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Este compartilhamento tem como objetivo impulsionar o conhecimento sobre esta nova doença chamada COVID 19.
COVID-19 Adaptation
Clinical Care for Severe Acute Respiratory Infection Toolkit

COVID-19 Adaptation

World Health Organization
Clinical care for severe acute respiratory infection: toolkit, update 2022. COVID-19 adaptation

WHO/2019-nCoV/SARI_toolkit/2022.1

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# Contents

FOREWORD ........................................................................................................................................... vii

ACKNOWLEDGEMENTS ................................................................................................................... viii

ABBREVIATIONS .............................................................................................................................. xi

EXECUTIVE SUMMARY .................................................................................................................... xvii

## 1. EPIDEMIOLOGY OF SARI ............................................................................................................ 1

Summary ............................................................................................................................................... 2
1.1 Differential diagnosis of SARI ...................................................................................................... 3
1.2 COVID-19 (SARS-CoV-2) fact sheet .......................................................................................... 5
1.3 Influenza virus fact sheet ........................................................................................................... 7
1.4 Middle East respiratory syndrome coronavirus (MERS-CoV) fact sheet .................................. 10
1.5 Risk factors for severe disease: influenza vs COVID-19 ............................................................ 11
References and resources .................................................................................................................. 12

## 2. SCREENING, TRIAGE, CLINICAL ASSESSMENT AND MANAGEMENT OF SARI .......... 15

Summary ............................................................................................................................................... 16
2.1 Screening for SARI ..................................................................................................................... 18
2.2 Triage for SARI .......................................................................................................................... 23
2.3 Clinical assessment of acutely ill patients - basic emergency care: ABCDE approach ............... 29
2.4 Classification of severity in patients with COVID-19 ............................................................... 32
2.5 Decision-making algorithm for patients presenting with acute respiratory infection (ARI) (influenza or COVID-19 suspected or known to be circulating) ................ 34
2.6 Decision-making algorithm for hospitalization of patients with pneumonia (influenza or COVID-19 known to be circulating) ........................................................................... 35
2.7 Decision-making algorithm for hospitalization and ICU admission for patients with SARI and severe pneumonia ........................................................................... 36
2.8 Checklist for admission .............................................................................................................. 37
2.9 Checklist for transfer ................................................................................................................. 38
2.10 Transfer of critically ill patients: air medevac for COVID-19 patients ..................................... 39
References and resources .................................................................................................................. 42

## 3. INFECTION PREVENTION AND CONTROL FOR PATIENTS WITH SARI ................. 45

Summary ............................................................................................................................................... 46
3.1 How to implement infection control measures for SARI .......................................................... 48
3.2 How to implement infection control measures for ARIs of potential concern ......................... 49
3.3 Personal protective equipment (PPE) ....................................................................................... 50
3.4 How to improve medical mask fit in health care settings .......................................................... 52
3.5 Steps to perform a particulate respirator seal check during the putting on of PPE .................. 53
3.6 Hand hygiene ............................................................................................................................ 54
3.7 The 5 moments for hand hygiene in health care facilities ....................................................... 56
8. SEPSIS AND SEPTIC SHOCK .............................................................. 155

8.1 Sepsis and septic shock definitions .............................................. 156
8.2 Sequential Organ Failure Assessment (SOFA) score ...................... 160
8.3 Quick Sequential Organ Failure Assessment (qSOFA) .................... 161
8.4 Paediatric Logistic Organ Dysfunction (PELOD-2) score ................ 162
8.5 Algorithm on targeted resuscitation in adults with shock ............... 164
8.6 Algorithms on initial resuscitation, and on fluid and vasoactive-inotrope management for children with septic shock .......... 165
8.7 Guide to the use of vasopressors in septic shock for adults and children .................. 167
8.8 The five rules for passive leg raise (PLR) ....................................... 169

References and resources ................................................................. 170

9. ACUTE RESPIRATORY DISTRESS SYNDROME (ARDS) ..................... 173

9.1 Memory aid: diagnosis and classification of ARDS in adults .......... 176
9.2 Memory aid: diagnosis and classification of pARDS in children ....... 177
9.3 Advanced non-invasive oxygen delivery in ARDS: algorithm to escalate supportive respiratory therapy .............................................. 178
9.4 Checklist for rapid sequence intubation procedure in adults and children ...................................................... 179
9.5 Considerations for intubation and mechanical ventilation in children ...................................................... 181
9.6 List of commonly used medicines and dosage in ICU with ventilated patients (adults, children) .................. 183
9.7 Choice of induction agents in adults ............................................. 187
9.8 Choice of induction agents in children ........................................ 188
9.9 Protocol to deliver lung protective ventilation (LPV) .................... 189
9.10 Memory aid: comparison of normal waveforms during volume and pressure-limited ventilation .............................................. 191
9.11 Memory aid: recognizing and interpreting abnormal pressure and flow waveforms during volume control ventilation .......... 192
9.12 Guide to distinguishing between the causes of high peak airway pressures: resistance versus compliance .............................................. 193
9.13 Troubleshooting high peak airway pressures, low tidal volumes, desaturation or haemodynamic instability in ventilated patients .......... 194
9.14 Respiratory care pocket card reference ........................................ 195
9.15 Adult ventilation order set (ARDS) ............................................. 204
9.16 Checklist for proning in severe ARDS ........................................ 205
9.17 Ventilator circuit types, filter and humidifier locations for SARI ........ 207

References and resources ................................................................. 212

10. MANAGE PAIN, SEDATION AND DELIRIUM .................................. 215

10.1 Numerical pain assessment scales .............................................. 218
10.2 Behavioural pain assessment scales ........................................... 219
10.3 Richmond Agitation-Sedation Scale (RASS) .................................. 225
10.4 Flowchart and worksheet for the Confusion Assessment Method of the ICU for adults (CAM-ICU) .................................................................................................................... 226
10.5 Flowchart and worksheet for the Confusion Assessment Method of the ICU for children (pCAM-ICU) .................................................................................................................... 228
10.6 Procedure for assessing attention: attention screening exam (ASE) for adults .................. 231
10.7 Guide to commonly used sedatives in adults ............................................................... 233
10.8 Guide to commonly used opioid analgesics in adults .................................................... 234
10.9 Guide to using neuromuscular blockers in adults ......................................................... 235
10.10 Guide to commonly used antipsychotics (haloperidol) in adults .................................. 236
10.11 Guide to paediatric analgesics, sedatives and neuromuscular blockers ....................... 237

References and resources .................................................................................................. 238

11. LIBERATION FROM INVASIVE MECHANICAL VENTILATION ........................................ 239
Summary ............................................................................................................................ 240
11.1 Algorithm for daily sedation interruption and daily spontaneous breathing trial (SBT) ........ 241
11.2 How to perform a cuff leak test .................................................................................. 242
11.3 Respiratory care pocket card reference ...................................................................... 244
11.4 Spontaneous breathing trial (SBT) order set .................................................................. 246
References and resources ................................................................................................. 247

12. BEST PRACTICES TO PREVENT COMPLICATIONS .......................................................... 249
Summary ............................................................................................................................ 250
12.1 Interventions to prevent complications in hospitalized and critically ill patients with COVID-19 ........................................ 251
12.2 Checklist for central venous catheter (CVC) insertion .................................................. 252
12.3 Checklist for preventing ventilator-associated pneumonia (VAP) ................................. 253
12.4 Checklist for preventing urinary tract infections (UTI) .................................................. 254
12.5 Procedure for providing enteral nutrition (EN) for adults ........................................... 255
12.6 Procedure for providing enteral nutrition (EN): paediatric considerations .................... 256
12.7 Algorithm for early mobility in the ICU ...................................................................... 258
12.8 Thromboembolic prophylaxis in COVID-19 ................................................................. 259
12.9 ABCDE bundle ............................................................................................................ 260
References and resources ................................................................................................. 263

13. QUALITY IN CRITICAL CARE ...................................................................................... 265
Summary ............................................................................................................................ 266
13.1 Checklist for ICU daily best practices ........................................................................ 268
13.2 Surviving Sepsis Campaign bundles .......................................................................... 269
13.3 Checklist for high-quality use of invasive mechanical ventilation for ARDS .................. 270
13.4 Process for selecting problem to focus on in the ICU and quality improvement programme .................................................. 271
13.5 Checklist for initiating, improving, evaluating and sustaining a quality improvement programme .................................................................................................................. 272
References and resources ................................................................................................. 273

14. ETHICAL CONSIDERATIONS ..................................................................................... 275
Summary ............................................................................................................................ 276
14.1 Ethical values and principles ...................................................................................... 277
14.2 Triage decision process flow ...................................................................................... 279
14.3 Hospital scarce resource decision-making................................................................... 280
References and resources ................................................................................................. 281
Foreword

This toolkit is intended for clinicians working with adult and paediatric patients with severe forms of acute respiratory infection, including severe pneumonia, acute respiratory distress syndrome, sepsis and septic shock in low- and middle-income countries.

Its main objective is to provide some of the necessary tools that can be used to care for patients with respiratory conditions from hospital entry to hospital discharge. It is a hands-on practical guide for health care professionals involved in critical care management during the COVID-19 pandemic and outbreaks of influenza (seasonal or avian influenza), Middle East respiratory syndrome coronavirus (MERS-CoV) or other emerging respiratory viral epidemics.

The toolkit is structured by topic and follows the different levels of care required to manage the respiratory conditions. Each topic starts with a summary and follows with the tools; complementary references and resources are listed at the end of each section. The tools provide a framework for users and are to be adapted to local conditions.

The child icon identifies tools to be used and adapted when caring for paediatric patients.

The adult icon identifies tools to be used and adapted when caring for adult patients.

Tools without an icon can be used and adapted when caring for adult and paediatric patients.

Accompanying the toolkit there are several links and QR codes that clinicians can use to access materials for use in clinical settings.
Acknowledgements

In 2015 and 2016, a major revision of the toolkit and associated materials was conducted to include the most recent internationally peer-reviewed publications at that time. In 2020, the toolkit was adapted for the COVID-19 pandemic and, in 2022, the updated version was developed to add new evidence and improved content with new algorithms, infographics and tables to facilitate the management of patients with severe acute respiratory infections (SARI) (including specifications for influenza virus and SARS-CoV-2 infection).

Confidentiality undertakings and declaration of interests forms were collected and reviewed from all the collaborators for this toolkit and no conflict of interests were identified.

2022 version

This latest update of the toolkit, with practical and simplified materials to manage patients with SARI, is the product of the contribution of many individuals, under the coordination of the case management team from the WHO Health Emergency Programme and guidance from Janet Diaz. Major contributions were provided by Vanessa Cramond (WHO), Janet Diaz (WHO), Bharath Kumar (WHO), Krutika Kuppalli (WHO), Marta Lado (WHO), Michael Lipnick (Zuckerberg San Francisco General Hospital, United States of America [USA]), Kobus Preller (WHO), Pryanka Relan (WHO), Alejandra Velez Ruiz Gaitan (WHO) and Archanah Seahwag (WHO consultant). Special thanks also go to our copyeditor Vivien Stone (Etchingham, United Kingdom of Great Britain and Northern Ireland [United Kingdom]) and for the design to Irene Lengui from L’IV Com Sàrl (Villars-sous-Yens, Switzerland).

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2020 version

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# Abbreviations

<table>
<thead>
<tr>
<th>Abbreviation</th>
<th>Description</th>
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<tbody>
<tr>
<td>ABCCs</td>
<td>airway, breathing, circulation, consciousness/convulsing</td>
</tr>
<tr>
<td>ABCDE</td>
<td>awakening, breathing coordination, delirium monitoring/management and early mobility (bundle)</td>
</tr>
<tr>
<td>ABG</td>
<td>air blood gases</td>
</tr>
<tr>
<td>AC</td>
<td>assist control</td>
</tr>
<tr>
<td>ACVPU</td>
<td>alert, confusion, verbal, pain, unresponsive</td>
</tr>
<tr>
<td>AGP</td>
<td>aerosol-generating procedure</td>
</tr>
<tr>
<td>Ag-RDT</td>
<td>antigen rapid diagnostic test</td>
</tr>
<tr>
<td>ALT</td>
<td>alanine aminotransferase</td>
</tr>
<tr>
<td>AMS</td>
<td>altered mental status</td>
</tr>
<tr>
<td>ANC</td>
<td>absolute neutrophil count</td>
</tr>
<tr>
<td>APRV</td>
<td>airway pressure release ventilation</td>
</tr>
<tr>
<td>ARDS</td>
<td>acute respiratory distress syndrome</td>
</tr>
<tr>
<td>ARI</td>
<td>acute respiratory infection</td>
</tr>
<tr>
<td>ASE</td>
<td>attention screening exam</td>
</tr>
<tr>
<td>AST</td>
<td>aspartate aminotransferase</td>
</tr>
<tr>
<td>AVPU</td>
<td>alert, verbal, pain, unresponsive (scale for assessing level of consciousness)</td>
</tr>
<tr>
<td>bCPAP</td>
<td>bubble continuous positive airway pressure</td>
</tr>
<tr>
<td>BEC</td>
<td>basic emergency care</td>
</tr>
<tr>
<td>BEE</td>
<td>basal energy expenditure</td>
</tr>
<tr>
<td>BID</td>
<td>twice a day</td>
</tr>
<tr>
<td>BiPAP</td>
<td>bilevel positive airway pressure</td>
</tr>
<tr>
<td>BMI</td>
<td>body mass index</td>
</tr>
<tr>
<td>BP</td>
<td>blood pressure</td>
</tr>
<tr>
<td>bpm</td>
<td>beats per minute</td>
</tr>
<tr>
<td>BPS</td>
<td>Behavioural Pain Scale</td>
</tr>
<tr>
<td>BS</td>
<td>British Standards</td>
</tr>
<tr>
<td>BSI</td>
<td>blood stream infection</td>
</tr>
<tr>
<td>BV filter</td>
<td>bacteria-virus filter</td>
</tr>
<tr>
<td>BVM</td>
<td>bag valve mask (Ambu)</td>
</tr>
<tr>
<td>CAM-ICU</td>
<td>confusion assessment method for the intensive care unit for adults</td>
</tr>
<tr>
<td>CAPA</td>
<td>coronavirus disease-associated pulmonary aspergillosis</td>
</tr>
<tr>
<td>CBC</td>
<td>cell blood count</td>
</tr>
<tr>
<td>CCB</td>
<td>calcium channel blockers</td>
</tr>
<tr>
<td>CCC</td>
<td>Clinical Care Committee</td>
</tr>
<tr>
<td>CDC</td>
<td>Centers for Disease Control and Prevention (United States of America)</td>
</tr>
<tr>
<td>Abbreviation</td>
<td>Description</td>
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<tr>
<td>--------------</td>
<td>-------------</td>
</tr>
<tr>
<td>CFR</td>
<td>case fatality ratio</td>
</tr>
<tr>
<td>CNS</td>
<td>central nervous system</td>
</tr>
<tr>
<td>CO</td>
<td>carbon monoxide/cardiac output</td>
</tr>
<tr>
<td>CO₂</td>
<td>carbon dioxide</td>
</tr>
<tr>
<td>COPD</td>
<td>chronic obstructive pulmonary disease</td>
</tr>
<tr>
<td>COVID-19</td>
<td>coronavirus disease</td>
</tr>
<tr>
<td>CPAP</td>
<td>continuous positive airway pressure</td>
</tr>
<tr>
<td>CPOT</td>
<td>Critical-Care Pain Observation Tool</td>
</tr>
<tr>
<td>CPP</td>
<td>cerebral perfusion pressure</td>
</tr>
<tr>
<td>CR</td>
<td>capillary refill</td>
</tr>
<tr>
<td>CRP</td>
<td>c-reactive protein</td>
</tr>
<tr>
<td>CPR</td>
<td>cardio-pulmonary resuscitation</td>
</tr>
<tr>
<td>CrAg</td>
<td>cryptococcal antigen</td>
</tr>
<tr>
<td>CRBSI</td>
<td>catheter-related blood stream infection</td>
</tr>
<tr>
<td>CSF</td>
<td>cerebrospinal fluid</td>
</tr>
<tr>
<td>CT</td>
<td>computerized tomography</td>
</tr>
<tr>
<td>CURB-65</td>
<td>confusion, urea, respiratory rate, blood pressure, 65 years (score)</td>
</tr>
<tr>
<td>CVC</td>
<td>central venous catheter</td>
</tr>
<tr>
<td>CVP</td>
<td>central venous pressure</td>
</tr>
<tr>
<td>CXR</td>
<td>chest X-ray</td>
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<tr>
<td>DBP</td>
<td>diastolic blood pressure</td>
</tr>
<tr>
<td>DKA</td>
<td>diabetic ketoacidosis</td>
</tr>
<tr>
<td>DM</td>
<td>diabetes mellitus</td>
</tr>
<tr>
<td>DVT</td>
<td>deep venous thrombosis</td>
</tr>
<tr>
<td>ECG</td>
<td>electrocardiogram</td>
</tr>
<tr>
<td>ECLS</td>
<td>extracorporeal life support</td>
</tr>
<tr>
<td>ECMO</td>
<td>extracorporeal membrane oxygenation</td>
</tr>
<tr>
<td>EIA</td>
<td>enzyme immunoassay</td>
</tr>
<tr>
<td>EIP</td>
<td>electromagnetic inductance plethysmography</td>
</tr>
<tr>
<td>EN</td>
<td>enteral nutrition</td>
</tr>
<tr>
<td>ENT</td>
<td>ear, nose and throat</td>
</tr>
<tr>
<td>EPAP</td>
<td>expiratory positive airway pressure</td>
</tr>
<tr>
<td>ESBL</td>
<td>extended spectrum beta-lactamase</td>
</tr>
<tr>
<td>ESI</td>
<td>emergency severity index</td>
</tr>
<tr>
<td>ESR</td>
<td>erythrocyte sedimentation rate</td>
</tr>
<tr>
<td>ETAT</td>
<td>emergency triage, assessment and treatment</td>
</tr>
<tr>
<td>ETCO₂</td>
<td>end-tidal carbon dioxide</td>
</tr>
<tr>
<td>ETT</td>
<td>endotracheal tube</td>
</tr>
<tr>
<td>EV-D68</td>
<td>Enterovirus D68</td>
</tr>
<tr>
<td>FFP2</td>
<td>filtering face pieces 92%</td>
</tr>
<tr>
<td>Abbreviation</td>
<td>Description</td>
</tr>
<tr>
<td>--------------</td>
<td>-------------</td>
</tr>
<tr>
<td>FiO₂</td>
<td>fraction of inspired oxygen</td>
</tr>
<tr>
<td>FLACC</td>
<td>face, legs, activity, cry, consolability</td>
</tr>
<tr>
<td>FM</td>
<td>face mask</td>
</tr>
<tr>
<td>GCS</td>
<td>Glasgow Coma Scale</td>
</tr>
<tr>
<td>GI</td>
<td>gastrointestinal</td>
</tr>
<tr>
<td>GMP</td>
<td>good manufacturing practices</td>
</tr>
<tr>
<td>Hb</td>
<td>haemoglobin</td>
</tr>
<tr>
<td>HBCAb</td>
<td>hepatitis B core antibody</td>
</tr>
<tr>
<td>HBsAg</td>
<td>hepatitis B surface antigen</td>
</tr>
<tr>
<td>HCAb</td>
<td>hepatitis C antibody</td>
</tr>
<tr>
<td>HCO₃⁻</td>
<td>blood bicarbonate</td>
</tr>
<tr>
<td>HDL</td>
<td>high density lipoprotein</td>
</tr>
<tr>
<td>HEPA</td>
<td>high-efficiency particulate air (filter)</td>
</tr>
<tr>
<td>HFNC</td>
<td>high-flow nasal cannula</td>
</tr>
<tr>
<td>HFNO</td>
<td>high-flow nasal oxygen</td>
</tr>
<tr>
<td>HICS</td>
<td>hospital incident command system</td>
</tr>
<tr>
<td>HIV</td>
<td>human immunodeficiency virus</td>
</tr>
<tr>
<td>HME</td>
<td>heat moisture exchanger</td>
</tr>
<tr>
<td>HR</td>
<td>heart rate</td>
</tr>
<tr>
<td>ICP</td>
<td>intracranial pressure</td>
</tr>
<tr>
<td>ICRC</td>
<td>International Committee of the Red Cross</td>
</tr>
<tr>
<td>ICU</td>
<td>intensive care unit</td>
</tr>
<tr>
<td>IHR</td>
<td>International Health Regulations</td>
</tr>
<tr>
<td>IITT</td>
<td>Interagency Integrated Triage Tool</td>
</tr>
<tr>
<td>ILI</td>
<td>influenza-like illness</td>
</tr>
<tr>
<td>IL-6</td>
<td>Interleukin-6</td>
</tr>
<tr>
<td>IL-6 RB</td>
<td>Interleukin-6 receptor blockers</td>
</tr>
<tr>
<td>IM</td>
<td>intramuscular</td>
</tr>
<tr>
<td>IMAI</td>
<td>Integrated Management of Adolescent and Adult Illness</td>
</tr>
<tr>
<td>IMV</td>
<td>invasive mechanical ventilation</td>
</tr>
<tr>
<td>iNO</td>
<td>inhaled nitric oxide</td>
</tr>
<tr>
<td>IO</td>
<td>intraosseous</td>
</tr>
<tr>
<td>IPC</td>
<td>infection prevention and control</td>
</tr>
<tr>
<td>ISO</td>
<td>International Organization for Standardization</td>
</tr>
<tr>
<td>IU</td>
<td>international units</td>
</tr>
<tr>
<td>IV</td>
<td>intravenous</td>
</tr>
<tr>
<td>JVP</td>
<td>jugular venous pressure</td>
</tr>
<tr>
<td>LAT</td>
<td>latex agglutination test</td>
</tr>
<tr>
<td>LDH</td>
<td>lactate dehydrogenase</td>
</tr>
<tr>
<td>LDL</td>
<td>low density lipoprotein</td>
</tr>
<tr>
<td>Abbreviation</td>
<td>Description</td>
</tr>
<tr>
<td>--------------</td>
<td>-------------</td>
</tr>
<tr>
<td>LFA</td>
<td>lateral flow immunochromatographic assay</td>
</tr>
<tr>
<td>LOS</td>
<td>length of stay</td>
</tr>
<tr>
<td>LPM/L/min</td>
<td>litres per minute</td>
</tr>
<tr>
<td>LPV</td>
<td>lung protective ventilation</td>
</tr>
<tr>
<td>LR</td>
<td>lactated Ringer's</td>
</tr>
<tr>
<td>LRT</td>
<td>lower respiratory tract</td>
</tr>
<tr>
<td>MAP</td>
<td>mean arterial pressure</td>
</tr>
<tr>
<td>MDR</td>
<td>multidrug-resistant</td>
</tr>
<tr>
<td>MERS-CoV</td>
<td>Middle East respiratory syndrome coronavirus</td>
</tr>
<tr>
<td>MEWS</td>
<td>modified Early Warning Score</td>
</tr>
<tr>
<td>MI</td>
<td>myocardial infarction</td>
</tr>
<tr>
<td>MIS-C</td>
<td>multisystem inflammatory syndrome in children</td>
</tr>
<tr>
<td>MRI</td>
<td>magnetic resonance image</td>
</tr>
<tr>
<td>MRSAR</td>
<td>methicillin-resistant Staphylococcus aureus</td>
</tr>
<tr>
<td>MSF</td>
<td>Médecins Sans Frontières</td>
</tr>
<tr>
<td>MV</td>
<td>minute ventilation</td>
</tr>
<tr>
<td>NAAT</td>
<td>nucleic acid amplification testing</td>
</tr>
<tr>
<td>NCD</td>
<td>noncommunicable disease</td>
</tr>
<tr>
<td>NEWS</td>
<td>National Early Warning Score (adults)</td>
</tr>
<tr>
<td>NG</td>
<td>nasogastric</td>
</tr>
<tr>
<td>NJ</td>
<td>nasojejunal</td>
</tr>
<tr>
<td>NIPPV</td>
<td>nasal intermittent positive pressure ventilation</td>
</tr>
<tr>
<td>NIV</td>
<td>non-invasive ventilation</td>
</tr>
<tr>
<td>NMB</td>
<td>neuromuscular blockers</td>
</tr>
<tr>
<td>NPA</td>
<td>nasopharyngeal airway</td>
</tr>
<tr>
<td>NRB</td>
<td>non-rebreather</td>
</tr>
<tr>
<td>NS</td>
<td>normal saline</td>
</tr>
<tr>
<td>NSAID</td>
<td>non-steroidal anti-inflammatory drugs</td>
</tr>
<tr>
<td>N95</td>
<td>filtering less 95%</td>
</tr>
<tr>
<td>OG</td>
<td>orogastric</td>
</tr>
<tr>
<td>OI</td>
<td>oxygenation index using SpO₂</td>
</tr>
<tr>
<td>OMD</td>
<td>Office of Medical Director</td>
</tr>
<tr>
<td>OPA</td>
<td>oropharyngeal airway</td>
</tr>
<tr>
<td>OSI</td>
<td>oxygen saturation index</td>
</tr>
<tr>
<td>PAD</td>
<td>pain, agitation and delirium</td>
</tr>
<tr>
<td>PALS</td>
<td>paediatric advanced life support</td>
</tr>
<tr>
<td>PaCO₂</td>
<td>partial pressure of carbon dioxide</td>
</tr>
<tr>
<td>PaO₂</td>
<td>partial pressure arterial oxygen</td>
</tr>
<tr>
<td>pARDS</td>
<td>paediatric acute respiratory distress respiratory</td>
</tr>
<tr>
<td>PBW</td>
<td>predicted body weight</td>
</tr>
<tr>
<td>Abbreviation</td>
<td>Description</td>
</tr>
<tr>
<td>--------------</td>
<td>-------------</td>
</tr>
<tr>
<td>pCAM-ICU</td>
<td>confusion assessment method for the intensive care unit for children</td>
</tr>
<tr>
<td>PCR</td>
<td>polymerase chain reaction</td>
</tr>
<tr>
<td>PDR</td>
<td>pressure driving</td>
</tr>
<tr>
<td>PEEP</td>
<td>positive end-expiratory pressure</td>
</tr>
<tr>
<td>PELOD</td>
<td>paediatric logistic organ dysfunction</td>
</tr>
<tr>
<td>PES</td>
<td>post-extubation stridor</td>
</tr>
<tr>
<td>PEWS</td>
<td>Paediatric Early Warning Score</td>
</tr>
<tr>
<td>P:F</td>
<td>ratio between arterial partial pressure of oxygen and the fraction of inspired oxygen</td>
</tr>
<tr>
<td>PIP</td>
<td>pressure inspiratory peak</td>
</tr>
<tr>
<td>PLR</td>
<td>passive leg raising</td>
</tr>
<tr>
<td>PLT</td>
<td>platelets</td>
</tr>
<tr>
<td>po</td>
<td>per os</td>
</tr>
<tr>
<td>POCT</td>
<td>point-of-care testing</td>
</tr>
<tr>
<td>PONV</td>
<td>post-operative nausea and vomiting</td>
</tr>
<tr>
<td>PPE</td>
<td>personal protective equipment</td>
</tr>
<tr>
<td>Pplat</td>
<td>plateau pressure</td>
</tr>
<tr>
<td>PPx</td>
<td>prophylaxis</td>
</tr>
<tr>
<td>PS</td>
<td>pressure support</td>
</tr>
<tr>
<td>PSA</td>
<td>pressure swing adsorption</td>
</tr>
<tr>
<td>PSI</td>
<td>pounds per square inch absolute</td>
</tr>
<tr>
<td>PSV</td>
<td>pressure support ventilation</td>
</tr>
<tr>
<td>PT</td>
<td>physical therapist</td>
</tr>
<tr>
<td>PTSD</td>
<td>post-traumatic stress disorder</td>
</tr>
<tr>
<td>QID</td>
<td>four times a day</td>
</tr>
<tr>
<td>qSOFA</td>
<td>quick sequential organ failure assessment</td>
</tr>
<tr>
<td>RASS</td>
<td>Richmond Agitation-Sedation Scale</td>
</tr>
<tr>
<td>RB</td>
<td>receptor blockers</td>
</tr>
<tr>
<td>RDT</td>
<td>Rapid diagnostic test</td>
</tr>
<tr>
<td>ROCM</td>
<td>rhino-orbito-cerebral mucormycosis</td>
</tr>
<tr>
<td>ROM</td>
<td>range of motion</td>
</tr>
<tr>
<td>RR</td>
<td>respiratory rate</td>
</tr>
<tr>
<td>RSBI</td>
<td>rapid shallow breathing index</td>
</tr>
<tr>
<td>RSI</td>
<td>rapid sequence intubation</td>
</tr>
<tr>
<td>RSV</td>
<td>respiratory syncytial virus</td>
</tr>
<tr>
<td>RT</td>
<td>respiratory therapist</td>
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<tr>
<td>RT-PCR</td>
<td>reverse transcription polymerase chain reaction</td>
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<tr>
<td>SARI</td>
<td>severe acute respiratory infection</td>
</tr>
<tr>
<td>SARS-CoV-2</td>
<td>severe acute respiratory syndrome coronavirus 2</td>
</tr>
<tr>
<td>SAT</td>
<td>spontaneous awakening trial</td>
</tr>
<tr>
<td>Abbreviation</td>
<td>Full Form</td>
</tr>
<tr>
<td>--------------</td>
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</tr>
<tr>
<td>SBP</td>
<td>systolic blood pressure</td>
</tr>
<tr>
<td>SBT</td>
<td>spontaneous breathing trial</td>
</tr>
<tr>
<td>SCI</td>
<td>spinal cord injury</td>
</tr>
<tr>
<td>ScvO$_2$</td>
<td>saturation of central venous blood</td>
</tr>
<tr>
<td>SD</td>
<td>standard deviation</td>
</tr>
<tr>
<td>S:F</td>
<td>ratio between saturation of oxygen and the fraction of inspired oxygen</td>
</tr>
<tr>
<td>SIMV</td>
<td>synchronized intermittent mandatory ventilation</td>
</tr>
<tr>
<td>SIRS</td>
<td>systemic inflammatory response syndrome</td>
</tr>
<tr>
<td>SmPC</td>
<td>Summary of Product Characteristics (European Union)</td>
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<td>SOFA</td>
<td>sequential organ failure assessment</td>
</tr>
<tr>
<td>SOP</td>
<td>standard operating procedure</td>
</tr>
<tr>
<td>SpO$_2$</td>
<td>oxygen saturation</td>
</tr>
<tr>
<td>TB</td>
<td>tuberculosis</td>
</tr>
<tr>
<td>TBI</td>
<td>traumatic brain injury</td>
</tr>
<tr>
<td>TMJ</td>
<td>temporomandibular joint</td>
</tr>
<tr>
<td>TNF-inhibitors</td>
<td>tumoral necrosis factor inhibitors</td>
</tr>
<tr>
<td>TOF</td>
<td>train-of-four</td>
</tr>
<tr>
<td>TT</td>
<td>triage team</td>
</tr>
<tr>
<td>TV</td>
<td>tidal volume</td>
</tr>
<tr>
<td>ULN</td>
<td>upper limit of normal</td>
</tr>
<tr>
<td>URT</td>
<td>upper respiratory tract</td>
</tr>
<tr>
<td>US</td>
<td>ultrasound</td>
</tr>
<tr>
<td>USP</td>
<td>United States Prescribing Information</td>
</tr>
<tr>
<td>UTI</td>
<td>urinary tract infection</td>
</tr>
<tr>
<td>VA</td>
<td>veno-arterial</td>
</tr>
<tr>
<td>VAP</td>
<td>ventilator-associated pneumonia</td>
</tr>
<tr>
<td>VAS</td>
<td>Visual Analogue Scale</td>
</tr>
<tr>
<td>VOC</td>
<td>Variant of Concern</td>
</tr>
<tr>
<td>VOI</td>
<td>Variant of Interest</td>
</tr>
<tr>
<td>VPSA</td>
<td>vacuum pressure swing adsorption</td>
</tr>
<tr>
<td>VTE</td>
<td>venous thromboembolism</td>
</tr>
<tr>
<td>VTM</td>
<td>viral transport medium</td>
</tr>
<tr>
<td>VV</td>
<td>veno-venous</td>
</tr>
<tr>
<td>WBC</td>
<td>white cell count</td>
</tr>
<tr>
<td>WHO</td>
<td>World Health Organization</td>
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</table>
Executive summary

Description of updates and new tools in the SARI toolkit update 2022

1. EPIDEMIOLOGY OF SARI

- Updated epidemiological information on COVID-19 and other viral infections that produce SARI with pandemic potential.

**UPDATED**

1.1 Differential diagnosis of SARI
1.2 COVID-19 (SARS-CoV-2) fact sheet
1.3 Influenza virus fact sheet
1.4 Middle East respiratory syndrome coronavirus (MERS-CoV) fact sheet

**NEW**

1.5 Risk factors for severe disease: influenza vs COVID-19

2. SCREENING, TRIAGE, CLINICAL ASSESSMENT AND MANAGEMENT OF SARI

- Detailed description to differentiate screening and triage for patients with SARI and recommended tools.
- Updated WHO classification of severity in COVID-19 as part of triage.
- Update of algorithms for the management of ARI and SARI in the community and in health facilities with consideration of influenza and COVID-19.
- Inclusion of medevac protocols for transfer of patients with SARI due to COVID-19.

**UPDATED**

2.5 Decision-making algorithm for patients presenting with acute respiratory infection (ARI) (influenza or COVID-19 suspected or known to be circulating)
2.6 Decision-making algorithm for hospitalization of patients with pneumonia (influenza or COVID-19 known to be circulating)
2.7 Decision-making support tool for hospitalization and ICU admission for patients with SARI and severe pneumonia

**NEW**

2.1 Screening for SARI
2.2 Triage for SARI
2.4 Classification of severity in patients with COVID-19
2.10 Transfer of critically ill patients: air medevac for COVID-19 patients
3. INFECTION PREVENTION AND CONTROL FOR PATIENTS WITH SARI

• Updated measures for IPC for SARI and specifically for COVID-19.
• Updates on PPE for SARI with special consideration of COVID-19.

UPDATED
3.1 How to implement infection control measures for SARI
3.3 Personal protective equipment (PPE)
3.6 Hand hygiene
3.9 Checklist for aerosol-generating procedures

NEW
3.2 How to implement infection control measures for ARIs of potential concern
3.4 How to improve medical mask fit in health care settings
3.7 The 5 moments for hand hygiene in health care facilities
3.8 The “Three Cs”: settings where transmission of the COVID-19 virus spreads more easily

4. MONITORING PATIENTS WITH ACUTE RESPIRATORY INFECTION

• Updates on the description of monitoring of patients with SARI: pulse oximetry, blood gases, capnometry.
• New table on monitoring patients with COVID-19 depending on disease severity and patient disposition (home, health facility, etc.).
• Bundle of recommendations for health care workers to care for and monitor patients with mild COVID-19 at home.

UPDATED
4.2 Pulse oximetry monitoring
4.3 Blood gas analysis monitoring
4.5 National Early Warning Score (NEWS) for adults

NEW
4.4 Capnometry (capnography)
4.7 Routine monitoring and care framework for COVID-19 patients
4.8 WHO Mild COVID-19 home care bundle for health care workers

5. DIAGNOSTIC TESTING FOR PATIENTS WITH ARI

• Specifications for diagnostic testing in COVID-19.
• Link to videos with explanation of different specimen collection systems (nasopharyngeal and pharyngeal swabs).

UPDATED
5.3 Specimen collection kit for upper respiratory tract specimens

NEW
5.1 Diagnostic testing for SARS-CoV-2 infection
5.2 Use of antigen-detection rapid diagnostic testing for SARS-CoV-2
## 6. OXYGEN THERAPY

- Definition of concepts related to oxygen therapy: oxygen concentration, fraction of inspired oxygen, saturation of oxygen.
- Indications for oxygen therapy.
- Explanation of the different oxygen delivery devices for low and high flows of oxygen.
- Specifications for oxygen therapy in children.
- Algorithms and flowcharts to understand how to titrate oxygen therapy according to need in neonates, children and adults.
- Awake prone positioning as an adjunctive therapy to oxygen in SARI/ARDS: indications and tips.
- Explanation and description of different oxygen supply sources: PSA, cryogenic liquid oxygen, concentrators, cylinders.
- Recommended template document to check oxygen therapy in patients in health care facilities.

### UPDATED

6.9 Checklist to troubleshoot warning signs during oxygen therapy delivery

### NEW

6.1 Indications for oxygen therapy
6.2 Memory aid: oxygen delivery devices
6.3 Memory aid: oxygen delivery in children
6.4 Algorithm to escalate respiratory support in adults and children with pneumonia
6.5 Flowchart on how to titrate oxygen in neonates
6.6 Flowchart on how to titrate oxygen in children
6.7 Flowchart on how to titrate oxygen in adults
6.8 Key tips on awake prone positioning
6.10 Oxygen supply calculations.
6.11 Memory aids: oxygen supply sources and distribution
6.12 Respiratory care order template for oxygen therapy

## 7. THERAPIES FOR SARI (COVID-19, INFLUENZA, BACTERIAL PNEUMONIA): ANTIMICROBIALS AND IMMUNOMODULATORS

- Update on antimicrobial therapies for SARI.
- Updated tables for antimicrobials in ARI adapted to influenza and COVID-19.
- Summary of COVID-19 therapeutics approved for use.
- Tables with descriptions of and administration instructions for COVID-19 therapeutics: corticosteroids, IL-6 RB; monoclonal antibodies.

### NEW

7.1 Memory aid: treatment for acute respiratory infections according to severity (when COVID-19 and influenza are circulating)
7.2 COVID-19 and therapeutics
7.3 Memory aid: invasive fungal infections in patients with COVID-19
8. SEPSIS AND SEPTIC SHOCK
• Updated definitions for sepsis and septic shock in adults and children.
• Description of different tools related to sepsis (SOFA, PELOD-2).
• Tables and algorithms for management of sepsis and shock, including use of vasopressors.

UPDATED
8.6 Algorithms on initial resuscitation, and on fluid and vasoactive-inotrope management for children with septic shock
8.7 Guide to the use of vasopressors in septic shock for adults and children

NEW
8.1 Sepsis and septic shock definitions
8.2 Sequential Organ Failure Assessment (SOFA) score
8.3 Quick Sequential Organ Failure Assessment (qSOFA)
8.4 Paediatric Logistic Organ Dysfunction (PELOD-2) score
8.5 Algorithm on targeted resuscitation in adults with shock

9. ACUTE RESPIRATORY DISTRESS SYNDROME (ARDS)
• Algorithms to escalate respiratory support in patients with ARDS including principles for intubation and mechanical ventilation in adults and children.
• New list of commonly used medicines in ventilated patients.
• Update of the protocol to deliver lung protective ventilation and description of different types of ventilation.
• New tools: respiratory pocket cards to support clinicians with the fundamentals of respiratory support.
• Checklist to help with proning techniques for patients with ARDS.
• New section to explain the locations for ventilator circuits, filters and humidifiers for patients with invasive and non-invasive ventilation.

UPDATED
9.1 Memory aid: diagnosis and classification of ARDS in adults
9.2 Memory aid: diagnosis and classification of pARDS in children

NEW
9.3 Advanced non-invasive oxygen delivery in ARDS: algorithm to escalate supportive respiratory therapy
9.4 Checklist for rapid sequence intubation procedure in adults and children
9.5 Considerations for intubation and mechanical ventilation in children
9.6 List of commonly used medicines and dosage in ICU with ventilated patients (adults, children)
9.7 Choice of induction agents in adults
9.8 Choice of induction agents in children
9.9 Protocol to deliver lung protective ventilation (LPV)
9.10 Memory aid: comparison of normal waveforms during volume and pressure-limited ventilation
9.11 Memory aid: recognizing and interpreting abnormal pressure and flow waveforms during volume control ventilation
9.12 Guide to distinguishing between the causes of high peak airway pressures: resistance versus compliance
9.13 Troubleshooting high peak airway pressures, low tidal volumes, desaturation or hemodynamic instability in ventilated patients
9.14 Respiratory care pocket card reference
9.15 Adult ventilation order set (ARDS)
9.16 Checklist for proning in severe ARDS
9.17 Ventilator circuit types, filter and humidifier locations for SARI

10. MANAGE PAIN, SEDATION AND DELIRIUM
   • Revised formatting of tables, but no new tools.

11. LIBERATION FROM INVASIVE MECHANICAL VENTILATION
   • Updates in the algorithm for daily sedation interruption and daily spontaneous breathing trial, and the cuff leak test protocol.
   NEW
   11.1 Algorithm for daily sedation interruption and daily spontaneous breathing trial (SBT)
   11.2 How to perform a cuff leak test
   11.3 Respiratory care pocket card reference
   11.4 Spontaneous breathing trial (SBT) order set

12. BEST PRACTICES TO PREVENT COMPLICATIONS
   • Update on identification and management of complications in critical patients (ICU): tables.
   • Addition of thromboembolic prophylaxis in COVID-19.
   NEW
   12.1 Interventions to prevent complications in hospitalized and critically ill patients with COVID-19
   12.8 Thromboembolic prophylaxis in COVID-19

13. QUALITY IN CRITICAL CARE
   • Revised formatting of the summary of principles for quality in critical care.
   • New checklists for ICU daily best practices.
   NEW
   13.1 Checklist for ICU daily best practices

14. ETHICAL CONSIDERATIONS
   • Updates in tables related to key ethical principles, triage decision process flow and algorithm for hospital scarce resources decision-making.
   NEW
   14.1 Ethical values and principles
   14.2 Triage decision process flow
   14.3 Hospital scarce resource decision-making
Epidemiology of SARI

Summary

A global case definition for severe acute respiratory infection (SARI) for surveillance purposes, applicable to all age groups, was first published by the World Health Organization (WHO) in 2011. The current WHO case definition is:

Patients with acute respiratory infection who have history of fever (or measured fever of ≥ 38 °C), cough and onset within the last 10 days (symptoms within 10 days) and require hospitalization.

The differential diagnoses for SARI include a wide spectrum of community-acquired pathogens, including respiratory viruses, bacteria and other less common micro-organisms. The ranking of differential diagnoses will vary by host factors (e.g. age, presence of chronic conditions, travel history, vaccination), environmental factors (e.g. geographic location, vectors), epidemiologic factors (e.g. the prevalence of the pathogen in the community) and pathogen factors (e.g. tropism for lungs). The differential diagnosis for SARI should also include endemic infections, such as malaria, arboviruses (dengue, chikungunya, yellow fever), tuberculosis (TB), or even human immunodeficiency virus (HIV) related opportunistic infections that can produce severe acute respiratory infections. See Tool 1.1 for a comprehensive differential diagnosis list that can be adapted to local settings.

Clinicians should be aware that in addition to the common viral and bacterial pathogens, an emerging respiratory virus may result in SARI and lead to a public health emergency of international concern. For example, influenza virus types A and B is a common respiratory virus that circulates worldwide and can cause seasonal influenza outbreaks and epidemics. Most people recover from fever and other symptoms within a week without requiring medical attention; however, hospitalization and death may mainly among high-risk groups. Worldwide, these annual epidemics are estimated to result in 3 to 5 million cases of severe illness, and 290 000 to 650 000 respiratory deaths. This estimate does not take in account deaths from other diseases such as cardiovascular disease, which can be influenza-related. Emerging respiratory viruses include: zoonotic influenza viruses, such as avian influenza, and coronaviruses, such as MERS-CoV and SARS-CoV-2. In addition, a new subtype of influenza A virus may also have pandemic potential. See Tools 1.2-1.4 for comprehensive information on these respiratory viruses.

Tools

1.1 Differential diagnosis of SARI
1.2 COVID-19 (SARS-CoV-2) fact sheet
1.3 Influenza virus fact sheet
1.4 Middle East respiratory syndrome coronavirus (MERS-CoV) fact sheet
1.5 Risk factors for severe disease: influenza vs COVID-19
1.1 Differential diagnosis of SARI

It is important to develop a differential diagnosis rapidly for all patients presenting with SARI. This will guide health care workers in the initial infection prevention and control (IPC), diagnostic and treatment measures.

The rate of co-infection among these pathogens is unknown. Therefore, a positive diagnostic test for another infection does not exclude the need for other micro-organism testing.

**VIRAL PATHOGENS**

<table>
<thead>
<tr>
<th>Common viral pathogens</th>
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<tbody>
<tr>
<td>• Respiratory syncytial virus (RSV)</td>
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<tr>
<td>• Parainfluenza virus</td>
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<tr>
<td>• Rhinoviruses</td>
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<tr>
<td>• Adenovirus, enterovirus (EV-D68)</td>
</tr>
<tr>
<td>• Human metapneumovirus</td>
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<tr>
<td>• Bocavirus</td>
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<tr>
<td>• Seasonal influenza, known subtype</td>
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</tbody>
</table>

<table>
<thead>
<tr>
<th>Less common viral pathogens</th>
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</thead>
<tbody>
<tr>
<td>• Varicella zoster</td>
</tr>
<tr>
<td>• Measles</td>
</tr>
<tr>
<td>• Hantavirus</td>
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</tbody>
</table>

<table>
<thead>
<tr>
<th>Virus that may constitute a public health emergency of international concern</th>
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</thead>
<tbody>
<tr>
<td>• SARS</td>
</tr>
<tr>
<td>• SARS-CoV-2</td>
</tr>
<tr>
<td>• MERS</td>
</tr>
<tr>
<td>• New subtypes of influenza</td>
</tr>
<tr>
<td>• Other zoonotic viral infections</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>If immunosuppressed (i.e. people living with HIV)</th>
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<tbody>
<tr>
<td>• Cytomegalovirus,</td>
<td></td>
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<tr>
<td>• Herpes simplex viruses</td>
<td></td>
</tr>
</tbody>
</table>
## Bacterial Pathogens

### Most Common Bacterial Pathogens
- *Streptococcus pneumoniae*
- *Hemophilus influenzae*
- *Moraxella catarrhalis*
- *Legionella pneumophila, non-pneumophila Legionella*
- *Chlamydia pneumonia*
- *Mycoplasma pneumoniae*
- *Klebsiella pneumonia*
- *Staphylococcus aureus*

### Less Common, Unless at Risk or in High-Prevalence Geographic Area
- *Mycobacterium tuberculosis*
- *Burkholderia pseudomallei*
- Rickettsial infections
  - *Coxiella burnetti* (Q fever)
- *Leptospira spp.*
- *Chlamydia psittaci*
- *Bordetella pertussis*
- *Salmonella sp.*

### Resistant Pathogens

Risk factors for multidrug-resistant pathogens: Intravenous antimicrobial therapy within < 90 days.

Resistant pathogens include:
- Methicillin-resistant *S. aureus* (MRSA)
- Non-fermenters such as *Pseudomonas aeruginosa, Acinetobacter baumannii*
- Extended spectrum beta-lactamase (ESBL) producers such as *E. coli, Klebsiella, Enterobacter*

## Other Endemic Infections

### Potential Endemic Infections
Can co-exist with respiratory infections in endemic areas.

- Malaria, dengue, chikungunya, tuberculosis, HIV
1.2 COVID-19 (SARS-CoV-2) fact sheet

COVID-19

Introduction

- The novel coronavirus SARS-CoV-2 is similar genetically to the SARS coronavirus.
- The first cases were reported in December 2019 in China, with SARS-CoV-2 identified in early January 2020.
- Since then, cases have been reported in virtually all countries, and the disease declared as a Public Health Emergency of International Concern by WHO on 30 January 2020 and described as a pandemic in March 2020.
- The latest epidemiology and case counts are available in the COVID-19 WHO situation reports: Coronavirus disease (COVID-19) weekly epidemiological and operational updates.
- The latest technical guidance can be found in: Country & Technical Guidance - Coronavirus disease (COVID-19).

Transmission

- The SARS-CoV-2 virus is a zoonotic virus. The intermediary animal host and source of the virus has not yet been identified but is currently under study.
- The SARS-CoV-2 virus is spread mainly via inhalation of respiratory droplets coming from coughing or sneezing of an infected person to a person who is in close contact (within 1 m). Those respiratory droplets can reach or can be introduced in the mouth, nose or eyes of a susceptible person and can result in infection. Crowded closed indoor spaces with poor ventilation can be favourable environments for the virus to spread easily among people.
- Additionally, indirect contact transmission involving contact of a susceptible host with a contaminated object or surface (fomite transmission) may also be possible.
- Nosocomial transmission can occur where there are inadequate IPC measures, including personal protective equipment (PPE) to be used during close contact with infected individuals.
- Aerosol-generating procedures (AGP), such as open suctioning of airways, sputum induction, cardiopulmonary resuscitation, endotracheal intubation and extubation, non-invasive ventilation (e.g. bilevel positive airway pressure [BiPAP], continuous positive airway pressure [CPAP]), bronchoscopy, and manual ventilation, may present additional risk in health care settings for transmission and infection of health care workers, requiring higher levels of respiratory protection.
- The median incubation period is about 5-6 days (range: 1-14 days). The most infectious period is 1-2 days before symptoms appear. The infectious period can last up to 5-9 days for mild patients, up to 3 weeks for severe patients, and immunosuppressed patients can be infectious for many months, with high virus levels being detected in the upper respiratory tract early in the disease course or pre-symptomatic phase.
- SARS-CoV-2, as other viruses, may present changes and/or mutations. There have been emergences of several variants of the virus that could pose an increased risk to global public health which prompted the characterization of specific Variants of Interest (VOI) and Variants of Concern (VOC). For the latest updated list of variants see: WHO Tracking SARS-CoV-2 variants.

Clinical features

- The most common clinical features include: fever, cough, malaise and shortness of breath and other symptoms included in the case definition: general weakness/fatigue, headache, myalgia, sore throat, coryza, anorexia/nausea/vomiting, diarrhoea, altered mental status. In children poor feeding, fussiness, vomiting and stiff neck can also be possible symptoms. The loss of taste and smell is a less common symptom, but seems to be quite specific to, although not exclusively associated with, COVID-19 compared with influenza.
- Clinical features range from mild, moderate ARI and, in some cases, SARI requiring oxygen, and critical disease with respiratory failure, sepsis, septic shock and ARDS with progressive multi-organ failure, thromboembolic disease, requiring intensive care interventions such as non-invasive or invasive mechanical ventilation, dialysis or vasopressors. Overall, the case fatality ratio (CFR) is around 1-2% of all infected patients. For more details per region and country, see the WHO Coronavirus (COVID-19) Dashboard.
- According to data, approximately 80% of symptomatic patients will have mild (40%) to moderate (40%) disease and recover. Moderate disease may include a mild form of pneumonia. Approximately 15% of mild/moderate cases progress to severe disease and an additional approximately 5% become critically ill; 20% of cases will remain asymptomatic. With COVID-19 vaccination, the proportion of patients with severe disease and mortality is decreasing. VOC may also impact the disease severity in COVID-19, see WHO Tracking SARS-CoV-2 variants.
Clinical features continued

- Severe/critical disease (20%) has a higher CFR and has been seen in older persons (> 60 years old) and those with chronic medical conditions, including noncommunicable diseases (NCDs) (e.g. hypertension, cardiac disease, diabetes, chronic lung disease, cerebrovascular disease, dementia, mental health disorders, chronic kidney disease) and some immunosuppressed conditions (e.g. cancer and HIV), obesity, smoking, pregnant or recently pregnant: women > 35 years old, obesity, with chronic medical conditions or pregnancy specific disorders (e.g. gestational diabetes and pre-eclampsia/eclampsia) and unvaccinated against COVID-19; with clinical deterioration occurring at around day 7 of illness.
- Complications such as thromboembolism, myocardial injury, arrhythmias, cardiomyopathy, heart failure and encephalopathy have been reported in severe/critical cases.
- Children appear to mainly have asymptomatic or mild disease, but there are reports of a small number of cases with multisystem inflammatory syndrome in children (MIS-C).
- Bilateral infiltrates and ground-glass changes are the most commonly reported signs on chest X-ray and CT imaging, with lymphopaenia frequently seen in blood tests and elevation of inflammatory markers (e.g. C-reactive protein [CRP], lactate dehydrogenase [LDH], ferritin, erythrocyte sedimentation rate [ESR], interleukin 6 [IL-6], D-dimer).

Prevention

Infection prevention and control and public health interventions
- For all individuals, proper hand washing techniques, respiratory hygiene, social distancing and limiting contact with symptomatic individuals are the main preventive measures (droplet and contact precautions). Universal masking and targeted continuous use of medical masks are recommended in specific transmission scenarios such as crowded indoor spaces with poor ventilation and inability to maintain the correct social distancing; WHO current guidance is provided in Mask use in the context of COVID-19.
- In health care settings, enhanced IPC measures are required when caring for patients with suspected, probable or confirmed COVID-19: including appropriate use of PPE (gown, gloves, medical mask, eye protection), and addition of airborne precautions (N95/FFP2/3) when performing AGP. See also WHO recommendations on mask use by health workers, in light of the Omicron variant of concern.

Vaccines and medications for prophylaxis
- Vaccines to protect from COVID-19 are in available and clinical trials and further developments are ongoing. See COVID-19 vaccines for the most up-to-date information.
- Prophylaxis: see WHO Living guideline: drugs to prevent COVID-19 for the most up-to-date recommendations for the use of drugs to prevent COVID-19.

Treatment

- Early recognition of those patients with (or at risk of) severe disease, and access to critical care interventions are key (Living guidance for clinical management of COVID-19).
- The most updated recommended therapeutics for COVID-19 can be found in Therapeutics and COVID-19: living guideline.
- Diagnosis and treatment of co-infections like respiratory viral, secondary bacterial infections or endemic diseases such as malaria or tuberculosis, which can cause febrile illness are important.
- For symptomatic patients: discharge from hospitals and/or removal of isolation measures can be done 10 days after symptom onset, plus at least 3 days without symptoms (without fever and respiratory symptoms).
- For asymptomatic patients: removal of isolation measures can be done 10 days after positive test for SARS-CoV-2.
# 1.3 Influenza virus fact sheet

## SEASONAL INFLUENZA VIRUSES

### Introduction

- Circulates worldwide causing outbreaks and seasonal epidemics.
- Some immunity already in the population, depending on age and vaccination status.
- Populations at higher risks of severe disease are patients > 65 years old, with chronic conditions: cardiac, respiratory, endocrine, neurological, renal, including metabolic and haematologic disorders, chronic liver disease and other immunosuppressed conditions, as well as children < 59 months, pregnant and post-partum period women.
- Health care workers are at high risk of acquiring influenza virus infection due to increased exposure to patients and risk further spread particularly to vulnerable individuals.
- For updated information on influenza see WHO Influenza [here](#) and Guidelines for the clinical management of severe illness from influenza virus infections [here](#).

### Human infection

Seasonal influenza epidemics are caused by influenza A and B viruses.
- These circulate worldwide and spread easily from person to person.
- Can cause annual epidemics that peak during winter in temperate climates or may circulate year-round in tropical regions.
- Epidemics can result in high levels of work/school absenteeism and productivity losses. Clinics and hospitals can be overwhelmed during peak illness periods worldwide, these annual epidemics are estimated to result in about 3 to 5 million cases of severe illness, and about 290 000 to 650 000 deaths from respiratory causes.
- In high-income countries most deaths associated with influenza occur among people age 65 or older.
- The effects of seasonal influenza epidemics in developing countries are not fully known, but research estimates that 99% of deaths in children under 5 years of age with influenza-related lower respiratory tract infections are found in low-resource countries.

### Transmission

- Influenza viruses are spread mainly via inhalation of respiratory droplets coming from coughing or sneezing of an infected person to a person who is in close contact (within 1 m). Those respiratory droplets can reach the mouth, nose or eyes of a susceptible person and can result in infection.
- Indirect contact transmission involving contact of a susceptible host with a contaminated object or surface (fomite transmission) may also be possible.
- The incubation period is about 2 days. Patients can be infectious 1 day before symptoms appear and up to 1 day after symptoms go away.
- Children shed virus longer than adults.
- The estimated attack rate is 5-20% and higher in densely populated communities and schools.

### Clinical features

- Uncomplicated ARI with high fever, cough and viral syndrome that commonly lasts for 1 week and does not require medical attention.
- Can also cause severe illness with pneumonia, sepsis, ARDS; seen more in patients at high risk (children < 59 months of age, older patients, pregnant woman and those with chronic medical conditions).
Prevention

Infection prevention and control and public health interventions

- For all individuals, proper hand washing techniques, respiratory hygiene, social distancing and limiting contact with symptomatic individuals are the main preventive measures (droplet and contact precautions).
- In healthcare settings, enhanced IPC measures are required when caring for patients with suspected, probable or confirmed influenza infection; including appropriate use of PPE (gown, gloves, medical mask, eye protection) with the addition of airborne precautions (N95/FFP2/3) when performing AGP.

Vaccines and medications for prophylaxis

- Annual vaccination is recommended for pregnant women, children aged 6-59 months, older age (≥65 years), individuals with chronic medical conditions and healthcare workers.
- No medication for prevention has been recommended.

Treatment

- Neuraminidase inhibitors (i.e. oseltamivir) are active against all circulating strains of seasonal influenza and should be given as soon as possible to patients with SARI, and those high-risk patients with uncomplicated ARI. See Guidelines for the clinical management of severe illness from influenza virus infections ( ).

Zoonotic Influenza

Introduction

- Depending on the origin host, humans can be infected with avian, swine and other zoonotic influenza viruses: such as avian influenza virus subtypes A(H5N1), A(H7N9) and A(H9N2), and swine influenza virus subtypes A(H1N1), A(H1N2) and A(H3N2).
- Human infections are rare and primarily acquired through direct contact with infected animals or contaminated environments; all these animal influenza type A viruses are distinct from human influenza viruses and have not acquired the ability of sustained transmission among humans.
- Aquatic birds are the primary natural reservoir for most subtypes of influenza A viruses. Most cause asymptomatic or mild infection in birds, where the range of symptoms depends on the virus properties.
- Human infections with these viruses need to be monitored closely. As the extent of virus circulation in animals is often not clear, epidemiological and virological surveillance and follow up of suspected human cases should remain high. Guidance on investigation of non-seasonal influenza and other emerging acute respiratory diseases can be found on the WHO website: Global Influenza Surveillance and Response System ( ).

Transmission

- Most human cases of influenza A(H5N1) and A(H7N9) virus infection have been associated with direct or indirect contact with infected live or dead poultry.
- Human infections with avian and other zoonotic influenza viruses, though rare, have been reported sporadically.
- Human infections are primarily acquired through direct contact with infected animals or contaminated environments, but do not result in efficient transmission of these viruses between people.

Clinical features

- Avian, swine and other zoonotic influenza virus infections in humans may cause disease ranging from mild upper respiratory tract infection (fever and cough), early sputum production and rapid progression to severe pneumonia, sepsis with shock, ARDS, and even death. Conjunctivitis, gastrointestinal symptoms, encephalitis and encephalopathy have also been reported to varying degrees depending on subtype.
### Prevention

**Infection prevention and control and public health interventions**
- Proper hand washing techniques, respiratory hygiene, social distancing and limiting contact with symptomatic individuals are the main preventive measures (droplet and contact precautions).
- Disease control in animals, avoid direct and prolonged exposure to infected animals.
- In health care settings, enhanced IPC measures are required when caring for patients with suspected, probable or confirmed zoonotic influenza; including appropriate use of PPE (gown, gloves, medical mask, eye protection), and the addition of airborne precautions (N95/FFP2/3) when performing AGP.

**Vaccines and medications for prophylaxis**
- Not available.

**Treatment**
- Early treatment with neuraminidase inhibitor, as soon as possible. See *Guidelines for the clinical management of severe illness from influenza virus infections* ( ).

### NEW INFLUENZA VIRUSES WITH PANDEMIC POTENTIAL

- Any new influenza virus is considered to have the potential to cause a pandemic if:
  - the virus has demonstrated the capacity to infect a human; and
  - the haemagglutinin (HA) gene or protein is not a variant or mutated form of those - A(H1) or A(H3) - circulating widely in the human population.
- A pandemic influenza virus may arise when genes from animal and human influenza viruses mix together to create a human-animal influenza reassortant virus (genetic reassortment); or genes in an animal influenza virus change allowing the virus to infect humans and transmit easily among them (genetic mutation).
- It is mandatory to notify a human influenza case caused by a new subtype to WHO, under the International Health Regulations (IHR) (2005). Because it is a new virus to which people have not been exposed, the population has no or little immunity and the virus is able to spread quickly between people and cause illness.
- Although we start with the assumption that the risk groups for infection and severe outcome are the same as in seasonal influenza, there might be differences. Historical knowledge from the 1918 and 2009 pandemics indicates that healthy, young adults can be disproportionately and more severely affected.
- Pandemic influenza might present differently from seasonal influenza and symptoms may be more severe and complications more frequent.
### 1.4 Middle East respiratory syndrome coronavirus (MERS-CoV) fact sheet

**MERS-CoV**

### Introduction
- Type of coronavirus whose primary reservoir is dromedary camels, with origination in bats (similar strains isolated from camels in Egypt, Oman, Qatar and Saudi Arabia).
- First case reported in March 2012 (Saudi Arabia).
- Since then, cases have been reported in 27 countries. 83% of the cases have been in Saudi Arabia. There was a large outbreak in the Republic of Korea in 2015; and moderate numbers have occurred in Jordan, Oman, Qatar and the United Arab Emirates.
- To date, there are 2578 laboratory-confirmed cases and 888 deaths (October 2021).

### Transmission
- Camel-human transmission route is unknown.
- Human-human transmission has been limited to health care settings when inadequate IPC measures occurred during close contact with infected individual (nosocomial transmission).
- No sustained community transmission reported.

### Clinical features
- Ranges from asymptomatic to mild ARI: fever, cough and shortness of breath. Pneumonia is common, but not always present. Gastrointestinal symptoms, including diarrhoea, have also been reported. In some cases, it can present as SARI with progressive organ failure, sepsis and ARDS.
- Some laboratory-confirmed cases of MERS-CoV infection have been reported as asymptomatic, following aggressive contact tracing of a laboratory-confirmed case.
- Approximately 35% of reported patients with MERS-CoV infection died.
- More severe disease in older people, immunosuppressed and those with chronic medical conditions.

### Prevention
**Infection prevention and control and public health interventions**
- Proper hand washing techniques, respiratory hygiene, social distancing and limiting contact with symptomatic individuals are the main preventive measures (droplet and contact precautions).
- When visiting areas where camels are present, use proper hand washing techniques.
- Avoid contact with sick camels.
- Avoid eating raw meat or unpasteurized milk.
- In health care settings, enhanced IPC measures are required when caring for patients with suspected, probable or confirmed MERS-CoV; including appropriate use of PPE (gown, gloves, medical mask, eye protection), and the addition of airborne precautions (N95/FFP2/3) when performing AGP.

### Vaccines and medications for prophylaxis
- Not available.

### Treatment
- Experimental protocols for treatment are available but no medication approved (unavailable).
### 1.5 Risk factors for severe disease: influenza vs COVID-19

#### INFLUENZA

<table>
<thead>
<tr>
<th>Chronic conditions, including:</th>
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<tbody>
<tr>
<td>• cardiac conditions (hypertension and cardiovascular disease)</td>
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<tr>
<td>• chronic lung conditions (asthma or chronic obstructive pulmonary disease [COPD]), endocrine disorders (diabetes)</td>
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<tr>
<td>• neurological disorders (stroke and neurodevelopmental conditions)</td>
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<tr>
<td>• chronic kidney disease</td>
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<td>• metabolic disorders</td>
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<tr>
<td>• haematologic disorders</td>
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<tr>
<td>• chronic liver disease</td>
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<tr>
<td>• other immunosuppressed conditions, including cancer and HIV/AIDS and chronic conditions requiring immunosuppressive therapy, such as chronic steroid treatment or chemotherapy</td>
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#### COVID-19

<table>
<thead>
<tr>
<th>Chronic conditions, including:</th>
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<tbody>
<tr>
<td>• cardiac conditions (hypertension and cardiovascular disease)</td>
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<tr>
<td>• chronic neurological disorders (e.g. stroke)</td>
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<tr>
<td>• dementia or mental health disorders</td>
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<tr>
<td>• chronic lung disease (e.g. COPD)</td>
</tr>
<tr>
<td>• endocrine disorders (diabetes)</td>
</tr>
<tr>
<td>• chronic kidney disease</td>
</tr>
<tr>
<td>• some immunosuppressed conditions (e.g. cancer, HIV)</td>
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#### Obesity

#### Smoking

In pregnant or recently pregnant: women > 35 years old, obesity, with chronic medical conditions or pregnancy specific disorders (e.g. gestational diabetes and pre-eclampsia/eclampsia)

#### Unvaccinated against COVID-19

#### Older persons (> 60 years old)

#### Young children (< 59 months)

**Note:**

- List of risk factors to be updated as evidence emerges.
References and resources


  • Clinical care
  • Country-level coordination, planning, and monitoring
  • Critical preparedness, readiness and response actions for COVID-19
  • Essential resource planning
  • Guidance for schools, workplaces and institutions
  • Health workers
  • Humanitarian operations, camps, refugees/migrants in non-camps and other fragile settings
  • Infection prevention and control/WASH
  • Maintaining essential health services and systems
  • Naming the coronavirus disease (COVID-19)
  • National laboratories
  • Risk communication and community engagement
  • Serology and early investigation protocols
  • Surveillance, rapid response teams, and case investigation
  • Travel, points of entry and border health
  • Virus origin/reducing animal-human transmission


1. Epidemiology of SARI


Screening, triage, clinical assessment and management of SARI
2 Screening, triage, clinical assessment and management of SARI

Summary

The screening and identification of patients with acute respiratory symptoms, mainly SARI, at all points of access to the health system, including primary health centres, clinics, hospital emergency units and community settings, is a priority when such symptoms may be due to pathogens of high concern, such as influenza and/or SARS-CoV-2.

At all first points of access to the health system, it is important to always apply appropriate IPC precautions to prevent the spread of illness to health care workers or other patients. See Chapter 3 for details.

1. It is fundamental to set up systems for reporting and referral of cases from the community to the appropriate destination for triage, clinical assessment, testing, and/or treatments as per local clinical care pathways and protocols.

2. After screening at the point of access, triage should be done with a validated acuity-based triage tool to identify and treat patients needing immediate care. Do not delay emergency care.

3. After triage, care for all patients should be delivered in designated areas, according to disease severity and acute care needs. Generally, patients with SARI need acute hospital care because of possible complications such as severe pneumonia, sepsis, organ dysfunction, and/or exacerbation of chronic disease or co-infections. These patients can progress to acute organ failure that may require critical care and admission to ICU for closer monitoring and supportive therapies that cannot be delivered on a general ward. Do not delay ICU admission for these patients.

For example, suspected COVID-19 patients with mild or moderate disease and without risk factors should be instructed to self-isolate and contact the COVID-19 information line or a health care provider for advice on testing and referral if needed. These patients can be isolated (or cohorted) at a health facility (if resources allow), community facility with rapid access to health advice, or at home (see WHO guidance) according to local care pathways.

Patients with suspected or confirmed COVID-19 with mild or moderate disease and with risk factors should be instructed to self-isolate and call the COVID-19 information line or a health care provider for advise on testing and referral if needed. These patients can be isolated (or cohorted) at a health facility (if resources allow), community facility with rapid access to health advice, or at home (see WHO guidance) according to local care pathways.
Patients with suspected or confirmed COVID-19 with severe or critical disease should be instructed to call the emergency hotline (such as the COVID-19 information line for emergency referrals) as soon as possible and be isolated and transferred to designated hospital for inpatient care.

See WHO guidance for more details:
- Home care for patients with suspected or confirmed COVID-19 and management of their contacts
- COVID-19 home care bundle for health care workers
- Coronavirus disease (COVID-19): home care for families and caregivers
- Living guidance for clinical management of COVID-19

Tools
2.1 Screening for SARI
   2.1.1 Case definition for suspected cases of COVID-19
   2.1.2 Clinical classification for respiratory infection due to influenza
2.2 Triage for SARI
   2.2.1 Setting up a triage/resuscitation area in a health care facility: rationale and requirements
   2.2.2 Examples of acuity-based triage tools: Interagency Integrated Triage Tool
2.3 Clinical assessment of acutely ill patients - basic emergency care: ABCDE approach
2.4 Classification of severity in patients with COVID-19
2.5 Decision-making algorithm for patients presenting with acute respiratory infection (ARI) (influenza or COVID-19 suspected or known to be circulating)
2.6 Decision-making algorithm for hospitalization of patients with pneumonia (influenza or COVID-19 known to be circulating)
2.7 Decision-making support tool for hospitalization and ICU admission for patients with SARI and severe pneumonia
2.8 Checklist for admission
2.9 Checklist for transfer
2.10 Transfer of critically ill patients: air medevac for COVID-19 patients
2.1 Screening for SARI

Screening is the process by which a patient is evaluated to see whether they meet a standardized case definition. It can be performed by different types of health workers and in many different areas such as the emergency unit, outpatient department/primary care clinic, in the community by a community health worker or by telemedicine.

During an epidemic, screening protocols should be established at all health care access points and during contact tracing activities. The screening questions may need to be adjusted for certain settings and guided by epidemiological considerations.

As a precaution, screening should always be done at a distance (> 1 m), with no physical contact.

For example, in the context of a COVID-19 outbreak, screening consists of a simple set of questions based on the WHO COVID-19 case definition (fever, cough, dyspnoea) (4). This screening has a simple binary outcome: yes or no.

A person screened positive becomes a suspected case and enters the COVID-19 care pathway. The patient should immediately be given a medical mask and referred to the next appropriate site (health facility for medical care and/or isolation, community facility for medical care or isolation, home for self-isolation, laboratory site for testing, and medical care).

A person screened negative can continue on the normal patient pathway (non-COVID-19) within the health care system without any special precaution.

COVID-19 screening helps ensure the safety of patients, staff and the community and is the first step in the COVID-19 care pathway.

![Screen for COVID-19 at first point of access to the health system](image)

All patients should be screened for COVID-19 using WHO case definitions at the first point they access the health system.

<table>
<thead>
<tr>
<th>Patients suspected to have COVID-19</th>
<th>Patients NOT suspected to have COVID-19</th>
</tr>
</thead>
<tbody>
<tr>
<td>Refer to appropriate facility or testing site as per local protocol</td>
<td>Management as per local protocol (routine management or referral as per reorganization of service delivery)</td>
</tr>
</tbody>
</table>

2.1.1 Case definition for suspected cases of COVID-19

A. Clinical and epidemiological criteria

**Clinical criteria:**
- Acute onset of fever AND cough; or
- Acute onset of any three or more of the following signs or symptoms: fever, cough, general weakness/fatigue, headache, myalgia, sore throat, coryza, dyspnoea, anorexia/nausea/vomiting, diarrhoea, altered mental status.

**Epidemiological criteria:**
- Residing in or working in an area with high risk of transmission of virus: closed residential settings, humanitarian settings such as camps and camp-like settings for displaced persons; any time within the 14 days prior to symptom onset; or
- Residing in or travel to an area with community transmission any time within the 14 days prior to symptom onset; or
- Working in any health care setting, including within health facilities or within the community; any time within the 14 days prior of symptom onset.

B. A patient with severe acute respiratory illness (SARI)

- acute respiratory infection with history of fever or measured fever of ≥ 38 °C;
- cough;
- onset within the last 10 days; and
- requires hospitalization.

C. A person NOT meeting the clinical AND epidemiological criteria

- Asymptomatic person not meeting epidemiological criteria with a positive SARS-CoV-2 antigen-RDT.

Checklist for COVID-19 neurological screening

COVID-19 may present with neurological symptoms with or without respiratory signs, and patients with COVID-19 are at increased risk for neurological manifestations.

How to use this screening checklist:
- In addition to standardized screening questions, the COVID-19 neurological screening checklist may be used to identify patients who may present with non-respiratory symptoms of COVID-19.
- Additionally, this checklist may be used while monitoring patients with COVID-19 for new or emerging neurological symptoms.

<table>
<thead>
<tr>
<th>Adult screening checklist</th>
</tr>
</thead>
<tbody>
<tr>
<td>Loss (ageusia) or decreased sense of taste (dysgeusia)</td>
</tr>
<tr>
<td>Loss (anosmia) or decreased sense of smell (hyposmia)</td>
</tr>
<tr>
<td>New onset headache</td>
</tr>
<tr>
<td>New onset dizziness (including pre-syncopal symptoms or syncope)</td>
</tr>
<tr>
<td>Confusion (delirium), behavioural changes including agitation, or psychotic symptoms</td>
</tr>
<tr>
<td>Decreased consciousness (e.g. somnolence, stupor, unresponsiveness)</td>
</tr>
<tr>
<td>New focal or diffuse muscle pain (myalgias) with or without weakness</td>
</tr>
<tr>
<td>Worsening of underlying neurological condition</td>
</tr>
<tr>
<td>Myoclonus</td>
</tr>
<tr>
<td>New onset seizures or increased seizure activity</td>
</tr>
<tr>
<td>New onset weakness (paresis, hemiparesis)</td>
</tr>
<tr>
<td>Neck stiffness (meningismus)</td>
</tr>
<tr>
<td>Trouble with speech</td>
</tr>
<tr>
<td>Numbness/tingling (sensory symptoms)</td>
</tr>
<tr>
<td>New onset of fatigue</td>
</tr>
<tr>
<td>Blurring/loss of vision</td>
</tr>
<tr>
<td>Problems passing urine or with bowel movements</td>
</tr>
<tr>
<td>Problems with balance in standing or walking</td>
</tr>
</tbody>
</table>
Special considerations in infants and children

- COVID-19 infection is frequently asymptomatic and requires a high index of suspicion when screening.
- Neurologic symptoms of COVID-19 infection in children include poor feeding, fussiness, vomiting, stiff neck.
- Multisystem inflammatory syndrome in children (MIS-C) due to COVID-19 may present with serious neurological symptoms.

This checklist was developed by members of the WHO Neurology and COVID-19 Forum Acute Clinical Care and Support Working Group. Working group members generated a list of symptoms based on their clinical and academic experience with COVID-19. In order to identify all relevant symptoms for inclusion in this checklist, an auditing process was carried out using meta-analytic data from a forthcoming WHO review (manuscript in progress) with a pre-defined threshold for inclusion and exclusion. Symptoms for which there are at least five cases reported in peer-reviewed literature in at least two different sites met the threshold for inclusion. After this process, each symptom was discussed within the working group and final decisions were reached by consensus.
Clinical classification for respiratory infection due to influenza

<table>
<thead>
<tr>
<th>Clinical syndrome</th>
<th>Definition</th>
</tr>
</thead>
<tbody>
<tr>
<td>Influenza-like infection (ILI)</td>
<td>Acute onset within the last 10 days following respiratory symptoms, measured fever of ≥ 38 °C and cough.</td>
</tr>
<tr>
<td>Acute respiratory infection (ARI)</td>
<td>Sudden onset of respiratory infection symptoms (cough, sore throat shortness of breath, or coryza).</td>
</tr>
<tr>
<td>Severe acute respiratory infection (SARI)</td>
<td>Acute respiratory infection with a history of fever or measured fever of ≥ 38 °C and cough with onset within the last 10 days and requires overnight hospitalization.</td>
</tr>
</tbody>
</table>

Signs and symptoms of uncomplicated influenza-like illness (ILI)

- Fever
- Cough
- Sore throat
- Rhinorrhoea or nasal congestion
- Headache
- Muscle pain or malaise
- Gastrointestinal illness such as diarrhoea or vomiting, but no evidence of dehydration
- No shortness of breath.

*Note:* Older people and immunosuppressed may present with atypical symptoms. Symptoms due to physiologic adaptations of pregnancy and adverse pregnancy events, such as dyspnoea, fever, gastrointestinal symptoms or fatigue, may overlap with ILI.

Signs and symptoms of complicated ARI (SARI)

- Respiratory distress: fast breathing, shortness of breath, accessory muscle use, cyanosis.
  - In children: central cyanosis, severe distress, grunting, severe chest indrawing or danger signs of lethargy, convulsions or inability to breastfeed or drink.
- Cardiovascular distress:
  - Adult: low blood pressure (SBP < 100); delayed capillary refill (> 3 sec < 65 years or > 4.5 sec in older people); fast and weak pulse.
  - Child: delayed capillary refill (> 3 sec); fast and weak pulse; or cool extremities or hypotension.
- Neurologic distress: alteration in mental status such as coma, lethargy, confusion, seizures, agitation.
- Dehydration:
  - In children: diarrhoea plus any of the following signs: lethargy, sunken eyes, very slow skin pinch, unable to drink or drinks poorly.
  - Persistent fever that is not responding after 3 days.

Source: Revision of clinical case definitions: influenza-like illness and severe acute respiratory infection (WHO, 2020).
2.2 Triage for SARI

Triage refers to the sorting of patients by priority after screening, based on specific criteria (e.g. severity) and can be performed at any point of access to the health care system, including in both pre-hospital and facility-based settings.

**Acuity-based triage** is the action of sorting and prioritizing patients based on the estimation of their severity. This is used as the basis for identification of those patients who require immediate medical intervention and those who can safely wait, or those who may need to be transported to a specific destination based on their condition. The concept of triage has been around for a long time and different triage tools have been created over the years. Acuity-based triage is the standard method of sorting patients in medical settings.

22.1 Setting up a triage/resuscitation area in a health care facility: rationale and requirements

**DESIGNATING A RESUSCITATION AREA IN THE EMERGENCY UNIT**

**Rationale:** Early recognition of conditions requiring time-sensitive management saves lives. A standardized approach in a designated resuscitation area ensures that the sickest patients in the emergency unit are clearly identified and receive necessary life-saving care. Dedicated resuscitation areas ensure that essential resources are accessible and providers are aware of critical patients as soon as possible.

**Use of the resuscitation area**

- Upon arrival to the emergency unit, all patient are triaged.
- Patients triaged as "red" are immediately transferred to the resuscitation area.
- Triage personnel alert emergency unit staff as patients are transferred to the resuscitation area (e.g. overhead announcement).
- Pre-hospital providers or bystanders remain until report is given to the receiving medical team.
- Patients in the resuscitation area are the top staff priority.
- The emergency unit ideally has other staff not assigned to the resuscitation area that continue to care for lower acuity patients.
- Initial assessment and resuscitation are followed by monitoring and re-evaluation.
- After initial resuscitation, team leader releases additional providers to care for other patients.
- Care plan (diagnostic, management and disposition) is developed before the team leaves the resuscitation area.

**Space**

- For rapid reception of patients, the resuscitation area is easily accessible to the main emergency unit entry areas including the ambulance entrance, main entrance and triage area.
- The resuscitation area is easily visible from the main nurses’ station and physicians’ work area, and staff are aware of its location and function.
- The resuscitation area has enough space to accommodate multiple providers and equipment.
- Easy access to radiology, operating theatre and ICU.

**Staff**

- Staff trained in resuscitation respond immediately to the resuscitation area.
- At the beginning of each shift, resuscitation area providers (doctors, nurses, technicians) are clearly identified and this information is communicated on a centrally visible whiteboard, chalkboard or monitor.

**Equipment and supplies**

The resuscitation area is equipped to handle critically ill patients at all times:
- Equipment to check and monitor vital signs.
- Medications cart equipped with key medications for critically ill patients (e.g. fluids, glucose, pain meds).
- Supplies (e.g. blood vials, IV kits, bandages, needles).
- Equipment for critical procedures (e.g. instruments, ultrasound if available).
- Equipment for airway emergencies (e.g. nasal and oral airways, intubation supplies, oxygen, bag valve masks).
- Crash/code cart with advanced life support medications and defibrillator.
- Essential utilities (e.g. electricity, lights, running water).
- Charting and documentation equipment and supplies.
- PPE for providers (e.g. gloves, gowns) and disposal area for sharps, infectious and non-infectious waste.
- Patient gowns and bed linens.
**2.2.2 Examples of acuity-based triage tools: Interagency Integrated Triage Tool**

Interagency Integrated Triage Tool (IITT) is a novel triage tool developed in collaboration between WHO, the International Committee of the Red Cross (ICRC) and Médecins Sans Frontières (MSF) to provide an integrated set of protocols for routine triage of adults and children. It can be utilized for facility-based triage, facility-based mass casualty triage and pre-hospital triage.

The tool was developed in 2019 and focuses on a three-tier triage system:
- high-acuity resuscitation area
- clinical treatment area
- low-acuity or waiting room.

Based on red, yellow and green criteria.
Interagency Integrated Triage Tool: > 12 years

1. CHECK FOR RED CRITERIA

- Unresponsive
- AIRWAY & BREATHING
  - Stridor
  - Respiratory distress* or central cyanosis
- CIRCULATION
  - Capillary refill >3 sec
  - Weak and fast pulse
  - Heavy bleeding
  - HR <50 or >120
- DISABILITY
  - Active convulsions
    - Any two of:
      - Altered mental status
      - Stiff neck
      - Hypoglycaemia
- OTHER
  - High-risk trauma*
  - Poisoning/ingestion or dangerous chemical exposure*
  - Threatened limb*
  - Snake bite
  - Acute chest or abdominal pain (>50 years old)
  - ECG with acute ischaemia (if done)
  - Violent or aggressive
- PREGNANT WITH ANY OF:
  - Heavy bleeding
  - Severe abdominal pain
  - Seizures or altered mental status
  - Severe headache
  - Visual changes
  - SBP ≥160 or DBP ≥110
  - Active labour
  - Trauma

2. CHECK FOR YELLOW CRITERIA

- AIRWAY & BREATHING
  - Any swelling/mass of mouth, throat or neck
  - Wheezing (no red criteria)
- CIRCULATION
  - Vomits everything or ongoing diarrhoea
  - Unable to feed or drink
  - Severe pallor (no red criteria)
  - Ongoing bleeding (no red criteria)
  - Recent fainting
- DISABILITY
  - Altered mental status or agitation (no red criteria)
  - Acute general weakness
  - Acute focal neurologic complaint
  - Acute visual disturbance
  - Severe pain (no red criteria)
- OTHER
  - New rash worsening over hours or peeling (no red criteria)
  - Visible acute limb deformity
  - Open fracture
  - Suspected dislocation
  - Other trauma/burns (no red criteria)
  - Known diagnosis requiring urgent surgical intervention
  - Sexual assault
  - Acute testicular/scrotal pain or priapism
  - Unable to pass urine
  - Exposure requiring time-sensitive prophylaxis (eg. animal bite, needlestick)
  - Pregnancy, referred for complications

Patients with high-risk vital signs or clinical concern need up-triage or immediate review by supervising clinician.

3. CHECK FOR HIGH-RISK VITAL SIGNS

<table>
<thead>
<tr>
<th>CRITERIA</th>
<th>VALUE</th>
</tr>
</thead>
<tbody>
<tr>
<td>HR &lt;60 or &gt;130</td>
<td></td>
</tr>
<tr>
<td>RR &lt;10 or &gt;30</td>
<td></td>
</tr>
<tr>
<td>Temp &lt;36° or &gt;39°C</td>
<td></td>
</tr>
<tr>
<td>SpO2 &lt;92%</td>
<td></td>
</tr>
<tr>
<td>AVPU other than A</td>
<td></td>
</tr>
</tbody>
</table>

NO

MOVE TO HIGH ACUITY RESUSCITATION AREA IMMEDIATELY

YES

MOVE TO CLINICAL TREATMENT AREA

NO

MOVE TO LOW ACUITY OR WAITING AREA

*See Reference Card

Developed by World Health Organization, The International Committee of the Red Cross, Médecins Sans Frontières
Interagency Integrated Triage Tool: < 12 years

**CHECK FOR RED CRITERIA**
- Unresponsive
  - AIRWAY & BREATHING
    - Stridor
    - Respiratory distress* or central cyanosis
  - CIRCULATION
    - Capillary refill >3 sec
    - Weak and fast pulse
    - Heavy bleeding
    - Cold extremities
    - Any two of:
      - Lethargy
      - Sunken eyes
      - Very slow skin pinch
      - Drinks poorly
  - DISABILITY
    - Active convulsions
    - Altered mental status (confused, restless, continuously irritable or lethargic) with stiff neck, hypothermia or fever
    - Hypoglycaemia (if known)
  - OTHER
    - Any infant <8 days old
    - Age <2 months and temp <36 or >39°C
    - High-risk trauma*
    - Threatened limb*
    - Acute testicular/scrotal pain or priapism
    - Snake bite
    - Poisoning/ingestion or dangerous chemical exposure*
    - Pregnant with adult red criteria

**CHECK FOR YELLOW CRITERIA**
- AIRWAY & BREATHING
  - Any swelling/mass of mouth, throat or neck
  - Wheezing (no red criteria)
- CIRCULATION
  - Unable to feed or drink
  - Vomits everything
  - Ongoing diarrhoea
  - Dehydration
  - Severe pallor (no red criteria)
- DISABILITY
  - Restless, continuously irritable or lethargy
  - Severe pain
- OTHER
  - Any infant 8 days to 6 months old
  - Malnutrition with visible severe wasting OR oedema of both feet
  - Trauma/burn (no red criteria)
  - Sexual assault
  - Known diagnosis requiring urgent surgical intervention
  - New rash worsening over hours or peeling (no red criteria)
  - Exposure requiring time-sensitive prophylaxis (e.g., animal bite)
  - Pregnancy (no red criteria)
  - Headache (no red criteria)

**CHECK FOR HIGH-RISK VITAL SIGNS**
- Temp <36° or >39°C
- SpO2 < 92%
- AVPU other than A

**MOVE TO HIGH ACUITY RESUSCITATION AREA IMMEDIATELY**

**MOVE TO CLINICAL TREATMENT AREA**

**MOVE TO LOW ACUITY OR WAITING AREA**

Patients with high-risk vital signs or clinical concern need up-triage or immediate review by supervising clinician.

*See Reference Card

Developed by World Health Organization, International Committee of the Red Cross, Médecins Sans Frontières
### High-Risk Trauma Criteria

<table>
<thead>
<tr>
<th>General Trauma</th>
<th>Road Traffic</th>
</tr>
</thead>
<tbody>
<tr>
<td>Fall from twice person’s height</td>
<td>High speed motor vehicle crash</td>
</tr>
<tr>
<td>Penetrating trauma excluding distal to knee/elbow with bleeding controlled</td>
<td>Pedestrian or cyclist hit by vehicle</td>
</tr>
<tr>
<td>Crush injury</td>
<td>Other person in same vehicle died at scene</td>
</tr>
<tr>
<td>Polytrauma (injuries in multiple body areas)</td>
<td>Motor vehicle crash without a seatbelt</td>
</tr>
<tr>
<td>Patient with bleeding disorder or on anticoagulation</td>
<td>Trapped or thrown from vehicle (including motorcycle)</td>
</tr>
<tr>
<td>Pregnant</td>
<td></td>
</tr>
</tbody>
</table>

#### Major Burns

<table>
<thead>
<tr>
<th>Criteria</th>
<th>Signs</th>
</tr>
</thead>
<tbody>
<tr>
<td>Greater than 15% body surface area</td>
<td>Inhalation injury</td>
</tr>
<tr>
<td>Circumferential or involving face or neck</td>
<td>Any burn in age &lt;2 or age &gt;70</td>
</tr>
</tbody>
</table>

#### Threatened Limb

A patient presenting with a limb that is:
- Pulseless
- Painful and one of the following: pale, weak, numb, or with massive swelling after trauma.

### Other High-Risk Criteria

#### Signs of Respiratory Distress

<table>
<thead>
<tr>
<th>Adult</th>
<th>Child</th>
</tr>
</thead>
<tbody>
<tr>
<td>Very fast or very slow breathing</td>
<td>Very fast breathing</td>
</tr>
<tr>
<td>Inability to talk or walk unaided</td>
<td>Inability to talk, eat or breastfeed</td>
</tr>
<tr>
<td>Confused, sleepy or agitated</td>
<td>Nasal flaring, grunting</td>
</tr>
<tr>
<td>Accessory muscle use (neck, intercostal, abdominal)</td>
<td>Accessory muscle use (e.g. head nodding, chest indrawing)</td>
</tr>
</tbody>
</table>

#### Ingestion/exposure

Use of clinical signs alone may not identify all those who need time-dependent intervention. Patients with high risk ingestion or exposure should initially be up-triaged to Red for early clinical assessment.

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2.3 Clinical assessment of acutely ill patients - basic emergency care: ABCDE approach

Basic emergency care (BEC): ABCDE approach to the acutely ill

Developed by WHO and ICRC in collaboration with the International Federation for Emergency Medicine, Basic emergency care (BEC): approach to the acutely ill and injured is an open-access training course for frontline health care providers who manage acute illness and injury with limited resources, including students, nurses, pre-hospital technicians, clinical officers and doctors who are working in field (pre-hospital) or hospital settings.

BEC integrates guidance from WHO Emergency triage assessment and treatment (ETAT) and the Integrated management of adolescent and adult illness (IMAI) district clinician manual and teaches a systematic approach to the initial assessment and management of four time-sensitive conditions - difficulty in breathing, shock, altered mental status and injury - where early intervention saves lives.

Because emergency care providers must respond to “undifferentiated” patients, those with acute symptoms for which the cause may not be known, BEC teaches a simple, systematic ABCDE approach to managing acute, potentially life-threatening conditions even before a diagnosis is known.

Patients who are acutely ill due to a SARI may present with any of three life-threatening conditions: difficulty in breathing, shock or altered mental status. The following “quick cards” from BEC summarize the initial approach to assessment and management of key findings from the ABCDE approach.

See: Basic emergency care (BEC): approach to the acutely ill and injured or WHO Emergency care or contact emergencycare@who.int for more information.
<table>
<thead>
<tr>
<th>ASSESSMENT FINDINGS</th>
<th>IMMEDIATE MANAGEMENT</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Airway</strong> Unconscious with limited or no air movement</td>
<td>If <strong>NO TRAUMA</strong>: head-tilt and chin-lift, use OPA or NPA to keep airway open, place in recovery position or position of comfort. If possible <strong>TRAUMA</strong>: use jaw thrust with c-spine protection and place OPA to keep the airway open (no NPA if facial trauma).</td>
</tr>
</tbody>
</table>
| Foreign body in airway | Remove visible foreign body. Encourage coughing.  
- If **unable** to cough: chest/abdominal thrusts/back blows as indicated.  
- If patient becomes unconscious: CPR. |
| Gurgling | Open airway as above, suction (avoid gagging). |
| Stridor | Keep patient calm and allow position of comfort.  
- For signs of anaphylaxis: give IM adrenaline.  
- For hypoxia: give oxygen. |
| **Breathing** Signs of abnormal breathing or hypoxia | Give oxygen. Assist ventilation with BVM if breathing NOT adequate. |
| Wheeze | Give salbutamol. For signs of anaphylaxis: give IM adrenaline. |
| Signs of tension pneumothorax (absent sounds/hyperresonance on one side WITH hypotension, distended neck veins) | Perform needle decompression, give oxygen and IV fluids. Will need chest tube. |
| Signs of opiate overdose (AMS and slow breathing with small pupils) | Give naloxone. |
| **Circulation** Signs of poor perfusion/shock | If **no pulse**, follow relevant CPR protocols.  
Give oxygen and IV fluids. |
| Signs of internal or external bleeding | Control external bleeding. Give IV fluids. |
| Signs of pericardial tamponade (poor perfusion with distended neck veins and muffled heart sounds) | Give IV fluids, oxygen.  
Will need rapid pericardial drainage. |
| **Disability** Altered mental status (AMS) | If **NO TRAUMA**, place in recovery position. |
| Seizure | Give benzodiazepine. |
| Seizure in pregnancy (or after recent delivery) | Give magnesium sulphate. |
| Hypoglycaemia | Give glucose if < 3.5 mmol/L or unknown. |
| Signs of opiate overdose (AMS with slow breathing with small pupils) | Give naloxone. |
| Signs of life-threatening brain mass or bleed (AMS with unequal pupils) | Raise head of bed, monitor airway.  
Will need rapid transfer for neurosurgical services. |
| **Exposure** Remove wet clothing and dry skin thoroughly | |
| Remove jewellery, watches and constrictive clothing | |
| Prevent hypothermia and protect modesty | |
| Snake bite | Immobilize extremity. Send picture of snake with patient. Call for anti-venom if relevant. |

**If cause unknown, remember trauma:** Examine the entire body and always consider hidden injuries [see also TRAUMA card]

**REMEMBER:** PATIENTS WITH ABNORMAL ABCDE FINDINGS MAY NEED RAPID HANDOVER/TRANSFER. PLAN EARLY.
NORMAL ADULT VITAL SIGNS

- **Pulse rate:** 60–100 beats per minute
- **Respiratory rate:** 10–20 breaths per minute
- **Systolic blood pressure:** > 90 mmHg
- **Oxygen saturation:** > 92%

Estimating systolic blood pressure (not reliable in children and the elderly):
- Carotid (neck) pulse \( \rightarrow \) SBP ≥ 60 mmHg
- Femoral (groin) pulse \( \rightarrow \) SBP ≥ 70 mmHg
- Radial (wrist) pulse \( \rightarrow \) SBP ≥ 80 mmHg

SPECIAL CONSIDERATIONS IN THE ASSESSMENT OF CHILDREN

- Children have bigger heads and tongues, and shorter, softer necks than adults. Position airway as appropriate for age.
- Always consider foreign bodies.

- Look for signs of increased work of breathing (e.g., chest indrawing, retractions, nasal flaring).
- Listen for abnormal breath sounds (e.g., grunting, stridor, or silent chest).

<table>
<thead>
<tr>
<th>AGE</th>
<th>RESPIRATORY RATE (breaths per minute)</th>
</tr>
</thead>
<tbody>
<tr>
<td>&lt; 2 months</td>
<td>40–60</td>
</tr>
<tr>
<td>2–12 months</td>
<td>25–50</td>
</tr>
<tr>
<td>1–5 years</td>
<td>20–40</td>
</tr>
</tbody>
</table>

- Signs of poor perfusion in children include: slow capillary refill, decreased urine output, lethargy, sunken fontanelle, poor skin pinch.
- Look for signs of anaemia and malnourishment (adjust fluids).
- Remember that children may not always report trauma and may have serious internal injury with few external signs.

<table>
<thead>
<tr>
<th>AGE (in years)</th>
<th>NORMAL HEART RATE (beats per minute)</th>
</tr>
</thead>
<tbody>
<tr>
<td>&lt;1</td>
<td>100–160</td>
</tr>
<tr>
<td>1–3</td>
<td>90–150</td>
</tr>
<tr>
<td>4–5</td>
<td>80–140</td>
</tr>
</tbody>
</table>

- Always check AVPU (alert, verbal, pain, unresponsive). Check for tone and response to stimulus.
- Hypoglycaemia is common in ill children. Look for lethargy or irritability.

INFANTS AND CHILDREN HAVE DIFFICULTY MAINTAINING TEMPERATURE

- Remove wet clothing and dry skin thoroughly. Place infants skin-to-skin when possible.
- For hypothermia, cover the head (but be sure mouth and nose are clear).
- For hyperthermia, unbundle tightly wrapped babies.

DANGER SIGNS IN CHILDREN

- Signs of airway obstruction (unable to swallow saliva/drooling or stridor).
- Increased breathing effort (fast breathing, nasal flaring, grunting, chest indrawing or retractions).
- Cyanosis (blue colour of the skin, especially at the lips and fingertips).
- Altered mental status (including lethargy or unusual sleepiness, confusion, disorientation).
- Moves only when stimulated or no movement at all (AVPU other than “A”).
- Not feeding well, cannot drink or breastfeed or vomiting everything.
- Seizures/convulsions.
- Low body temperature (hypothermia).

ESTIMATED WEIGHT in KILOGRAMS for CHILDREN 1–10 YEARS OLD:

\[ \text{[age in years + 4] \times 2} \]

### 2.4 Classification of severity in patients with COVID-19

#### Population

<table>
<thead>
<tr>
<th>Disease severity</th>
<th>Non-severe</th>
<th>Severe</th>
<th>Critical</th>
</tr>
</thead>
<tbody>
<tr>
<td>Patients with confirmed covid-19</td>
<td>Absence of signs of severe or critical disease</td>
<td>Oxygen saturation &lt;90% on room air</td>
<td>Requires life sustaining treatment</td>
</tr>
<tr>
<td></td>
<td>Signs of pneumonia</td>
<td>Acute respiratory distress syndrome</td>
<td></td>
</tr>
<tr>
<td></td>
<td>Signs of severe respiratory distress</td>
<td>Sepsis</td>
<td></td>
</tr>
<tr>
<td></td>
<td></td>
<td>Septic shock</td>
<td></td>
</tr>
</tbody>
</table>

Source: A living WHO guideline on drugs for COVID-19 (BMJ, 2020)

#### Mild disease

Symptomatic patients meeting the case definition for COVID-19 without evidence of viral pneumonia or hypoxia.

#### Moderate disease

**Pneumonia**

Adolescent or adult with clinical signs of pneumonia (fever, cough, dyspnoea, fast breathing) but no signs of severe pneumonia, including SpO₂ ≥ 90% on room air.

Child with cough or difficulty breathing + fast breathing and/or chest indrawing and no signs of severe pneumonia.

Fast breathing: < 2 months: ≥ 60 breaths/min; 2-11 months: ≥ 50; 1-5 years: ≥ 40.

The diagnosis can be made on clinical grounds; chest imaging (radiograph, CT scan, ultrasound) may assist in diagnosis and identify or exclude pulmonary complications.

**Caution:** The oxygen saturation threshold of 90% to define severe COVID-19 is arbitrary and should be interpreted cautiously. For example, clinicians must use their judgment to determine whether a low oxygen saturation is a sign of severity or is normal for a given patient with chronic lung disease. Similarly, a saturation between 90-94% on room air may be abnormal (in patient with normal lungs) and can be an early sign of severe disease, mainly if patient is on a downward trend. Generally, if there is any doubt, the panel suggested erring on the side of considering the illness as severe.

#### Severe disease

Adolescent or adult with clinical signs of pneumonia (fever, cough, dyspnoea) plus one of the following: respiratory rate > 30 breaths/min, severe respiratory distress, or SpO₂ < 90% on room air.

Child: with clinical signs of pneumonia (cough or difficulty breathing + fast breathing or chest wall indrawing) + at least one of the following:

- SpO₂ < 90%
- Very severe chest indrawing, grunting, central cyanosis, or presence of any other general danger sign (inability to breastfeed or drink, lethargy or unconsciousness or convulsions).

The diagnosis can be made on clinical grounds; chest imaging (radiograph, CT scan, ultrasound) may assist in diagnosis and identify or exclude pulmonary complications.
Critical disease

ARDS

Onset: within 1 week of a known clinical insult (i.e. pneumonia) or new or worsening respiratory symptoms.

Chest imaging: radiograph, CT scan or lung ultrasound: bilateral opacities, not fully explained by volume overload, lobar or lung collapse, or nodules.

Origin of pulmonary infiltrates: respiratory failure not fully explained by cardiac failure or fluid overload. Need objective assessment (e.g. echocardiography) to exclude hydrostatic cause of infiltrates/oedema if no risk factors present.

Oxygenation impairment in adults:

Air blood gases (ABG) available

- Mild ARDS: 200 mmHg < PaO₂/FiO₂ ≤ 300 mmHg (with PEEP or CPAP ≥ 5 cmH₂O)
- Moderate ARDS: 100 mmHg < PaO₂/FiO₂ ≤ 200 mmHg (with PEEP ≥ 5 cmH₂O)
- Severe ARDS: PaO₂/FiO₂ ≤ 100 mmHg (with PEEP ≥ 5 cmH₂O).

Oxygen impairment in children: note OI and OSI. Use OI when available. If PaO₂ not available, wean FiO₂ to maintain SpO₂ > 97% to calculate OSI or SpO₂/FiO₂ ratio:

- Bilevel (NIV or CPAP) ≥ 5 cmH₂O via full face mask: PaO₂/FiO₂ ≤ 300 mmHg or SpO₂/FiO₂ ≤ 264
- Mild ARDS (invasively ventilated): 4 ≤ OI < 8 or 5 ≤ OSI < 7.5
- Moderate ARDS (invasively ventilated): 8 ≤ OI < 16 or 7.5 ≤ OSI < 12.3
- Severe ARDS (invasively ventilated): OI ≥ 16 or OSI ≥ 12.3.

Abbreviation: OSI = Oxygenation Index

ABG not available (Kigali modification)

- SpO₂/FiO₂ < 315 suggests ARDS (including non-ventilated patients)

Sepsis

Adults: acute life-threatening organ dysfunction caused by a dysregulated host response to suspect or proven infection. Signs of organ dysfunction include: altered mental status (delirium), difficult or fast breathing, low oxygen saturation, reduced urinary output, fast heart rate, weak pulse, cold extremities or low blood pressure, skin mottling, laboratory evidence of coagulopathy, thrombocytopenia, acidosis, high lactate or hyperbilirubinaemia.

Children: suspected or proven infection and ≥ 2 age-based systemic inflammatory response syndrome (SIRS) criteria, of which one must be abnormal temperature or white blood cell count.

Septic shock

Adults: persistent hypotension despite volume resuscitation, requiring vasopressor to maintain MAP ≥ 65 mmHg and serum lactate level ≥ 2 mmol/L.

Children: any hypotension (SBP < 5th centile or 2SD below normal for age) or two or three of the following: altered mental status; bradycardia or tachycardia (HR < 90 beats/min [bpm] or > 160 bpm in infants and heart rate < 70 bpm or > 150 bpm in children); prolonged capillary refill (> 2 sec) or weak pulse; fast breathing; mottled or cool skin or petechial or purpuric rash; high lactate; reduced urine output; hyperthermia or hypothermia.

Acute thrombosis

Acute venous thromboembolism (i.e. pulmonary thromboembolism), acute coronary syndrome, acute stroke.

MIS-C

Preliminary case definition: children and adolescents 0-19 years of age with fever ≥ 3 days AND two of the following: rash or bilateral non purulent conjunctivitis or muco-cutaneous inflammation signs (oral, hands or feet); hypotension or shock; features of myocardial dysfunction, pericarditis, valvulitis, or coronary abnormalities (including ECHO findings or elevated troponin/NT-proBNP); evidence of coagulopathy (PT, PTT, elevated D-dimers); acute gastrointestinal problems (diarrhoea, vomiting or abdominal pain); AND elevated markers of inflammation such as ESR, C-reactive protein, or procalcitonin AND no other obvious microbial cause of shock syndrome AND evidence of COVID-19 (RT-PCR, antigen test or serology positive), or likely contact with patients with COVID-19.

(See scientific brief, 15 May 2020, WHO: Multisystemic inflammatory syndrome in children and adolescents temporarily related to COVID-19.)

Abbreviations: BP = blood pressure; bpm = beats/min; CPAP = continuous positive airway pressure; CT = computed tomography; FiO₂ = fraction of inspired oxygen; MAP = mean arterial pressure; NIV = non-invasive ventilation; OI = Oxygenation Index using SpO₂; OSI = Oxygen Saturation Index; PaO₂ = partial pressure of oxygen; PEEP = positive end-expiratory pressure; SBP = systolic blood pressure; SD = standard deviation; SIRS = systemic inflammatory response syndrome; SOFA = sequential organ failure assessment; SpO₂ = oxygen saturation.
2.5 Decision-making algorithm for patients presenting with acute respiratory infection (ARI) (influenza or COVID-19 suspected or known to be circulating)

**Fever (> 38 °C) or history of fever + cough + shortness of breath within 14 days**

**Screening for COVID-19**
- Travel/residence in a location with COVID-19 community transmission
- Contact with a confirmed/probable COVID-19 case
- Patient with severe acute respiratory illness (fever + cough or shortness of breath) and requiring hospitalization and no alternative diagnosis that fully explains the clinical presentation

**Screening for flu if influenza is circulating**

**Test according to national protocols**

- **No risk factors**
  - Isolation in community facility/home with rapid health advice
  - Seek usual care according to local guidance

- **≥ 1 risk factors**
  - Isolation in hospital or community facility/home with monitoring by health worker
  - If influenza circulating: test and treat (oseltamivir)

**Yes**

- **SpO₂ < 90%**
- **SpO₂ 90-94% with instability**
- **Signs of severe pneumonia**
- **Presence of emergency signs**

**Emergency signs**
- Obstructed or absent breathing
- Severe respiratory distress
- Cyanosis
- Shock
- Coma and/or convulsions

**Yes**

- Isolation and treatment in the hospital AND consider ICU
- If COVID-19 positive, refer to Therapeutics and COVID-19: living guideline and Living guidance for clinical management of COVID-19
- If influenza circulating: treat (oseltamivir), see Guidelines for the clinical management of severe illness from influenza virus infections

**Risk factors associated with severe disease**
- Age more than 60 years (increasing with age).
- Underlying noncommunicable diseases (NCDs): diabetes, hypertension, cardiac disease, chronic lung disease, cerebrovascular disease, dementia, mental disorders, chronic kidney disease, immunosuppression (including HIV), obesity and cancer.
- In pregnant or recently pregnant: women > 35 years old, obesity, with chronic medical conditions or pregnancy specific disorders (e.g. gestational diabetes and pre-ectampsia/ectampsia).
- Smoking.
- Unvaccinated against COVID-19.
- In influenza: young children (< 59 months).
2.6 Decision-making algorithm for hospitalization of patients with pneumonia (influenza or COVID-19 known to be circulating)

**WHO criteria for severe pneumonia**

**Children:** Pocket book of hospital care for children (WHO, 2013).
- Cough or difficulty breathing with at least one of the following:
  - Central cyanosis or oxygen saturation (\(\text{SpO}_2\) < 90%)
  - Severe respiratory distress (e.g. grunting, very severe chest indrawing)
  - General danger sign (e.g. inability to breastfeed or drink, lethargy or unconscious, convulsions).
- Any or all of the following may also be present:
  - Fast breathing (< 2 months ≥ 60 breaths/min; 2-11 months ≥ 50; 1-5 years ≥ 40; 5-15 years ≥ 30).
  - Chest indrawing.

**Adults:** IMAI district clinician manual: hospital care for adults and adolescents (WHO, 2011).
- \(\text{RR} > 30\) breaths/min
- \(\text{SpO}_2 < 90\%\)
- Signs of severe respiratory distress (e.g. inability to speak, use of accessory muscles).
2.7 Decision-making support tool for hospitalization and ICU admission for patients with SARI and severe pneumonia

Patients should be admitted to ICU based on severity of clinical condition and resource availability. In hospitals where oxygen therapy is only available in the ICU, admit all SARI patients to the ICU. In hospitals where oxygen therapy can also be delivered on wards, admit less severe SARI patients to wards but with increased monitoring. During outbreaks, a surge of patients may exhaust resources; less severe cases may need to be managed outside the ICU.

In adults, the CURB-65 score is a validated tool that, when combined with clinical judgments can be used to predict mortality and aid in determining admission for adult patients with pneumonia. This is adapted from the British Thoracic Society guidelines for the management of community acquired pneumonia in adults (BTS, 2009).

<table>
<thead>
<tr>
<th>CURB-65 score: one point for each feature present</th>
<th>Total points</th>
</tr>
</thead>
<tbody>
<tr>
<td>Confusion</td>
<td>1</td>
</tr>
<tr>
<td>Urea &gt; 7 mmol/L</td>
<td>1</td>
</tr>
<tr>
<td>RR ≥ 30/min</td>
<td>1</td>
</tr>
<tr>
<td>Blood pressure (SPB &lt; 90 or DPB ≤ 60 mmHg)</td>
<td>1</td>
</tr>
<tr>
<td>Age ≥ 65 years</td>
<td>1</td>
</tr>
</tbody>
</table>

Level of severity, risk of death and management according to score

- **Score 0-1**
  - Low severity
  - Risk of death < 3%
  - Consider home-based care

- **Score 2-3**
  - Moderate severity
  - Risk of death 3-15%
  - Consider hospitalization

- **Score 4-5**
  - High severity
  - Risk of death > 15%
  - Consider ICU
2.8 Checklist for admission

Once you have decided to admit a patient with severe acute respiratory infection (SARI) to the hospital, consider using this checklist to ensure the following have been done in preparation for admission.

This is adapted from the IMAI district clinician manual: hospital care for adults and adolescents (WHO, 2011).

Checklist for admission

☐ Essential diagnostic tests obtained, e.g. complete blood cell count, chemistry panel, glucose, chest radiograph, upper respiratory tract specimens for viral testing (COVID-19 and during influenza season include influenza testing), blood sample for culture (when possible, before first dose of antimicrobials), but do not delay antimicrobials.

☐ Emergency treatments given and patient’s response checked, e.g. oxygen therapy, insertion of peripheral IV (use appropriate antisepsis for the skin to prevent catheter-related infections), initial fluid therapy (and vasopressors if in shock).

☐ For patients with SARI and sepsis or septic shock: administer appropriate antimicrobials immediately for the suspected or confirmed pathogen, ideally within 1 hour of recognition (see Chapter 7).

☐ Give steroids (if suspected or confirmed severe or critical COVID-19).

☐ Documentation completed.

☐ Determined the level of care the patient needs, e.g. ICU, high dependency unit, general ward.

☐ Determined IPC measures and proper PPE health care workers need to manage the patient.

☐ Verbal communication with ward staff completed to ensure continuity of care.

☐ Patient prepared for safe transfer.
2.9 Checklist for transfer

Transport of the critically ill patient can be risky as complications during this process can be life threatening and may be related to clinical, organizational, or equipment issues.

Consider using this checklist to ensure the safe transport of the patient to the designated unit.

This is adapted from the IMAI district clinician manual: hospital care for adults and adolescents (WHO, 2011).

---

Checklist for transfer

☐ Patient is stabilized.
☐ Adequate IPC measures and proper PPE needed, e.g. medical mask for patients with ARI.
☐ Everything secured: airway, NG tube, IV lines, monitors, endotracheal tubes, ventilator.
☐ Enough drugs: e.g. vasopressors, sedatives.
☐ Enough oxygen: adequate oxygen saturation (SpO₂).
☐ Enough IV fluids: blood pressure adequate.
☐ Health care workers (e.g. transporters, receiving staff) and receiving unit/ward prepared.
2.10 Transfer of critically ill patients: air medevac for COVID-19 patients

Air ambulance services are facing multiple challenges when transporting highly infectious patients for several hours in enclosed spaces. The preference is to treat patients on site rather than transporting them from an outbreak area. However, an evacuation may be needed if patients require care beyond what can be provided in the current facility/location. Moving patients closer to hospital facilities is likely to increase the chance of a good outcome and relocating them will take the load off overburdened and under-resourced health care facilities during an outbreak. While there is no single standardized scoring system on emergency aeromedical evacuations, the patient’s clinical condition, age, local resources and locations have been found to be the criteria that inform decision-making.

This overview is meant to provide a simplified guidance to clinicians that will facilitate air medevac and does not deliver detailed recommendations for necessary equipment, medications, flight physiology specific ventilation, waste management, IPC considerations, etc.

Pre-flight patient assessment

Presenting signs and symptoms of COVID-19 vary. When considering a medevac transportation, pre-clarification of the severity of the disease is crucial.

<table>
<thead>
<tr>
<th>Pre-flight physiological considerations</th>
<th>Altitude exposure imposes hypoxia and gas expansion in body cavities.</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>• Consider supplemental oxygen, ventilation and low-altitude flight paths.</td>
</tr>
<tr>
<td></td>
<td>• Prepare for any deterioration and for conditions induced by the aerospace environment.</td>
</tr>
<tr>
<td></td>
<td>• Plan for early endotracheal intubation:</td>
</tr>
<tr>
<td></td>
<td>- Pre-emptive intubation is recommended for patients with the potential of respiratory decompensation during aeromedical transport.</td>
</tr>
<tr>
<td></td>
<td>- Anaemia: fit to fly if Hb ≥ 95g/L. If, due to chronic disease and compensation, Hb ≥ 80 g/L is acceptable. If lower or if concurrent lung or cardiac disease, consider transfusion.</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Pre-flight checklist</th>
<th>• Allocate key roles.</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>• Flight crew must include a specialized intensive care flight physician and nurse, that are knowledgeable about flight physiology, experienced in inter-facility air-transfer and the management of critical care patients/ intubated patients.</td>
</tr>
<tr>
<td></td>
<td>• Precise SOPs (e.g. correct use and handling of PPE/environmental cleaning and decontamination); every crew member and service staff (e.g. aircraft cleaning) strictly adhere to SOPs.</td>
</tr>
<tr>
<td></td>
<td>• Medevac personnel to ensure that patient receives instruction on in-flight IPC measures.</td>
</tr>
<tr>
<td></td>
<td>• Written informed consent and signed liability waiver forms from patient and/or relatives along with outline of intention to transfer must occur before the transfer.</td>
</tr>
<tr>
<td></td>
<td>• Clear communication and agreement between coordination mechanism, transferring and receiving facility:</td>
</tr>
<tr>
<td></td>
<td>- Formal agreement from the receiving facility.</td>
</tr>
<tr>
<td></td>
<td>- Ensure full and up-to-date medical records on patient’s clinical condition, treatment received, intention to transfer, mode and timeline of transfer and communicate them to flight crew and receiving facility.</td>
</tr>
<tr>
<td></td>
<td>• Equipment check: monitoring devices (ECG, blood pressure, pulse oximetry, capnography, defibrillators, pacing devices, aeronautical oxygen systems, oxygen, suction, face mask, filter, breathing circuit, respiratory support device, equipment to manage sedation, intubation, ventilation, cardiovascular support, infusion devices, medications including resuscitation drugs, point-of-care testing (POCT) for e.g. arterial blood gases analysis, etc. All equipment must be certified for aeronautical use.</td>
</tr>
<tr>
<td></td>
<td>• Check resuscitation medications.</td>
</tr>
<tr>
<td></td>
<td>• Be prepared to deepen anaesthesia and administer additional catecholamines during transport.</td>
</tr>
</tbody>
</table>
In-flight checklist
- Precise SOPs (e.g., correct use and handling of PPE/environmental cleaning and decontamination); every crew member and service staff (e.g., aircraft cleaning) strictly adhere to SOPs.
- Medevac personnel to ensure that patient receives instruction on in-flight IPC measures.
- Avoid aerosol-generating procedures such as non-invasive ventilation, high-flow oxygen therapy, tracheal suction or nebulization.
- Clear SOPs in place to ensure safe sharps/waste management.

Post-flight checklist
- SOPs on decontamination.
- SOPs on safe equipment handling.
- SOPs on safe waste disposal.
- SOPs on personnel monitoring.

Medevac algorithm

- Is it possible to delay the medevac until the patient is treated and fully stabilized? Are there resources in place to do so?
- Is it possible to delay the medevac until the patient is treated and fully stabilized? Are there resources in place to do so?
- Is it possible to delay the medevac until the patient is treated and fully stabilized? Are there resources in place to do so?
- Is it possible to delay the medevac until the patient is treated and fully stabilized? Are there resources in place to do so?

- Proceed to endotracheal intubation and mechanical ventilation
- Proceed to endotracheal intubation and mechanical ventilation
- Proceed to endotracheal intubation and mechanical ventilation
- Proceed to endotracheal intubation and mechanical ventilation

- Patient is stable?
- Unconscious?
- GCS < 11
- SatO₂ < 90% or PaO₂/FiO₂ < 300

- Perform medevac to the closest city/airport where adequate treatment capacities are available

Notes: GCS - Glasgow Coma Scale; PaO₂/FiO₂ - ratio of partial pressure arterial oxygen and fraction of inspired oxygen; SatO₂ - oxygen saturation.
**Minimize risk of in-flight transmission**

The risk of transmission in the confined environment of a helicopter or aircraft, potentially over long distances with prolonged transport times can be minimized by isolating the patient during transportation.

<table>
<thead>
<tr>
<th>Rotary wing</th>
<th>Fixed wing</th>
</tr>
</thead>
<tbody>
<tr>
<td>• The cockpit should be isolated through use of the aircraft blind.</td>
<td>• Use of barriers such as screens or curtains may provide some level of protection for personnel positioned in the cockpit, and their effectiveness is reliant on airflow and the movement of airborne particles within the aircraft.</td>
</tr>
<tr>
<td></td>
<td>• Air-conditioning (if applicable) should be selected in non-recirculating mode.</td>
</tr>
<tr>
<td></td>
<td>‒ Pressurized aircraft: if available, aircraft recirculation should be deselected.</td>
</tr>
<tr>
<td></td>
<td>‒ If cabin air recirculation is selected, then HEPA filtration is preferred. Aircraft ventilation should remain on at all times during transport of respiratory patients, including during ground delays.</td>
</tr>
</tbody>
</table>
References and resources


3. Infection prevention and control for patients with SARI
Infection prevention and control for patients with SARI

Summary

Administrative and engineering measures and PPE work in harmony to prevent the spread of infection and keep health care workers and patients safe. For health care facility readiness for COVID-19 see Severe acute respiratory infections treatment centre ( ).

When providing health care for any patients in a health care facility, one must implement standard precautions, which should include hand hygiene according to the WHO 5 moments, respiratory hygiene, use of appropriate PPE according to a risk assessment, safe injection practices and sharps management, waste management, safe handling and cleaning of soiled linens, environmental cleaning, and safe handling, cleaning and disinfection of patient care equipment. Standard precautions apply to all patients regardless of their diagnosis or presumed infection status.

When caring for patients with certain types of ARI, such as avian influenza, MERS, SARS-CoV-2, seasonal influenza or another novel viral infection, it is also recommended to use droplet and contact precautions.

When carrying out certain high-risk procedures such as aerosol-generating procedures like open suctioning of airways, sputum induction, cardiopulmonary resuscitation, endotracheal intubation and extubation, non-invasive ventilation (e.g. BiPAP, CPAP), bronchoscopy and manual ventilation, one should use airborne precautions in addition to contact and droplet precautions.

In the context of the current COVID-19 epidemic, there are some specificities of SARS-CoV-2 in its transmission that have implications for the infection prevention precautions. The virus SARS-CoV-2 can spread from an infected person’s mouth or nose in small liquid particles when they cough, sneeze, speak, sing or breathe. These particles range from larger respiratory droplets to smaller aerosols.

- Current evidence suggests that the virus spreads mainly between people who are in close contact with each other, typically within 1 m (short range). A person can be infected when aerosols or droplets containing the virus are inhaled or come directly into contact with the eyes, nose or mouth.
- The virus can also spread in poorly ventilated and/or crowded indoor settings, where people tend to spend longer periods of time. This is because aerosols remain suspended in the air or travel farther than 1 m (long range).
- People may also become infected by touching surfaces that have been contaminated by the virus when touching their eyes, nose or mouth without cleaning their hands (fomite transmission).

These different ways of transmission of COVID-19 can be prevented by following basic transmission-based precautions (Coronavirus disease (COVID-19): How is it transmitted?) ()

- Social distance: keep at least 1 m from others.
- Wear a mask (Mask use in the context of COVID-19) ( )
• Ventilation: avoid crowded places, poorly ventilated, indoor locations and avoid prolonged contact with others.
• Spend more time outdoors than indoors; and avoid the three Cs: Crowded places; Close-contact settings; Confined and enclosed spaces.
• Avoid touching surfaces, especially in public settings or health facilities, in case people infected with COVID-19 have touched them. Clean surfaces regularly with standard disinfectants.
• Clean hands with soap and water, or an alcohol-based handrub.
• Respiratory hygiene: cover nose and mouth during coughing or sneezing with tissue or flexed elbow.
• Vaccination against COVID-19 whenever possible (COVID-19 vaccines).

Tools
3.1 How to implement infection control measures for SARI
3.2 How to implement infection control measures for ARIs of potential concern
3.3 Personal protective equipment (PPE)
3.4 How to improve medical mask fit in health care settings
3.5 Steps to perform a particulate respirator seal check during the putting on of PPE
3.6 Hand hygiene
3.7 The 5 moments for hand hygiene
3.8 The “Three Cs”: settings where transmission of the COVID-19 virus spreads more easily
3.9 Checklist for aerosol-generating procedures
3.1 How to implement infection control measures for SARI

These algorithms are adapted from the WHO guidelines, *Infection prevention and control of epidemic- and pandemic-prone acute respiratory infections in health care* (WHO, 2014).

Decision-tree for IPC measures for patients known or suspected to have an acute respiratory infection

---

**Patient**

Patient enters triage with symptoms of acute febrile respiratory illness  
Plus clinical and epidemiological clues for ARI of potential concern

- Report to public health authorities

**Infection control measures**

- Health care workers should perform adequate hand hygiene, use medical masks and, if splashes into eyes are possible, eye protection (goggles/face shield)
- Paediatric patients with clinical symptoms and signs indicating specific diagnosis (e.g. croup for parainfluenza, acute bronchiolitis for respiratory syncytial virus), especially during seasonal outbreaks, may require isolation precautions as soon as possible
- Encourage respiratory hygiene (i.e. use of medical mask or tissues when coughing or sneezing followed by hand hygiene) by patients in the waiting room
- Ensure patients maintain a minimum 1 m distance from other patients

- Health care workers should use PPE (medical mask, eye protection, long-sleeved gown and gloves) and adequate hand hygiene before and after PPE use
- If required, use separate, adequately ventilated or airborne precaution room
- If no separate room available, cohort patients with same laboratory-confirmed etiological diagnosis, maintain at least 1 m distance between beds
- If etiology cannot be laboratory confirmed and no separate room, adopt special measures

- Patient diagnosed with ARI of potential concern
  - IPC precautions to remain in place for the duration of symptomatic illness

- Other diagnosis
  - Reassess IPC precaution

---

*ARIs of potential concern include SARS, COVID-19, new influenza virus causing human infection (e.g. human cases of avian influenza) and novel pathogens causing ARIs that can cause outbreaks with high morbidity and mortality. Clinical and epidemiological clues include severe disease in a previously healthy host, exposure to household member or close contact with severe ARI, cluster of cases, travel, exposure to ill animals or laboratory.*

*Airborne precaution rooms include both mechanically and naturally ventilated rooms with ≥ 12 air changes per hour and controlled direction of airflow.*

*The term “special measures” means allowing patients with epidemiological and clinical information suggestive of a similar diagnosis to share a room, but with a spatial separation of at least 1 m.*
3.2 How to implement infection control measures for ARIs of potential concern

Acute respiratory infections of potential concern include SARS, COVID-19, new influenza viruses causing human infection (e.g. human cases of avian influenza) and novel pathogens causing ARIs that can cause outbreaks with high morbidity and mortality.

<table>
<thead>
<tr>
<th>Instructions for patients</th>
<th>Give suspect patient a medical mask and direct the patient to a separate area; an isolation room if available. If not possible to separate the patient, try to keep at least 1 m distance between suspected patients and other patients in the waiting room. All patients in the waiting room should have a mask and should practise respiratory hygiene (cover their nose and mouth during coughing or sneezing with a tissue or flexed elbow) and perform hand hygiene after contact with respiratory secretions (such as coughing, sneezing or blowing nose).</th>
</tr>
</thead>
<tbody>
<tr>
<td>Apply droplet precautions</td>
<td>Droplet precautions prevent transmission of respiratory viruses through respiratory droplets that are expelled when an infected person speaks, coughs, or sneezes. Place patients in single rooms, or group those with the same etiological diagnosis together, maintaining at least 1 m distance between beds. If an etiological diagnosis is not possible, group patients with similar clinical diagnosis and based on epidemiological risk factors, with a spatial separation. Limit patient movement within the institution and ensure that patients wear medical masks when being taken outside their rooms. When providing care in close contact with a patient with respiratory symptoms (e.g. coughing or sneezing), use a medical mask and eye protection (face mask or goggles), because sprays of secretions may occur. Note: The risk of COVID-19 spreading is especially high in places where the three Cs overlap: Crowded places, Close-contact settings (especially where people have conversations very near each other) and Confined and enclosed spaces with poor ventilation (see Tool 3.7).</td>
</tr>
<tr>
<td>Apply contact precautions</td>
<td>Contact precautions prevent direct or indirect transmission from contact with contaminated surfaces or equipment (i.e. contact with contaminated oxygen tubing/interfaces). PPE for contact precautions are a long-sleeved gown and gloves. Hand hygiene followed by putting on PPE for contact and droplet precautions (medical mask and eye protection) should be applied when entering the room. Remove PPE when leaving the room, and practise hand hygiene following PPE removal. If possible, use either disposable or dedicated equipment (e.g. stethoscopes, blood pressure cuffs, pulse oximeters and thermometers). If equipment needs to be shared among patients, clean and disinfect items between each patient use. Ensure that health workers refrain from touching their eyes, nose and mouth with potentially contaminated gloved or ungloved hands. Avoid contaminating environmental surfaces that are not directly related to patient care (e.g. door handles and light switches). Avoid medically unnecessary movement and transport of patients. Practise hand hygiene frequently.</td>
</tr>
<tr>
<td>Apply airborne precautions when performing an aerosol-generating procedure</td>
<td>Ensure that health workers performing, or in the room during, an aerosol-generating procedure (e.g. open suctioning of respiratory tract, tracheal intubation, non-invasive ventilation, tracheostomy, bronchoscopy, cardiopulmonary resuscitation) use the appropriate PPE, including gloves, long-sleeved gowns, eye protection, and fit-tested particulate respirators (N95 or equivalent, or higher level of protection). A scheduled fit test should not be confused with a user’s seal check before each use. Whenever possible, use adequately ventilated single rooms when performing aerosol-generating procedures, meaning negative pressure rooms with a minimum of 6-12 air changes per hour (e.g. equivalent to 40-80 L/s/patient for a 4 × 2 × 3 m² room) or at least 160 L/s/patient in facilities with natural ventilation. Avoid the presence of unnecessary individuals in the room. Care for the patient in the same type of room after mechanical ventilation begins.</td>
</tr>
</tbody>
</table>
3.3 Personal protective equipment (PPE)

For specifications for COVID-19 review *Personal protective equipment for COVID-19* and WHO recommendations on mask use by health care workers ( ).

Remember, PPE use should be guided by risk assessment concerning anticipated contact with blood and other bodily fluids, including respiratory droplets and secretions, during patient care and presence of non-intact skin. For example, if there is a risk of splash to the body and face then use hand hygiene, gloves, gown, medical mask and eyewear. A “how to guide” for putting on and taking off PPE is shown below.

**HOW TO GUIDE - PUTTING ON PPE FOR CONTACT/DROPLET PRECAUTIONS**

1. Perform hand hygiene
   - Alcohol based handrub
     Rub hands for 20–30 seconds.
   - Water and soap
     Wash hands for 40–60 seconds.

2. Put on the gown

3. Put on the mask
   - Medical mask.

4. Put on eye protection
   - Put on face shield or goggles.

5. Put on gloves
   - Ensure glove is placed over the cuff of the gown.

Full PPE
HOW TO GUIDE - TAKING OFF PPE FOR CONTACT/DROPLET PRECAUTIONS

Ensure that infectious waste containers are available for safe disposal of PPE. Separate containers should be available for reusable items.

Order is important

1. Remove gloves

2. Remove the gown
   Ensure gown is pulled away from the body during removal and that clothing does not become contaminated and dispose of them safely.

3. Perform hand hygiene
   Alcohol based handrub
   Rub hands for 20-30 seconds.
   or
   Water and soap
   Wash hands for 40-60 seconds.

4. Remove eye protection
   Remove face shield or goggles.

5. Remove the mask
   Ensure you are taking the mask off from the straps, avoid touching the mask.

6. Perform hand hygiene
   Alcohol based handrub
   Rub hands for 20-30 seconds.
   or
   Water and soap
   Wash hands for 40-60 seconds.
3.4 How to improve medical mask fit in health care settings

How to improve medical mask fit in health care settings

When linking ear loops behind the head

1. Clean hands thoroughly before putting on and before and after taking off your mask.
2. Attach a clean connector to link ear loops together.
3. Place the medical mask colour-side facing outward, attach ear loops behind ears.
4. Attach ear loops using connector behind head tightly.
5. Adjust the wire at the bridge of the nose and ensure there are no gaps between the mask and your face at the sides of your nose, cheeks, and under your chin.

When using knot-and-tuck method

1. Fold the mask horizontally.
2. Make a knot on both ear loops as close to the edge of the mask as possible.
3. Push the extra material under the mask inward to ensure no gaps on both sides.

*Find a clean practical connector to link your ear loops, it can be adjustable rope or silicone.

If a surface is used to fold and manipulate the mask, clean the surface first using a cloth wipe with soap and water, followed by disinfection using a cloth wipe soaked in 70-90% alcohol OR 0.1% sodium hypochlorite (or comparable hospital grade disinfectant) and allow for least 1 minute contact time before surface is used.
3.5 Steps to perform a particulate respirator seal check during the putting on of PPE

1. Cup the respirator in your hand with the nosepiece at your fingertips allowing the headbands to hang freely below your hand.

2. Position the respirator under your chin with the nosepiece up.

3. Pull the top strap over your head, resting it high at the back of your head. Pull the bottom strap over your head and position it around the neck below the ears.

4. Place fingertips of both hands at the top of the metal nosepiece. Mould the nosepiece (using two fingers of each hand) to the shape of your nose. (Pinching the nosepiece using one hand may result in less effective respirator performance.) Cover the front of the respirator with both hands, being careful not to disturb the position of the respirator.

A. Positive seal check
Exhale sharply. A positive pressure inside the respirator = no leakage. If leakage, adjust position and/or tension straps. Reset the seal. Repeat the steps until respirator is sealed properly.

B. Negative seal check
Inhale deeply. If no leakage, negative pressure will make respirator cling to your face. Leakage will result in loss of negative pressure in the respirator due to air entering through gaps in the seal.
3.6 Hand hygiene

Hand hygiene must be performed before and after any contact with patients, after contact with contaminated items or surfaces, before and after PPE use and according to the WHO 5 moments (see Tool 3.6).

Use an alcohol-based product if hands are not visibly soiled.

Wash hands with soap and water when they are visibly soiled or contaminated with proteinaceous material.

Below is an example of hand washing with soap and water. The same rubbing technique can be used with alcohol-based product.

This entire procedure should take 40-60 seconds for water and soap (or 20-30 seconds for alcohol-based handrub).
How to Handrub?

RUB HANDS FOR HAND HYGIENE! WASH HANDS WHEN VISIBLY SOILED

Duration of the entire procedure: 20–30 seconds

1a. Apply a palmful of the product in a cupped hand, covering all surfaces;

1b. Rub hands palm to palm;

2. Right palm over left dorsum with interlaced fingers and vice versa;

3. Palm to palm with fingers interlaced;

4. Backs of fingers to opposing palms with fingers interlocked;

5. Rotational rubbing of left thumb clasped in right palm and vice versa;

6. Rotational rubbing, backwards and forwards with clasped fingers of right hand in left palm and vice versa;

7. Once dry, your hands are safe.
3.7 The 5 moments for hand hygiene in health care facilities

Your 5 Moments for Hand Hygiene

1. BEFORE TOUCHING A PATIENT
   - Wash: Clean your hands before touching a patient when approaching a patient.
   - Dry: To protect the patient against harmful germs carried on your hands.

2. BEFORE CLEAN/ASEPTIC PROCEDURE
   - Wash: Clean your hands immediately before performing a clean/aseptic procedure.
   - Dry: To protect the patient against harmful germs, including the patient’s own, from entering their body.

3. AFTER BODY FLUID EXPOSURE/RISK
   - Wash: Clean your hands immediately after an exposure to body fluids (and after gown removal).
   - Dry: To protect yourself and the health-care environment from harmful patient germs.

4. AFTER TOUCHING A PATIENT
   - Wash: Clean your hands after touching a patient and their immediate surroundings, when leaving the patient's room.
   - Dry: To protect yourself and the health-care environment from harmful patient germs.

5. AFTER TOUCHING PATIENT SURROUNDINGS
   - Wash: Clean your hands after touching any object or surface in the patient’s immediate surroundings, even if the patient has not been touched.
   - Dry: To protect yourself and the health-care environment from harmful patient germs.
3.8 The “Three Cs”: settings where transmission of the COVID-19 virus spreads more easily

The risk of COVID-19 spreading is especially high in places where these “3 Cs” overlap.
3.9 Checklist for aerosol-generating procedures

In health facilities where people are receiving treatment for COVID-19, there is an increased risk of infection during medical procedures called aerosol-generating procedures. These can produce very small droplets that can stay suspended in the air for long periods of time and spread beyond conversational distances (typically 1 m). Therefore, health workers performing these procedures or in settings where these procedures are performed should take specific airborne protection measures, including using appropriate PPE such as respirators.

Consider using this checklist when performing aerosol-generating procedures, such as tracheal intubation, non-invasive ventilation, tracheotomy, cardiopulmonary resuscitation and manual ventilation before intubation, bronchoscopy, aspiration, or open suctioning of respiratory tract secretions.

Note: There is limited research available regarding the risk of non-invasive ventilation and high-flow oxygen therapy, but experts suggest using airborne precautions during these procedures.

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Checklist for aerosol-generating procedures

☐ Perform hand hygiene before and after patient contact and before and after PPE use.

☐ All health workers present should wear the following PPE:
  • A facial particulate respirator (e.g. European Union FFP2 or United States of America National Institute for Occupational Safety and Health-certified N95). Perform a seal check.
  • Eye protection (e.g. goggles or a face shield).
  • A clean, non-sterile, long-sleeved gown.
  • Gloves (some of these procedures require sterile gloves).

☐ Make sure adequately ventilated room:
  • For mechanically ventilated rooms: Ensure ≥ 6-12 air changes per hour plus control of airflow direction, ideally 12 air changes per hour for new constructions, with a recommended negative pressure differential of ≥ 2.5 Pa (0.01 inch water gauge) to ensure that air flows from the corridor into the patient room.
  • For natural ventilation: The recommended average natural ventilation rate is 160 L/s/patient.

☐ Social distancing - avoid unnecessary individuals in the room.
References and resources


WHO IPC posters:

• Contact and droplet precautions - COVID-19 personal protective equipment (PPE): https://www.who.int/docs/default-source/infection-prevention-and-control/contact-droplet-covid-19-precautions.pdf?sfvrsn=74c1a87c_2

• How to guide - Putting on PPE for contact/droplet precautions: https://www.who.int/docs/default-source/infection-prevention-and-control/ppe-en.pdf?sfvrsn=4b45270e_2

• Screening for acute respiratory infection: https://www.who.int/docs/default-source/infection-prevention-and-control/screening-respiratory-infection.pdf?sfvrsn=2c899e42_2
Monitoring patients with acute respiratory infection
Summary

Vital signs, including temperature, heart rate, blood pressure, respiratory rate, oxygen saturation and mental status with AVPU, are standard metrics that can be used when assessing and monitoring patients with ARI in hospital and pre-hospital settings.

Pulse oximetry is used to monitor oxygen saturation (SpO₂). It is an essential tool that should be available at all first access health points, where patients with ARI are assessed to inform triage, clinical assessment and appropriate referral and/or treatment plans. Pulse oximeter is also an essential tool to monitor patients for signs of clinical deterioration or improvement that may require change in clinical management interventions.

In patients with mild or moderate COVID-19 with risk factors for severe disease who do not need hospitalization, monitoring oxygenation with a pulse oximeter at home can be beneficial to identify patients that may develop complications and require urgent referral to health facilities for additional treatments. See Living guidance for clinical management of COVID-19 Section 10. Management of moderate COVID-19: pneumonia treatment.

Patients with severe or critical COVID-19 should be cared for in a hospital and monitored frequently because of their dynamic clinical condition and need for timely (and titrated) interventions. The acute condition of the patient will dictate the moment when advanced monitors are deployed, as well as the frequency of checks. For example, in an ICU compared with a general ward, haemodynamic and respiratory parameters are monitored more frequently (sometimes continuously), along with a more complete assessment that includes frequent physical examinations, laboratory tests, and intake and output.

All parameters that are monitored should be captured in a standardized charting system (paper based or electronic health record) so that trends can be monitored over time and response to treatments assessed.

There are published monitoring scoring tools that can be used to assess and identify deteriorating patients that may be progressing to critical illness. Two examples includes in this toolkit are the National Early Warning Score (NEWS) and the Paediatric Early Warning Score (PEWS). These tools may be useful to identify when patients fail to respond to treatments or deteriorate and should trigger the need for escalation of care and/or new treatment intervention. Regardless of scoring tool used, monitoring vital signs alone must be paired with a systematic approach to interpretation of the data (including clinical history and physical examination) and modification of monitoring and treatment plans accordingly.
Tools

4.1 AVPU scale: a simple tool for assessing level of consciousness
4.2 Pulse oximetry monitoring
4.3 Blood gas analysis monitoring
4.4 Capnometry (capnography)
4.5 National Early Warning Score (NEWS) for adults
4.6 Paediatric Early Warning Score (PEWS)
4.7 Routine monitoring and care framework for COVID-19 patients
4.8 WHO Mild COVID-19 home care bundle for health care workers
4.9 Memory aid: key criteria used to assess vital signs in children
4.10 Memory aid: key physiological aspects to assess in pregnant women
4.1 AVPU scale: a simple tool for assessing level of consciousness

This scale is a simple way to assess a patient’s mental status. Each letter corresponds to the patient’s level of consciousness.

| A | ALERT: Patient is aware of the examiner, responds to the environment on their own, follows commands, opens eyes spontaneously, and tracks objects. |
| V | VERBAL: Patient does not open eyes spontaneously, but does in response to a verbal stimulus and reacts meaningfully to the verbal stimulus. |
| P | PAIN: Patient does not open eyes spontaneously, but does in response to pain and may move, moan, or cry in response to pain. |
| U | UNRESPONSIVE: Patient does not respond to verbal or painful stimuli. |
4.2 Pulse oximetry monitoring

A pulse oximeter measures oxygen saturation of haemoglobin in the blood by comparing the absorbance of light of different wavelengths across a translucent part of the body.

Pulse oximetry is the best non-invasive method available for detecting hypoxaemia and titrating oxygen delivery accordingly. Using clinical signs may mislead the diagnosis of some patients with hypoxaemia (e.g. some COVID-19 patients with “silent hypoxaemia”). Pulse oximetry should be performed on all patients with SARI.

Pulse oximetry may have some limitations and may produce inaccurate results during select clinical conditions (e.g. carbon monoxide poisoning, methaemoglobinemia, and low perfusion) or when using low-cost oximeters that do not meet the technical specifications for clinical use (see WHO technical specifications for oxygen concentrators and WHO-UNICEF Technical specifications and guidance for oxygen therapy devices).

Note: Oxygen partial pressures in the atmosphere are lower at higher altitudes. Therefore, patients at facilities in higher altitudes may require higher flow rates for longer duration for adequate therapy compared with patients at sea level. If the altitude is higher than 1000 m, then a correction factor should be calculated to define ARDS: $\text{PaO}_2/\text{FiO}_2 \times \text{barometric pressure}/760$.
Pulse oximeter displaying normal SpO₂ reading

This image shows a pulse oximeter with a normal oxygen saturation (SpO₂) reading (pulse rate = 102 bpm; SpO₂ = 97%) and a plethysmographic (pulse) wave (i.e. the waveform corresponds with arterial pulsation and rate) indicating a good arterial trace and a valid reading.

Note: Not all pulse oximeters may display a plethysmograph or signal quality, and without this feature, interpretation of the data must be done cautiously.

Pulse oximeter displaying abnormal SpO₂ reading

In this image (pulse rate = 150 bpm; SpO₂ = 82%), the pulse oximeter has a good plethysmographic wave, indicating a valid arterial trace. Therefore, the SpO₂ reading, which is abnormally low (82%), is likely accurate and indicates that the patient is hypoxaemic. Oxygen should be given. Note the increased heart rate, which is common in seriously ill patients.

Pulse oximeter showing a poor plethysmography (pulse) wave

In this image, the SpO₂ reading is 83% (pulse rate 55 bpm) with a poor pulse wave. The poor pulse wave raises concern that the reading of 83% may not be correct. Multiple factors may cause a poor waveform, including patient movement, poor probe placement, or poor perfusion (shock).

4.3 Blood gas analysis monitoring

Blood gas analysis can be used to measure the pH, partial pressure of oxygen (PaO₂), and partial pressure of carbon dioxide (PaCO₂) in arterial, venous or capillary blood.

The pH is a direct indicator of overall acid-base status in arterial, capillary and venous blood. The probable cause of pH disturbances can be inferred only from the PaCO₂ and blood bicarbonate (HCO₃⁻) concentration (or the base excess or deficit). In the absence of blood gas analysis, there is no accurate surrogate for assessing acid-base status.

Acidosis is a process that lowers the extracellular fluid pH (pH < 7.35). This can be caused by a fall in the serum bicarbonate concentration (metabolic acidosis with low pH and HCO₃⁻) or an elevation of PaCO₂ (respiratory acidosis with low pH and high PaCO₂). Acidosis (metabolic or respiratory) is commonly seen when there is major disturbance of the circulation or oxygen delivery, as in severe hypoxaemia due to SARI, ARDS, sepsis and septic shock.

Arterial blood gas analysis can be used for monitoring changes in response to therapy, including ventilator changes. Venous blood and capillary blood may be easier to monitor than arterial blood but should not be used for oxygen level determination. The CO₂ level in arterial, capillary or venous blood helps in assessing and monitoring alveolar ventilation, but peripheral venous values can be inaccurate. While SpO₂ can often be used as a surrogate for PaO₂, end-tidal CO₂ (EtCO₂) measurement has limitations when deployed as a surrogate for PaCO₂ (see Tool 4.4). Direct measurement of PaCO₂ is an important diagnostic tool in patients with SARI and ARDS to help guide protocolized lung protective ventilation.

Blood gas analysis provides information on oxygenation, ventilation and circulation, but can also inform about electrolyte concentrations (particularly sodium and potassium) which can be measured in the same blood sample by some analysers, such as point of care. Electrolyte abnormalities are common in critically ill patients with SARI and it may be useful for them to be measured for diagnosis and monitoring, besides pH, HCO₃⁻, PaCO₂ and PaO₂.

✓ BENEFITS

• Measures pH, HCO₃⁻, PaCO₂, PaO₂.
• Informs about ventilation and acidosis.
• May be used to measure lactate, haemoglobin, potassium (e.g. point of care analysers).

⚠️ LIMITATIONS

• Invasive arterial puncture.
• Special (heparinized) syringe.
• Rapid transfer (on ice if > 20 minutes) to laboratory.
• Blood gas analyser machine (e.g. I-STAT).
• CO level not detected on all machines.

4.4 Capnometry (capnography)

Capnometry is the non-invasive measurement of EtCO₂ partial pressure. This is achieved by shining infrared light through exhaled gas and detecting absorption, from which EtCO₂ partial pressure can be derived.

Capnography is the non-invasive measurement of EtCO₂ in exhaled breath expressed as the CO₂ concentration over time. The relationship of CO₂ concentration to time is graphically represented by the CO₂ waveform (capnogram).

Capnography can be done in line with the breathing circuit, at the hub of the endotracheal tube (i.e. mainstream) or by continuously sampling gas from the exhaled breath of a patient or from a breathing circuit (e.g. nasal, nasal-oral cannula) to a detector (i.e. sidestream). Mainstream systems are configured for intubated patients. Sidestream systems are configured for both intubated and non-intubated patients. Capnography can be used to spot check for correct endotracheal tube placement and continuously monitor for circuit continuity and adequacy of ventilation.¹

CO₂ monitors are either quantitative or qualitative. Quantitative devices measure the precise EtCO₂ as a number (capnometry) or a number and a waveform (capnography). Qualitative devices (e.g. colorimetric detectors) report the range in which the EtCO₂ falls as opposed to a precise value. For example, the colour in the colorimetric end-tidal CO₂ (EtCO₂) detector may change when the endotracheal tube is not correctly placed. EtCO₂ measurements can be used to monitor and manage ventilation and circulation.

**✓ BENEFITS**

- Measures EtCO₂ tension in expired air from sampling gas in respiratory circuit.
- In normal lungs, EtCO₂ is about 3-5 mmHg less than PaCO₂.
- Confirms endotracheal intubation.
- Assess perfusion (during CPR).

**⚠ LIMITATIONS**

- Inaccurate if there is no discernible plateau: e.g. airflow obstruction.
- Underestimates PaCO₂ when there is decreased lung perfusion:
  - pulmonary emboli
  - hypotension
  - high PEEP
  - severe ARDS
  - emphysema.

¹ If there is significant dead space in the circuit (e.g. large filters or heat moisture exchangers) or the patient’s lungs (e.g. ARDS), capnography may significantly underestimate PaCO₂ and will have limited utility for guiding titration of ventilation.
4.5 National Early Warning Score (NEWS) for adults

The NEWS score was developed by the Royal College of Physicians (United Kingdom of Great Britain and Northern Ireland) to improve the assessment of acute-illness severity of patients in hospital and pre-hospital settings.

For the NEWS update (NEWS 2) in Chart 4 some specific areas were reviewed:

- Determining how the NEWS could be better used to identify patients likely to have sepsis who were at immediate risk of serious clinical deterioration and required urgent clinical intervention.
- Highlighting that a NEWS score of 5 or more is a key threshold for an urgent clinical alert and response.
- NEWS 2:
  - Improves the recording of oxygen use and the NEWS scoring of recommended oxygen saturations in patients with hypercapnic respiratory failure (most often due to COPD).
  - Recognizes the importance of new onset confusion, disorientation, delirium or any acute reduction in the Glasgow Coma Scale (GCS) score as a sign of potentially serious clinical deterioration by including new confusion as part of the AVPU scoring scale (i.e. ACVPU).

Please refer to all materials, including posters and training materials, on their website (ex.). Local adaptation and validation may be necessary.

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**Chart 1: NEWS scoring system**

<table>
<thead>
<tr>
<th>Physiological parameter</th>
<th>Score 0</th>
<th>Score 1</th>
<th>Score 2</th>
<th>Score 3</th>
</tr>
</thead>
<tbody>
<tr>
<td>Respiration rate (per minute)</td>
<td>8</td>
<td>9-11</td>
<td>12-20</td>
<td>21-24</td>
</tr>
<tr>
<td>SpO2 Scale 1 (%)</td>
<td>91</td>
<td>92-93</td>
<td>94-95</td>
<td>≥ 96</td>
</tr>
<tr>
<td>SpO2 Scale 2 (%)</td>
<td>83</td>
<td>84-85</td>
<td>86-87</td>
<td>88-92</td>
</tr>
<tr>
<td>Air or oxygen?</td>
<td>Oxygen</td>
<td>Oxygen</td>
<td>Oxygen</td>
<td>Oxygen</td>
</tr>
<tr>
<td>Systolic blood pressure (mmHg)</td>
<td>90</td>
<td>91-100</td>
<td>101-110</td>
<td>111-219</td>
</tr>
<tr>
<td>Pulse (per minute)</td>
<td>40</td>
<td>41-50</td>
<td>51-90</td>
<td>91-110</td>
</tr>
<tr>
<td>Consciousness</td>
<td>Alert</td>
<td>Alert</td>
<td>Alert</td>
<td>Alert</td>
</tr>
<tr>
<td>Temperature (°C)</td>
<td>35.0</td>
<td>35.1-36.0</td>
<td>36.1-38.0</td>
<td>38.1-39.0</td>
</tr>
</tbody>
</table>

Source: Royal College of Physicians (2017).
Chart 2: NEWS thresholds and triggers

<table>
<thead>
<tr>
<th>NEW score</th>
<th>Clinical risk</th>
<th>Response</th>
</tr>
</thead>
<tbody>
<tr>
<td>Aggregate score 0-4</td>
<td>Low</td>
<td>Ward-based response</td>
</tr>
<tr>
<td>Red score</td>
<td>Low-medium</td>
<td>Urgent ward-based responsea</td>
</tr>
<tr>
<td>Score of 3 in any individual parameter</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Aggregate score 5-6</td>
<td>Medium</td>
<td>Key threshold for urgent responseb</td>
</tr>
<tr>
<td>Aggregate score 7 or more</td>
<td>High</td>
<td>Urgent or emergency responseb</td>
</tr>
</tbody>
</table>

a Response by a clinician or team with competence in the assessment and treatment of acutely ill patients and in recognizing when the escalation of care to a critical care team is appropriate.

b The response team must also include staff with critical care skills, including airway management.

Source: Royal College of Physicians (2017).

Chart 3: Clinical response to NEWS trigger thresholds

<table>
<thead>
<tr>
<th>NEW score</th>
<th>Monitoring frequency</th>
<th>Clinical response</th>
</tr>
</thead>
<tbody>
<tr>
<td>0</td>
<td>Minimum 12 hourly</td>
<td>• Continue routine NEWS monitoring</td>
</tr>
<tr>
<td>Total 1-4</td>
<td>Minimum 4-6 hourly</td>
<td>• Inform registered nurse, who must assess the patient • Registered nurse decides whether increased frequency of monitoring and/or escalation of care is required</td>
</tr>
<tr>
<td>3 in single parameter</td>
<td>Minimum 1 hourly</td>
<td>• Registered nurse to immediately inform the medical team caring for the patient • Registered nurse to request urgent assessment by a clinician or team with core competencies in the care of acutely ill patients • Provide clinical care in an environment with monitoring facilities</td>
</tr>
<tr>
<td>Total 5 or more</td>
<td>Minimum 1 hourly</td>
<td>• Registered nurse to inform medical team caring for the patient, who will review and decide whether escalation of care is necessary</td>
</tr>
<tr>
<td>Urgent response threshold</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Total 7 or more</td>
<td>Continuous monitoring of vital signs</td>
<td>• Registered nurse to immediately inform the medical team caring for the patient - this should be at least at specialist registrar level • Emergency assessment by a team with critical care competencies, including practitioner(s) with advanced airway management skills • Consider transfer of care to a level 2 or 3 clinical care, i.e. higher dependency unit or ICU • Clinical care in an environment with monitoring facilities</td>
</tr>
</tbody>
</table>

Source: Royal College of Physicians (2017).
Chart 4: NEWS 2 observation chart

<table>
<thead>
<tr>
<th>NEWS key</th>
<th>FULL NAME</th>
<th>DATE OF BIRTH</th>
<th>DATE OF ADMISSION</th>
</tr>
</thead>
<tbody>
<tr>
<td>D</td>
<td>Alert</td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>Confusion</td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>V</td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>P</td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>U</td>
<td></td>
<td></td>
</tr>
<tr>
<td>E</td>
<td>Temperature</td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>38.1–39.0°C</td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>37.1–38.0°C</td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>36.1–37.0°C</td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>35.1–36.0°C</td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>≤35.0°C</td>
<td></td>
<td></td>
</tr>
<tr>
<td>NEWS TOTAL</td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Source: Royal College of Physicians (2017).
## 4.6 Paediatric Early Warning Score (PEWS)

This score was published in Critical Care in 2011 (see Parshuram et al., 2011), has been used in Canada and the United Kingdom of Great Britain and Northern Ireland, and has been shown to be clinically effective in low-resource settings (see Brown et al., 2019).

As in the adult scoring system, it is used to alert staff on general paediatric wards that a child is becoming critically unwell. The scoring system may need calibration or adjustment if used in a different environment to that for which it was developed. A score of 8 or more has a sensitivity of 83% for an impending emergency, including a possible cardiopulmonary arrest, and indicates that the child is critically ill and should be evaluated immediately by a physician and that a higher level of care should be considered. The seven items in the lefthand column should be scored and added together.

<table>
<thead>
<tr>
<th>Item</th>
<th>Age group</th>
<th>0</th>
<th>1</th>
<th>2</th>
<th>4</th>
</tr>
</thead>
<tbody>
<tr>
<td>HR (bpm)</td>
<td>0 to &lt; 3 m</td>
<td>&gt; 110 and &lt; 150</td>
<td>≥ 150 or ≤ 110</td>
<td>≥ 180 or ≤ 90</td>
<td>≥ 190 or ≤ 80</td>
</tr>
<tr>
<td></td>
<td>3 to &lt; 12 m</td>
<td>&gt; 100 and &lt; 150</td>
<td>≥ 150 or ≤ 100</td>
<td>≥ 170 or ≤ 80</td>
<td>≥ 180 or ≤ 70</td>
</tr>
<tr>
<td></td>
<td>1-4 yr</td>
<td>&gt; 90 and &lt; 120</td>
<td>≥ 120 or ≤ 90</td>
<td>≥ 150 or ≤ 70</td>
<td>≥ 170 or ≤ 60</td>
</tr>
<tr>
<td></td>
<td>&gt; 4-12 yr</td>
<td>&gt; 70 and &lt; 110</td>
<td>≥ 110 or ≤ 70</td>
<td>≥ 130 or ≤ 60</td>
<td>≥ 150 or ≤ 50</td>
</tr>
<tr>
<td></td>
<td>&gt; 12 yr</td>
<td>&gt; 60 and &lt; 100</td>
<td>≥ 100 or ≤ 60</td>
<td>≥ 120 or ≤ 50</td>
<td>≥ 140 or ≤ 40</td>
</tr>
<tr>
<td>SBP (mmHg)</td>
<td>0 to &lt; 3 m</td>
<td>&gt; 60 and &lt; 80</td>
<td>≥ 80 or ≤ 60</td>
<td>≥ 100 or ≤ 50</td>
<td>≥ 130 or ≤ 45</td>
</tr>
<tr>
<td></td>
<td>3 to &lt; 12 m</td>
<td>&gt; 80 and &lt; 100</td>
<td>≥ 100 or ≤ 80</td>
<td>≥ 120 or ≤ 70</td>
<td>≥ 150 or ≤ 60</td>
</tr>
<tr>
<td></td>
<td>1-4 yr</td>
<td>&gt; 90 and &lt; 110</td>
<td>≥ 110 or ≤ 90</td>
<td>≥ 125 or ≤ 75</td>
<td>≥ 160 or ≤ 60</td>
</tr>
<tr>
<td></td>
<td>&gt; 4-12 yr</td>
<td>&gt; 90 and &lt; 120</td>
<td>≥ 120 or ≤ 90</td>
<td>≥ 140 or ≤ 80</td>
<td>≥ 170 or ≤ 70</td>
</tr>
<tr>
<td></td>
<td>&gt; 12 yr</td>
<td>&gt; 100 and &lt; 130</td>
<td>≥ 130 or ≤ 100</td>
<td>≥ 150 or ≤ 85</td>
<td>≥ 190 or ≤ 75</td>
</tr>
<tr>
<td>CR time</td>
<td>&lt; 3 seconds</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>RR (breaths/min)</td>
<td>0 to &lt; 3 m</td>
<td>&gt; 29 and &lt; 61</td>
<td>≥ 61 or ≤ 29</td>
<td>≥ 81 or ≤ 19</td>
<td>≥ 91 or ≤ 15</td>
</tr>
<tr>
<td></td>
<td>3 to &lt; 12 m</td>
<td>&gt; 24 or &lt; 51</td>
<td>≥ 51 or ≤ 24</td>
<td>≥ 71 or ≤ 19</td>
<td>≥ 81 or ≤ 15</td>
</tr>
<tr>
<td></td>
<td>1-4 yr</td>
<td>&gt; 19 or &lt; 41</td>
<td>≥ 41 or ≤ 19</td>
<td>≥ 61 or ≤ 15</td>
<td>≥ 71 or ≤ 12</td>
</tr>
<tr>
<td></td>
<td>&gt; 4-12 yr</td>
<td>&gt; 19 or &lt; 31</td>
<td>≥ 31 or ≤ 19</td>
<td>≥ 41 or ≤ 14</td>
<td>≥ 51 or ≤ 10</td>
</tr>
<tr>
<td></td>
<td>&gt; 12 yr</td>
<td>&gt; 11 or &lt; 17</td>
<td>≥ 17 or ≤ 11</td>
<td>≥ 23 or ≤ 10</td>
<td>≥ 30 or ≤ 9</td>
</tr>
<tr>
<td>Respiratory effort</td>
<td>Normal</td>
<td>Milde increase</td>
<td>Moderate increase</td>
<td>Severe increase/any apnoea</td>
<td></td>
</tr>
<tr>
<td>SpO2 (%)</td>
<td>&gt; 94%</td>
<td>91% to 94%</td>
<td>≤ 90%</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Oxygen</td>
<td>Room air</td>
<td>&lt; 4 L/min or</td>
<td>≥ 4 L/min or</td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td></td>
<td>&lt; 50%</td>
<td>≥ 50%</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Notes: CR time - capillary refill time; HR - heart rate; RR - respiratory rate; SBP - systolic blood pressure; SpO2 - peripheral oxygen saturation.

Source: Parshuram et al. (2011).
<table>
<thead>
<tr>
<th>SEVERITY OF ILLNESS</th>
<th>MILD</th>
<th>MODERATE</th>
<th>SEVERE</th>
<th>CRITICAL</th>
</tr>
</thead>
<tbody>
<tr>
<td>Patient disposition</td>
<td>Home</td>
<td>Home(^b)</td>
<td>Inpatient ward</td>
<td>High dependency, step down or ICU</td>
</tr>
<tr>
<td>Vital signs</td>
<td></td>
<td></td>
<td></td>
<td>ICU</td>
</tr>
<tr>
<td>Temperature</td>
<td>Self-monitoring (initial assessment, daily and as needed)</td>
<td>Self-monitoring (initial assessment, daily and as needed)</td>
<td>Every 8-12 h</td>
<td>On admission and every 6-8 h</td>
</tr>
<tr>
<td>Oxygen saturation (SpO(_2))</td>
<td>No(^c)</td>
<td>Self-monitoring (initial assessment, daily and as needed)</td>
<td>No(^d)</td>
<td>Every 8-12 h</td>
</tr>
<tr>
<td>Respiration rate (RR)</td>
<td>No(^d)</td>
<td>Self-monitoring on initial assessment, daily and as needed</td>
<td>No(^d)</td>
<td>Every 8-12 h</td>
</tr>
<tr>
<td>Heart rate (HR) and regularity</td>
<td>No(^d)</td>
<td>Self-monitoring (initial assessment, daily and as needed) (if SpO(_2) is measured)</td>
<td>No(^d)</td>
<td>Every 8-12 h (if SpO(_2) is measured)</td>
</tr>
<tr>
<td>Blood pressure (BP)</td>
<td>No(^d)</td>
<td>N/A with telemedicine</td>
<td>No(^d)</td>
<td>Every 8-12 h</td>
</tr>
<tr>
<td>Assessment measures</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Symptoms’ assessment</td>
<td>Self-monitoring (initial assessment, daily and as needed)</td>
<td>Self-monitoring (initial assessment, daily and as needed)</td>
<td>Self-monitoring (initial assessment, daily and as needed)</td>
<td>Self-monitoring (initial assessment, daily and as needed)</td>
</tr>
<tr>
<td>Physical examination</td>
<td>No(^d)</td>
<td>No(^d)</td>
<td>No(^d)</td>
<td>No(^d)</td>
</tr>
<tr>
<td>Mental status (AVPU)</td>
<td>Self-monitoring (on initial assessment, daily and as needed)</td>
<td>Initial assessment, daily and as needed</td>
<td>Self-monitoring (initial assessment, daily and as needed)</td>
<td>Initial assessment, daily and as needed</td>
</tr>
</tbody>
</table>

To be adapted by context and local care pathway.
### Routine monitoring and care framework continued

<table>
<thead>
<tr>
<th>SEVERITY OF ILLNESS</th>
<th>MILD Without risk factors</th>
<th>MILD With ≥ 1 risk factor</th>
<th>MODERATE Without risk factor</th>
<th>MODERATE With ≥ 1 risk factor</th>
<th>SEVERE</th>
<th>CRITICAL</th>
</tr>
</thead>
<tbody>
<tr>
<td>Advanced monitoring tools: Intake and output measurements</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>Every 6-8 h</td>
<td>Every 4 h</td>
</tr>
<tr>
<td>3- or 5-lead electrocardiogram (ECG)</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>As needed</td>
<td>Continuous cardiac rhythm monitoring if needed</td>
</tr>
<tr>
<td>End-tidal CO₂ (EtCO₂)</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>As needed</td>
<td>Continuously or as frequent as possible in patients at risk of airway obstruction or hypoventilation</td>
</tr>
<tr>
<td>Arterial blood gases (ABGs)</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>As needed</td>
<td>As needed</td>
</tr>
<tr>
<td>Ventilator parameters</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
</tr>
<tr>
<td>Chest X-ray</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>As needed</td>
<td>As needed</td>
</tr>
<tr>
<td>12-lead ECG</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>As needed</td>
<td>As needed</td>
</tr>
</tbody>
</table>

N/A - not applicable/essential, but health care provider in charge of patient follow-up should consider if necessary.

**Notes:**

- **Patients with ≥ 1 risk factor and mild disease** who are not hospitalized should use pulse oximetry monitoring at home as part of a package of care (telemedicine, home hospitalization programmes) including patient and provider education and appropriate follow-up and report to health worker on a daily basis. The rest of the vital signs, such as temperature, RR, HR, BP, are recommended to be reported to the health care provider who follows up the patient at home if the equipment is available at home.

- **Patients without risk factors and moderate disease** can be initially managed at home with self-monitoring and reporting to a health worker if any complication or emergency signs occur.

- **Patients with ≥ 1 risk factor and moderate disease** should, preferably, be referred to a health care facility for monitoring and treatment. This is not possible, patients can be initially managed at home, preferably monitored by a health worker (telemedicine, home hospitalization programmes) at least once daily; and patients should record vital signs as detailed in the table above and report to the health worker on a daily basis.

- **The need for monitoring of vital signs (SpO₂, HR, RR, BP) and physical examination** should be assessed by health care provider who follows up the patient at home with telemedicine or home hospitalization programmes (ensure the equipment is available).

**Risk factors for severe disease**

Older age (> 60 years), hypertension, diabetes, cardiac disease, chronic lung disease, cerebrovascular disease, dementia, mental disorders, chronic kidney disease, immunosuppression (including HIV), obesity, cancer, pregnancy and post-partum period (up to 6 weeks) and unvaccinated for COVID-19. Risk factors in pregnant or recently pregnant: women > 35 years old, obesity, with chronic medical conditions or pregnancy specific disorders (e.g. gestational diabetes and pre-eclampsia/eclampsia).

Patients with any emergency sign at home, such as obstructed or absent breathing, severe respiratory distress, cyanosis, shock, coma and/or convulsions should seek immediate medical attention and be referred to a health facility.

4.8 WHO Mild COVID-19 home care bundle for health care workers

WHO Mild COVID-19 HOME CARE BUNDLE FOR HEALTH CARE WORKERS*

- Instruct patient to stay in isolation, preferably in separate room with adequate ventilation. Ensure good flow of fresh air and open windows where possible. Minimize close contact with others (households and/or visitors). If within 1 m of others, patient should wear a mask, and caregivers should wear PPE. Wash your hands regularly.
- Encourage patient to stay hydrated, eat well and take rest when needed but to try to resume activities at appropriate pace. Support patient’s psychosocial needs, such as through listening carefully to their needs and concerns and addressing them.
- Advise patient to monitor for worsening of symptoms, such as chest pain, fast or difficulty breathing (at rest or while speaking), fast heart rate, palpitations, confusion, altered mental status, or any other emergency signs. If present, instruct patient to call for emergency help according to national protocols.
- If patient is at risk for severe disease, monitor oxygen saturation with pulse oximeter, at least twice a day. If SpO₂ is <90%, instruct patient to call for emergency help. If SpO₂ is between 90-94%, call for urgent help, as this range may be an early sign for deterioration in someone with previously normal lungs. Oral corticosteroids may be prescribed at this time.

EMERGENCY SIGNS:
Obstructed or absent breathing, severe respiratory distress, cyanosis, shock, coma and/or convulsions.

Risk factors for severe disease includes: older age (> 60 years), hypertension, diabetes, cardiac disease, chronic lung disease, cerebrovascular disease, dementia, mental disorders, chronic kidney disease, immunosuppression (including HIV), obesity, cancer and unvaccinated against COVID-19. In pregnant or recently pregnant: women > 35 years old, obesity, with chronic medical conditions or pregnancy specific disorders (e.g. gestational diabetes and pre-eclampsia/eclampsia).

See Therapeutics and COVID-19: living guideline ( ) and Living guidance for clinical management of COVID-19 ( ).
WHO Mild COVID-19
HOME CARE BUNDLE
FOR HEALTH CARE WORKERS*

Severe disease
Adolescent or adult with clinical signs of pneumonia (fever, cough, dyspnoea) plus one of the following:
respiratory rate > 30 breaths/min; severe respiratory distress; or SpO₂ < 90% on room air at rest.

Child with clinical signs of pneumonia (cough or difficulty in breathing) + fast breathing or chest wall indrawing
+ at least one of the following:
• SpO₂ < 90%
• Very severe chest wall indrawing, grunting, central cyanosis or presence of any other general danger sign (inability
to breastfeed or drink, lethargy or unconsciousness or convulsions).

Critical COVID-19
Patient presenting with acute respiratory distress syndrome, sepsis, septic shock, acute thrombosis or other conditions
that normally require life-sustaining therapies.

CAUTION: The oxygen saturation threshold of 90% to define severe COVID-19 and should be
interpreted cautiously. For example, clinicians must use their judgment to determine whether
a low oxygen saturation is a sign of severity or is normal for a given patient with chronic lung disease.
Similarly, a saturation between 90-94% on room air may be abnormal (in patient with normal lungs)
and can be an early sign of severe disease, if patient is on a downward trend. Generally, if there is any
doubt, err on the side of considering the illness as severe.

Supplemental oxygen and humidification at home should be medically prescribed and supervised by a
health worker. Use only concentrators that are approved by the local authorities. Follow the instructions for
use and avoid flammable sources close by.

Criteria for discharging patients from isolation (i.e. discontinuing transmission-based precautions)
without requiring retesting:
• For symptomatic patients: 10 days after symptom onset, plus at least 3 additional days without symptoms
(including without fever and without respiratory symptoms).
• For asymptomatic cases: 10 days after positive test for SARS-CoV-2.

ADDITIONAL REFERENCES
WHO patient leaflet for the self-management of symptoms
WHO Healthy at Home
https://www.who.int/campaigns/connecting-the-world-to-combat-coronavirus/healthyathome

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https://www.who.int/publications/i/item/clinical-management-of-covid-19
https://www.who.int/teams/health-care-readiness-clinical-unit/covid-19

*This is a derivative product related to
the WHO COVID-19 Clinical management:
living guidance, Therapeutics and
COVID-19: living guideline, WHO Home
care for patients with suspected or
confirmed COVID-19 and OpenWHO.org.
Advice for health workers that are caring
for COVID-19 patients at home.
4.9 Memory aid: key criteria used to assess vital signs in children

<table>
<thead>
<tr>
<th>Age</th>
<th>&lt; 1 month</th>
<th>1 month - 1 year</th>
<th>1-5 years</th>
<th>5-12 years</th>
<th>&gt; 12 years</th>
</tr>
</thead>
<tbody>
<tr>
<td>Normal RR/min</td>
<td>30-40</td>
<td>30-40</td>
<td>20-30</td>
<td>20-25</td>
<td>12-20</td>
</tr>
<tr>
<td>RR/min in severe distress</td>
<td>&gt; 60 or &lt; 20</td>
<td>&gt; 50 or &lt; 10</td>
<td>&gt; 40</td>
<td>&gt; 40</td>
<td>&gt; 40</td>
</tr>
<tr>
<td>Normal heart rate (HR)/min</td>
<td>120-180</td>
<td>120-180</td>
<td>100-140</td>
<td>90-140</td>
<td>90-140</td>
</tr>
<tr>
<td>Normal SBP (mmHg)</td>
<td>60</td>
<td>80</td>
<td>90 + (2 × age)</td>
<td>120</td>
<td></td>
</tr>
<tr>
<td>Lower limit SBP (mmHg)</td>
<td>50</td>
<td>70</td>
<td>70 + (2 × age)</td>
<td>90</td>
<td></td>
</tr>
<tr>
<td>Normal urine output</td>
<td>1.2 mL/kg/hr</td>
<td>1 mL/kg/hr</td>
<td>0.5-1 mL/kg/hr</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Key tips for assessing a sick child

**Blood pressure measurement in children**
- Cuff should cover to ¾ of the upper arm, calf or thigh.
- Cuffs that are too small or too large give falsely high readings.
- Child should be at rest and not distressed (this will falsely elevate the reading).

**To perform capillary refill (CR) assessment**
- Press the nail bed of finger or thumb (peripheral CR) or over the sternum (central CR) for 3 seconds. Release and count in seconds the time taken for the return of colour (perfusion).

**Weight estimates in children**
It is always best to weigh children rather than estimate their weight.

*In an emergency*, weight can be estimated in visibly well-nourished children.
- Term infants: 2.5-4.5 kg.
- 6 months of age: 5-7 kg.
- After 1 year of age: (age in years + 4) × 2 kg.

**Criteria to define severe malnutrition**
- Clinical signs of severe malnutrition: visible ribs and no fat on the buttocks, thighs, arms or shoulders.
- Mid-upper arm circumference < 11.5 cm.
- Severe wasting: < 70% weight-for-length or -3SD on charts.
- Bilateral pedal oedema.

**Signs of respiratory distress**
- Fast RR.
- Nasal flaring, grunting.
- Intercostal recession and tracheal tug.
- Very severe: indrawing of the lower chest wall; central cyanosis of the lips and tongue; inability to breastfeed or drink; lethargy.
4.10 Memory aid: key physiological aspects to assess in pregnant women

<table>
<thead>
<tr>
<th>Measurement</th>
<th>% change</th>
<th>Absolute change</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cardiac output</td>
<td>30-50%</td>
<td>(2 L/min)</td>
</tr>
<tr>
<td>Heart rate</td>
<td>15-20%</td>
<td>(12 bpm)</td>
</tr>
<tr>
<td>Stroke volume</td>
<td>20-30%</td>
<td>(18 mL)</td>
</tr>
<tr>
<td>Mean arterial pressure</td>
<td>0-5%</td>
<td></td>
</tr>
<tr>
<td>Central venous pressure</td>
<td>No change</td>
<td></td>
</tr>
<tr>
<td>Systemic vascular resistance</td>
<td>20-30%</td>
<td>(320 dynes/cm²)</td>
</tr>
<tr>
<td>Left ventricular stroke work index</td>
<td>No change</td>
<td></td>
</tr>
<tr>
<td>Mean pulmonary artery pressure</td>
<td>No change</td>
<td></td>
</tr>
<tr>
<td>Pulmonary capillary wedge pressure</td>
<td>No change</td>
<td></td>
</tr>
<tr>
<td>Pulmonary vascular resistance</td>
<td>30%</td>
<td>(40 dynes/cm²)</td>
</tr>
</tbody>
</table>

Source: Adapted from Hegewald and Crapo (2011).

**Immune system**
- May increase susceptibility to intracellular pathogens such as viruses.
- Changes persist following the end of pregnancy.

**Cardiovascular**
- Blood volume increases by 40-50% causing dilutional anaemia and decreased oncotic pressure.
- Cardiac output increases by 30-50%.
- Heart rate increases by 10-20 bpm.
- Blood pressure decreases by 5-10 mmHg systolic and 10-15 mmHg diastolic. But after 24 weeks’ gestation, gradually increases to non-pregnant level by term.
- Systemic vascular resistance decreases by 20%.

**Respiratory**
- Increased tidal volume (TV) and minute ventilation. Chronic compensated respiratory alkalosis.
- No change in RR, tachypnoea is not a normal variant of pregnancy!
- No change in vital capacity.
- Increased oxygen consumption to 20-40% above non-pregnant levels.
- Decreased oxygen reserve (this makes pregnant patient more susceptible to effects of respiratory compromise).
Maternal–fetal dyad
- Fetus completely dependent on placenta for oxygen, nutrition and waste removal.
- Placenta is dependent on maternal blood cardiac output (500-800 mL of blood or 17% cardiac output goes to uterus every minute).
- With maternal compromise, blood flow will shunt away from uterus and this can occur before discernible maternal haemodynamic changes.
- If maternal oxygen or blood pressure decreases, the placenta will not be able to maintain adequate perfusion or oxygenation and the fetus will become distressed.

Key tips for complications in pregnant women

Tips regarding preterm labour
- Tocolytics may worsen maternal status by decreasing blood pressure, tachycardia, arrhythmias or causing pulmonary oedema.
- Antenatal corticosteroids promote fetal lung maturation if there is need to deliver fetus preterm (weeks 24–34). Can use betamethasone 12 mg IM every 24 hours for two doses or dexamethasone 6 mg IM every 12 hours for four doses.

Tips for managing respiratory distress
- Keep SpO2 ≥ 92-95%.
- Do not delay intubation for worsening respiratory distress. Be prepared for difficult airway!

Tips for managing hypotension
- Ensure adequate resuscitation but avoid fluid overload.
- Do not lay flat. Position with lateral tilt (elevate either hip 10–12 cm) to augment venous return to heart.
- Cautious vasopressor use as risk of reducing uterine perfusion, must monitor fetus.

Tips regarding antimicrobial therapy
- For suspected influenza virus infection, it is safe to treat with oseltamivir and give as soon as possible.
- Also give antibiotics - penicillins, cephalosporins and macrolides are appropriate in pregnancy.
- Avoid fluoroquinolones and doxycycline if possible.
References and resources


Silcock DJ, Corfield AR, Gowen PA, Rooney KD. Validation of the National Early Warning Score in the prehospital setting. Resuscitation. 2015;89:31-5.


Key resources for supporting the management of SARI

**Basic emergency care (BEC): approach to the acutely ill and injured (2018)**
Developed by WHO and ICRC, in collaboration with the International Federation for Emergency Medicine, *Basic emergency care (BEC): approach to the acutely ill and injured* is an open-access training course for frontline health care providers who manage acute illness and injury with limited resources. Produced in response to requests from multiple countries and international partners, the BEC package includes a participant workbook and electronic slide decks for each module. Integrating the guidance from WHO Emergency triage, assessment and treatment (ETAT) for children and the Integrated management of adult/adolescent illness (IMAI), BEC teaches a systematic approach to the initial assessment and management of time-sensitive conditions where early intervention saves lives.

This is for use by doctors, nurses and other health workers caring for children at first level referral hospitals with basic laboratory facilities and essential medicines. These guidelines focus on the management of the major causes of childhood mortality in most developing countries including pneumonia, and also cover common procedures, patient monitoring and supportive care on the wards.

**Oxygen therapy for children (2016)**
This is a bedside manual for health workers to guide the provision of oxygen therapy for children. The manual focuses on the availability and clinical use of oxygen therapy in children in health facilities to guide health workers, biomedical engineers and administrators. It addresses detection of hypoxaemia, use of pulse oximetry, clinical use of oxygen, delivery systems and monitoring of patients on oxygen therapy. The manual also addresses the practical use of pulse oximetry, and oxygen concentrators and cylinders.

The manual is written for clinicians working at the district hospital (first-level referral care) who diagnose and manage sick adolescents and adults in resource-constrained settings. It aims to support clinical reasoning, and to provide an effective clinical approach and protocols for the management of common and serious or potentially life-threatening conditions at district hospitals. The target audience thus includes doctors, clinical officers, health officers and senior nurse practitioners. It has been designed to be applicable in both high- and low-HIV prevalence settings.

**Technical specifications for oxygen concentrators (2015)**
This provides an overview of oxygen concentrators and technical specifications to aid in selection, procurement and quality assurance. It highlights the minimum performance requirements and technical characteristics for oxygen concentrators and related equipment that are suitable for the use in health facilities.

The purpose of this document is to increase access to quality products to ensure the supply of oxygen, especially in low- and middle-income countries and low-resource settings within countries from all income groupings. This project is one of many related to improving oxygen supply that other stakeholders are working on. These efforts aim to support ministries of health to ensure oxygen supply is available, as well as raise awareness of the importance of appropriate selection, procurement, maintenance and use of medical devices, both capital equipment and single-use devices.

WHO Priority medical devices list for the COVID-19 response and associated technical specifications (November 2020)

This document describes the medical devices required for the clinical management of COVID-19, selected and prioritized according to the latest available evidence and interim guidelines. It includes: oxygen therapy, pulse oximeters, patient monitors, thermometers, infusion and suction pumps, X-ray, ultrasound and CT scanners as well as PPE. In order to facilitate access to quality assured priority medical devices, the document also includes technical and performance characteristics, related standards, accessories and consumables. It is intended for policy-makers and planning officers in ministries of health, procurement and regulatory agencies, intergovernmental and international agencies as well as the medical device industry.
Diagnostic testing for patients with ARI
5 Diagnostic testing for patients with ARI

Summary

The differential diagnoses for SARI include a wide spectrum of community-acquired pathogens, including respiratory viruses, bacteria and other less common micro-organisms. The ranking of differential diagnoses will vary by host factors (e.g. age, presence of chronic conditions, travel history, vaccination), environmental factors (e.g. geographic location, vectors), local epidemiology (e.g. the prevalence of the pathogen in the community, endemic infections) and pathogen factors (e.g. tropism for lungs). See Chapter 1 for more details. Diagnostic testing should be conducted based on the differential diagnosis and pre-test clinical likelihood of disease.

In patients presenting with ARI, the collection of upper respiratory tract samples is recommended to guide further management. Testing should be done as soon as possible and inform treatment plans. Use appropriate IPC precautions when collecting specimens. See Infection prevention and control during health care when coronavirus disease (COVID-19) is suspected or confirmed ( ).

Collect specimens from the upper respiratory tract (URT: nasopharyngeal and oropharyngeal) AND, where clinical suspicion remains and URT specimens are negative, collect specimens from the lower respiratory tract when readily available (LRT: expectorated sputum, endotracheal aspirate or bronchoalveolar lavage in ventilated patient) for SARS-CoV-2 testing by RT-PCR or antigen detection rapid diagnostic tests and bacterial stains/cultures.

If patient presents with signs or symptoms that meet criteria for SARI, also collect:
- Samples of blood and sputum for bacterial culture, to diagnose potential bacterial cause of pneumonia and sepsis, ideally before antimicrobial therapy. However, do not delay empiric and appropriate antimicrobial treatment.

Other considerations for areas with endemic infections should align with local protocols:
- In malaria-endemic areas, patients with fever should be tested for malaria and treated appropriately.
- In areas where arbovirus infection is endemic, then testing for dengue, chikungunya or yellow fever, may be considered in patients with undifferentiated febrile illness, particularly when thrombocytopenia is present.
- In areas where there is high prevalence of TB, TB testing can also be considered.
- Co-infection with SARS-CoV-2 may also occur and a positive diagnostic test for dengue does not exclude the testing for COVID-19.

See WHO Diagnostic testing for SARS-CoV-2 infection ( ).
See WHO TB guidelines: recent updates ( ).
See WHO Malaria testing ( )
Tools
5.1 Diagnostic testing for SARS-CoV-2 infection
5.2 Use of antigen-detection rapid diagnostic testing for SARS-CoV-2
5.3 Specimen collection kit for upper respiratory tract specimens
  5.3.1 Nasopharyngeal swab technique
  5.3.2 Posterior pharyngeal swab or throat swab technique
  5.3.3 Tracheal aspirate technique
5.4 Guideline for specimen storage
5.5 Material for specimen transportation
5.6 Guideline for specimen transportation
5.7 Guide for blood culture collection in patients with SARI
5.1 Diagnostic testing for SARS-CoV-2 infection

Diagnostic testing for SARS-CoV-2 infection

Countries need to test for SARS-CoV-2 according to the national strategy, using available and approved diagnostic tests. WHO recommends testing of all SARS-CoV-2 suspected cases.

Types of tests:

- **Nucleic acid amplification testing**
  - Detects genetic material of the virus
  - Uses upper respiratory specimens* to diagnose acute SARS-CoV-2 infection.
  - Nucleic acid amplification testing (NAAT), for example RT-PCR, is the reference method for detection of acute SARS-CoV-2 infection.
  - Results: usually available within 24 hours. Testing takes 30 minutes to 4 hours (depending on the test), but transport to the testing laboratory can add hours to days.

- **Antigen**
  - Detection rapid diagnostic testing – detects viral protein(s)
  - Uses upper respiratory specimens* to diagnose acute SARS-CoV-2 infection.
  - Performance is best within first 5-7 days of symptoms.
  - Results: within 15-30 minutes, not requiring laboratory infrastructure.

- **Antibody testing**
  - Detects antibodies against the virus
  - Uses serum/plasma or whole blood specimens to detect antibodies generated by prior SARS-CoV-2 infection or vaccination.
  - SARS-CoV-2 antibodies are usually detectable 1-2 weeks after infection or vaccination.
  - Results: within 24 hours; point of care tests within 10-30 mins.

*Some NAA tests and some Ag-POC are designed to work on upper respiratory tract samples or saliva
For more information: https://www.youtube.com/watch?v=Phs5dJu_GQ0

For further information see WHO Diagnostic testing for SARS-CoV-2 infection and video on testing for COVID-19.
5.2 Use of antigen-detection rapid diagnostic testing for SARS-CoV-2

Use of antigen-detection rapid diagnostic testing

WHO recommends that all suspected cases be tested for SARS-CoV-2

Antigen-detection diagnostic testing
Uses upper respiratory specimen or saliva to test for SARS-CoV-2 infection by detecting viral proteins (e.g. nucleoprotein).

Can be used outside of clinical and laboratory settings, including in communities (*)

1. CASE FINDING
As a primary case-detection test for testing all suspected cases.
Can be used to detect SARS-CoV-2 in many settings, including in health facilities, testing centres, care homes, prisons, schools, communities where there is ongoing transmission.

2. CONTACT TRACING
To identify infection among asymptomatic contacts of cases. (1)

3. OUTBREAK INVESTIGATIONS
To confirm suspected outbreaks of COVID-19 among symptomatic individuals, especially in closed or semi-closed settings including schools, care homes, cruise ships, prisons, work places and dormitories, etc.

4. MONITOR TRENDS
To monitor trends in disease incidence in communities, and particularly among frequently exposed workers, including health and care workers, irrespective of symptoms.

*AgRDT testing should be performed by trained individuals. WHO training materials can be found here: https://extranet.who.int/tdg/content/sars-cov-2-antigen-rapid-diagnostic-test-training-package

1 Ag-RDTs can be used to test asymptomatic contacts of confirmed cases, even if the Ag-RDT is not specifically authorized for this use.

For further information see WHO guidance September 2020: https://www.who.int/publications/i/item/antigen-detection-in-the-diagnosis-of-sars-cov-2-infection-using-rapid-immunoassays
5.3 Specimen collection kit for upper respiratory tract specimens

It is best to compile a specimen collection kit before starting to take specimens.

Here is an inventory of all items that should be in the specimen collection kit for URT specimens.

**Required items**

- PPE (gloves, medical or FFP2/N95 mask, gown, face shield/goggles)
- ice packs/cooler box
- field collection forms
- an alcohol-resistant pen or marker for labelling samples
- sterile Dacron or rayon swabs
- 1–2 mL viral transport medium (VTM)*
- specimen collection containers.

* When antigen rapid diagnostic tests (Ag-RDT) are conducted, sampling/collection material used should follow the manufacturer’s instructions. Often VTM is not suitable for RDT. However, having VTM samples is critical for subsequent culture and often for PCR (important for influenza surveillance); thus if Ag-RDT is to be done, an additional sample should be taken.

**Technique**

1. Disinfect bottles.
2. Swab with rigid (plastic) shaft for throat and nasal specimens.
3. Use tongue depressors for throat swabs.
4. Use sterile saline (0.9% NS) for nasopharyngeal aspiration.
5. Use sputum or mucus trap for nasopharyngeal aspiration (also require negative pressure).

**Swabs**

The type of swab used is very important.

Only sterile Dacron or rayon swabs with aluminum or plastic shafts should be used. This is because calcium alginate or cotton swabs, or swabs with wooden sticks, may contain substances that inactivate some viruses and inhibit PCR testing.
5.3.1 Nasopharyngeal swab technique

Required materials
- swab with flexible (aluminium or plastic) shaft.

Technique
1. Apply standard, contact and droplet precautions.
2. Insert swab into one nostril and back into the nasopharynx.
3. Leave swab in place for a few seconds.
4. Then slowly remove swab while rotating it over surface of posterior nasopharynx.
5. Withdraw swab from collection site; insert into transport tube or container with VTM.
7. After collection, immediately transport specimen to the laboratory for viral PCR testing and/or viral antigen detection. If transport to the laboratory is delayed, place specimen on ice or in refrigeration.

In case of nasopharyngeal swab in infants and young children:
- Use a swab of appropriate size: measure the distance from the nose to the ear (philtrum to the tragus).
- Insert the swab half to full amount of that distance, stopping if you encounter resistance.
- Insert the swab horizontally, below the inferior turbinate, not diagonally up the nose.

How to collect oropharyngeal and nasopharyngeal specimens for the diagnosis of COVID-19 (© WHO)
5.3.2 Posterior pharyngeal swab or throat swab technique

**Required materials**
- swab with rigid (plastic) shaft
- tongue depressor.

**Technique**
1. Apply standard, contact and droplet precautions.
2. Ask the subject to open his or her mouth and say “ah” to elevate the uvula.
3. Depress the tongue to hold out of way with tongue depressor.
4. Swab the posterior pharynx and do not touch tongue with swab.
5. Insert into transport tube or container with VTM. Break applicator tip to ensure closure of vial.
7. Immediately transport specimen to the laboratory for viral PCR testing and/or viral antigen detection. If transport to the laboratory is delayed, place specimen on ice or in refrigeration.

For further information see *Optimal insertion depth for nasal mid-turbinate and nasopharyngeal swabs* (Callesan et al., 2021) and video on collecting oropharyngeal and nasopharyngeal specimens for the diagnosis of COVID-19.

© WHO
5.3.3 Tracheal aspirate technique

Intended for patients intubated and receiving invasive mechanical ventilation (IMV).

**Required materials**
- suction outlet (portable or wall)
- sterile suction catheter
- specimen mucus trap (i.e. Lucken’s tube)
- sterile saline (0.9% NS)
- IPC for airborne precautions (N-95 particulate mask)
- a sterile suction catheter (not a closed, inline system)
- suction tubing
- airway emergency equipment.

**Technique**
1. Apply standard, contact, droplet and airborne precautions.
2. Prepare patient: pre-oxygenate with 100% fraction of inspired oxygen (FiO₂). Give adequate sedation.
3. Attach mucous trap to catheter and suction outlet. Turn on suction to make sure functioning. Then turn it off.
4. When you are ready, disconnect ventilator tubing from endotracheal tube.
5. Without applying suction, insert sterile suction catheter apparatus into endotracheal tube, about 2–3 cm beyond tip.
6. Apply suction and collect sample into the mucous trap. Hold trap upright to prevent secretions from going into the pump. Slowly withdraw catheter. Replace ventilator tubing.
7. If inadequate sample, instil 3-5 mL of sterile saline, give two insufflations/deep breaths and apply suction.
8. After collection, immediately transport specimen to laboratory for viral testing and bacteriology.
9. Store in refrigerator (2-8 °C) for maximum 24 hours.
10. If delay, store in freezer < -20 °C.
5.4 Guideline for specimen storage

Viral transport medium is used immediately after the collection of samples for viral isolation and testing. It prevents the specimen from drying out and prevents bacterial and fungal growth.

Although you should send specimens in VTM to the laboratory as soon as possible, it is important to properly store them before you send them to a laboratory if there is a delay.

Do not freeze samples in the standard freezer. It is very important to avoid freeze-thaw cycles because this will destroy some types of virus. It is better to keep a sample on ice even for a week, than to allow the sample to freeze and thaw multiple times.

Viral transport medium information

Possible suppliers
Local laboratory and commercial supplier.

Description
It is usually supplied in the form of 1-3 mL of VTM in sterile container.

Stock management
It is important that clinicians liaise with the laboratory to make sure that there is sufficient stock of VTM available at facility, and that it is stored in an area which is accessible to clinicians when needed.

Conservation
If VTM must be stored for long periods, this should be done in a freezer at -20 °C. For short periods of time VTM may be stored in a fridge at 4-6 °C.
5.5 Material for specimen transportation

When you are ready to pack specimens, no more than 500 mL should be in the specimen container. For transportation from the field to the laboratory, you must use three packaging layers. This is done to protect specimens from damage during transportation.

**Required materials**
- primary waterproof container (e.g. Falcon tube)
- absorbent material:
  - bubble wrap
  - secondary recipient
  - cooler box
  - ice packs
  - sample identification form.

**Packing and labelling of infectious substances not refrigerated**

For SARS-CoV-2, specimens should be classified as Category A, UN2814, “infectious substance, affecting humans”. Clinical samples will be classified as UN3373, “Biological Substance Category B”.

For more details review:
- *Laboratory biosafety guidance related to coronavirus disease (COVID-19)*
- *Guidance on regulations for the transport of infectious substances 2021-2022*
5.6 Guideline for specimen transportation

1 Envelop the cryo-tube with blotting paper.

Place all components in a waterproof secondary recipient container and close in order to be watertight.

Place the primary waterproof container in bubble wrap or a shock-absorbing material.

Place all components in a waterproof secondary recipient container and close in order to be watertight.

Place ice packs in the cooler box. Put the filled secondary container in the cooler box. The recipient container should be in a vertical position.

Insert the sample identification form in a zip bag and place the zip bag in the cooler box, next to the secondary recipient container.

Close the cooler box in order to be watertight. Write expeditor and addressee on the external part of the cooler box. Put infectious substance label if necessary.

Source: Adapted from influenza sentinel surveillance training. Institute Pasteur of Madagascar, CDC and WHO.
5.7 Guide for blood culture collection in patients with SARI

Blood cultures should be obtained before starting antimicrobial therapy in all patients with sepsis in the hospital. The Surviving Sepsis Campaign guidelines caution that this should not delay antimicrobial treatment by more than 45 minutes.

This technique is adapted from the United States Centers for Disease Control and Prevention (CDC) website.

Required materials
- PPE (gloves and mask)
- alcohol swabs
- chlorhexidine swabs (associated with less contamination than standard povidone-iodine)
- blood culture bottles (two bottles per set, one anaerobe and one aerobe)
- two sterile needles (adult: 22 gauge; paediatric: 25 gauge)
- two syringes (adult 20 mL; paediatric 5 mL)
- tourniquet
- sterile gauze pad
- adhesive tape
- patient labels
- plastic zip lock bag for transport.

Technique
1. Check patient ID, explain procedure.
2. Hand washing.
3. Disinfect bottle tops with 70% isopropyl alcohol (alcohol pad) in a circular motion, allow to dry.
5. Puncture the vein with clean needle. Use sterile gloves if you plan to palpate vein after cleaning site.
6. For adults, collect 10-20 mL; for children, collect 3-5 mL for each blood culture set.
7. Remove needle from vein, divide blood into two blood culture bottles, by placing same needle perpendicularly into the bottle. Do not overfill bottles. If not enough for both bottles, preferably start filling the aerobic bottle. There are systems with bottles that can tap the blood directly from the vein. If possible, this system is preferred.
8. Gently rotate bottle to mix blood and broth.
9. Two blood cultures (by separate stick) per septic episode is sufficient (except in endocarditis).
10. Place label and put into plastic bag and send to the laboratory.

Contaminated blood culture
If skin is not adequately cleansed before obtaining the culture, or the procedure for taking the culture is not done carefully and cleanly, bacteria from the skin may be injected into the bottle, producing contamination and a false positive blood culture. This may lead to misdiagnosis and prolonged antimicrobial use.
References and resources


Oxygen therapy
6 Oxygen therapy

Summary

The administration of supplemental oxygen is necessary when a patient has low blood oxygen levels, termed hypoxaemia. Untreated, acute hypoxaemia can lead to tissue hypoxia (low level of oxygen at the cellular level), organ dysfunction and death. Delivery of oxygen to tissues also relies on adequate cardiac output and haemoglobin to carry oxygen to the tissues.

Oxygen delivery ($DO_2$) = cardiac output (CO) × oxygen content in arterial blood ($CaO_2$)

- $CaO_2 = 1.34 \times (Hb) \times SpO_2 + (0.003 \times PaO_2)$
- $CO = SV \times HR$

Oxygen therapy improves oxygen delivery to the tissues by increasing oxygen content in the blood. Oxygen content in the blood is frequently measured by pulse oximetry as oxygen saturation ($SpO_2$).

Oxygen therapy provides patients with an oxygen concentration greater than that in ambient air (> 0.21). While oxygen therapy can be generated from various sources and applied via various delivery mechanisms, the fundamentals of oxygen therapy remain the same.

1. **Oxygen concentration (%) $O_2$** from the oxygen source (i.e. liquid oxygen, pressure swing adsorption [PSA], bedside concentrator) is the purity of oxygen produced by the device. It can range widely depending on the quality of the source but generally should range from 0.82-1.0 (82-100%).

2. **Fraction of inspired oxygen (%) $FiO_2$** is the oxygen concentration that is inspired by the patient, usually a result of mixing the oxygen source and ambient air and can range from 0.21-1.0 (21-100%). This varies depending on the delivery device and the patient’s respiratory drive.

3. **Oxygen saturation (%) $SpO_2$** and oxygen partial pressure ($PaO_2$) are the measured oxygen levels in blood, the former measured by pulse oximeter and the latter by blood gas analyser.

**Pulse oximeters** should be available in all clinical areas where oxygen is delivered. Blood gas analysers and end tidal carbon dioxide ($EtCO_2$) monitors should ideally be available at least in ICU or critical care units to measure ventilatory parameters ($EtCO_2$, pH, $PaCO_2$) (see Chapter 4: Monitoring patients with acute respiratory infection).
Oxygen delivery devices should be selected based on the oxygen needs of the patient and include nasal cannulas, conventional face masks, Venturi face masks and face masks with reservoir bag. If patients require higher levels of oxygen flow (> 10-15 L/min) to reach SpO₂ targets, and/or have other signs of acute respiratory failure, other respiratory support options, that can deliver higher flows and/or positive pressure support should be considered. These include:
- high-flow nasal oxygen (HFNO);
- non-invasive ventilation devices (CPAP, BiPAP delivered via oronasal mask, full face mask, helmet or nasal interface);
- invasive mechanical ventilation (IMV) delivered via endotracheal tube. Do not delay intubation if there are urgent indications.

Note: Use of HFNO, CPAP, BiPAP and IMV for treatment of severe/critical COVID-19 may consume significant quantities of oxygen.

Rational oxygen use delivers the least amount of oxygen necessary to achieve SpO₂ goals. Although avoiding hypoxaemia is the primary goal of oxygen therapy, it is important to remember that giving a patient more oxygen than they need to meet the SpO₂ goal can be harmful and more rapidly deplete oxygen supplies. Therefore, titration up and down of oxygen therapy and trials of oxygen weaning should be performed on a regular basis (see how to titrate oxygen with different devices: Tools 6.5-6.7).

Oxygen sources should routinely be checked for quality, including oxygen concentration and flow or pressure output.

Tools
- 6.1 Indications for oxygen therapy
- 6.2 Memory aid: oxygen delivery devices
- 6.3 Memory aid: oxygen delivery in children
- 6.4 Algorithm to escalate respiratory support in adults and children with pneumonia
- 6.5 Flowchart on how to titrate oxygen in neonates
- 6.6 Flowchart on how to titrate oxygen in children
- 6.7 Flowchart on how to titrate oxygen in adults
- 6.8 Key tips on awake prone positioning
- 6.9 Checklist to troubleshoot warning signs during oxygen therapy delivery
- 6.10 Oxygen supply calculations
- 6.11 Memory aids: oxygen supply sources and distribution
- 6.12 Respiratory care order template for oxygen therapy
6.1 Indications for oxygen therapy

Give oxygen immediately to any patient (adult or child) with:

- Respiratory distress
- Sepsis with hypoperfusion or shock
- Alteration of mental status
- Hypoxaemia

- $\text{SpO}_2 < 90\%$ (if patient is haemodynamically stable)
- $\text{SpO}_2 < 94\%$ (if patient with any emergency signs, with or without respiratory distress)
- $\text{SpO}_2 < 92–95\%$ (if pregnant)

*Note:* Emergency signs - obstructed or absent breathing, severe respiratory distress, central cyanosis, shock, coma and/or convulsions - may require airway management in addition to oxygen.

Supplemental oxygen therapy should be provided immediately to achieve the following $\text{SpO}_2$ targets:

<table>
<thead>
<tr>
<th>$\text{SpO}_2$ targets</th>
</tr>
</thead>
<tbody>
<tr>
<td>$\text{SpO}_2 \geq 90%$ in adults and children</td>
</tr>
<tr>
<td>$\text{SpO}_2 \geq 92-95%$ in pregnant patients</td>
</tr>
<tr>
<td>$\text{SpO}_2 \geq 94%$ if child or adult with signs of multi-organ failure, including shock, alteration of mental status, severe anaemia, and/or during resuscitation. Once stabilized, resume target $\text{SpO}_2 \geq 90%$.</td>
</tr>
</tbody>
</table>
6.2 Memory aid: oxygen delivery devices

### Non-invasive O₂ delivery devices

<table>
<thead>
<tr>
<th>Device</th>
<th>O₂ flow</th>
<th>FiO₂</th>
</tr>
</thead>
<tbody>
<tr>
<td>Nasal cannula</td>
<td>1–5 L/min</td>
<td>0.23–0.35</td>
</tr>
<tr>
<td>Face mask</td>
<td>5–10 L/min</td>
<td>0.30–0.50</td>
</tr>
<tr>
<td>Venturi face mask</td>
<td>2–15 L/min</td>
<td>0.24–0.6</td>
</tr>
<tr>
<td>Face mask reservoir</td>
<td>10–15 L/min</td>
<td>0.5–0.95</td>
</tr>
</tbody>
</table>

### Advanced non-invasive O₂ delivery devices

<table>
<thead>
<tr>
<th>Device</th>
<th>O₂ flow</th>
<th>FiO₂</th>
</tr>
</thead>
<tbody>
<tr>
<td>High-flow nasal oxygen</td>
<td>10–60 L/min</td>
<td>0.23–1.0</td>
</tr>
<tr>
<td>BiPAP/CPAP</td>
<td>10–80 L/min</td>
<td>0.21–1.0</td>
</tr>
</tbody>
</table>

* O₂ flow ranges differ for neonates, children and adults; see Tool 6.2 for ranges by age.
* Delivered O₂ concentration depends on multiple factors including the concentration of the oxygen source and the patient’s respiratory pattern (e.g. peak inspiratory flow and minute ventilation).
* O₂ consumption for BiPAP/CPAP is widely variable depending on device used and the leak of the system.

Source: USAID-STAR-UCSF OpenCriticalCare.org Project (illustrations Holly Sullivan)
## Oxygen delivery devices (single use)

<table>
<thead>
<tr>
<th>Device Description</th>
<th>Typical flow rate range</th>
<th>FlO₂ delivered (range)</th>
<th>Considerations</th>
<th>Advantages/disadvantages</th>
<th>Can be used with</th>
</tr>
</thead>
<tbody>
<tr>
<td>Nasal cannula, adult and paediatric (single use)</td>
<td>1-6 L/min</td>
<td>24-44% oxygen</td>
<td>Increases by approximately 4% with each litre of oxygen per minute</td>
<td>Technically, it is possible to deliver higher flows with this device, however, the oxygen source should deliver the desired flow; it can dry the nasal mucosa and disturb sleeping patterns; humidification might be required according to clinical guidance. For paediatric patients with a flow &gt; 4 L/min humidification is necessary (WHO)</td>
<td>Advantages: Easy to use; Patient can eat and talk. Disadvantages: Can be easily dislodged and is not as effective in patients with deviated septum or polyps</td>
</tr>
<tr>
<td>Venturi mask; adult, paediatric</td>
<td>2-15 L/min</td>
<td>24-60% oxygen, according to the type of mask</td>
<td>Allows precise measurement of FlO₂ delivered</td>
<td>Utilizes different sized ports to change the FlO₂ delivered (24-50%) Some brands relate a colour to a flow rate and FlO₂ delivered, e.g. blue = 2-4 L/min = 24%; white = 4-6 L/min = 28%; yellow = 8-10 L/min = 35%; red = 10-12 L/min = 40%; green = 12-15 L/min = 60%.</td>
<td>Advantages: Precise measurement of FlO₂ delivered; Does not dry mucous membranes. Disadvantages: Is confining for some patients; It interferes with talking and eating.</td>
</tr>
<tr>
<td>Mask with reservoir bag; adult</td>
<td>&gt; 10 L/min</td>
<td>80-95% oxygen FlO₂; depends on the patient’s pattern of breathing</td>
<td>Non-rebreather mask with reservoir bag</td>
<td></td>
<td>Advantages: Delivers high concentration of oxygen. Disadvantages: Oxygen flow should be &gt; 10 L/min; less can cause the bag to collapse during inspiration.</td>
</tr>
</tbody>
</table>

**Disclaimer:** This table is intended to provide information from the technical point of view about oxygen delivery devices, including flow rate ranges, achievable FlO₂, and possible oxygen sources that can be used with each device. Clinical decisions should determine the methods of administration of oxygen therapy and device selection.

**Source:** Priority medical devices list for the COVID-19 response and associated technical specifications (WHO, 2020).
6.3 Memory aid: oxygen delivery in children

Nasal cannula is the preferred method of delivering oxygen to infants and children < 5 years of age with hypoxaemia who require oxygen therapy.

<table>
<thead>
<tr>
<th>Age of child</th>
<th>Nasal cannula size</th>
<th>Maximal oxygen flow rates</th>
</tr>
</thead>
<tbody>
<tr>
<td>Neonates</td>
<td>Neonatal</td>
<td>0.5-1.0 L/min by nasal cannula</td>
</tr>
<tr>
<td>Infants</td>
<td>Infant</td>
<td>1-2 L/min by nasal cannula</td>
</tr>
<tr>
<td>Pre-school aged</td>
<td>Child</td>
<td>1-4 L/min by nasal cannula</td>
</tr>
<tr>
<td>School-aged</td>
<td>Child to adult</td>
<td>1-5 L/min by nasal cannula</td>
</tr>
</tbody>
</table>

Practical considerations

- **Fitting:** The distal prong should fit well into the nostril (premature infants: 1 mm; infants weighing up to 10 kg: 2 mm). The prongs should be secured with a piece of tape on the cheeks near the nose as shown. Care should be taken to keep the nostrils clear of mucus to avoid blockage.

- **Humidification:**

  When oxygen is delivered at standard flow rate (0.5-1 L/min for a neonate, 1-2 L/min for an infant, 1-4 L/min for an older child) through nasal prongs, humidification is not necessary.

  Higher flow rates without effective humidification may cause drying of nasal mucosa with associated bleeding, airway obstruction and may cause bronchoconstriction in some conditions (e.g. asthma, COPD).

- Humidification is essential when cold oxygen is delivered from a cylinder, compared with concentrators (mainly in tropical countries), as normally concentrators provide oxygen at room temperature.

- Bubble humidifiers reduce the dryness of the oxygen supplied by bubbling the gas through water at room temperature. They must be filled with clean water at least once a day (distilled water or tap water that has been boiled and cooled). The water level in the humidifier should be checked twice daily and topped up as necessary. The humidifier equipment must be washed and disinfected regularly to prevent bacterial colonization.
6.4 Algorithm to escalate respiratory support in adults and children with pneumonia

Assess patient
Are any of the following signs of severe pneumonia present? Clinical signs of pneumonia and any of the following:
- Adults: RR > 30/min, severe respiratory distress, or SpO₂ < 90%
- Child: very severe chest indrawing, grunting, central cyanosis, or presence of any other general danger signs (inability to breastfeed or drink, lethargy or unconsciousness or convulsions).

Continue monitoring and reassess (~q2–6 hr)
Is the patient meeting treatment goals?
- SpO₂ (>90%)
- Improving work of breathing
- pH > 7.30
- PaCO₂ < 45
- Adequate mental status
If not, consider alternative therapies, escalation, or transfer to a higher level of care

Assess response
Increased distress or SpO₂ < 90%?

Initiate
Nasal cannula
0.5–1 LPM neonates; 1–2 LPM infants; 1–4 LPM adults and school children

Assess response
Increased distress or SpO₂ < 90%?

Consider intubation and mechanical ventilation
Use lung protective ventilation or highest level of support available (e.g., bag mask ventilation for respiratory failure) and transfer to a ward or facility where mechanical ventilation can be done safely

Titration and SpO₂ goals
- SpO₂ ≥ 90% in adults and children
- SpO₂ ≥ 92–95% in pregnant patients
- SpO₂ ≥ 94% if child or adult with signs of multi-organ failure
- Avoid SpO₂ > 98% to avoid ill effects of hypoxemia and to preserve O₂ supply

Adjuncts
- Ensure adequate PPE and IPC
- Place medical mask over nasal cannula, face mask, non-rebreather mask with reservoir, or high-flow nasal cannula to decrease risk of droplet and aerosolization
- Prone positioning
- Reposition airway (shoulder roll, jaw thrust)
- Elevate the head of the bed
- Provide chest physiotherapy
- Oral and/or nasal suctioning

CPAP/NIPPV contraindications
- Aspiration risk
- Inability to protect airway or remove mask
- Hemodynamic instability
- Abnormal mental status
- Need for emergent intubation
- Anatomic barriers to mask seal (relative)
- Insufficient respiratory drive/effort

Non-rebreather mask + reservoir
10–15 LPM
May consider adding nasal cannula to augment FiO₂ especially if devices that deliver FiO₂ 1.0 are limited.

Assess response
Increased distress or SpO₂ < 90%?

Simple face mask
5–10 LPM

Assess response
Increased distress or SpO₂ < 90%?

Are emergency signs present?

Consider 1–2 hr trial of one of the following based on patient & local context

• Oral/NIV ventilation
• High-flow nasal oxygen

6.5 Flowchart on how to titrate oxygen in neonates

- Start nasal cannula oxygen at:
  0.5-1 L/min neonates
  1-2 L/min infants

  If continued distress or SpO₂ < 90%
  (or < 94% if emergency signs)

- Consider face mask with reservoir
  only if HFNO, CPAP, bCPAP, BiPAP
  unavailable

- Start oxygen at 10-15 L/min and titrate
  to ensure bag inflates

  If continued distress or SpO₂ < 90%
  (or < 94% if emergency signs)

Find higher level of care and consider one
of the following if adequate O₂ supply:

- HFNO: 0-10 kg 2 L/kg/min
  10-20 kg 1 L/kg/min

- CPAP: 5-10 cmH₂O

- Bubble CPAP: start flow at 5 L/min;
  immerse expiratory limb (cm depth in H₂O
  equals CPAP "pressure"; look for bubbles;
  titrate to 10 L/min if needed to generate
  bubbles

- BiPAP: PS Δ5-15/PEEP 5-15
  i.e. IPAP 10-30/EPAP 5-15
  (IPAP = PEEP + PS; EPAP = PEEP)

If continued respiratory distress or SpO₂ < 90% on 15 L/min, consider intubation and invasive mechanical ventilation (clinical management decisions should be made based on individual patient characteristics, local resources and expertise).

Wean O₂ flow and avoid SpO₂ 100% to avoid ill effects of hyperoxia and excess O₂ consumption.

Source: USAID-STAR-UCSF OpenCriticalCare.org Project (Illustrations Holly Sullivan)
6.6 Flowchart on how to titrate oxygen in children

A medical mask should be placed over nasal cannula and HFNO for patients with suspected or confirmed highly infectious respiratory illness (e.g. COVID-19).

- Start oxygen at 1-5 L/min
- Use nasal prongs
- Assess response

If continued distress or SpO₂ < 90% (or < 94% if emergency signs)

- Use face mask
- Increase oxygen to 5-10 L/min
- Assess response

If continued distress or SpO₂ < 90% (or < 94% if emergency signs)

- Use face mask with reservoir
- Start oxygen at 10-15 L/min and titrate to ensure bag inflates

If continued distress or SpO₂ < 90% (or < 94% if emergency signs)

Transfer to higher level care and consider:

- HFNO: 10-20 kg 1 L/kg/min
  20-40 kg 0.5-1 L/kg/min (max 30)
  > 40 kg 0.5-1 L/kg/min (max 60)
- CPAP: 5-10 cmH₂O
- BiPAP: PS Δ5-15/PEEP 5-15
  i.e. IPAP 10-30/EPAP 5-15
  (IPAP = PEEP + PS; EPAP = PEEP)

If continued respiratory distress or SpO₂ < 90% on 15 L/min, consider intubation and invasive mechanical ventilation (clinical management decisions should be made based on individual patient characteristics, local resources and expertise).

Wean O₂ flow and avoid SpO₂ 100% to avoid ill effects of hyperoxia and excess O₂ consumption.

Source: USAID-STAR-UCSF OpenCriticalCare.org Project (illustrations Holly Sullivan) (P)
6.7 Flowchart on how to titrate oxygen in adults

A medical mask should be placed over nasal cannulas for patients with suspected or confirmed highly infectious respiratory illness (e.g. COVID-19)

- Preferably, if available, use Venturi mask instead of face mask
- Titrate oxygen 2-15 L/min by colour

Heated humidification systems should be used with HFNO and BiPAP/CPAP

- Start oxygen at 1-5 L/min
- Use nasal prongs
- Assess response

If continued distress or SpO₂ < 90% (or < 94% if pregnant or emergency signs)

- Use face mask
- Increase oxygen to 5-10 L/min
- Assess response

If continued distress or SpO₂ < 90% (or < 94% if pregnant or emergency signs)

- Use face mask with reservoir
- Start oxygen at 10-15 L/min and titrate to ensure bag inflates

If continued distress or SpO₂ < 90% (or < 94% if pregnant or emergency signs)

Transfer to higher level care and consider:
- HFNO: 30-60 L/min (may also adjust FiO₂)
- CPAP: 10-15 cmH₂O
- BiPAP: PS Δ5-15/PEEP 5-15 i.e. IPAP 10–30/EPAP 5–15 (IPAP = PEEP + PS; EPAP = PEEP)

If continued respiratory distress or SpO₂ < 90% on 15 L/min, consider intubation and invasive mechanical ventilation (clinical management decisions should be made based on individual patient characteristics, local resources and expertise).

Wean O₂ flow and avoid SpO₂ 100% to avoid ill effects of hyperoxia and excess O₂ consumption.

6.8 Key tips on awake prone positioning

Prone positioning (lying on their front) in awake spontaneously breathing patients, may improve hypoxaemia. This prone positioning has been often used in patients with moderate-severe ARDS requiring invasive mechanical ventilation and is associated with improved oxygenation and reduced mortality in these patients.

The WHO COVID-19 Clinical Guideline Development Group conditionally recommends:
Awake prone positioning of patients severely ill and hospitalized with COVID-19 requiring supplemental oxygen (including high-flow nasal cannula) or non-invasive ventilation.
- **Benefits:** observational studies of awake prone patients with severe COVID-19 suggest decreased mortality and need for intubation (very low certainty evidence).
- **Harms:** include possible patient discomfort and pain (very low certainty evidence).

Source: USAID-STAR-UCSF OpenCriticalCare.org Project (illustrations Holly Sullivan)

Indications and contraindications for awake prone positioning

<table>
<thead>
<tr>
<th>Characteristics of patients appropriate for prone position</th>
<th>Contraindications to prone positioning</th>
</tr>
</thead>
<tbody>
<tr>
<td>• Awake and alert.</td>
<td>• Need for immediate intubation.</td>
</tr>
<tr>
<td>• Capable of communicating and moving independently.</td>
<td>• Haemodynamically unstable (tachycardia, hypotension).</td>
</tr>
<tr>
<td>• Patient must be able get help if they have discomfort or pain.</td>
<td>• Spinal instability.</td>
</tr>
<tr>
<td>• Patient must be able to supinate independently, if needed.</td>
<td>• Altered mental status or reduced ability to protect the airway.</td>
</tr>
<tr>
<td>• Haemodynamically stable.</td>
<td>• Unable to readily call for help if needed.</td>
</tr>
<tr>
<td>• Able to protect their airway.</td>
<td>• Caution if nausea or vomiting.</td>
</tr>
<tr>
<td>• Able to be closely monitored by workers with experience with prone positioning.</td>
<td>• Not enough human resources in the unit to monitor.</td>
</tr>
</tbody>
</table>
Proning positioning tips

**Characteristics of patients appropriate for prone position**
- Patients should attempt to prone on a regular basis (e.g. every 4 hours) and maintain the prone position for as long as possible. (Many patients are unable to maintain the prone position for more than 1-2 hours.)
- Patients should be able to stop proning at any time and return to the supine position as needed.

**Rotation and timed position changes**
- Regimens vary, and target being in awake prone position 8-12 hours/day, broken into shorter periods over the day. For example, some institutional protocols describe rotational protocols, with patients changing position on a regular schedule (e.g. every 1 hour changing position, with positions rotating from prone, to lying on right side, to sitting straight upright, to lying on left side, to prone again, etc.).

**Patient comfort: frequent limitations for patients are low back pain, nausea and vomiting**
- For nausea or vomiting, immediately assist the patient to an upright position or recovery position. Gently suction or wipe the airway, if the patient cannot clear spontaneously.
- For low back pain, patients may find comfort using padding (i.e. pillows, blankets) under the pelvis.
- If possible, tilt the bed slightly in reverse Trendelenburg position to reduce pressure on the eyes and face.
6.9 Checklist to troubleshoot warning signs during oxygen therapy delivery

✔ If respiratory distress and hypoxaemia fail to improve despite increasing oxygen, use a systematic approach to manage your patient. Consider using this checklist. Repeat the quick check BEC with ABCDE approach (see Chapter 2: Screening, triage, clinical assessment and monitoring of SARI).
## Checklist to troubleshoot warning signs during oxygen therapy delivery

### Equipment

**Is the measurement correct?**
- Repeat measurement:
  - Place pulse oximeter correctly; try another measurement location (e.g. a different finger or the ear or nose if adapter available).
  - Check pulse oximeter plethysmograph or signal quality.
  - Use another pulse oximeter or get an arterial blood gas (if appropriate).

**Is there technical difficulty in delivering treatments?**
- Is the mask of the appropriate size for the patient (neonate, infant, child, adult)?
- Check that the oxygen source is working:

<table>
<thead>
<tr>
<th>Cylinder</th>
<th>Yes</th>
<th>No</th>
</tr>
</thead>
<tbody>
<tr>
<td>Does the cylinder contain oxygen (or another gas)?</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Does the cylinder contain sufficient oxygen?</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Is the pressure above 200 psi?</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Is there any leakage in the circuit?</td>
<td></td>
<td></td>
</tr>
<tr>
<td>• tubing</td>
<td></td>
<td></td>
</tr>
<tr>
<td>• connections</td>
<td></td>
<td></td>
</tr>
<tr>
<td>• masks</td>
<td></td>
<td></td>
</tr>
<tr>
<td>If using the mask with reservoir bag, does it fill up?</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Concentrator</th>
<th>Yes</th>
<th>No</th>
</tr>
</thead>
<tbody>
<tr>
<td>Is it connected to the electricity source and the power is on?</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Is the oxygen purity measured (or displayed in the screen) above 82%?</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Is there any leakage in the circuit?</td>
<td></td>
<td></td>
</tr>
<tr>
<td>• tubing</td>
<td></td>
<td></td>
</tr>
<tr>
<td>• connections</td>
<td></td>
<td></td>
</tr>
<tr>
<td>• masks</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Are the flows and pressures correct for the type of concentrator used?</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

### Patient and treatments

**Is there an alternate or additional diagnosis?**
- Does the patient have acute heart failure?
- Does the patient have pleural effusions?
- Does the patient have pulmonary embolism?

**Is the patient getting appropriate therapy for the correct diagnosis?**
- Ensure underlying etiology is being appropriately managed (e.g. antimicrobials given for pneumonia).

**Is our treatment causing harm?**
- Consider complications and modify management accordingly (e.g. too much fluid leading to pulmonary oedema? Allergic reaction to medication?).
- Does the patient have hypoxemia that is refractory to high-flow oxygen (e.g. significant shunt from ARDS)?
- Consider initiation of mechanical ventilator support for management of respiratory failure.

**If the patient’s mental status deteriorates despite SpO2 > 90%, consider the following:**
- Manage airway, assist ventilation if needed - do not wait for arterial blood gas results if the patient requires assisted ventilation on clinical grounds.
- Check arterial blood gas, if available, to evaluate ventilation. CO2 retention causing acute respiratory acidosis will not be detected with SpO2 alone.
- Consider alternate causes of altered mental status and treat appropriately (e.g. acute central nervous system [CNS] event, electrolyte abnormalities, low glucose).
6.10 Oxygen supply calculations

The ability to administer oxygen to patients with SARI depends on several factors including the supply of oxygen, the device(s) and flow rates being used to deliver oxygen, and the number of patients with SARI who are being treated.

At the facility level, there are many considerations required to estimate oxygen demand, including:

1. **Oxygen source type:** This may include pressure swing absorption (PSA), vacuum pressure swing adsorption (VPSA) plants, bulk liquid tank, bedside oxygen concentrators, distribution manifolds with high-pressure cylinders, or a combination of sources.

2. **Distribution system:** Oxygen may travel from the source to the patient’s bedside in two main ways: pipe network or bedside high-pressure gas cylinders (or bedside concentrators). Piping networks should be designed to fit in infrastructure; sectioning of the pipes, alarms and monitoring components should be carefully planned. Accessories for monitoring and regulation of the medical gas should be included (e.g. pressure regulators and flowmeters).

3. **Delivery equipment and devices:** This refers to biomedical equipment connected to the network (e.g. patient ventilators, or direct interfaces such as nasal cannula).

Forecasting oxygen demand can be done using different methodologies and includes considerations like number of beds, hypoxaemia rates per ward, bed occupancy rates, and distance from the main source. Currently, various tools are publicly available to estimate oxygen needs at facility, subnational or national level.

Useful links:
- USAID Open Critical Care.org Project oxygen supply and demand calculator (🔗)
- UNICEF: Oxygen system planning tool (🔗)
- PATH: Oxygen delivery toolkit (🔗)
6.11 Memory aids: oxygen supply sources and distribution

Overview of oxygen supply sources

<table>
<thead>
<tr>
<th>Description</th>
<th>PSA bedside concentrators</th>
<th>PSA or VSA O₃ generator plants</th>
<th>Cryogenic liquid</th>
</tr>
</thead>
<tbody>
<tr>
<td>Different flow rates, typical: 5, 8 or 10 L/min - medical use</td>
<td>Various sizes and configurations: single and duplex 2-200+ Nm³/hr</td>
<td>Produced mainly for heavy industry; serves medical sector where GMP allows</td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Requirements</th>
<th>PSA bedside concentrators</th>
<th>PSA or VSA O₃ generator plants</th>
<th>Cryogenic liquid</th>
</tr>
</thead>
<tbody>
<tr>
<td>• Situated onsite, bedside.</td>
<td>• Various own-operate models.</td>
<td>• Third party responsible for production and supply chain.</td>
<td></td>
</tr>
<tr>
<td>• Continuous and reliable electrical source.</td>
<td>• Often situated onsite.</td>
<td>• Plants must be offshore. Bulk liquid tanks with passive</td>
<td></td>
</tr>
<tr>
<td>• Device-specific spares needed.</td>
<td>• Continuous and reliable electrical source during plant</td>
<td>storage (specialized materials).</td>
<td></td>
</tr>
<tr>
<td>• Time technical maintenance (preventive every 6 months).</td>
<td>and booster operations.</td>
<td>Capital expenditure (CAPEX) and operating expenses (OPEX)</td>
<td></td>
</tr>
<tr>
<td>• Need for PC measures as is situated bedside.</td>
<td>• Detailed technical and financial planning for long-term</td>
<td>are very right, borne by third party.</td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Additional considerations</th>
<th>PSA bedside concentrators</th>
<th>PSA or VSA O₃ generator plants</th>
<th>Cryogenic liquid</th>
</tr>
</thead>
<tbody>
<tr>
<td>• Difficult to optimize for all scale needs.</td>
<td>• Need ≥ 4 technicians for 24/7 operation.</td>
<td>• Goods and service contract.</td>
<td></td>
</tr>
<tr>
<td>• Not suitable for high-flow or higher pressure needs (e.g., patient ventilators).</td>
<td>• Continuous supply at all atmospheric pressures.</td>
<td>• Product can be used via high-pressure gas cylinders or</td>
<td></td>
</tr>
<tr>
<td>• Depending on the capacity and oxygen therapy, flow could be split among patients.</td>
<td>• Supply can be piped bedside and/or plant can fill cylinders to be used bedside or transported elsewhere.</td>
<td>piped bedside from bulk tank.</td>
<td></td>
</tr>
</tbody>
</table>

Primary system: PSA plan

Primary system: cryogenic liquid oxygen

Source: Oxygen access scale up (WHO, 2020).
Cylinders

There is significant variability in terminology (i.e. letter system) and the colour coding systems to label oxygen cylinders. Always check with the cylinder manufacturer and consider the dimensions of the cylinder for accurate capacity information. More information on oxygen cylinder standards can be found in the WHO-UNICEF technical specifications and guidance for oxygen therapy devices (https://www.bochealthcare.co.uk/en/images/cylinder_data_med309965_2011_tcm409-54065.pdf, accessed 12 June 2019).

<table>
<thead>
<tr>
<th>Cylinder size</th>
<th>D</th>
<th>E</th>
<th>F</th>
<th>G</th>
<th>J</th>
</tr>
</thead>
<tbody>
<tr>
<td>Nominal content/oxygen capacity (L)</td>
<td>340 L</td>
<td>680 L</td>
<td>1360 L</td>
<td>3400 L</td>
<td>6800 L</td>
</tr>
<tr>
<td>Nominal content (m^3)</td>
<td>0.34 m^3</td>
<td>0.68 m^3</td>
<td>1.36 m^3</td>
<td>3.4 m^3</td>
<td>6.8 m^3</td>
</tr>
<tr>
<td>Water capacity (L)</td>
<td>2.3 L</td>
<td>4.7 L</td>
<td>9.4 L</td>
<td>23.6 L</td>
<td>47.2 L</td>
</tr>
<tr>
<td>Dimensions (height × diameter) (mm)</td>
<td>535 × 102</td>
<td>865 × 102</td>
<td>930 × 140</td>
<td>1320 × 178</td>
<td>1520 × 229</td>
</tr>
<tr>
<td>Approximate full weight (kg)</td>
<td>3.9</td>
<td>6.5</td>
<td>17</td>
<td>39</td>
<td>78</td>
</tr>
<tr>
<td>Nominal service pressure (kPa/bar/psi)</td>
<td>13 700 kPa (137 bar/1987 psi)</td>
<td>13 700 kPa (137 bar/1987 psi)</td>
<td>13 700 kPa (137 bar/1987 psi)</td>
<td>13 700 kPa (137 bar/1987 psi)</td>
<td>13 700 kPa (137 bar/1987 psi)</td>
</tr>
<tr>
<td>Health facility use</td>
<td>Emergency and ambulance transport</td>
<td>Emergency and ambulance transport</td>
<td>Stand-alone</td>
<td>Stand-alone</td>
<td>Manifold connection and stand-alone</td>
</tr>
</tbody>
</table>

### Duration of cylinders

<table>
<thead>
<tr>
<th>Rate of oxygen administration for one patient</th>
<th>Cylinder size</th>
<th></th>
<th></th>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>D 340 L</td>
<td>E 680 L</td>
<td>F 1360 L</td>
<td>G 3400 L</td>
<td>J 6800 L</td>
</tr>
<tr>
<td>2 L/min</td>
<td>2 hr 50 min</td>
<td>5 hr 40 min</td>
<td>11 hr 20 min</td>
<td>28 hr 20 min</td>
<td>56 hr</td>
</tr>
<tr>
<td></td>
<td>8.5 tanks</td>
<td>4 tanks</td>
<td>2 tanks</td>
<td>1 tanks</td>
<td>0.5 tanks</td>
</tr>
<tr>
<td>5 L/min</td>
<td>1 hr 8 min</td>
<td>2 hr 16 min</td>
<td>4 hr 30 min</td>
<td>11 hr 20 min</td>
<td>23 hr</td>
</tr>
<tr>
<td></td>
<td>21 tanks</td>
<td>10 tanks</td>
<td>2 tanks</td>
<td>1 tanks</td>
<td></td>
</tr>
<tr>
<td>8 L/min</td>
<td>42 min</td>
<td>1 hr 24 min</td>
<td>2 hr 50 min</td>
<td>7 hr</td>
<td>14 hr</td>
</tr>
<tr>
<td></td>
<td>34 tanks</td>
<td>17 tanks</td>
<td>8 tanks</td>
<td>4 tanks</td>
<td></td>
</tr>
<tr>
<td>10 L/min</td>
<td>34 min</td>
<td>1 hr 8 min</td>
<td>2 hr 16 min</td>
<td>5 hr 50 min</td>
<td>11 hr</td>
</tr>
<tr>
<td></td>
<td>42 tanks</td>
<td>21 tanks</td>
<td>10 tanks</td>
<td>4 tanks</td>
<td></td>
</tr>
</tbody>
</table>

**Top row:** duration of cylinder  
**Bottom row:** number of cylinders required/24 hr

### Concentrators

Source: USAID-STAR-UCSF OpenCriticalCare.org Project.

![Concentrators diagram](image)
There are multiple devices available and approved, with different power efficiencies and a range of minimum and maximum oxygen output between a minimum of 0.125 L/min to a maximum of 10 L/min. Few portable oxygen concentrators can deliver more than 10 L/min, and many oxygen concentrators may claim a wide range of flows but only with low oxygen concentrations. Always check manufacturers’ specifications and WHO technical specifications for oxygen concentrators (WHO, 2015).

Oxygen concentrators are easily movable by an individual and can be located some distance from the flowmeter assembly. This is useful in situation where different patients must share the same oxygen source using a flowmeter stand (mainly for children).
### 6.12 Respiratory care order template for oxygen therapy

**Oxygen therapy order set v1.2**

<table>
<thead>
<tr>
<th>Hospital name/Logo</th>
<th>Oxygen therapy order set v1.2</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Surname/Family name</strong></td>
<td><strong>Name</strong></td>
</tr>
<tr>
<td><strong>Date</strong></td>
<td><strong>/ /</strong></td>
</tr>
</tbody>
</table>

**CHOOSE A DELIVERY DEVICE**

- **Nasal cannula**
  - Titrating flow rate _____ L/min to maintain oxygen saturation (SpO₂) ≥ 95%
  - Set flow rate to: _____ L/min (typical range 0.5-1 LPM)
  - Attach room temperature bubble humidifier and change sterile distilled H₂O (per protocol every) or [_____ hours]
  - Consider 0% - 4LPM for infants, especially in paediatrics (may evidence benefit is lacking)

- **Simple face mask**
  - Titrating flow rate from 5-10 L/min to maintain oxygen saturation (SpO₂) by pulse oximeter to goal ≥ 90% - 95%
  - Set flow rate to: _____ L/min (typical range 5-10 LPM)
  - Attach room temperature bubble humidifier and change sterile distilled H₂O (per protocol every) or [_____ hours]
  - Consider especially in paediatrics (any evidence of benefit is lacking)

- **Face mask with reservoir bag**
  - Titrating flow rate from 10-15 L/min to maintain oxygen saturation (SpO₂) by pulse oximeter to goal ≥ 90% - 94%
  - Set flow rate to: _____ L/min (typical range 10-15 LPM)
  - Attach room temperature bubble humidifier and change sterile distilled H₂O (per protocol every) or [_____ hours]

- **Venturi mask**
  - Select Venturi device or settings (FiO₂ 0.4-0.6) to maintain oxygen saturation (SpO₂) by pulse oximeter to goal = 91% - 96%
  - Set flow rate to: _____ L/min (0.4-0.6)
  - Note: Do not use humidifiers with Venturi masks as moisture may affect accuracy of FiO₂

- **High-flow nasal cannula (HFNO)**
  - Titrating flow rate with 1.0 FiO₂ to maintain oxygen saturation (SpO₂) by pulse oximeter to goal ≥ 90% - 95%
  - Set flow rate to: _____ L/min and rate of expiratory FiO₂ = 1.0

**Pulse oximetry**

- In addition to routine monitors, check oxygen saturation (SpO₂) continuously or every [_____ hours] minutes

---

*SpO₂ goal = 90% in adults and children; SpO₂ goal = 92% - 95% in pregnant patients; SpO₂ goal = 94% if child or adult with signs of multi-organ failure, including shock, alteration of mental status, severe anaemia and/or ongoing resuscitation. (Once stabilized, resume target SpO₂ = 90%.)

---

**Date (time):**

**Name:**

**Signature:**

**Contact #:**

---

**Source:** USAID-STAR-UCSF OpenCriticalCare.org Project.
References and resources


Therapies for SARI (COVID-19, influenza, bacterial pneumonia): antimicrobials and immunomodulators
7 Therapies for SARI (COVID-19, influenza, bacterial pneumonia): antimicrobials and immunomodulators

Summary

Patients with SARI should receive optimized supportive care, as soon as possible, and this includes oxygen therapy. In addition, based on the differential diagnosis and results of diagnostic tests (when available) appropriate antimicrobial and/or immunomodulator therapy should be given.

In patients with mild confirmed COVID-19, empirical antibiotic or antifungal therapy or prophylaxis is not recommended. In general, the use of empiric antibiotics or antifungals should be discouraged in COVID-19 cases, as their use may lead to higher bacterial or fungal resistance rates, which will impact the burden of disease and deaths in a population during the pandemic and beyond. If patient is at high risk for severe COVID-19, see WHO Therapeutics and COVID-19: living guideline for current recommendations.

In patients with moderate COVID-19 with non-severe pneumonia, antibiotics or antifungal should also not be prescribed unless there is clinical suspicion of a bacterial or fungal infection. A recent systematic review of critical patients hospitalized with COVID-19 reported only 8% of patients experiencing bacterial/fungal co-infection during hospital admission. If patient is at high risk for severe COVID-19, see WHO Therapeutics and COVID-19: living guideline for current recommendations.

In patients with severe and critical COVID-19 (severe pneumonia, ARDS, sepsis, septic shock), the following treatments should be administered immediately (within 1 hour): empiric antimicrobial therapy if suspect bacterial co-infection, initiation of fluid bolus and/or vasopressors for hypotension (see Chapter 8 for details) AND COVID-19 therapeutics: see WHO Therapeutics and COVID-19: living guideline for most recent recommendations; in addition to treatment with corticosteroids and either IL-6 receptor blockers or baricitinib.

When seasonal influenza A or B viruses are confirmed or suspected to be circulating in the community, treat patients at risk and those with SARI and patients as soon as possible with oseltamivir AND antimicrobials for all likely pathogens (if bacteria are considered) as soon as possible (within 1 hour). Patients with suspected or confirmed avian influenza should also be treated. Oseltamivir is a neuraminidase inhibitor antiviral drug and is active against all currently circulating influenza viruses that infect humans. See Guidelines for the clinical management of severe illness from Influenza virus infections.

In settings endemic for other infections, such as TB and malaria, patients should be screened as per usual local protocols, and if screened positive, appropriate PPE should be donned and appropriate treatment started. For example, in areas where TB is prevalent and a patient is suspected to have COVID-19 and/or TB, appropriate PPE should be donned for possible COVID-19 and TB immediately. In malaria-endemic areas, patients with fever should be tested for the presence of malaria and treated as appropriate.

A positive test for another pathogen does not rule out a SARS-CoV-2 infection and a positive test for SARS-CoV-2 does not rule out co-infection with another pathogen or other etiology for the patient’s symptoms.
Tools

7.1 Memory aid: treatment for acute respiratory infections according to severity (when COVID-19 and influenza are circulating)

7.2 COVID-19 and therapeutics
   7.2.1 Corticosteroids for COVID-19
   7.2.2 Interleukin-6 receptor blockers for COVID-19: tocilizumab, sarilumab
   7.2.3 Monoclonal antibodies for COVID-19: casirivimab and imdevimab

7.3 Memory aid: invasive fungal infections in patients with COVID-19

7.4 Treatment for influenza infection fact sheet
7.1 Memory aid: treatment for acute respiratory infections according to severity (when COVID-19 and influenza are circulating)

Testing for different pathogens causing SARI should follow the national protocols.

<table>
<thead>
<tr>
<th>Adults</th>
<th>Mild disease</th>
<th>Pneumonia</th>
<th>Severe pneumonia</th>
</tr>
</thead>
</table>
|                 | Patients with uncomplicated upper respiratory tract viral infection may have non-specific symptoms such as fever, fatigue, cough (with or without sputum production), anorexia, malaise, muscle pain, sore throat, dyspnoea, nasal congestion or headache. Rarely, patients may also present with diarrhoea, nausea and vomiting. The elderly, immunosuppressed and pregnant women may present with atypical symptoms (mainly COVID-19). | Fever, cough, dyspnoea, fast breathing, but no signs of severe pneumonia, including SpO₂ ≥ 90% on room air. | Clinical signs of pneumonia (fever, cough, dyspnoea) plus one of the following:  
> respiratory rate > 30 breaths/min  
> severe respiratory distress  
> SpO₂ < 90% on room air. |

| Children        | Patients with uncomplicated upper respiratory tract viral infection may have non-specific symptoms such as fever, fatigue, cough (with or without sputum production), anorexia, malaise, muscle pain, sore throat, dyspnoea, nasal congestion or headache. Rarely, patients may also present with diarrhoea, nausea and vomiting. | Cough or difficulty breathing, plus fast breathing* and/or chest indrawing but no signs of severe pneumonia. | Pneumonia plus at least one of the following signs of severe pneumonia:  
> SpO₂ < 90%  
> Very severe chest wall indrawing, grunting, central cyanosis, or presence of any other general danger sign (inability to breastfeed or drink, lethargy or unconsciousness, or convulsions). |

| Treatment       | • Isolation in hospital, community facility or home care (if required for highly infectious pathogen).  
• Give antipyretics for fever. Monitor and refer immediately if signs of decompensation.  
If COVID-19 and risk factors: consider if therapies are required (see Therapeutics and COVID-19: living guideline  
If influenza circulating and risk factors for severe disease: consider if oseltamivir should be prescribed (see Guidelines for the clinical management of severe illness from influenza virus infections). | • Isolation in hospital, community facility or home care depending on risk factors (if required for highly infectious pathogen).  
• Give antipyretics for fever.  
• Give appropriate antibiotic if required if suspicion of bacterial source or co-infection.  
Monitor (including saturation of oxygen with pulse oximeter twice daily for patients of high risk of severity) and refer immediately if signs of decompensation.  
If COVID-19 and risk factors: consider if therapies are required (see Therapeutics and COVID-19: living guideline).  
If influenza circulating and risk factors for severe disease: consider if oseltamivir should be prescribed (see Guidelines for the clinical management of severe illness from influenza virus infections). | • Isolation and treatment in a hospital (if required for highly infectious pathogen), consider intensive care.  
• Give antipyretics for fever.  
• Give recommended antibiotic.  
• Manage airway as appropriate.  
• Give oxygen if:  
  – SpO₂ < 90% and haemodynamically stable  
  – SpO₂ < 94% and with any emergency signs (obstructed or absent breathing, severe respiratory distress, central cyanosis, shock, coma or convulsions) with or without respiratory distress  
  – SpO₂ < 92-95% if pregnant.  
• If COVID-19 suspected, give steroids and IL-6 Rb and/or other medications (see Therapeutics and COVID-19: living guideline).  
• If influenza circulating treat (oseltamivir) (see Guidelines for the clinical management of severe illness from influenza virus infections).  
Monitor for signs of decompensation. |

* Fast breathing is defined by age:  
  < 2 months: ≥ 60 breaths/min; 2-11 months: ≥ 50 breaths/min; 1-5 years: ≥ 40 breaths/min.

Empiric antibiotics for severe pneumonia and sepsis in adults

For severe pneumonia, and sepsis in adults, give empirical broad-spectrum IV antimicrobials within the first hour for suspected pathogens. This is crucially important. Refer to national or institutional recommendations. Common choices include:

### Empiric antibiotic treatment for community-acquired bacterial pneumonia in adults

<table>
<thead>
<tr>
<th></th>
<th>Adults</th>
<th>Total treatment duration</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Mild to moderate cases</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>FIRST CHOICE</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Amoxicillin (oral):</td>
<td>1 g given every 8 hours</td>
<td>5 days</td>
</tr>
<tr>
<td>Phenoxyymethylpenicillin (oral):</td>
<td>500 mg given every 6 hours</td>
<td></td>
</tr>
<tr>
<td><strong>SECOND CHOICE</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Amoxicillin+clavulanic acid (oral):</td>
<td>875 mg + 125 mg given every 8 hours</td>
<td></td>
</tr>
<tr>
<td>Doxycycline (oral):</td>
<td>100 mg given every 12 hours</td>
<td></td>
</tr>
</tbody>
</table>

| **Severe cases**   |        |                          |
| **FIRST CHOICE**   |        |                          |
| Ceftriaxone (IV/IM): | 2 g given once a day (IV), 1 g given once a day (IM) | 5 days (consider longer treatment and/or investigate for complications if the patient is not clinically stable at Day 5) |
| Cefotaxime (IV/IM): | 2 g given every 8 hours |                          |
| if CURB-65 ≥ 2 CONSIDER ADDING Clarithromycin b (oral or IV): | 500 mg given every 12 hour |                          |
| **SECOND CHOICE**  |        |                          |
| Amoxicillin+clavulanic acid (IV): | 1 g + 200 mg given every 8 hours |                      |
| if CURB-65 ≥ 2 CONSIDER ADDING Clarithromycin b (oral or IV): | 500 mg given every 12 hour |                          |

**Notes:**
- IM: intramuscular; IV: intravenous; IU: international units.
- Doxycycline is contraindicated in pregnant women.
- The rationale of adding clarithromycin to beta-lactam is to cover for possible atypical bacteria. Azithromycin could be used as an alternative when clarithromycin is not available but there are increasing concerns about its potential for the emergence and spread of antibiotic resistance because of its long half-life. Erythromycin could also be considered but it is associated with higher toxicity (diarrhoea is frequently associated with its use).
- All dosages are for normal renal function.

**ACCESS** antibiotics are indicated in green, **WATCH** antibiotics in yellow and **RESERVE** antibiotics in red.
### Empiric antibiotic treatment for community-acquired sepsis of bacterial origin in adults

<table>
<thead>
<tr>
<th>Most probable source of infection</th>
<th>Empiric antibiotic treatment</th>
<th>Total treatment duration</th>
</tr>
</thead>
<tbody>
<tr>
<td>Clinical sepsis of unknown origin*</td>
<td>Ceftriaxone (IV): 2 g given once a day or Cefotaxime (IV): 2 g given every 8 hours and Gentamicin (IV): 5 mg/kg given once a day or Amikacin (IV): 15 mg/kg given once a day</td>
<td>7 days (but duration depends on the patient’s underlying disease and clinical progression)</td>
</tr>
</tbody>
</table>

**Notes:**
- IV: intravenous.
- *If the source of the infection is determined please follow infection-specific guidance.
- Ceftriaxone or cefotaxime are alternative options. The choice can be made based on local availability.
- Gentamicin and amikacin are alternative options. The choice can be made based on local availability. In addition, amikacin is still effective against isolates producing extended-spectrum β-lactamases (ESBL) and is considered an appropriate carbapenem-sparing option in settings where ESBL-producing isolates are very prevalent.
- All dosages are for normal renal function.

ACCESS antibiotics are indicated in green, WATCH antibiotics in yellow and RESERVE antibiotics in red.

### Empiric antibiotics for SARI and severe bacterial pneumonia in children

**Children**

#### Severe pneumonia
(pneumonia with any danger sign, which requires referral to facility/hospital, admission and injectable therapy)

<table>
<thead>
<tr>
<th></th>
<th><strong>Ampicillin</strong> (IV/IM):</th>
</tr>
</thead>
<tbody>
<tr>
<td>50 mg/kg dose given every 12 hours (1st week of life)</td>
<td>50 mg/kg dose given every 8 hours (&gt; 1st week of life)</td>
</tr>
<tr>
<td><strong>Gentamicin</strong> (IV/IM):</td>
<td></td>
</tr>
<tr>
<td>Neonates: 5 mg/kg dose given once a day</td>
<td>Children: 7.5 mg/kg dose given once a day</td>
</tr>
</tbody>
</table>

**Ampicillin can be replaced by**

<table>
<thead>
<tr>
<th></th>
<th><strong>Amoxicillin</strong> (IV/IM):</th>
</tr>
</thead>
<tbody>
<tr>
<td>50 mg/kg dose given every 12 hours</td>
<td></td>
</tr>
</tbody>
</table>

**or**

<table>
<thead>
<tr>
<th></th>
<th><strong>Benzylpenicillin</strong> (IV/IM):</th>
</tr>
</thead>
<tbody>
<tr>
<td>30 mg (50,000 IU)/kg given every 6 hours</td>
<td></td>
</tr>
</tbody>
</table>

**If no clinical response to ampicillin and gentamicin change to second line:**

<table>
<thead>
<tr>
<th></th>
<th><strong>Cefotaxime</strong> (IV/IM):</th>
</tr>
</thead>
<tbody>
<tr>
<td>50 mg/kg dose given every 8 hours</td>
<td></td>
</tr>
</tbody>
</table>

**or**

<table>
<thead>
<tr>
<th></th>
<th><strong>Ceftriaxone</strong> (IV/IM):</th>
</tr>
</thead>
<tbody>
<tr>
<td>80 mg/kg dose given once a day</td>
<td></td>
</tr>
</tbody>
</table>

**Note:**

ACCESS antibiotics are indicated in green, WATCH antibiotics in yellow and RESERVE antibiotics in red.
### Empiric antibiotics for hospital-acquired bacterial pneumonia (non VAP)

<table>
<thead>
<tr>
<th>Adults</th>
<th>Children</th>
<th>Total treatment duration</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>FIRST CHOICE</strong></td>
<td><strong>Amoxicillin+clavulanic acid</strong> (IV/oral): 40-50 mg/kg/dose of amoxicillin component, given every 12 hours or 30 mg/kg dose given every 8 hours</td>
<td>7 days$^d$</td>
</tr>
<tr>
<td>Amoxicillin+clavulanic acid $^a$ (IV): 1 g + 20 mg given every 8 hours</td>
<td><strong>Amoxicillin+clavulanic acid</strong> (IV/oral): 40-50 mg/kg/dose of amoxicillin component, given every 12 hours or 30 mg/kg dose given every 8 hours</td>
<td>7 days$^d$</td>
</tr>
<tr>
<td>or Ceftriaxone (IV/IM): 2 g given once a day (IV), 1 g given once a day (IM)</td>
<td>or Ceftriaxone (IV/IM): 2 g given once a day (IV), 1 g given once a day (IM)</td>
<td>7 days$^d$</td>
</tr>
<tr>
<td>or Cefotaxime (IV/IM): 2 g given every 8 hours</td>
<td>or Cefotaxime (IV/IM): 2 g given every 8 hours</td>
<td>7 days$^d$</td>
</tr>
<tr>
<td>or Piperacillin+tazobactam $^b$ (IV): 4 g + 500 mg given every 6 hours</td>
<td>or Piperacillin+tazobactam (IV): 100 mg/kg/dose of piperacillin component, given every 8 hours</td>
<td>7 days$^d$</td>
</tr>
<tr>
<td>or Ceftriaxone (IV/IM): 80 mg/kg dose given once a day</td>
<td>or Cefotaxime (IV/IM): 50 mg/kg dose given every 8 hours</td>
<td>7 days$^d$</td>
</tr>
<tr>
<td>or Cefotaxime (IV/IM): 50 mg/kg dose given every 8 hours</td>
<td>or Cefotaxime (IV/IM): 50 mg/kg dose given every 8 hours</td>
<td>7 days$^d$</td>
</tr>
<tr>
<td>or Piperacillin+tazobactam (IV): 100 mg/kg/dose of piperacillin component, given every 8 hours</td>
<td>or Piperacillin+tazobactam (IV): 100 mg/kg/dose of piperacillin component, given every 8 hours</td>
<td>7 days$^d$</td>
</tr>
</tbody>
</table>

**Notes:**

$^a$ Amoxicillin+clavulanic acid can be used within 5 days of hospital admission and if no prior antibiotic exposure or risk for resistance.

$^b$ Piperacillin+tazobactam offers anti-pseudomonal coverage (which the other options do not). Risk of Pseudomonas aeruginosa is higher in patients with recent antibiotic exposure and especially in patients with known previous respiratory colonization and underlying lung diseases.

$^c$ Where possible use dispersible tablets. Oral syrup must be refrigerated as clavulanic acid is rapidly metabolized in high ambient temperatures.

$^d$ Reassess the diagnosis and consider longer treatment if the patient is not clinically stable at Day 7.

**ACCESS** antibiotics are indicated in green, **WATCH** antibiotics in yellow and **RESERVE** antibiotics in red.

**VAP:** ventilator-associated pneumonia.
## 7.2 COVID-19 and therapeutics

<table>
<thead>
<tr>
<th>Population</th>
<th>Disease severity</th>
<th>Recommendations</th>
</tr>
</thead>
<tbody>
<tr>
<td>Patients with confirmed covid-19</td>
<td>Non-severe</td>
<td>Absence of signs of severe or critical disease</td>
</tr>
<tr>
<td></td>
<td>Severe</td>
<td>Oxygen saturation &lt;90% on room air</td>
</tr>
<tr>
<td></td>
<td>Critical</td>
<td>Requires life sustaining treatment</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Signs of pneumonia</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Signs of severe respiratory distress</td>
</tr>
</tbody>
</table>

### Interventions

#### Strong recommendations in favour

- **Molnupiravir**
  - For those with highest risk of hospital admission

- **Sotrovimab**
  - For those with highest risk of hospital admission

- **Casirivimab and imdevimab**
  - For those with highest risk of hospital admission

- **Casirivimab and imdevimab**
  - Evidence of limited efficacy against Omicron BA.1 variant

#### Weak or conditional recommendations in favour

- **Ruxolitinib and tofacitinib**
  - Should be considered only if neither baricitinib nor IL-6 receptor blockers are available

#### Weak or conditional recommendations against

- **Remdesivir**
  - Next update will incorporate important new evidence

- **Ivermectin**
  - Should be considered only in the context of a clinical trial

#### Strong recommendations against

- **Convalescent plasma**
  - Should be considered only in the context of a clinical trial

- **Convalescent plasma**

- **Hydroxychloroquine**

- **Lopinavir-ritonavir**

---

Source: A living WHO guideline on drugs for COVID-19 (BMJ, 2020)
7.2.1 Corticosteroids for COVID-19

In patients with severe or critical COVID-19 WHO makes a strong recommendation to use systemic corticosteroids rather than no corticosteroids. See updated recommendations on therapeutics for patients with COVID-19: Therapeutics and COVID-19: living guideline.

The following recommendations on the use of corticosteroids are based on clinical trials in which the benefit was observed in patients who were mechanically ventilated or required supplemental oxygen at enrolment. No benefit was seen in patients who did not require supplemental oxygen at enrolment.

**Indication: severe or critical COVID-19**

<table>
<thead>
<tr>
<th>Route*</th>
<th>Systemic corticosteroids may be administered orally or intravenously.</th>
</tr>
</thead>
<tbody>
<tr>
<td>Type*</td>
<td>Daily regimen of dexamethasone 6 mg once daily is equivalent to:</td>
</tr>
<tr>
<td></td>
<td>• 150 mg of hydrocortisone daily (e.g. 50 mg every 8 hours) or</td>
</tr>
<tr>
<td></td>
<td>• 40 mg of prednisone daily or</td>
</tr>
<tr>
<td></td>
<td>• 32 mg of methylprednisolone daily (e.g. 8 mg every 6 hours or 16 mg every 12 hours).</td>
</tr>
<tr>
<td>Duration</td>
<td>Up to 5-14 days or until hospital discharge.</td>
</tr>
<tr>
<td>General monitoring</td>
<td>Monitor glucose levels (even if no diabetes diagnosed previously).</td>
</tr>
<tr>
<td>Special populations to monitor closely for complications</td>
<td>Patients receiving other immunosuppressants/immunomodulators.</td>
</tr>
<tr>
<td></td>
<td>Patients with these conditions:</td>
</tr>
<tr>
<td></td>
<td>- diabetes (specially with diabetic ketoacidosis)</td>
</tr>
<tr>
<td></td>
<td>- severe immunodeficiency disorders</td>
</tr>
<tr>
<td></td>
<td>- haematological and other malignancies</td>
</tr>
<tr>
<td></td>
<td>- low number of white blood cells</td>
</tr>
<tr>
<td></td>
<td>- organ transplantations</td>
</tr>
<tr>
<td></td>
<td>- iron overload states</td>
</tr>
<tr>
<td></td>
<td>- severe burns</td>
</tr>
<tr>
<td></td>
<td>- injection drug use</td>
</tr>
<tr>
<td></td>
<td>- malnutrition</td>
</tr>
<tr>
<td></td>
<td>- open wound following trauma.</td>
</tr>
<tr>
<td>Examples of possible complications</td>
<td>• Hyperglycaemia or decompensated diabetes</td>
</tr>
<tr>
<td></td>
<td>• Immunosuppression</td>
</tr>
<tr>
<td></td>
<td>• Superinfections: bacterial, fungal, viral, parasites</td>
</tr>
<tr>
<td></td>
<td>• Poor wound healing.</td>
</tr>
</tbody>
</table>

**Notes:**
* If intestinal dysfunction is suspected, clinicians must consider administering systemic corticosteroids intravenously rather than orally.
* Different corticosteroid preparations have different potencies and doses vary for different steroids. Care should be taken not to confuse doses for different products.

Interleukin-6 receptor blockers for COVID-19: tocilizumab, sarilumab

In patients with severe or critical COVID-19 WHO makes a strong recommendation for treatment with IL-6 receptor blockers (tocilizumab or sarilumab) in combination with other corticosteroids.


IL-6 receptor blockers should be prescribed and supervised by doctors who are experienced in the use of biologics and who have fully familiarized themselves with the efficacy and safety profile of these products.

**Tocilizumab**

<table>
<thead>
<tr>
<th>Indication (patient criteria)</th>
<th>Patients with severe or critical COVID-19 requiring supplemental oxygen and/or mechanical ventilation AND corticosteroid treatment (see Therapeutics and COVID-19: living guideline). Treatment should be started as early as possible in the patient’s critical illness.</th>
</tr>
</thead>
<tbody>
<tr>
<td>Dose and route</td>
<td>Tocilizumab 8 mg/kg (maximum dose of 800 mg) intravenous infusion administered over 60 minutes as a single dose.</td>
</tr>
<tr>
<td></td>
<td>• Avoid IV push or bolus.</td>
</tr>
<tr>
<td></td>
<td>• If clinical response is determined to be inadequate after 12-48 hours, a second dose may be considered.</td>
</tr>
<tr>
<td></td>
<td>• Renal dose adjustment is not warranted.</td>
</tr>
<tr>
<td></td>
<td>• Do not exceed maximum dose of 800 mg.</td>
</tr>
<tr>
<td>Dose of tocilizumab (mg) to be given:</td>
<td></td>
</tr>
<tr>
<td>Patient body weight</td>
<td>Band dose</td>
</tr>
<tr>
<td>&lt; 30 kg</td>
<td>12 mg/kg</td>
</tr>
<tr>
<td>31-45 kg</td>
<td>8 mg/kg</td>
</tr>
<tr>
<td>46-55 kg</td>
<td>400 mg</td>
</tr>
<tr>
<td>56-65 kg</td>
<td>480 mg</td>
</tr>
<tr>
<td>66-80 kg</td>
<td>600 mg</td>
</tr>
<tr>
<td>81-90 kg</td>
<td>680 mg</td>
</tr>
<tr>
<td>&gt; 91 kg</td>
<td>800 mg</td>
</tr>
</tbody>
</table>

**Available formulations**

Volume tocilizumab 20 mg/mL:
80 mg in 4-mL vial
200 mg in 10-mL vial
400 mg in 20-mL vial.
### Tocilizumab continued

**Preparation of infusion and administration**

1. Ensure area of preparation is well ventilated, clear and clean.
2. Ensure appropriate PPE is worn: gloves, mask and goggles.
3. Collect the appropriate numbers of tocilizumab vials from the fridge (between 2-8 °C) and a 100-mL bag of 0.9% sodium chloride for IV infusion.
4. Tocilizumab solution should be clear to opalescent, colourless to pale yellow and free of visible particles for administration.
5. Withdraw the volume of sodium chloride 0.9% from the 100-mL IV infusion bag equivalent to that which you will be injecting of tocilizumab, calculated as required for the patient dose (see table above).
   * For patients < 30 kg, dilute to 50 mL in 0.9% or 0.45% sodium chloride Injection for intravenous infusion.
6. Discard the withdrawn sodium chloride 0.9% inside the syringe and needle into a sharp bin.
7. Using an appropriate volume IV syringe, draw up the calculated mL of tocilizumab 20 mg/mL solution required from the vial(s) (see table above).
8. Slowly add the tocilizumab solution to the sodium chloride 0.9% IV infusion to make a final volume of 100 mL.
9. Dispose the tocilizumab needle and syringe in a sharp bin.
10. Mix the tocilizumab solution by gently inverting several times. Do not shake the solution.
11. Complete and apply an IV infusion label to the tocilizumab infusion.
12. Obtain second check for the IV tocilizumab infusion from a colleague and sign for preparation and administration.
13. Record baseline vital signs before, and saturation of oxygen (SpO₂) during and after administration: heart rate, blood pressure, temperature, respiratory rate.
14. Connect the tocilizumab IV infusion to the patient IV line and administer over 1 hour via volumetric infusion pump. Do not use the same IV line for other medications while the tocilizumab is being administered.
15. Complete patient observations again and monitor the patient for signs of hypersensitivity to tocilizumab during and after the administration.
16. Acute infusion reactions can occur during the administration of tocilizumab or within 24 hours of infusion.
17. For mild reactions (such as flushing or chills), the infusion rate can be slowed down and the patient continually monitored.
18. For severe reactions (such as hives, difficulty breathing, chest pain, high or low blood pressure, swelling of hand and face, fever, chills or anaphylaxis) or when mild reactions that do not disappear despite slowing infusion, stop the infusion and inform the doctor immediately for additional treatment.
19. Once tocilizumab infusion is complete, take down the infusion and flush the giving set with 20 mL of sodium chloride 0.9% over 15 minutes to ensure all the tocilizumab has been given.
20. Dispose of the infusion and giving set in a sharp bin.
Tocilizumab continued

<table>
<thead>
<tr>
<th>Monitoring for potential serious adverse events</th>
<th>Please refer to USPI, SmPC or local labelling for important safety issues and warnings.</th>
</tr>
</thead>
<tbody>
<tr>
<td>• Monitor the patient for signs of hypersensitivity to tocilizumab during and after administration: 15 minutes after starting the infusion, then every 30 minutes during the infusion and for 1 hour after the end of the infusion (15 min, 45 min, 1 hr, 1 hr 15 min, 1 hr 45 min).</td>
<td></td>
</tr>
<tr>
<td>• Laboratory monitoring is recommended due to potential consequences of treatment-related changes, at baseline, 72 hours, and after infusion of IL-6 RB:</td>
<td></td>
</tr>
<tr>
<td>– <strong>Neutrophil count</strong> (it is not recommended to initiate treatment in patients with neutropenia).</td>
<td></td>
</tr>
<tr>
<td>– <strong>Platelets count</strong> (it is not recommended to initiate treatment in patients with &lt; 50 000/mL).</td>
<td></td>
</tr>
<tr>
<td>– <strong>Transaminases</strong> (it is not recommended to initiate treatment in patients with elevated transaminases ALT or AST above 1.5× ULN. Discontinue infusion or do not give second dose in patients who develop persistent elevated ALT or AST above 3× ULN or who develop ALT or AST above 5× ULN).</td>
<td></td>
</tr>
<tr>
<td>– <strong>Lipid profile</strong> (LDL, HDL cholesterol, triglycerides) (possibility of elevated lipid profile after treatment).</td>
<td></td>
</tr>
<tr>
<td>• At baseline:</td>
<td></td>
</tr>
<tr>
<td>– <strong>Screening for HIV, Hep B (HBsAg, HBCAb) and C (HC Ab)</strong> (Discuss the results with microbiology or infectious diseases physicians if unsure how to interpret.) Individual clinical assessment of these patients is needed.</td>
<td></td>
</tr>
<tr>
<td>* <strong>Delays in the result of these tests should not restrain clinicians from starting treatment when indicated.</strong></td>
<td></td>
</tr>
<tr>
<td>• Before, during and after IL-6 receptor blockers infusion: patients should be regularly clinically assessed for bacterial infection. Monitor the appearance of sepsis produced by other pathogens different from COVID-19 (caution is recommended when considering the use in patients with a history of recurring or chronic infections or underlying conditions which may predispose patients to infections).</td>
<td></td>
</tr>
<tr>
<td>• In areas of high prevalence of TB or immunosuppressed patients, monitoring and assessment for presumptive TB is important before and after administration.</td>
<td></td>
</tr>
<tr>
<td>• Observe for hypersensitivity reactions during and after administration as mentioned in preparation and infusion section.</td>
<td></td>
</tr>
</tbody>
</table>

| Safety profile/adverse effects | • The most common side-effects (occurring in up to 1 patient in 10) with tocilizumab are upper respiratory tract infections (nose and throat infection), nasopharyngitis (inflammation of the nose and throat), headache, hypertension (high blood pressure) and abnormal liver function tests. The most serious side-effects are serious infections, complications of diverticulitis and hypersensitivity (allergic) reactions. |

<table>
<thead>
<tr>
<th>High-risk groups that require special precautions for use</th>
<th>• <strong>Age &gt; 70 years.</strong></th>
</tr>
</thead>
<tbody>
<tr>
<td>• Patients with recurring chronic infections or underlying conditions that predispose to infections, e.g. interstitial lung disease, diabetes, diverticulitis or patients taking steroids. Additionally there appears to be an increase risk of serious infections with increased body weight.</td>
<td></td>
</tr>
<tr>
<td>• Patients receiving IL-6 RB on long-term regimens for conditions other than COVID-19 (risk of fatal infections such as active TB, bacterial, viral and other opportunistic infections).</td>
<td></td>
</tr>
<tr>
<td>• Women of childbearing potential must use effective contraception during and up to 3 months after treatment.</td>
<td></td>
</tr>
<tr>
<td>• Pregnant women - only use in pregnancy only if the potential benefit justifies the potential risk to the fetus.</td>
<td></td>
</tr>
</tbody>
</table>
Tocilizumab continued

**Examples of possible complications**
- Acute severe infections: TB, bacterial, invasive fungal, viral and other opportunistic superinfections.
- Immunosuppression.
- Elevated liver function tests and lipid profile.
- Gastrointestinal perforation.
- Hypersensitive reactions, anaphylaxis.
- Important risks include:
  - Serious infection
  - Complications of diverticulitis
  - Serious hypersensitivity reactions
  - Neutropenia
  - Hepatotoxicity
  - Thrombocytopenia and the potential risk of bleeding
  - Elevated lipid levels and the potential risk of cardiovascular and cerebrovascular events
  - Malignancies
  - Demyelinating disorders
  - Immunogenicity.

**Not recommended**
- On immunosuppressive therapy (excluding steroids)
- HIV with CD4 < 200.
- ANC < 200.
- Suspected active severe bacterial, viral, fungal or TB infection (other than COVID-19).
- Patients who may be at increased risk of gastrointestinal (GI) perforations: history of diverticulitis or bowel perforation.
- Patients with elevated ALT or AST above 10 times the upper limit of the reference range.
- Paediatric population < 2 years old.

**Contraindications**
- Tocilizumab is contraindicated in those with prior hypersensitivity to the medication or any known ingredient in it.

**Notes:**
Avoid concurrent use of live vaccines during treatment as clinical safety has not been established.
Further guidance for use can be obtained from the product information and training packages provided by the license holder.
Report all adverse events to the national pharmacovigilance centre or the manufacturer.
ALT - alanine aminotransferase; AST - aspartate aminotransferase; SmPC - Summary of Product Characteristics (European Union); ULN - upper limit of normal; USPI - United States Prescribing Information.
Sarilumab

**Indication (patient criteria)**

Patients with severe or critical COVID-19 requiring supplemental oxygen and/or mechanical ventilation AND corticosteroid treatment (see Therapeutics and COVID-19: living guideline ).

Treatment should be started as early as possible in the patient’s critical illness.

**Dose and route**

Sarilumab will be administered as an IV infusion at a dose of 400 mg over at least 60 minutes using a dedicated IV line.

- Avoid IV push or IV bolus. (Subcutaneous administration has not been studied for COVID-19.)
- If a clinical response is determined to be inadequate after 12-48 hours, a second dose may be considered.
- Renal dose adjustment is not warranted.

**Available formulations**

Single dose 200 mg pre-filled syringe 200 mg/1.14 mL.

- Do not use sarilumab with autoinjector directly for SC injection, follow instructions to dilute them for IV infusion (see below).

**Preparation of infusion and administration**

1. Ensure area of preparation is well-ventilated, clear and clean.
2. Ensure appropriate PPE is worn: gloves, mask and goggles.
3. Collect the contents of 2 prefilled syringes (200 mg/1.14 mL) of sarilumab, which should be diluted in 100 mL of 0.9% sodium chloride to prepare 400 mg of sarilumab IV bag.

Note: The needle attached to the sarilumab syringe is approx. 12.5 mm long (half of normal). Take care to ensure it fully penetrates the port and reaches the fluid.

4. After mixing it is best practice to ensure the drug is not trapped in the injection port. A flush of 25 mL of normal saline can be given.
5. Invert bag 10 times to mix, do not shake.
6. Ensure the product solution is clear and free from any precipitation. Label as local policy.
7. The infusion should be started within 4 hours of preparation and can be given via central line or peripheral line.
8. Do not infuse concomitantly in the same IV line with other medications.
9. The inline filter and the IV infusion pump must be able to deliver as little as 0.17 mL/min (10 mL/hr).
10. Infusion speed must be set at 10 mL/h for 15 minutes, then increased to 130 mL/hr for the next 45 minutes. Infuse over 60 minutes.
11. After completion of the infusion, 25 mL of 0.9% saline should be used to flush the drug through the giving set.
12. IV infusion sets that have been evaluated in sarilumab compatibility include:

<table>
<thead>
<tr>
<th>Infusion set</th>
<th>Examples</th>
</tr>
</thead>
<tbody>
<tr>
<td>Standard infusion set with PVC tubing containing DEHP (PVS+DEHP) with 0.2 um PES inline filter</td>
<td>Baxter, product code 2C6571 or similar</td>
</tr>
<tr>
<td>DEHP-free infusion set made from polyethylene lined PVC tubing (PE-line PVC) with 0.2 um PES inline filter</td>
<td>Alaris, product no. 11532269 or similar</td>
</tr>
<tr>
<td>DEHP-Free PVC infusion set made from PVC tubing containing TOTAM (PVC-TOTM) with 0.2 um PES inline filter</td>
<td>Hospira, product no. 14255-28 or similar</td>
</tr>
<tr>
<td>Infusion set made from polyurethane (PU) with 0.2 um PES inline filter</td>
<td>B Braun, product 8700095P or similar</td>
</tr>
</tbody>
</table>
Sarilumab continued

| Preparation of infusion and administration | 13. Monitor for infusion-related reactions: chills, nausea, headache, wheezing, itching, flushing, pyrexia, dizziness. If infusion-related reactions are mild, stop infusion and treat symptoms. Reduce infusion rate by at least 50% when re-starting infusion. |
| Infusion instructions | 14. For severe infusion-related reactions, stop the infusion and inform the doctor immediately. |
| 15. Once the sarilumab infusion is complete, take down the infusion and flush the giving set with 20 mL of sodium chloride 0.9% at same rate as infusion. |
| 16. Document the administration. |
| 17. Dispose the infusion and giving set in a sharp bin. |

| Monitoring for potential serious adverse events | Please refer to USPI, SmPC or local labelling for important safety issues and warnings. |
| 1. Monitor the patient for signs of hypersensitivity to sarilumab during and after administration: 15 minutes after starting the infusion, then every 30 minutes during the infusion and for 1 hour after the end of the infusion (15 min, 45 min, 1 hr, 1 hr 15 min, 1 hr 45 min). |
| 2. Laboratory monitoring is recommended due to potential consequences of treatment-related changes, at baseline, 72 hours, and after infusion of IL-6 RB: |
| Neutrophil count (it is not recommended to initiate treatment in patients with neutropenia). |
| Platelets count (it is not recommended to initiate treatment in patients with < 50,000/mL). |
| Transaminases (it is not recommended to initiate treatment in patients with elevated transaminases ALT or AST above 1.5x ULN. Discontinue infusion or do not give second dose in patients who develop persistent elevated ALT or AST above 3x ULN or who develop ALT or AST above 5x ULN. |
| SmPC recommends dose to be modified if ALT > 1-3 upper limit. |
| Lipid profile (LDL, HDL cholesterol, triglycerides) (possibility of elevated lipid profile after treatment). |
| At baseline: |
| Screening for HIV, Hep B (HBsAg, HBCAb) and C (HC Ab)* (Discuss the results with microbiology or infectious diseases physicians if unsure how to interpret.) |
| * Delays in the result of these tests should not restrain clinicians from starting treatment when indicated. |
| Before, during and after IL-6 receptor blockers infusion: patients should be regularly carefully assessed for bacterial infection. Monitor the appearance of sepsis produced by other pathogens different from COVID-19 (caution is recommended when considering the use in patients with a history of recurring or chronic infections or underlying conditions which may predispose patients to infections). |
| In areas of high prevalence of TB or immunosuppressed patients, monitoring and assessment for presumptive TB is important before and after administration. |

| Safety profile/adverse effects | Transient and/or reversible elevations in liver enzyme levels that appear to be dose dependent and rare occurrences of neutropenia and thrombocytopenia. Additional adverse effects, such as serious infections (e.g. TB, bacterial or fungal infections), and bowel perforation, have been reported, but only with long-term use of sarilumab. |

| High-risk groups that require special precautions for use | Age > 70 years. |
| 1. Patients with recurring chronic infections or underlying conditions that predispose to infections. |
| 2. Patients receiving IL-6 RB on long-term regimens for conditions other than COVID-19 (risk of fatal infections such as active TB, bacterial, viral and other opportunistic infections). |
| 3. Women of childbearing potential must use effective contraception during and up to 3 months after treatment. |
| 4. Pregnant women - only use in pregnancy only if the potential benefit justifies the potential risk to the fetus. |
Sarilumab continued

### Examples of possible complications
- Acute severe infections: TB, bacterial, invasive fungal, viral and other opportunistic superinfections.
- Immunosuppression
- Elevated liver function tests and lipid profile.
- Gastrointestinal perforation.
- Hypersensitive reactions, anaphylaxis.

### Not recommended
- On immunosuppressive therapy (excluding steroids).
- HIV with CD4 < 200.
- ANC < 200.
- Suspected active severe bacterial, viral, fungal or TB infection (other than COVID-19).
- Patients who may be at increased risk of gastrointestinal (GI) perforations: history of diverticulitis or bowel perforation.
- Patients with elevated ALT or AST above 10 times the upper limit of the reference range.
- Paediatric population < 2 years old.

### Contraindications
- Sarilumab is contraindicated in those with prior hypersensitivity to the medication or any known ingredient in it.

**Notes:**
Avoid concurrent use of live vaccines during treatment as clinical safety has not been established. Further guidance for use can be obtained from the product information and training packages provided by the license holder. Report all adverse events to the national pharmacovigilance centre or the manufacturer.

ALT - alanine aminotransferase; AST - aspartate aminotransferase; SmPC - Summary of Product Characteristics (European Union); ULN - upper limit of normal; USPI - United States Prescribing Information.
723 Monoclonal antibodies for COVID-19: casirivimab and imdevimab

For patients with non-severe COVID-19 (who do not meet the criteria for severe or critical infection) at the highest risk of developing severe disease there is a conditional recommendation to use a combination of neutralizing monoclonal antibodies (casirivimab and imdevimab).

For patients with severe and critical COVID-19 with seronegative status there is a conditional recommendation to use a combination of neutralizing monoclonal antibodies (casirivimab and imdevimab). See updated recommendations on therapeutics for patients with COVID-19: Therapeutics and COVID-19: living guideline.

See the following posters/clinical tools regarding administration of casirivimab and imdevimab:

- Preparation of intravenous casirivimab and imdevimab for COVID-19
- Preparation and administration of subcutaneous casirivimab and imdevimab for COVID-19
- Safety and monitoring in patients receiving casirivimab and imdevimab for COVID-19

WHO has noted that there is a predicted lack of efficacy for casirivimab and imdevimab with the Omicron variant; updated recommendations may be warranted when sufficient evidence addressing this is available.
Casirivimab and imdevimab

Indication (patient criteria)
1. Patients with confirmed non-severe COVID-19 at highest risk for infection:
   - Those with risk beyond 10% for being hospitalized with COVID-19. Those at highest risk for progression of infection:
     - Older age.
     - Risk factors: hypertension, diabetes, cardiac disease, chronic lung disease, cerebrovascular disease, dementia, mental disorders, chronic kidney disease, immunosuppression, obesity, cancer.
     - Not vaccinated.
2. Severe or critical COVID-19:
   - Seronegative for SARS-CoV-2.
   - Serologic testing should be conducted with tests that detect the presence of the SARS-CoV-2 spike protein antibodies and have performance characteristics similar to the reference standard test used to characterize seronegative patients in the RECOVERY trial (i.e. Oxford fluorescent-based ELISA assay for serum IgG against the SARS-CoV-2 spike protein), with an arbitrary cut-off determined by a panel of positive controls.
   - Some lateral flow assay may be suitable and can usually be performed in several minutes.
3. No medical contraindication:
   - No allergy to casirivimab and imdevimab or components: L-histidine, L-histidine monohydrochloride monohydrate, polysorbate 80, sucrose.
   (See Therapeutics and COVID-19: living guideline (1)).

Dose and route
Casirivimab and imdevimab can be administered as 1200-8000 mg (600-4000 mg each antibody) demonstrating efficacy at all doses. (Adults and paediatric patients ≥ 12 years of age and weighing ≥ 40 kg.)

<table>
<thead>
<tr>
<th>Drug</th>
<th>Dose (non-severe disease)</th>
<th>Dose (severe and critical disease)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Casirivimab</td>
<td>600 mg IV/SC or 1200 mg IV</td>
<td>1200-4000 mg IV</td>
</tr>
<tr>
<td>Imdevimab</td>
<td>600 mg IV/SC or 1200 mg IV</td>
<td>1200-4000 mg IV</td>
</tr>
<tr>
<td>Total dose</td>
<td>1200-2400 mg IV or 1200 mg SC*</td>
<td>2400-8000 mg IV</td>
</tr>
</tbody>
</table>

Table 1. Dose of casirivimab and imdevimab
Casirivimab and imdevimab continued

Dose and route
Notes:
* If administered subcutaneously (SC), maximum total dose is 1200 mg.
IV - intravenous infusion is preferred.
SC - subcutaneous is an alternative route of administration. It can be given when intravenous infusion is not feasible and would lead to a delay in treatment.

Preferred: Casirivimab and imdevimab intravenous single infusion using a dedicated IV line with a sterile low protein binding inline or add-on 0.2 micron filter via pump (preferred) or gravity + 1 hour post-infusion close monitoring (covered below in more detail).

Alternative: Casirivimab and imdevimab by consecutive subcutaneous injections (syringes) + 1 hour post-injection close monitoring (covered below in more detail).

Renal or hepatic dose adjustment is not currently warranted for either drug.

Storage conditions
- Store in a refrigerator at 2-8 °C in the original carton to protect from light.
- Do not freeze.
- Do not shake.

Available formulations
The vials of casirivimab and imdevimab come in two different sizes (20 mL or 6 mL).

a) Each 20-mL vial contains 11.1 mL of product
   Concentration: 1332 mg per 11.1 mL (120 mg/mL).

b) Each 6-mL vial contains 2.5 mL of product
   Concentration: 300 mg per 2.5 mL (120 mg/mL).

Preparation of intravenous perfusion

1. Wear PPE: gloves, gown, protective eyewear and respiratory mask.
2. Remove casirivimab and imdevimab vials from refrigerated storage.
3. Prepare in a well-ventilated area in clean room.
4. Allow to equilibrate to room temperature for approximately 20 minutes before preparation.
   Do not shake the vials or expose to direct heat.
5. Inspect the casirivimab and imdevimab vials to ensure there is no discoloration or particulate matter prior to administration. If either is observed, the vials should be discarded.
   The solution in each vial should be clear to slightly opalescent (colourless to pale yellow).
6. Obtain a prefilled intravenous infusion bag containing 50 mL, 100 mL, 150 mL or 250 mL of 0.9% of sodium chloride injection or 5% dextrose injection.
7. To allow for space in the infusion bag for the addition of the casirivimab and imdevimab, use a syringe and aseptic non-touch technique to withdraw the total volume dose equivalent from the infusion bag and discard.
   For example, if preparing a total dose of casirivimab and imdevimab of 1200 mg infusion, remove 10 mL of fluid from the infusion bag prior to injecting the monoclonal antibody (see Table 2 below).
8. Using a separate syringe for each vial, withdraw the appropriate amount of casirivimab and imdevimab from each respective vial and inject into a prefilled intravenous infusion bag containing either 0.9% of sodium chloride injection or 5% dextrose injection.
   If using one vial to prepare more than one bag, then prepare all bags at the same time.
9. Gently invert infusion bag by hand ten times to mix.
   Do not shake.
10. The product is preservative free, so the diluted infusion solution should be administered immediately.
    If not possible to immediately administer, the diluted casirivimab and imdevimab infusion solution can be stored in the refrigerator at 2-8 °C for up to 36 hours or at room temperature up to 25 °C for no more than 4 hours. If refrigerated, remove approximately 30 minutes prior to patient infusion to allow for equilibration to room temperature.
### Preparation of intravenous perfusion

**Table 2. Recommended dilution instructions for 600 mg of casirivimab and 600 mg of imdevimab for intravenous infusion (total dose: 1200 mg)**

<table>
<thead>
<tr>
<th>Size of prefilled 0.9% sodium chloride or 5% dextrose infusion bag</th>
<th>Preparing casirivimab and imdevimab using individual vials</th>
</tr>
</thead>
<tbody>
<tr>
<td>50 mL</td>
<td>Add:</td>
</tr>
<tr>
<td>100 mL</td>
<td>• 5 mL of casirivimab (use two 6-mL vials of casirivimab or 5-mL of one 20-mL vial) and • 5 mL of imdevimab (use two 6-mL vials of imdevimab or 5-mL of one 20-mL vial)</td>
</tr>
<tr>
<td>150 mL</td>
<td>Inject 5 mL of casirivimab + 5 mL of imdevimab into a prefilled 0.9% sodium chloride or 5% dextrose infusion bag and administer as instructed.</td>
</tr>
<tr>
<td>250 mL</td>
<td></td>
</tr>
</tbody>
</table>

**Table 3. Recommended dilution instructions for 1200 mg of casirivimab and 1200 mg of imdevimab for intravenous infusion (total dose: 2400 mg)**

<table>
<thead>
<tr>
<th>Size of prefilled 0.9% sodium chloride or 5% dextrose infusion bag</th>
<th>Preparing casirivimab and imdevimab using individual vials</th>
</tr>
</thead>
<tbody>
<tr>
<td>50 mL</td>
<td>Add:</td>
</tr>
<tr>
<td>100 mL</td>
<td>• 10 mL of casirivimab (use four 6-mL vials of casirivimab or 10-mL from a 20-mL vial) and • 10 mL of imdevimab (use four 6-mL vials of imdevimab or 10-mL from a 20-mL vial)</td>
</tr>
<tr>
<td>150 mL</td>
<td>Inject 10 mL of casirivimab + 10 mL of imdevimab into a prefilled infusion bag of 0.9% sodium chloride or 5% dextrose and administer as instructed.</td>
</tr>
<tr>
<td>250 mL</td>
<td></td>
</tr>
</tbody>
</table>

**Table 4. Recommended dilution instructions for 4000 mg of casirivimab and 4000 mg of imdevimab for intravenous infusion (total dose: 8000 mg)**

<table>
<thead>
<tr>
<th>Size of prefilled 0.9% sodium chloride or 5% dextrose infusion bag</th>
<th>Preparing casirivimab and imdevimab using individual vials</th>
</tr>
</thead>
<tbody>
<tr>
<td>150 mL</td>
<td>Add:</td>
</tr>
<tr>
<td>250 mL</td>
<td>• 33.3 mL of casirivimab (use three 20-mL vials of casirivimab) and • 33.3 mL of imdevimab (use three 20-mL vials of imdevimab)</td>
</tr>
<tr>
<td>500 mL</td>
<td>Inject 33.3 mL of casirivimab + 33.3 mL of imdevimab into a prefilled infusion bag of 0.9% sodium chloride or 5% dextrose and administer as instructed.</td>
</tr>
</tbody>
</table>
Casirivimab and imdevimab continued

Preparation of subcutaneous injections

1. Wear PPE: gloves, gown, protective eyewear and respiratory mask.
2. Remove casirivimab and imdevimab vials from refrigerated storage.
3. Prepare in a well-ventilated area in clean room.
4. Allow to equilibrate to room temperature for approximately 20 minutes before preparation. **Do not shake the vials or expose to direct heat.**
5. Inspect the casirivimab and imdevimab vials to ensure there is no discolouration or particulate matter prior to administration. If either is observed, the vials should be discarded. **The solution in each vial should be clear to slightly opalescent (colourless to pale yellow).**
6. Casirivimab and imdevimab should be administered consecutively by subcutaneous injection using the appropriate number of syringes. **Obtain 3-mL or 5-mL polypropylene luer lock syringes with luer connection and 21-gauge 1½ inch transfer needles.**
7. Withdraw appropriate dose of casirivimab and imdevimab into each syringe. Replace the transfer needle with a 25-27 gauge needle for subcutaneous injection.
8. The product is preservative free, so should be administered immediately. **If not possible to immediately administer, store the prepared syringes of casirivimab and imdevimab at room temperature up to 25 °C for no more than 4 hours.**

Administration of intravenous infusion

Casirivimab and imdevimab infusion should be administered by a qualified health care professional using aseptic non-touch technique.

1. Materials needed:
   a) Polyvinyl chloride (PVC), polyethylene (PE)-lined PVC or polyurethane (PU) infusion set;
   b) In-line or add-on low protein binding 0.2 micron polyethersulfone (PES) filter.
2. Attach the infusion set to the intravenous bag.
3. Prime the infusion set.
4. Administer the entire infusion solution in the bag via pump (preferred) or gravity through an intravenous line containing a sterile inline or add-on low protein binding 0.2 micron filter.
5. Do not administer the infusion solution with another medication.
6. After infusion is complete, flush tubing with either 0.9% sodium chloride injection or 5% dextrose injection.
7. Discard the completed infusion bag.
8. Clinically monitor patient during administration of medication and observe patient for 1 hour after infusion is complete.

| Table 5. Administration rate for 600 mg casirivimab and 600 mg imdevimab for intravenous infusion (total 1200 mg) |
|---|---|---|
| Size of prefilled 0.9% sodium chloride or 5% dextrose infusion bag | Maximum infusion rate | Minimum infusion time |
| 50 mL | 150 | 20 minutes |
| 100 mL | 300 | 20 minutes |
| 150 mL | 500 | 20 minutes |
| 250 mL | 500 | 30 minutes |

**Note:** Infusion should not be administered > 4 hours.
Casirivimab and imdevimab continued

**Administration of intravenous infusion**

<table>
<thead>
<tr>
<th>Size of prefilled 0.9% sodium chloride or 5% dextrose infusion bag</th>
<th>Maximum infusion rate</th>
<th>Minimum infusion time</th>
</tr>
</thead>
<tbody>
<tr>
<td>50 mL</td>
<td>150</td>
<td>20 minutes</td>
</tr>
<tr>
<td>100 mL</td>
<td>300</td>
<td>20 minutes</td>
</tr>
<tr>
<td>150 mL</td>
<td>450</td>
<td>20 minutes</td>
</tr>
<tr>
<td>250 mL</td>
<td>500</td>
<td>30 minutes</td>
</tr>
</tbody>
</table>

Note: Infusion should not be administered > 4 hours.

**Administration of subcutaneous injection**

1. Administer the subcutaneous injection of casirivimab and imdevimab consecutively, each at a different injection site into the upper thigh, back of the upper arm or abdomen to space apart each injection. **Avoid the waistline and 2 inches (5 cm) around the navel.**
   - Do NOT inject into skin that is tender, damaged, bruised or scarred.
   - Clinically monitor patient after the injections and observe for 1 hour.

2. **General monitoring**
   - Casirivimab and imdevimab can be administered as an intravenous infusion or subcutaneous injection.
   - It should only be administered in health care settings by a qualified health care provider who has access to immediate emergency medical services that can treat severe infusion reactions.
   - Patient should be clinically monitored during dose administration.
   - Patient should be observed for 1 hour after intravenous or subcutaneous dosing is complete, with vital signs (blood pressure, heart rate, respiratory rate, temperature and oxygen saturation) checked at 15 minutes, 30 minutes and 1 hour post infusion.

3. **Special populations to monitor closely for complications**
   - There are limited data regarding the use of casirivimab and imdevimab in pregnant patients with COVID-19.
   - Casirivimab and imdevimab should be used in pregnancy only if the potential benefit justifies the potential risk to the mother and fetus.
   - There are no available data on the presence of casirivimab and imdevimab in human milk or animal milk, the effects on breastfed infants, or the effects of the drug on milk production.

4. **Contraindications**
   - Casirivimab and imdevimab is contraindicated in those with prior hypersensitivity to the medication or any known ingredient in casirivimab and imdevimab.

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7. Therapies for SARI (COVID-19, influenza, bacterial pneumonia): antimicrobials and immunomodulators 143
Casirivimab and imdevimab continued

### Examples of possible complications

- Hypersensitivity reactions including anaphylaxis have been observed with casirivimab and imdevimab.
- Infusion-related reactions (IV).
- Injection site related reactions (SQ/SC).
- Hypersensitivity reactions occurring more than 24 hours after the infusion of casirivimab and imdevimab have been reported. If signs or symptoms of a clinically significant hypersensitivity reaction or anaphylaxis occur, immediately discontinue administration, and initiate appropriate medications and/or supportive therapy.
- Infusion-related reactions occurring during infusion and up to 24 hours post infusion have been observed and may be severe or life threatening.

**These reactions may include:** fever, difficulty breathing, reduced oxygenation, chills, fatigue, irregular heart beat, chest pain or discomfort, weakness, nausea, headache, angioedema, throat irritation, bronchospasm, hypertension, hypotension, throat irritation, rash, pruritis, muscle aches, pre-syncope, syncope, dizziness and diaphoresis.

- If an infusion-related reaction occurs consider slowing or stopping the infusion and administer appropriate medications and/or supportive care.

### Reporting of adverse events

Report all adverse events to WHO and the manufacturer (OfMonth):

www.roche.com/products/local_safety_reporting.htm

**Notes:**

As a precautionary measure, vaccination for SARS-CoV-2 should be deferred for ≥ 90 days in people who have received casirivimab and imdevimab. The antibody treatment may interfere with vaccine-induced immune responses.

If using the 20-mL multidose vial of casirivimab or imdevimab and there is remaining product, it can be returned to the refrigerator and stored for a maximum for 48 hours per the open vial policy.
7.3 Memory aid: invasive fungal infections in patients with COVID-19

Secondary invasive fungal infections such as mucormycosis, candidemia, aspergillosis or cryptococcosis with life-threatening outcomes are increasingly being observed in settings where corticosteroids are being used inappropriately (higher than recommended doses, for longer than recommended duration or in non-severe COVID-19 patients).

**Mucormycosis**

- **Risk factors**
  - diabetes mellitus (poorly controlled or complicated) with or without diabetic ketoacidosis (DKA)
  - prolonged corticosteroid use
  - haematological malignancies
  - haematopoietic stem cell transplant
  - solid organ transplantation
  - iron chelation therapy
  - iron overload states
  - burns
  - other immunosuppressive conditions or medications (e.g. TNF- inhibitors or other immunosuppressive medication, cancer - chemotherapy, chronic immunosuppression - HIV).

- **Prevention**
  - Use the recommended dose and duration of corticosteroids based on WHO recommendations and clinical practice guidelines for the indication. See WHO Therapeutics and COVID-19: living guideline [1].
  - Follow strict adherence to protocol of low-dose steroid and strict glycaemic control.
  - Follow strict adherence to quality of oxygen humidification protocol and biosafety:
    - Appropriate hygiene should be followed when oxygen is being administered.
    - Use of sterile water for humidifiers during the oxygen therapy is recommended.
    - Disposable items should not be reused and where this avoidable they should be properly sterilized.
    - Use of medical masks.
    - Educate patients on early signs and symptoms of some of the most important secondary fungal infections in COVID-19 recovering patients (e.g. facial pain or orbital pain-diplopia, nasal blockage or excessive discharge of blood or brown/black discharge, loosening of teeth, etc. all could be signs of rhino-orbital-cerebral mucormycosis [ROCM]).
    - Include ENT surgeon and ophthalmologist in the post-COVID-19 clinic to pick up the cases early.
    - Environmental control in hospital.
    - Restrict antibiotics only when bacterial infection is suspected/proven, as the use of antibiotics also increases likelihood of fungal infection.

- **Signs and symptoms**
  - facial swelling (pain and numbness can be present)
  - nasal or sinus congestion
  - black lesions on nasal bridge or upper inside of mouth
  - blackish and foul-smelling nasal discharge
  - loosening of a tooth or a TMJ-like symptoms
  - eye pain-diplopia
  - headache or cranial nerve palsy
  - fever
  - for respiratory or gastrointestinal mucormycosis, there is no specific symptom or sign; however, suspect the disease if there is chest pain, shortness of breath, cough haemoptysis, blackish discoloration in the skin with necrosis, abdominal pain, nausea and vomiting, gastrointestinal bleeding, mental status changes or coma.

- **Diagnosis**
  - early diagnosis is key!
  - imaging (CT or MRI)
  - endoscopic or bronchoscopic sample collection
  - histopathology or culture PLUS patient risk factors and clinical presentation.
<table>
<thead>
<tr>
<th>Treatment</th>
<th>Primary regimen:</th>
</tr>
</thead>
<tbody>
<tr>
<td>Liposomal (lipid) amphotericin B 5-10 mg/kg IV daily for 3-6 weeks:</td>
<td></td>
</tr>
<tr>
<td>• early administration is key - delay of 6 days was associated with two-fold increase in mortality in 12 weeks;</td>
<td></td>
</tr>
<tr>
<td>• monitor renal function regularly.</td>
<td></td>
</tr>
</tbody>
</table>

Surgical debridement:

• early aggressive surgical resection and debridement is key for local control of disease and can reduce morbidity and mortality.

Control underlying disease/risk (e.g. DM-DKA - discontinue corticosteroids, etc.).

Alternative regimens:

• amphotericin B deoxycholate 1-1.5 mg/kg IV daily (administered in at least 1000 mL of 5% dextrose over 2 hrs for 3-6 weeks):
  - during amphotericin B deoxycholate infusion, pre or peri-infusion normal saline and other symptomatic therapy may minimize the infusion related toxicity; monitor renal function regularly.
• posaconazole:
  - posaconazole IV 300 mg IV over 90 mins every 12h Day 1, then 300 mg IV daily;
  - delayed-release tablet 300 mg PO/12h on Day 1, then 300 mg PO daily;
  - suspension 200 mg QID, then 400 mg PO/12h after stabilization of disease;
• isavuconazonium sulfate (prodrug of isavuconazole) 372 mg PO/IV q8h x 6 doses, later 372 mg PO/IV daily.

Step-down or salvage therapy:

• if the patient is stable, posaconazole or isavuconazole tablet for 3-6 months after 3-6 weeks amphotericin B therapy;
• if there is disease progression while the patient is on amphotericin B therapy, liposomal amphotericin B dose may be increased to 10 g/kg/d or replace with posaconazole or isavuconazole;
• if the patient has renal compromise, start with isavuconazole or posaconazole.

NOTE 1: High doses of liposomal and deoxycholate amphotericin B need careful administration and monitoring and management - and reference detailed guidance. It should be given with saline fluid loading to reduce renal toxicity, and potassium and magnesium monitoring and supplements.

NOTE 2: As alternative regimen - posaconazole or isavuconazole injection is preferred.

NOTE 3: Step-down or salvage therapy:

- if the patient is stable, posaconazole or isavuconazole tablet for 3-6 months after 3-6 weeks amphotericin B therapy;
- if there is disease progression while the patient is on amphotericin B therapy, liposomal amphotericin B dose may be increased to 10 mg/kg/d or replace with posaconazole or isavuconazole;
- if the patient has renal compromise, start with isavuconazole or posaconazole.
# Candidemia

## Risk factors
- diabetes mellitus (poorly controlled or complicated)
- prolonged corticosteroid use
- haematological malignancies
- active solid malignancy
- recent chemotherapy
- HIV infection
- haemodialysis
- long ICU stays
- use of broad-spectrum antibiotics or antifungal agents
- central venous catheters
- parenteral nutrition
- immunosuppressive agents
- neutropenia
- severe pancreatitis
- Candida spp. colonization
- recent abdominal surgery
- extensive burns
- COVID-19 infection.

## Prevention
Control of risk factors (above) mainly in nosocomial transmission.

Use the recommended dose and duration of corticosteroids based on WHO recommendations and clinical practice guidelines. See WHO Therapeutics and COVID-19: living guideline (https://www.who.int/publications/m/item/who-therapeutics-and-covid-19-

Consider gradual withdrawal practices with corticosteroid regimens:
- if a patient has received more than 3 weeks' treatment;
- has recently received repeated courses of steroids;
- has taken a short course within 1 year of stopping long-term therapy;
- has other possible causes of adrenal suppression.

Follow strict adherence to protocol of low-dose steroid and strict glycaemic control.

## Clinical presentations
- Sepsis.
- Candidemia: one or more positive blood cultures for Candida spp.
- Less frequent: infective endocarditis; endophthalmitis.

## Diagnosis
- One or more positive blood cultures for Candida spp. (they could be negative in 50% cases).
- Beta-D-glucan (poor specificity).
- Candida mannan and anti-mannan detection.
- Clinical signs: suspected dissemination (e.g. fever, multiple colonized or infected sites, even if negative blood cultures).

## Treatment
*First-line: echinocandins:*
- caspofungin: 70 mg IV × 1, then 50 mg/day IV;
- micafungin: 100 mg/day IV;
- anidulafungin: 100 mg/day IV;

Duration: 14 days timed from clearance of bloodstream and resolution of symptoms.

*Alternative regimens: amphotericin B, voriconazole, isavuconazole.*
- amphotericin B (lipid formulation): 3–5 mg/kg/day IV (especially if azole and echinocandin resistance or intolerance is of concern);
- voriconazole, 6 mg/kg IV/PO × 2 doses on Day 1, then 3 mg/kg twice daily isavuconazole.

NOTE 1: The clinical presentation of candidemia is indistinguishable from most bacterial bloodstream infections - and the blood cultures can be falsely negative – this could lead to delays in appropriate therapy. Empiric treatment should be considered after consultation with infectious disease specialist if the patient is critically ill with unexplained sepsis and has some of the clinical risk factors, and/or colonized in none-sterile sites - or elevated markers (e.g. serum beta glucans levels, etc.).

NOTE 2: Fluconazole resistance is common depending on strain and varies with geographic location.
Aspergillosis

Risk factors
- prolonged neutropenia (acute leukaemia, myelodysplastic syndromes)
- prolonged high-dose corticosteroids
- allogenic stem cell transplant
- solid organ transplant
- haematologic malignancy
- immunosuppressive medications
- chronic granulomatous disease, e.g. TB, sarcoidosis
- chronic lung diseases, e.g. asthma, cystic fibrosis
- antimicrobial drug exposure
- inherited immunodeficiencies
- ICU care and mechanical ventilation
- COVID-19 infection, especially those treated in ICU.

Prevention
Use the recommended dose and duration of corticosteroids based on WHO recommendations and clinical practice guidelines. See WHO Therapeutics and COVID-19: living guideline ( ).

Control of risk factors.
Consider gradual withdrawal practices with corticosteroid regimens:
- if a patient has received more than 3 weeks’ treatment;
- has recently received repeated courses of steroids;
- has taken a short course within 1 year of stopping long-term therapy;
- has other possible causes of adrenal suppression.

Follow strict adherence to protocol of low-dose steroid and strict glycaemic control.
Follow strict adherence to quality of oxygen humidification protocol and biosafety:
- appropriate hygiene should be followed when oxygen is being administered;
- use of sterile water for humidifiers during the oxygen therapy is recommended;
- disposable items should not be reused and where this avoidable it should be properly sterilized.

Restrict antibiotics only when bacterial infection is suspected/proven, as the use of antibiotics also increases likelihood of fungal infection.

Signs and symptoms
- Chest pain, fever, cough, shortness of breath, haemoptysis.
- Invasive pulmonary infection - coronavirus disease-associated pulmonary aspergillosis (CAPA).
- Dissemination can occur: CNS, skin, GI tract, etc.
- Pulmonary deterioration - worsening respiratory status and need for increased respiratory support, hemoptysis, chest pain, sepsis (fever, tachycardia, tachypnoea, hypotension, change in mental status).

Diagnosis
- Imaging - multiple nodule, thick-walled cavities; in immunosuppressed patients halo sigh or air-crescent sign.
- Histopathology or culture PLUS patient risk factors and clinical presentation.
- Galactomannan antigen: serum, BAL fluid more sensitivity but reduced specificity for invasive disease.
- Beta-D-glucan assay in serum.

Treatment

Primary regimen:
- voriconazole 6 mg/kg IV/PO every 12 hrs on Day 1, then 4 mg/kg IV/PO every 12 hrs.

Alternative regimens:
- liposomal amphotericin B 3-5mg/kg IV daily.
- posaconazole:
  - delayed-release tablets 300 mg PO/12h on Day 1, then 300 mg PO daily;
  - suspension 200 mg QID, then 400 mg PO/12h after stabilization of disease;
  - posaconazole IV 300 mg IV over 90 mins every 12h Day 1, then 300 mg IV daily.
- isavuconazolium sulfate (prodrg of isavuconazole) 372 mg PO/IV q8h × 6 doses, later 372 mg PO/IV daily.

NOTE 1: High doses of liposomal and deoxycholate amphotericin B need careful administration and monitoring and management – and reference detailed guidance. It should be given with saline fluid loading to reduce renal toxicity, and potassium and magnesium monitoring.

NOTE 2: Posaconazole suspension should be taken with food. Not FDA approved treatment, but clinically used.

NOTE 3: As alternative regimen - posaconazole or isavuconazole injection is preferred.

NOTE 4: Step-down or salvage therapy:
- if the patient is stable, posaconazole or isavuconazole tablet for 3-6 months after 3-6 weeks amphotericin B therapy;
- if there is disease progression while the patient is on amphotericin B therapy, liposomal amphotericin B dose may be increased to 10 mg/kg/d or replace with posaconazole or isavuconazole;
- if the patient has renal compromise, start with isavuconazole or posaconazole.
Cryptococcosis

| Risk factors | • conditions associated with reduced cell mediated immunity (e.g. lymphoma)  
|             | • chronic corticosteroid use  
|             | • HIV infection (CD4 cell count < 200)  
|             | • allogenic stem cell transplant  
|             | • solid organ transplantation. |

| Prevention | Control of risk factors. |

| Signs and symptoms | • Meningoencephalitis (common presentation) with symptoms (fever, malaise, headache, meningism) but could be asymptomatic.  
|                    | – Encephalopathic presentation: lethargy, seizure, dementia, personality changes or altered mental status.  
|                    | – Meningitis (fever, malaise, headache, meningism is seen in 25% of patients).  
|                    | • Less frequent: pulmonary (pneumonia presentation) or disseminated disease. |

| Diagnosis | • Direct microscopy, culture, serology (CrAg, LAT, EIA, LFA), molecular identification (PCR).  
|           | • Lumbar puncture: analysis of CSF: CrAg, India staining, culture. |

| Treatment | Primary regimen:  
|           | • induction phase: liposomal amphotericin B 3-4 mg/kg IV daily (or amphotericin B or amphotericin B lipid complex) AND fluconazole 800-1200 mg/day IV/PO for 2 weeks; or Duration > 2 weeks or until CSF sterilization achieved (if takes more than 2 weeks).  
|           | • consolidation phase: fluconazole 800 mg/daily for 10 weeks.  
|           | Alternative regimens:  
|           | • amphotericin B deoxycholate AND fluycytosine followed by fluconazole  
|           | • fluconazole AND fluycytosine  
|           | • amphotericin B deoxycholate AND fluconazole  
|           | • fluycytosine (5-FC) 25 mg/kg PO four times daily if no renal impairment. |
7.4 Treatment for influenza infection fact sheet

- See Guidelines for the clinical management of severe illness from influenza virus infections, March 2022.
- Oseltamivir is recommended to be used when influenza is suspected or known to be circulating.
- If testing for influenza is not possible, empiric treatment is indicated.
- Oseltamivir is not proven to be effective for COVID-19.

<table>
<thead>
<tr>
<th>Populations</th>
<th>Dosing*</th>
</tr>
</thead>
<tbody>
<tr>
<td>Adults</td>
<td></td>
</tr>
<tr>
<td>Mild illness</td>
<td>75 mg orally, twice daily for 5 days</td>
</tr>
<tr>
<td>With severe illness or severe immunocompromising conditions</td>
<td>75 mg orally, twice daily for 5 days</td>
</tr>
<tr>
<td></td>
<td>Consider higher dose: 150 mg orally, twice daily</td>
</tr>
<tr>
<td>Children ≥ 1 year old</td>
<td></td>
</tr>
<tr>
<td>&lt; 15 kg</td>
<td>30 mg orally, twice daily for 5 days</td>
</tr>
<tr>
<td>15 to &lt; 23 kg</td>
<td>45 mg orally, twice daily for 5 days</td>
</tr>
<tr>
<td>23 to &lt; 40 kg</td>
<td>60 mg orally, twice daily for 5 days</td>
</tr>
<tr>
<td>≥ 40 kg</td>
<td>75 mg orally, twice daily for 5 days</td>
</tr>
<tr>
<td>Children &lt; 1 year old</td>
<td></td>
</tr>
<tr>
<td>14 days to 1 year</td>
<td>3 mg/kg orally, twice daily for 5 days</td>
</tr>
</tbody>
</table>

**Note:**
- The route of administration can be either via NG or OG tube if the patient cannot take medication orally (see safety profile). Where the clinical course remains severe or progressive, despite ≥ 5 days of antiviral treatment, treatment should be continued without a break until virus infection is resolved or there is satisfactory clinical improvement.

**Safety considerations and side-effects**

**Safety profile:** oseltamivir has not been associated with increased adverse effects in adult outpatients. However, oseltamivir has not been evaluated in severely ill patients, pregnancy or paediatric populations. Oseltamivir should be used with caution:

- In patients with kidney disease: reduce dose to 75 mg daily if creatinine clearance is 10-30 mL/min.
- In patients with liver disease the safety and efficacy has not been evaluated, so dose reduction is not recommended now.
- For pregnant or nursing mothers, oseltamivir is recommended as therapy in pandemic influenza (H1N1) virus as there is a high risk of severe illness in pregnant women and there is no evidence of adverse effects or birth defects.

**Side-effects:** side-effects are generally minor:

- Gastrointestinal tract: nausea (mitigated by taking with food), vomiting.
- Rare neuropsychiatric adverse events - association seen primarily in one country, causality has not been established.
### Oral formulations

<table>
<thead>
<tr>
<th>Formulations</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>Capsules</td>
<td>30 mg, 45 mg, 75 mg each</td>
</tr>
<tr>
<td></td>
<td>Store at room temperature (15-30 °C)</td>
</tr>
<tr>
<td>Liquid suspension</td>
<td>White powder mixed with 23 mL of drinking water fruit flavoured</td>
</tr>
<tr>
<td></td>
<td>Refrigeration required</td>
</tr>
<tr>
<td></td>
<td>Use within 10 days</td>
</tr>
<tr>
<td></td>
<td>Oral dispenser included (must confirm dosage and volume when administering)</td>
</tr>
<tr>
<td>Oral suspension</td>
<td>If commercial suspension unavailable a suspension may be prepared from oseltamivir capsules</td>
</tr>
</tbody>
</table>

**Note:**
- Preparation of oral oseltamivir suspension:
  - The inhouse suspension should be made at 15 mg/mL for patients > 1 year; and 10 mg/mL for ≤ 1 year.
  - The suspension can be made from oseltamivir phosphate capsules using sterile water at the bedside.
References and resources


CDC websites:
https://www.cdc.gov/fungal/diseases/mucormycosis/diagnosis.html
https://www.cdc.gov/fungal/diseases/mucormycosis/symptoms.html
https://www.cdc.gov/fungal/diseases/mucormycosis/health-professionals.html


EMEA. CHMP assessment report on novel influenza (H1N1) outbreak, Tamiflu (oseltamivir), Relenza (zanamivir). (EMEA/H/A-5.3/1172).


Sepsis and septic shock
Sepsis and septic shock

Summary


To treat patients with septic shock, it is crucial to deliver early, targeted resuscitation using crystalloid fluid, vasopressors and, in some cases, inotropes and/or blood transfusion. Fluid resuscitation with crystalloid fluids remains the most common intervention for septic shock; it should be given as a challenge to improve targets of perfusion, and promptly stopped when no longer responsive, to avoid harms of excess fluid. Resuscitation strategies for children with septic shock should be modified if the child has severe malaria with anaemia or severe malnutrition or is being cared for in settings without ICU capacity, specifically invasive mechanical ventilation. Resuscitation targets for adults and children include improved blood pressure and other markers of tissue perfusion.

### Markers of tissue perfusion in adults and children

<table>
<thead>
<tr>
<th>Marker</th>
</tr>
</thead>
<tbody>
<tr>
<td>Capillary refill &lt; 2 sec</td>
</tr>
<tr>
<td>Absence of skin mottling</td>
</tr>
<tr>
<td>Strong peripheral pulses</td>
</tr>
<tr>
<td>Warm and dry extremities</td>
</tr>
<tr>
<td>Regular urine output</td>
</tr>
<tr>
<td>Normal mental status</td>
</tr>
<tr>
<td>Normalization of lactate</td>
</tr>
<tr>
<td>In children: improved heart rate</td>
</tr>
<tr>
<td>(Tachycardia is an early sign of septic shock and low blood pressure is a late finding)</td>
</tr>
</tbody>
</table>

Refer to the shock quick cards for initial approach and management of patients with septic shock; from the WHO-ICRC Basic emergency care (BEC): approach to the acutely ill and injured.
Tools

8.1  Sepsis and septic shock definitions
8.2  Sequential Organ Failure Assessment (SOFA) score
8.3  Quick Sequential Organ Failure Assessment (qSOFA)
8.4  Paediatric Logistic Organ Dysfunction (PELOD-2) score
8.5  Algorithm on targeted resuscitation in adults with septic shock
8.6  Algorithms on initial resuscitation, and on fluid and vasoactive-inotrope management for children with septic shock
   8.6.1  Initial resuscitation algorithm for children
   8.6.2  Fluid and vasoactive-inotrope management algorithm for children
8.7  Guide to the use of vasopressors in septic shock for adults and children
8.8  Five rules for passive leg raise (PLR)
8.1 Sepsis and septic shock definitions

For further information see: Surviving Sepsis Campaign (October 2021)

**Sepsis adults**

*Life-threatening organ dysfunction caused by a dysregulated host response to suspected or proven infection*

**Signs of organ dysfunction include:**
- Altered mental status
- Difficulty or fast breathing
- Low oxygen saturation
- Reduced urinary output
- Fast heart rate
- Weak pulse
- Cold extremities
- Low blood pressure
- Skin mottling

Laboratory evidence of:
- coagulopathy
- thrombocytopenia
- acidosis
- high lactate
- hyperbilirubinaemia

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**Sepsis children**

*Suspected or proven infection and ≥ two age-based systemic inflammatory response syndrome (SIRS) criteria, of which one must be abnormal temperature or white blood cell count*

**SIRS criteria include:**
- Abnormal temperature < 36 °C or > 38.5 °C
- Heart rate > 2 SD above normal for age or bradycardia if < 1 year of age
- Respiratory rate > 2 SD above normal for age
- Abnormal white blood cell count or > 10% immature neutrophils

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**Septic shock adults**

*Sepsis with circulatory and cellular/metabolic abnormalities profound enough to substantially increase mortality*

**Criteria:**
- Persisting hypotension despite volume resuscitation, requiring vaspressors to maintain MAP ≥ 65 mmHg
  - or
- Serum lactate level > 2 mmol/L (18 mg/dL)
Septic shock children

Sepsis with circulatory and cellular/metabolic abnormalities profound enough to substantially increase mortality

Criteria: two or three of the following

- Altered mental status
- Tachycardia or bradycardia
  - HR < 90 bpm or > 160 bpm in infants and
  - HR < 70 bpm or > 150 bpm in children
- Prolonged capillary refill (> 2 sec) or feeble pulse
- Tachypnoea
  - Mottled or cool skin or petechial or purpuric rash
- High lactate
- Oliguria
- Hypothermia or hypothermia
  - or
- Any hypotension (SBP < 5th centile or 2 SD below normal for age)

WHO paediatric emergency, triage, assessment and treatment (ETAT): definition of shock for children

Shock is the presence of all three clinical criteria:

- Delayed capillary refill ≥ 3 sec
- Cold extremities
- Weak and fast pulse
  - or
- Frank hypotension for age:

<table>
<thead>
<tr>
<th>Age</th>
<th>&lt; 1 m</th>
<th>1-12 m</th>
<th>1-12 yr</th>
<th>&gt; 12 yr</th>
</tr>
</thead>
<tbody>
<tr>
<td>SBP</td>
<td>&lt; 50</td>
<td>&lt; 70</td>
<td>70+ (2 × age)</td>
<td>&lt; 90</td>
</tr>
</tbody>
</table>

For more information see: WHO Paediatric emergency triage, assessment and treatment (ETAT) ( )
8.2 Sequential Organ Failure Assessment (SOFA) score

The SOFA score is commonly used to describe and quantify organ failure and can also be used to predict outcome. The SOFA score has been proposed for use in triage strategies because it helps to quantify the principle of utility.

The SOFA score ranges from 0 to 24 and includes points related to six organ systems:
- respiratory (hypoxaemia defined by low PaO$_2$/FiO$_2$);
- coagulation (low platelets);
- liver (high bilirubin);
- cardiovascular (hypotension);
- central nervous system (low level of consciousness defined by Glasgow Coma Scale [GCS]);
- renal (low urine output or high creatinine).

Sepsis is defined by an increase in the sepsis-related SOFA score of ≥ 2 points. (Assume the baseline score is 0 if data are not available.) To use the SOFA scoring system for triage, add the points for each clinical characteristic at presentation and then at 48 hours. Both the initial and 48-hour scores are predictive of mortality. The maximum score is 24.

<table>
<thead>
<tr>
<th>Variables</th>
<th>0</th>
<th>1</th>
<th>2</th>
<th>3</th>
<th>4</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Respiratory</strong></td>
<td>PaO$_2$/FiO$_2$, mmHg</td>
<td>&gt; 400</td>
<td>≤ 400</td>
<td>≤ 300</td>
<td>≤ 200$^a$</td>
</tr>
<tr>
<td><strong>Coagulation</strong></td>
<td>Platelets $\times 10^3/\mu$L$^3$</td>
<td>&gt; 150</td>
<td>≤ 150</td>
<td>≤ 100</td>
<td>≤ 50</td>
</tr>
<tr>
<td><strong>Liver</strong></td>
<td>Bilirubin, mg/dL$^b$</td>
<td>&lt; 1.2</td>
<td>1.2-1.9</td>
<td>2.0-5.9</td>
<td>6.0-11.9</td>
</tr>
<tr>
<td><strong>Cardiovascular Hypotension</strong></td>
<td>No hypotension</td>
<td>Mean arterial pressure &lt; 70 mmHg</td>
<td>Dopamine ≤ 5 or Dobutamine (any dose)</td>
<td>Dopamine &gt; 5, Epinephrine ≤ 0.1, or Norepinephrine ≤ 0.1$^c$</td>
<td>Dopamine &gt; 15, Epinephrine &gt; 0.1, or Norepinephrine &gt; 0.1$^c$</td>
</tr>
<tr>
<td>Central nervous system</td>
<td>GCS</td>
<td>15</td>
<td>13-14</td>
<td>10-12</td>
<td>6-9</td>
</tr>
<tr>
<td>Renal</td>
<td>Creatinine, mg/dL or urine output, mL/day</td>
<td>&lt; 1.2</td>
<td>1.2-1.9</td>
<td>2.0-3.4</td>
<td>3.5-4.9 or &lt; 500 mL/day</td>
</tr>
</tbody>
</table>

Notes:
- bpm - beats per minute; FiO$_2$ - fraction of inspired oxygen; MAP - mean arterial pressure; PaO$_2$ - partial pressure of oxygen; SBP - systolic blood pressure; SOFA - sequential organ failure assessment.
- $^a$ Values are with respiratory support.
- $^b$ To convert bilirubin from mg/dL to µmol/L, multiply by 17.1.
- $^c$ Adrenergic agents administered for at least 1 hour (doses given are in µg/kg per minute).
- $^d$ To convert creatinine from mg/dL to µmol/L, multiply by 88.4.
### 8.3 Quick Sequential Organ Failure Assessment (qSOFA)

The qSOFA score (also known as quickSOFA) is a bedside prompt that may identify adult patients with suspected infection who may be at greater risk for a poor outcome outside the ICU. In a patient with suspected infection, the presence of $\geq$ two of the following qSOFA criteria may be associated with an increased risk of death:

*Note: Although the presence of a positive qSOFA should alert clinicians to the possibility of sepsis in all resource settings, given the poor sensitivity of the qSOFA, there is a strong recommendation by the Surviving Sepsis Campaign (October 2021) against its use as a single screening tool.*

For further information see the latest Surviving Sepsis Campaign guidelines (October 2021).

#### qSOFA Calculator

<table>
<thead>
<tr>
<th>Criteria</th>
<th>Yes</th>
<th>No</th>
</tr>
</thead>
<tbody>
<tr>
<td>Altered mental status (GCS &lt; 15, AVPU scale)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Respiratory rate (RR) 22 breaths per minute</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Systolic blood pressure (SBP) ≤ 100 mmHg</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Source: Adapted from www.qsofa.org

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8.4 Paediatric Logistic Organ Dysfunction (PELOD-2) score

Multiple organ dysfunction syndrome is a frequent cause of death in adult and paediatric ICUs. The PELOD-2 score was developed to describe the severity of age-specific organ dysfunction in children and has since been validated in many settings. This descriptive score relies on **10 variables that correspond to five different organ dysfunctions**. Any increased organ dysfunction in the PELOD-2 score is closely related to an increased risk of mortality, but neurologic and respiratory dysfunctions are the most critical.

In the population in which the PELOD-2 was developed, a score of 10 was associated with ~10% probability of mortality, while a score of 20 was associated with > 90% probability of mortality. However, the predicted risk of death is population specific and varies with resource availability.

All variables must be collected, but measurements can be done only if justified by the patient’s clinical status. If a variable is not measured, it should be considered normal. If a variable is measured more than once in 24 hours, the worst value is used in calculating the score.
### Organ dysfunctions and variables

<table>
<thead>
<tr>
<th>Organ dysfunctions and variables</th>
<th>Points by severity levels</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>0</td>
</tr>
<tr>
<td><strong>Neurologic</strong></td>
<td></td>
</tr>
<tr>
<td>GCS</td>
<td>≥ 11</td>
</tr>
<tr>
<td>Pupil reaction</td>
<td>Both reactive</td>
</tr>
<tr>
<td><strong>Cardiovascular</strong></td>
<td></td>
</tr>
<tr>
<td>Lactate (mmol/L)</td>
<td>&lt; 5.0</td>
</tr>
<tr>
<td><strong>Cardiovascular</strong></td>
<td></td>
</tr>
<tr>
<td><strong>Mean arterial pressure (mmHg)</strong></td>
<td></td>
</tr>
<tr>
<td>0 to &lt; 1 months</td>
<td>≥ 46</td>
</tr>
<tr>
<td>1-11 months</td>
<td>≥ 55</td>
</tr>
<tr>
<td>12-23 months</td>
<td>≥ 60</td>
</tr>
<tr>
<td>24-59 months</td>
<td>≥ 62</td>
</tr>
<tr>
<td>60-143 months</td>
<td>≥ 65</td>
</tr>
<tr>
<td>&gt;144 months</td>
<td>≥ 67</td>
</tr>
<tr>
<td><strong>Renal</strong></td>
<td></td>
</tr>
<tr>
<td>Cr (µmol/L)</td>
<td>≤ 69</td>
</tr>
<tr>
<td>0 to &lt; 1 months</td>
<td>≤ 22</td>
</tr>
<tr>
<td>1-11 months</td>
<td>≤ 34</td>
</tr>
<tr>
<td>12-23 months</td>
<td>≤ 50</td>
</tr>
<tr>
<td>24-59 months</td>
<td>≤ 58</td>
</tr>
<tr>
<td>60-143 months</td>
<td>≤ 92</td>
</tr>
<tr>
<td>&gt;144 months</td>
<td></td>
</tr>
<tr>
<td><strong>Respiratory</strong></td>
<td></td>
</tr>
<tr>
<td>PaCO₂ (mmHg) FiO₂</td>
<td>≥ 61</td>
</tr>
<tr>
<td>Haematologic</td>
<td></td>
</tr>
<tr>
<td>WBC (× 10⁹/L)</td>
<td>≥ 142</td>
</tr>
<tr>
<td>Platelets (× 10⁹/L)</td>
<td></td>
</tr>
</tbody>
</table>

Notes:

- **Neurologic dysfunction**: Glasgow Coma Score: use the lowest value. If the patient is sedated, record the estimated Glasgow Coma Score before sedation. Assess only patients with known or suspected acute central nervous system disease. Pupillary reactions: nonreactive pupils must be > 3 mm. Do not assess after iatrogenic pupillary dilation.
- **Cardiovascular dysfunction**: heart rate and mean arterial pressure: do not assess during crying or iatrogenic agitation.
- **Respiratory dysfunction**: PaO₂ used: use arterial measurement only. PaO₂/FiO₂ ratio is considered normal in children with cyanotic heart disease. PaCO₂ can be measured from arterial, capillary or venous samples. Invasive ventilation; the use of mask ventilation is not considered invasive ventilation. FiO₂: fraction of inspired oxygen.
8.5 Algorithm on targeted resuscitation in adults with shock

Shock features in adults:
- SBP < 100 mmHg, HR > 90/min
- Signs of hypoperfusion:
  - Delayed capillary refill > 3 sec
  - Cold extremities
  - RR > 22/min
  - Decreased urinary output (< 0.5 mL/kg/hr)
  - Altered mental status
  - Elevated lactate

General clinical management:
1. Immediately obtain body fluid cultures
2. Give empiric antimicrobials (within 1 hr)
3. Check laboratory: CBC, biochemistry, lactate, CRP
4. Check serum glucose, replete as needed
5. Monitor and replete electrolytes as needed
6. Treat severe acidemia (pH < 7.2) with bicarbonate
7. Monitor fluid balance
8. Provide symptom relief (paracetamol)
9. Begin early nutrition (if tolerating oral)
10. Counsel patient and family

Fluid responsiveness:
When fluid therapy improves blood pressure or markers of perfusion (i.e., test blood pressure after fluid bolus or positive leg raise)

Markers of good perfusion:
- SBP > 90-100 mmHg or age appropriate for child
- Strong pulse
- Normal RR
- Normal mental status (AVPU)
- Urine output > 0.5 mL/kg/hr for adult or > 1 mL/kg/hr for child
- Brisk capillary refill
- Normalization of lactate

Fluid overload:
- Dyspnea, drop in SpO2, increase RR
- Elevated JVP > 12 cm
- Peripheral oedema
- Hepatomegaly in children

Patient with shock features

**SpO2 < 94%**

Monitor for fluid overload

**Yes**

Fluid resuscitation
- Use crystalloid: Ringer’s lactate or normal saline
- Invert IV or IO line safely
- Give initial fluid bolus

Stop fluid challenges
- Switch to maintenance fluids (IV or oral per patient)
- Check vitals every 1 hr

Reassess vital signs every 10 min for fluid responsiveness, markers of perfusion and fluid losses
Are targets of perfusion met (shock resolved)?

**No**

Stop fluid challenges
- Repeat fluid challenge and reassess targets of perfusion

Refractory shock

If markers of perfusion are stable/improving, reduce inotrope and vasopressor dose

Initiate vasopressor perfusion:
- Epinephrine/norepinephrine: start 0.1 mcg/kg/min, titrate up by 0.1 mcg/kg/min every 10-15 min
- Maximum: 1.0 mcg/kg/min

**Notes:**
AVPU - alert, verbal, pain, unresponsive; HR - heart rate; JVP - jugular venous pressure; RR - respiratory rate;
SBP - systolic blood pressure; SpO2 - oxygen saturation.
Source: Modified from Optimized supportive care for Ebola virus disease (WHO, 2019) [1]
8.6 Algorithms on initial resuscitation, and on fluid and vasoactive-inotrope management for children with septic shock

This initial resuscitation algorithm for children from the Surviving Sepsis Campaign, is based on recently published paediatric sepsis and septic shock guidelines and has been adapted for use in health care systems with and without intensive care.

8.6.1 Initial resuscitation algorithm for children

*See fluid and vasoactive algorithm. Note: Fluid bolus should be omitted from bundle if a) fluid overload is present or b) it is a low-resource setting without hypotension. Fluid in mL/kg should be dosed as ideal body weight.

**Hydrocortisone may produce benefit or harm.

Notes: ECLS - extracorporeal life support; VA - veno-arterial; VV - veno-venous.
Source: Surviving Sepsis Campaign paediatric patients ( )
Fluid in mL/kg should be dosed as ideal body weight.

**Abnormal Perfusion with or without Hypotension**

- If signs of fluid overload are absent, administer fluid bolus, 10-20 mL/kg.
- Repeat assessment of hemodynamic response to fluid and consider fluid boluses, 10-20 mL/kg, until shock resolves or signs of fluid overload develop.
- Assess cardiac function.
- Consider epinephrine if there is myocardial dysfunction or epinephrine/norepinephrine if shock persists after 40-60 mL/kg (or sooner if signs of fluid overload develop).

**Abnormal Perfusion WITHOUT Hypotension**

- Do NOT give fluid bolus unless there are signs of dehydration with ongoing fluid losses (eg, diarrhea).
- Start maintenance fluids.
- Monitor hemodynamics closely.
- Consider vasoactive-inotropic support (if available).

**Abnormal Perfusion WITH Hypotension**

- If signs of fluid overload are absent, administer fluid bolus, 10-20 mL/kg.
- Assess hemodynamic response to fluid and repeat fluid boluses, 10-20 mL/kg, until hypotension resolves or signs of fluid overload develop.
- Assess cardiac function (if available)
- Consider epinephrine/norepinephrine if hypotension persists after 40 mL/kg or sooner if signs of fluid overload develop.

**Shock resolved, perfusion improved**

- Do not give more fluid boluses.
- Consider maintenance fluids.
- Monitor for signs/symptoms of recurrent shock.

*Hypotension in healthcare systems WITHOUT intensive care is defined as either: SBP < 50 mm Hg in children aged < 12 months, SBP < 60 mm Hg in children aged 1 to 5 years, SBP < 70 mm Hg in children aged > 5 years.

Presence of all 3 World Health Organization criteria: cold extremities, prolonged capillary refill > 3 seconds, weak/fast pulse

Source: Rhodes et al. (2020); Surviving Sepsis Campaign paediatric patients (Weiss et al. 2020).
8.7 Guide to the use of vasopressors in septic shock for adults and children

In adults, the Surviving Sepsis Campaign guidelines recommend vasopressors to be started if MAP < 65 mmHg. Administer vasopressors at a strictly controlled rate, titrate to maintain MAP 65 mmHg, reduce as the MAP improves and discontinue promptly when no longer needed. Dose initiation and titration should be individualized. The MAP goal can be individualized based on other clinical history (i.e. consider higher MAP target > 80 mmHg in patients with chronic hypertension).

Also, target other markers of perfusion such as:

<table>
<thead>
<tr>
<th>Markers of tissue perfusion in adults and children</th>
</tr>
</thead>
<tbody>
<tr>
<td>Capillary refill &lt; 2 sec</td>
</tr>
<tr>
<td>Absence of skin mottling</td>
</tr>
<tr>
<td>Strong peripheral pulses</td>
</tr>
<tr>
<td>Warm and dry extremities</td>
</tr>
<tr>
<td>Regular urine output</td>
</tr>
<tr>
<td>Normal mental status</td>
</tr>
<tr>
<td>Normalization of lactate</td>
</tr>
<tr>
<td>In children: improved heart rate</td>
</tr>
<tr>
<td>(tachycardia is an early sign of septic shock and low blood pressure is a late finding)</td>
</tr>
</tbody>
</table>

Vasopressors: norepinephrine is recommended as the first-line agent; however, epinephrine can be used as an alternative. Dopamine is not recommended because of the risk of tachyarrhythmias and concern of worse outcome.

Inotropes: dobutamine when there are persistent signs of hypoperfusion and clinical evidence of myocardial dysfunction (i.e. by ECHO, ScvO₂ < 70%) after adequate MAP and fluid status achieved.

Vasopressin is recommended as a second-line agent for refractory distributive shock.

Note: Vasopressor and inotrope selection should be informed by the patient’s physiology including heart rate and assessment of cardiac function.

In children, the Surviving Sepsis Campaign guidelines recommend vasopressors if clinical signs of shock persist after fluid resuscitation and should not be delayed. These agents should be administered at a strictly controlled rate and titrated to achieve targets of adequate tissue perfusion.

Recommendation of epinephrine or norepinephrine as the first-line vasoactive infusion guided by clinician preference, individual patient physiology, and local system factors. If shock persists, add a second agent, and vasopressin can be added in children requiring high-dose vasopressors.

Note: Children can move between various shock states and vasopressors should be adjusted accordingly.
## Dosing of vasopressors in adults and children

<table>
<thead>
<tr>
<th>Route</th>
<th>Norepinephrine</th>
<th>Dobutamine</th>
<th>Epinephrine</th>
<th>Vasopressin</th>
</tr>
</thead>
<tbody>
<tr>
<td>Central vein preferred</td>
<td>Initial: 0.05 µg/kg/min</td>
<td>Initial: 2-5 µg/kg/min</td>
<td>Initial: 0.05 µg/kg/min</td>
<td>Initial: 0.01-0.04 units/min</td>
</tr>
<tr>
<td></td>
<td>Range: increase by 0.1 µg/kg/min increments; consider refractory if &gt; 1 µg/kg/min</td>
<td>Range: increase by 2.5 µg/kg/min increments; maximum 20 µg/kg/min</td>
<td>Range: increase by 0.1 µg/kg/min increments; consider refractory if &gt; 1 µg/kg/min</td>
<td>Fixed dose</td>
</tr>
<tr>
<td>Peripheral vein if necessary</td>
<td>Same dosing</td>
<td>Same dosing</td>
<td>Same dosing</td>
<td>Same dosing</td>
</tr>
</tbody>
</table>

Note: Thus, in septic shock, inotropes should be used in combination with vasopressors to maintain MAP at goal in adults, and children with low systemic vascular resistance.

*Requires close nursing care to check infusion site. If necrosis, stop infusion and consider injection of 1 mL phentolamine (vasodilator) solution subcutaneously. Phentolamine dose (adults): 5-10 mg in 10 mL of NS.

## Side-effects of vasopressors and inotropes

**Side-effects of vasopressors**

- **Tachyarrhythmias**
- **Ischaemia to organs**
- **Cool and cyanotic extremities**
- **Soft tissue necrosis (with peripheral administration if the vasopressor is extravasated)**

**Side-effects of inotropes**

- **Tachyarrhythmias**
- **Hypotension (due to peripheral vasodilation)**
8.8 The five rules for passive leg raise (PLR)

In acute circulatory failure, passive leg raising (PLR) is a test that predicts whether cardiac output (CO) will increase with volume expansion. By transferring a volume of around 300 mL of venous blood from the lower body towards the right heart, PLR mimics a fluid challenge. However, no fluid is infused and the haemodynamic effects are rapidly reversible.

Best method for passive leg raising - the five rules to be followed

1. Check the trunk is at 45°
2. Use the bed adjustment and avoid touching the patient (pain, awakening)
3. Assess PLR effects by directly measuring CO (not with blood pressure only)
4. Use a real-time measurement of CO
5. Reassess CO in the semi-recumbent position (should return to baseline)

Notes: CO - cardiac output; PLR - passive leg raise.
References and resources


8. Sepsis and septic shock


Acute respiratory distress syndrome (ARDS)
Acute respiratory distress syndrome (ARDS)

Summary

When to suspect ARDS
• signs of severe or worsening respiratory distress
• hypoxaemia (SpO₂ < 90%) despite escalating oxygen therapy or SpO₂/FiO₂ < 315
• new bilateral opacities on chest images
• pulmonary oedema, cardiac failure or fluid overload not the primary cause.

Intubation and invasive mechanical ventilation may be indicated for many patients with ARDS (particularly those with moderate or severe ARDS) and hypoxaemic respiratory failure.

Lung protective ventilation (LPV) reduces mortality in patients with ARDS. A lung protective ventilation strategy includes:
• low tidal volumes (TV) (target 6 mL/kg predicted body weight or less);
• low plateau airway pressure (Pplat) (target Pplat ≤ 30 cmH₂O);
• moderate positive end-expiratory pressure (PEEP) (8–14 cmH₂O needed in most ARDS cases);
• driving pressure (Pplat-PEEP) ≤15 cmH₂O to titrate TV when PEEP≥15 and Pplat ≥30.

In ARDS patients who do not require intubation, i.e. mild ARDS, use of high-nasal flow oxygen (HNFO) or non-invasive ventilation (NIV) using CPAP or BiPAP may be safe when: significant hypercapnia is absent, mental status is normal, and haemodynamics are stable. These therapies require a monitored setting, with experienced personnel and capacity for performing emergent endotracheal intubation. A time-limited trial (~1 hr) of CPAP or BiPAP may be appropriate for select patients.

Do not delay the decision to intubate and start invasive mechanical ventilation when indications are present, including:
• severe hypoxaemia refractory to supplemental oxygen use
• severe hypercapnia refractory to non-invasive ventilation (if available)
• haemodynamic instability
• need for airway protection
• presence of emergency signs.
HFNO, BiPAP and CPAP may consume significant quantities of oxygen. See *Oxygen sources and distribution for COVID-19 treatment centres* ( ).

Use airborne precautions when conducting aerosol-generating procedures. See *Transmission of SARS-CoV-2: implications for infection prevention precautions* ( ).

**Tools**

9.1 Memory aid: diagnosis and classification of ARDS in adults
9.2 Memory aid: diagnosis and classification of pARDS in children
9.3 Advanced non-invasive mechanical ventilation in ARDS: algorithm to escalate supportive respiratory therapy
9.4 Checklist for rapid sequence intubation procedure in adults and children
9.5 Considerations for intubation and mechanical ventilation in children
9.6 List of commonly used medicines and dosage in ICU with ventilated patients (adults, children)
9.7 Choice of induction agent in adults
9.8 Choice of induction agent in children
9.9 Protocol to deliver lung protective ventilation (LPV)
  9.10.1 ARDS-net PEEP FiO\textsubscript{2} grid to guide PEEP
  9.10.2 Goals of Pplat and pH in lung protective ventilation
9.10 Memory aid: comparison of normal waveforms during volume and pressure-limited ventilation
9.11 Memory aid: recognizing and interpreting abnormal pressure and flow waveforms during volume control ventilation
9.12 Guide to distinguishing between causes of high peak airway pressures: resistance versus compliance
9.13 Troubleshooting high peak airway pressures, low tidal volumes, desaturation or haemodynamic instability in ventilated patients
9.14 Respiratory care pocket card reference
9.15 Adult ventilation order set (ARDS)
9.16 Checklist for proning in severe ARDS
9.17 Ventilation circuit types, filter and humidifier locations for SARI
9.1 Memory aid: diagnosis and classification of ARDS in adults

Berlin definition of acute respiratory distress syndrome (ARDS), 2012
ARDS Definition Task Force et al. (2012).

<table>
<thead>
<tr>
<th>Timing</th>
<th>Within 1 week of a known clinical insult or new or worsening respiratory symptoms</th>
</tr>
</thead>
<tbody>
<tr>
<td>Chest imaging(^a)</td>
<td>Bilateral opacities - not fully explained by effusions, lobar/lung collapse or nodules</td>
</tr>
<tr>
<td>Origin of oedema</td>
<td>Respiratory failure not fully explained by cardiac failure or fluid overload</td>
</tr>
<tr>
<td></td>
<td>Need objective assessment (e.g. echocardiography) to exclude hydrostatic oedema if no risk factor present</td>
</tr>
<tr>
<td>Oxygenation(^b)</td>
<td>Mild: 200 &lt; PaO(_2) / FiO(_2) ≤ 300 with PEEP or CPAP ≥ 5 cm H(_2)O</td>
</tr>
<tr>
<td></td>
<td>Moderate: 100 &lt; PaO(_2) / FiO(_2) ≤ 200 with PEEP ≥ 5 cm H(_2)O</td>
</tr>
<tr>
<td></td>
<td>Severe: PaO(_2) / FiO(_2) ≤ 100 with PEEP ≥ 5 cm H(_2)O</td>
</tr>
</tbody>
</table>

Notes:
\(^a\) Chest radiograph or computed tomography scan;
\(^b\) If altitude is higher than 1000 m, the correction factor should be calculated as follows: \(\text{PaO}_2 / \text{FiO}_2 \times (\text{barometric pressure} / 760)\);
\(^c\) This may be delivered non-invasively in the mild ARDS group;
CPAP - continuous positive airway pressure; FiO\(_2\) - fraction of inspired oxygen; PaO\(_2\) - partial pressure arterial oxygen; PEEP - positive end-expiratory pressure.

A recent publication suggests a modified definition for resource-constrained environments, that excludes the need for CPAP or PEEP, arterial blood analysis and chest radiograph (Kigali modifications).

Kigali modifications of Berlin definition, 2016
Riviello et al. (2016).

<table>
<thead>
<tr>
<th>Chest imaging</th>
<th>1. Bilateral opacities - not fully explained by effusions, lobar/lung collapse or nodules by chest radiograph or ultrasound</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>2. Ultrasound findings defined as presence of B-lines or consolidations without associated effusions found in at least one area on each side of the chest</td>
</tr>
<tr>
<td></td>
<td>The protocol requires six areas of each side of chest (two anterior, two lateral, two posterolateral) to be examined.</td>
</tr>
<tr>
<td>Oxygenation</td>
<td>SpO(_2) / FiO(_2) ≤ 315, no PEEP or CPAP requirement</td>
</tr>
</tbody>
</table>

176 Clinical Care for Severe Acute Respiratory Infection—Toolkit—Update 2022
## 9.2 Memory aid: diagnosis and classification of pARDS in children

**Paediatric acute respiratory distress syndrome (pARDS) definition**

<table>
<thead>
<tr>
<th>Age</th>
<th>Exclude patients with perinatal related lung disease</th>
</tr>
</thead>
<tbody>
<tr>
<td>Timing</td>
<td>Within 7 days of known clinical insult</td>
</tr>
<tr>
<td>Origin of oedema</td>
<td>Respiratory failure not fully explained by cardiac failure or fluid overload</td>
</tr>
<tr>
<td>Chest imaging</td>
<td>Chest imaging findings of new infiltrates(s) consistent with acute pulmonary parenchymal disease</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Oxygenation</th>
<th>Non-invasive mechanical ventilation</th>
<th>Invasive mechanical ventilation</th>
</tr>
</thead>
<tbody>
<tr>
<td>pARDS (no severity stratification)</td>
<td>Mild</td>
<td>Moderate</td>
</tr>
<tr>
<td>Full face mask bilevel ventilation of CPAP $\geq 5$ cm H$_2$O</td>
<td>$4 \leq$ OI $&lt; 8$</td>
<td>$8 \leq$ OI $&lt; 16$</td>
</tr>
<tr>
<td>PF ratio $\leq 300$, SF ratio $\leq 264$</td>
<td>$5 \leq$ OSI $&lt; 7.5$</td>
<td>$7.5 \leq$ OSI $&lt; 12.3$</td>
</tr>
</tbody>
</table>

**Notes:**
- CPAP - continuous positive airway pressure; OI - Oxygenation Index ($[\text{FiO}_2 \times \text{mean airway pressure} \times 100]/\text{PaO}_2$); OSI - oxygen saturation index ($[\text{FiO}_2 \times \text{mean airway pressure} \times 100]/\text{SpO}_2$); PF ratio - $\text{PaO}_2/\text{FiO}_2$ ratio; SF ratio - $\text{SpO}_2/\text{FiO}_2$ ratio.
9.3 Advanced non-invasive oxygen delivery in ARDS: algorithm to escalate supportive respiratory therapy

**Emergency signs**
- Obstructed or inadequate breathing
- Central cyanosis
- Severe respiratory distress (i.e., significant tachypnoea, accessory muscle use)
- Signs of shock
- Seriously reduced level of consciousness
- Seizures
- Acidosis (pH < 7.30)
- Severe hypoxaemia, P:F < 100

**Do not delay the decision to intubate and start invasive mechanical ventilation when indications are present, including:**
- Severe hypoxaemia refractory to supplemental oxygen use
- Severe hypercapnia refractory to non-invasive ventilation (if available)
- Haemodynamic instability
- Need for airway protection
- Presence of emergency signs

**Notes:** See Antonelli et al., 2001; Bellani et al., 2021; Bellani et al., 2017; Carteaux et al., 2016; Seghal et al., 2015; Suttapanit et al., 2020; Thillie et al., 2013.
9.4 Checklist for rapid sequence intubation procedure in adults and children

This tool can be used before performing endotracheal intubation. Intubation and IMV can be indicated, for adults and children, in case of hypoxaemia refractory to supplemental oxygen, depressed level of consciousness (AVPU) and severe shock.

1. Appropriate infection prevention precautions
   - If suspect COVID-19 or any other droplet/airborne infection, use airborne precautions.
   - Minimize personnel in patient’s room during aerosol-generating procedures.
   - Mask ventilation should be avoided if possible to minimize aerosolization.
     Note: If mask ventilation is necessary, then an inline bacterial viral filter should be utilized and for adults a two-handed, two-person technique should be used.

2. Equipment needed
   - Suction: working Yankauer sucker under right side of pillow.
   - Self-inflating resuscitation bag, 15 L/min oxygen, PEEP valve (pre-oxygenation and post-intubation) endotracheal tube (ETT): correct size, cuff checked and lubricated +/- stylet two working laryngoscopes with blades (direct laryngoscopes or videolaryngoscope).
   - 10 mL syringe. Tube tie or tape.
   - Gum elastic bougie on trolley top. Oropharyngeal airway on trolley top.
   - Confirm laryngeal mask airway and surgical airway are available.
   - Capnometry or capnography set up.
   - Stethoscope.
   - Ventilator checks complete.
   - Ensure appropriate bacterial viral filter placement and in-line/closed suction setup in place.
   - Alternate oxygen source (cylinder/flowmeter).

3. Drugs needed
   - IV access patent and accessible.
   - Induction agents: hypnotic/analgesic/neuromuscular blockers bolus and maintenance infusions prepared.
   - Vasopressor and atropine prepared.

4. Team role description
   - Lead provider 1: airway management and drug administration. Provider 2: assistance and drug administration.
   - Provider 3: cricoid pressure (controversial).
   - Provider 4 (respiratory therapist): airway management and ventilation assistance.
   - Team members available immediately outside patient room.
   - Team members available to provide spotting for correct PPE donning and doffing.

Note: See Gelb et al., 2018.
Rapid sequence intubation (RSI)

Definition: RSI is a protocol designed for the quick intubation of the trachea.

Target: Patients suspected of having an increased risk of aspirating stomach contents into the lungs.

Technique: Quicker form of the process normally used to “induce” a state of general anaesthesia.

It uses drugs to rapidly allow an ETT to be placed between the vocal cords, by blocking the patient’s involuntary reflexes and muscle tone in the oropharynx and larynx. Once the ETT has been passed between the vocal cords, a cuff is inflated around the tube in the trachea and the patient can then be artificially ventilated. Correct ETT position can be verified by direct visualization through the vocal cords; capnography (persistent CO₂ return; may show CO₂ transiently if in esophagus); high SpO₂; bilateral breath sounds on chest auscultation; and correct position on X-ray.

5. Protocol

☐ Pre-oxygenate for 5 minutes with 100% FiO₂

Ideally this is done with a device capable of reliably delivering 100% FiO₂ (HNFO, CPAP or BiPAP).

Note: Children, infants and obese patients have reduced functional residual capacity; they can desaturate quickly on induction.

Note: If the mask’s seal is broken, or for example the mask is removed so that the patient can speak clearly, the process must start again for an additional 5 minutes.

☐ Anticipate shock

Benzodiazepines, thiopental, inhalational agents and propofol cause myocardial depression and vasodilation; this can unmask or worsen shock. Induction agents such as etomidate, ketamine or opiates can also cause haemodynamic instability in patients with high pre-induction sympathetic tone.

☐ Anticipate instability and consider use ketamine for induction if available.

☐ Anticipate instability by pre-loading with volume (10-20 mL/kg 0.9% isotonic crystalloid) and/or starting/increasing inotropic support.

Ensure a blood pressure raising agent that can be given as a bolus is immediately available in a syringe bedside (e.g. ephedrine, phentylephrine, diluted epinephrine). Check the blood pressure as often as possible - if an automatic blood pressure machine is used, set at frequent intervals (e.g. every 1 minute) during induction and intubation.

☐ Use induction agent ± opiate and neuromuscular relaxant in all patients to optimize the view and make intubation easier.

☐ Confirm correct ETT placement with end-tidal CO₂ as the gold standard. When end-tidal CO₂ measurement is not available, several additional signs and techniques are frequently used to assess endotracheal tube placement, including:
  - direct visualization of cords
  - improving SpO₂
  - bilateral equal air on auscultation
  - condensation in the endotracheal with exhalation
  - external palpation of the endotracheal tube balloon in the trachea
  - ultrasound of the endotracheal tube in the trachea
  - chest X-ray position of ETT tip 1-2 cm above the carina (in adults), or T3 posterior.
9.5 Considerations for intubation and mechanical ventilation in children

Specifications for children

Decompress the stomach to prevent diaphragmatic splinting:
- use airway adjuncts to reduce stomach inflation;
- in bag-mask ventilation, place NG tube early and regularly aspirate with large bore syringe to decompress stomach.

Consider atropine in neonates and children to prevent bradycardia caused by vagal stimulation during laryngoscopy or with the use of succinylcholine.

Anatomical differences between children and adults

Anatomical differences between children and adults can make ventilation more difficult.
- **Lower chest wall rigidity** of children implies an earlier respiratory failure in infants in any pathology that causes ↓ compliance of lung, e.g. viral pneumonitis.
- **Smaller airway diameter** of children implies an upper airway resistance.
- **Larger abdomen** of children implies a ↓ functional residual capacity → ↑ atelectasis at end expiration and atelectrauma.
- **Larger tongue, anterior larynx, narrow cricoid ring, larger occiput** require positioning of the airway (e.g. use of neck rolls) to optimize visualization on laryngoscopy:
  - neonates and infants in neutral position
  - older children in “sniffing morning air” position.

Tips: Anticipate a difficult airway, particularly if stridor or a small posteriorly placed jaw are present. Pre-oxygenate, have a range of ETT and blades and the most experienced operator available.

Choice of endotracheal tubes size for paediatric patients

<table>
<thead>
<tr>
<th></th>
<th>Term infant</th>
<th>Estimate at 6 months</th>
<th>Children ≥ 1 year (kg)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Diameter (size) of ETT (cuffed preferred)</td>
<td>3–3.5</td>
<td>3.5–4</td>
<td>(Age/4) + 4 (uncuffed); (Age/4) + 3.5 (cuffed)</td>
</tr>
<tr>
<td>Length oral ETT at lips (confirm on X-ray)</td>
<td>8–9</td>
<td>10</td>
<td>(Age/2) + 12 cm</td>
</tr>
<tr>
<td>Length nasal ETT at nose (confirm on X-ray)</td>
<td>10–11</td>
<td>12</td>
<td>(Age/2) +15 cm</td>
</tr>
<tr>
<td>Suction catheter size</td>
<td>$2 \times \text{ETT} = 6$</td>
<td>$2 \times \text{ETT} = 8$</td>
<td>$2 \times \text{ETT}$</td>
</tr>
</tbody>
</table>
### Normal physiologic parameters and equipment

<table>
<thead>
<tr>
<th>Age</th>
<th>kg</th>
<th>HR</th>
<th>MAP</th>
<th>RR</th>
<th>Blade</th>
<th>ETT mm</th>
<th>ETT@Lips</th>
</tr>
</thead>
<tbody>
<tr>
<td>0-1 m</td>
<td>&lt;1</td>
<td>140</td>
<td>30</td>
<td>&lt;60</td>
<td>Miller 0</td>
<td>2.5</td>
<td>7 cm</td>
</tr>
<tr>
<td>0-1 m</td>
<td>1-2</td>
<td>140</td>
<td>30</td>
<td>&lt;60</td>
<td>Miller 0</td>
<td>3.0</td>
<td>8 cm</td>
</tr>
<tr>
<td>0-1 m</td>
<td>2-3</td>
<td>130-140</td>
<td>30</td>
<td>&lt;60</td>
<td>Mil0/Mil1</td>
<td>3.5</td>
<td>9 cm</td>
</tr>
<tr>
<td>0-1 m</td>
<td>&gt;3</td>
<td>130-140</td>
<td>40</td>
<td>&lt;60</td>
<td>Mil0/Mil1</td>
<td>3.5-4.0</td>
<td>10 cm</td>
</tr>
<tr>
<td>1-6 m</td>
<td>4-6</td>
<td>130</td>
<td>50</td>
<td>24-30</td>
<td>Mil1/Mil1.5</td>
<td>3.5-4.0</td>
<td>12 cm</td>
</tr>
<tr>
<td>6 m-1 yr</td>
<td>6-10</td>
<td>130</td>
<td>60</td>
<td>22-26</td>
<td>Wis 1.5</td>
<td>4.0</td>
<td>13 cm</td>
</tr>
<tr>
<td>1-2 yr</td>
<td>10-12</td>
<td>120</td>
<td>60</td>
<td>20-24</td>
<td>Wis 1.5</td>
<td>4.5</td>
<td>14 cm</td>
</tr>
<tr>
<td>2-4 yr</td>
<td>12-16</td>
<td>110</td>
<td>60</td>
<td>18-22</td>
<td>Wis 1.5/Mac 2</td>
<td>5.0</td>
<td>15 cm</td>
</tr>
<tr>
<td>4-6 yr</td>
<td>16-20</td>
<td>90-110</td>
<td>70</td>
<td>16-20</td>
<td>Mil 2/Mac 2</td>
<td>5.5</td>
<td>16 cm</td>
</tr>
<tr>
<td>6-8 yr</td>
<td>20-30</td>
<td>90</td>
<td>70</td>
<td>16-20</td>
<td>Mil 2/Mac 2</td>
<td>6.0</td>
<td>17 cm</td>
</tr>
<tr>
<td>9-12 yr</td>
<td>30-45</td>
<td>80</td>
<td>70-80</td>
<td>12-18</td>
<td>Mil/Mac 2-3</td>
<td>6.5-7.0</td>
<td>18 cm</td>
</tr>
<tr>
<td>&gt;12 yr</td>
<td>&gt;50</td>
<td>75</td>
<td>70-80</td>
<td>10-16</td>
<td>Mil/Mac 2-3</td>
<td>7.0</td>
<td>20-22 cm</td>
</tr>
</tbody>
</table>

### Neonatal and paediatric general estimates

- **Neonatal “1-2-3(kg)/7-8-9 (ETT@Lips) rule”**
- For preterm and term newborns: MAP equals the number of weeks post conceptual age (PCA)
- By day of life 5, MAP = number of weeks PCA + 5

**Notes:** ETT - endotracheal tube; HR - heart rate; LMA - laryngeal mask airway; MAP - median arterial pressure; RR - respiratory rate.
9.6 List of commonly used medicines and dosage in ICU with ventilated patients (adults, children)

**INDUCTION AGENT**

**NEUROMUSCULAR BLOCKING AGENT**

**SEDATIVE/HYPNOTIC**

**CARDIOVASCULAR SUPPORT AGENT**

**ANALGESIC**

### Medications

<table>
<thead>
<tr>
<th>Medication</th>
<th>Dosage</th>
</tr>
</thead>
<tbody>
<tr>
<td>ACETAMINOPHEN</td>
<td>See Paracetamol</td>
</tr>
</tbody>
</table>
| ADENOSINE               | Adult: 6 mg IV push; then 12 mg IV q1min × 2 prn  
Peds: 0.1 mg/kg IV push (max 6 mg/dose), may repeat 0.2 mg/kg IV (max 12 mg/dose) |
| ADRENALINE (EPINEPHRINE)| Adult: Arrest: 1 mg q3-5min IV prn; ETT 2-2.5 mg q3-5min prn (dilute in 5-10 mL NS or sterile water)  
Anaphylaxis/hypotension: 0.05-0.1 mg IV q5min prn; 0.2-0.5 mg IM q5min prn  
Infusion: 0.5-20 mcg/min IV  
Racemic: 2.25% solut 0.5 mL via neb  
Peds: Arrest: 5 mcg/kg IV (max 1 mg) q3-5min prn; 100 mcg/kg ETT q3-5 min prn  
Anaphylaxis: Children > 6 mo < 30 kg: 5 mcg/kg IM; > 30 kg 300 mcg IM  
Severe Hypotension: 0.5-10 mcg/kg IV  
Infusion: 0.02-1 mcg/kg/min IV  
Racemic: 2.25% solut 0.25-0.5 mL via neb |
| ALBUTEROL               | Adult & Peds: (bronchodilation) Nebulized: 2.5 mg in 3 mL every 20 min or continuous (5-20 mg/hr)   |
| AMIODARONE              | Adult: 150-300 mg IV (dependent on rhythm) then 1 mg/min × 6hr, then 0.5 mg/min ×1hr  
Peds: 5 mg/kg IV (max 300 mg) over 30 min, may repeat × 2; Infusion: 5-15 mcg/kg/min IV |
| ATRACURIUM              | Adult & Peds: 0.4-0.5 mg/kg IV (t½ = ~20 min)                                                   |
| ATROPINE                | Adult: Arrest/bradycardia: 0.5 mg IV q3-5min max 3 mg; ETT 1-2 mg q3-5 min prn  
Peds: Arrest/brady: 0.02 mg/kg (max 0.5 mg) IV, repeat x 1 q5min prn; ETT 0.04-0.06 mg/kg; repeat × 1 prn |
| CALCIUM CHLORIDE        | Adult: Arrest, CCB toxicity: 1-2 gm IV slowly; repeat q10min prn  
Peds: Arrest, CCB toxicity: 20 mg/kg IV (max 2 g); repeat q10min prn |
| CARBOPROST (HEMABATE)   | Adult: 250 mcg IM, repeat q15min prn. Max 2 mg (see PPH for full details)                      |
| CISATRACURIUM           | Adult: 0.1-0.2 mg/kg IV (t½ = ~25 min); Infusion 0.5-10 mcg/kg/min IV  
Peds: 0.1-0.15 mg/kg IV; Infusion 0.5-4 mcg/kg/min IV |
| CODEINE                 | Adult: 15-60 mg PO/IM/SQ; repeat q4h prn  
Peds: not recommended in children < 12 yr |
| DANTROLENE             | Adult & Peds: 2.5 mg/kg IV, repeat 1 mg/kg prn (max of 10 mg/kg)                                |
| DEXMEDETOMIDINE         | Adult & Peds: Load: 0.5-1 mcg/kg IV (over 10 min), Infusion: 0.2-1.5 mcg/kg/hr IV               |
| DEXAMETHASONE           | Adult & Peds: Airway oedema: 0.5 mg/kg IV q6hr  
PONV: Adults 4-8 mg IV; Peds 0.1 mg/kg IV |
| DIAZEPAM                | Adult: 5-10 mg IV  
Peds: 0.2-0.3 mg/kg IV |

9. Acute respiratory distress syndrome (ARDS) 183
<table>
<thead>
<tr>
<th>Drug</th>
<th>Adult Dose</th>
<th>Pediatric Dose</th>
<th>Notes</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>DICLOFENAC</strong></td>
<td>Adult: 50-100 mg PO</td>
<td>Peds: 0.5 mg/kg IV/IM; 1 mg/kg PO/PR</td>
<td></td>
</tr>
<tr>
<td><strong>DIPHENHYDRAMINE</strong></td>
<td>Adult: 25-50 mg IV/IM/PO q4-6hr</td>
<td>Peds: 0.5-1 mg/kg IV q4-6 hours; max 50 mg</td>
<td></td>
</tr>
<tr>
<td><strong>DOBUTAMINE</strong></td>
<td>Adult &amp; Peds: 0.5-20 mcg/kg/min IV infusion</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>DOPAMINE</strong></td>
<td>Adult &amp; Peds: 0.5-20 mcg/kg/min IV infusion</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>EPINEPHRINE</strong></td>
<td>See Adrenaline</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>EPHEDRINE</strong></td>
<td>Adult: 5-10 mg IV prn</td>
<td>Peds: 0.1-0.2 mg/kg (max 25 g/dose) IV prn</td>
<td></td>
</tr>
<tr>
<td><strong>ERGOMETRINE</strong></td>
<td>Adult: 0.5 mg IV/IM slow</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>ESMOLOL</strong></td>
<td>Adult &amp; Peds: Bolus: 0.5 mg/kg IV prn; Infusion: 50-300 mcg/kg/min IV</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>ETOMIDATE</strong></td>
<td>Adult &amp; Peds: 0.2-0.3 mg/kg IV</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>FENTANYL</strong></td>
<td>Adult: Analgesia: 25-100 mcg IV prn; Infusion 25-200 mcg/hr (or higher)</td>
<td>Peds: Analgesia: 0.5-1 mcg/kg IV prn; 1-2 mcg/kg intranasal prn; Infusion: 0.5-5 mcg/kg/hr IV</td>
<td></td>
</tr>
<tr>
<td><strong>GLYCOPHYRROLATE</strong></td>
<td>Adult: Reversal: 0.1-0.2 mg IV</td>
<td>Peds: Reversal: 0.015 mg/kg IV; Antisialagogue: 4 mcg/kg IM</td>
<td></td>
</tr>
<tr>
<td><strong>HYDRALAZINE</strong></td>
<td>Adult: 10-20 mg IV</td>
<td>Peds: 0.1-0.2 mg/kg IV</td>
<td></td>
</tr>
<tr>
<td><strong>HYDROCODONE</strong></td>
<td>Adult: 20-40 mg PO</td>
<td>Peds: 0.2 mg/kg PO</td>
<td></td>
</tr>
<tr>
<td><strong>HYDROCORTISONE</strong></td>
<td>Adult: 100 mg IV; stress dose 50 mg IV q6hr</td>
<td>Peds: (stress dose) 1-2 mg/kg IV</td>
<td></td>
</tr>
<tr>
<td><strong>HYDROMORPHONE</strong></td>
<td>Adult: 0.5-2 mg IV prn</td>
<td>Peds: IV: 5-10 mcg/kg IV prn</td>
<td>PO/PR: 50-80 mcg/kg q3-6h prn</td>
</tr>
<tr>
<td><strong>INTRALIPID</strong></td>
<td>Adult &amp; Peds: LAST: 1.5 mL/kg followed by infusion 0.25 mL/kg/min up to 0.5 mL/kg/min ; use ideal body weight; 12 mL/kg in ped</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>KETAMINE</strong></td>
<td>Adult: Induction: 0.5-2 mg/kg IV, 4-10 mg/kg IM; Analgesia: 0.2-0.8 mg/kg IV, 2-4 mg/kg IM; Infusion 2-15 mcg/kg/min IV</td>
<td>Peds: Induction: 1-3 mg/kg IV, 5-8 mg/kg IM, 5-10 mg/kg PR; Analgesia: 0.2-0.5 mg/kg IV, 2-4 mg/kg IM; Infusion: 2-10 mcg/kg/min IV</td>
<td></td>
</tr>
<tr>
<td><strong>KETOROLAC</strong></td>
<td>Adult: 30-60 mg IV/IM, then 15-30 mg IV/IM q6h prn</td>
<td>Peds: 0.5 mg/kg (max 30 mg) IV q6h prn; 1 mg/kg IM</td>
<td></td>
</tr>
<tr>
<td><strong>LABETALOL</strong></td>
<td>Adult: 10-20 mg IV, double dose q15min prn to max 300 mg; Infusion 0.5-2 mg/min (or higher)</td>
<td>Peds: 0.1 mg/kg IV q5-10min</td>
<td></td>
</tr>
<tr>
<td><strong>LIDOCAINE</strong></td>
<td>Adult: Arrest: 1-1.5 mg/kg IV, 0.5-0.75 mg/kg q5-10min prn (max 3 mg/kg), ETT 2-3.75 mg/kg, infusion 1-4 mg/min; analgesia: 1-2 mg/kg IV, infusion: 0.5-3 mg/kg/hr IV</td>
<td>Peds: Arrest: 1 mg/kg IV, repeat x1 prn, ETT 2.3 mg/kg infusion 20-50 mcg/kg/min IV; Analgesia: 1 mg/kg IV; Infusion: 1.5-2 mg/kg/hr IV</td>
<td></td>
</tr>
<tr>
<td><strong>LORAZEPAM</strong></td>
<td>Adult: 1-4 mg IV prn</td>
<td>Peds: 0.1 mg/kg IV prn (max 4 mg/dose)</td>
<td></td>
</tr>
<tr>
<td>Medication</td>
<td>Description</td>
<td></td>
<td></td>
</tr>
<tr>
<td>----------------------------</td>
<td>-----------------------------------------------------------------------------</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>
| **MAGNESIUM SULFATE**      | Adult: Asthma: 2 gm IV over 20 min; eclampsia/pre-eclampsia: load 4-6 gm IV, infusion 1-2 gm/hr IV; torsade de pointes: 1-2 gm IV, infusion 0.5-1 gm/hr IV  
  Peds: Asthma: 25-75 mg/kg (max 2 gm) IV over 20 min; torsade de pointes: 25-50 mg/kg/dose (max 2 gm) IV |
| **MEPERIDINE**             | See Pethidine                                                                |
| **METARAMINOL**            | Adult & Peds: 0.5 mg IV bolus, repeat q2-3min prn (avoid in children < 12) |
| **METHADONE**              | Adult: Analgesia: 2.5-10 mg PO/IM/IV/SQ (based on opioid tolerance), repeat q8-12h prn  
  Peds: Analgesia: 0.05-0.1 mg/kg PO/IM/IV/SQ; (t½ = 18-24 hrs) |
| **METHOHEXITAL**           | Adult: Induction: 1-1.5 mg/kg IV  
  Peds: Induction: 1-3 mg/kg IV, 20-30 mg/kg PR |
| **METHYLERGONOVINE/METHERGINE** | Adult: 0.2 mg IM; repeat q5-10min max 2 doses |
| **METHYLPRIDNISOLONE**     | Adult: Asthma: 40-80 mg IV; anaphylaxis: 125 mg IV  
  Peds: Asthma: 1 mg/kg IV; anaphylaxis: 1-2 mg/kg IV |
| **METOCLOPRAMIDE**         | Adult: 10-20 mg IV/PO, repeat 5-10 mg q6h prn  
  Peds: 0.1-0.15 mg/kg IV/PO q6h prn |
| **MIDAZOLAM**              | Adult: 0.5-4 mg IV  
  Peds: 0.1-0.2 mg/kg IV, 0.5 mg/kg PO/PR |
| **MISOPROSTOL**            | Adult: 1 mg PR                                                                |
| **MORPHINE SULFATE**       | Adult: 2.5-10 mg IV/IM  
  Peds: 0.05-0.1 mg/kg IV/IM |
| **NALOXONE**               | Adult: Excessive sedation: 0.02-0.2 mg q4-8; opioid overdose: 0.1-2 mg IV/IM q2-3min prn, 2 mg nebulized, 4 mg intranasal  
  Peds: Excessive sedation: 0.5-1 mcg/kg IV q2-3min prn; opioid overdose: 10 mcg/kg IV/IM q2-3min prn; 4 mg intranasal |
| **NEOSTIGMINE**            | Adult & Peds: 0.03-0.07 mg/kg IV (max 5 mg)  
  Add atropine IV 0.5-1 mg (adults), 20 mcg/kg (peds) or glycopyrrolate (see “glycopyrrolate”) |
| **NITROGLYCERIN**          | Adult: Infusion: 10-200 mcg/min IV  
  Peds: 0.5-20 mcg/kg/min IV infusion IV |
| **NOREPINEPHRINE**         | Adult: Infusion: 0.05-2 mcg/kg/min or 0.5-20 mcg/min IV  
  Peds: Infusion: 0.05-2 mcg/kg/min IV |
| **ONDANSETRON**            | Adult: 4-8 mg IV, repeat q4-8h prn  
  Peds: 0.15 mg/kg IV; repeat q6-8h prn |
| **OXYCOCONE**              | Adult: 5-15 mg (or higher depending on opioid tolerance), repeat q3-4h prn  
  Peds: 0.1 mg/kg PO; repeat q3-4h prn |
| **OXYTOCIN**               | Adult: 3 U stat IV over 30 sec, consider repeat dosing and infusion |
| **PANCURONIUM**            | Adult: 0.04-0.1 mg/kg IV  
  Peds: 0.05-0.15 mg/kg IV (t½ = ~110 min) |
| **PARACETAMOL (ACETAMINOPHEN)** | Adult: 500-1000 mg IV/PO, repeat q4-6 prn (max 2-4 gm/day)  
  Peds: PO/IV: 10-15 mg/kg, repeat q6h prn, PR: 40 mg/kg × 1, max: 75 mg/kg/24 hr |
| **PETHIDINE (MEPERIDINE)** | Adult: Shivering/analgesia: 12.5-50 mg IV  
  Peds: 0.5-1 mg/kg IV, max 400 mg daily |
<p>| <strong>PHENOYBARBITAL/PHENOBARBITONE</strong> | Adult &amp; Peds: Status epilepticus: 15-20 mg/kg IV, may repeat 5-10 mg/kg in 10 min prn x 1 |</p>
<table>
<thead>
<tr>
<th>Medication</th>
<th>Adult Dose/Route</th>
<th>Pediatric Dose/Route</th>
<th>Notes/Indications</th>
</tr>
</thead>
<tbody>
<tr>
<td>PHENYLEPHRINE</td>
<td>40-100 mcg IV q1-2min prn; infusion 10-200 mcg/min</td>
<td></td>
<td></td>
</tr>
<tr>
<td>PITOCIN</td>
<td>See Oxytocin</td>
<td></td>
<td></td>
</tr>
<tr>
<td>PROCHLORPERAZINE</td>
<td>5-10 mg IV/IM/PO q3-6h prn (max 40 mg/day)</td>
<td>0.1-0.15 mg/kg PO/IM/IV q6-8h prn (max 10 mg/dose)</td>
<td></td>
</tr>
<tr>
<td>PROMETHAZINE</td>
<td>12.5-25 mg PO/PR q4-6h prn</td>
<td>0.2-0.5 mg/kg PO/PR q6-8h</td>
<td>Max 25 mg/dose (do not give if &lt; 2 yr)</td>
</tr>
<tr>
<td>PROPOFOL</td>
<td>Induction: Dose variable, Adults: 1-2.5 mg/kg; Children 2-4 mg/kg Infusion: 10-250 mcg/kg/min</td>
<td></td>
<td></td>
</tr>
<tr>
<td>RANITIDINE</td>
<td>50 mg IV; 150-300 mg PO</td>
<td>1 mg/kg IV; 2.5 mg/kg PO</td>
<td></td>
</tr>
<tr>
<td>REMIFENTANIL</td>
<td>Adult &amp; Peds: Bolus: 0.5-1 mcg/kg IV; Infusion: 0.05-0.5 mcg/kg/min IV</td>
<td></td>
<td></td>
</tr>
<tr>
<td>ROCURONIUM</td>
<td>0.6-1.2 mg/kg IV (t½ = ~60 min)</td>
<td>0.9-1.2 mg/kg IV</td>
<td></td>
</tr>
<tr>
<td>ROVACURONIUM</td>
<td>Adult &amp; Adolescents: 1 patch q72h</td>
<td>6 mcg/kg IV (max 0.3 mg)</td>
<td></td>
</tr>
<tr>
<td>SODIUM CITRATE (Bicitra)</td>
<td>15-30 mL PO q6h prn</td>
<td>1-1.5 mL/kg q6-8h prn (max 30 mL/dose)</td>
<td></td>
</tr>
<tr>
<td>SODIUM BICARBONATE</td>
<td>50-100 mL 8.4% IV prn</td>
<td>1-2 mEq/kg IV</td>
<td></td>
</tr>
<tr>
<td>SUCCINYLCHOLINE/SUXAMETHONIUM</td>
<td>(induction) 0.6-2 mg/kg IV (high end for RSI) IM: 3-4 mg/kg; max 5 mL at injection site (t½ = ~6-8 min)</td>
<td>1-2 mg/kg IV; 3-4 mg/kg IM</td>
<td></td>
</tr>
<tr>
<td>SUFENTANIL</td>
<td>Analgesia: 0.5-2 mcg/kg IV Infusion: 0.05-2 mcg/kg/hr</td>
<td></td>
<td></td>
</tr>
<tr>
<td>SUGAMMADEX</td>
<td>2 TOF twitches: 2 mg/kg; 0 TOF, 1-2 PTC: 4 mg/kg; Immediate emergent reversal: 16 mg/kg</td>
<td></td>
<td></td>
</tr>
<tr>
<td>TERBUTALINE</td>
<td>(tolcylisis) 5-10 mcg/kg IV q15min (max 250 mcg)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>THIOPENTAL/THIOPENTONE</td>
<td>(induction) 3-6 mg/kg</td>
<td></td>
<td></td>
</tr>
<tr>
<td>TRAMADOL</td>
<td>25-100 mg PO q4-6h prn</td>
<td>not recommended in children &lt; 12 yr</td>
<td></td>
</tr>
<tr>
<td>TRANEXAMIC ACID</td>
<td>1 g IV over 10 min, repeat × 1 after 30 min prn</td>
<td></td>
<td></td>
</tr>
<tr>
<td>VASOPRESSIN</td>
<td>(shock) 0.03-0.05 units/minute drip</td>
<td>Infusion: 0.0002-0.002 units/kg/min IV</td>
<td></td>
</tr>
<tr>
<td>VECURONIUM</td>
<td>(induction) 0.1 mg/kg IV (t½ = ~65 min)0.8-1.7 mcg/kg/min drip</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Notes: CCB - calcium channel blocker; ETT - endotracheal tube; IV - intravenous; IM - intramuscular; min - minute; PO - per os/oral; PONV - post-operative nausea and vomiting; PR - per rectal; prn - pro re nata; PTC - post-tetanic count; SQ - subcutaneous; TOF - train of four.
Source: Adapted from USAID-STAR-UCSF Project (2022).
9.7 Choice of induction agents in adults

<table>
<thead>
<tr>
<th>Choice of induction agents in adult patients</th>
<th>Intravenous dose</th>
<th>Notes</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Opiates</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Fentanyl</td>
<td>50-250 mcg</td>
<td>Can cause ↓ blood pressure</td>
</tr>
<tr>
<td></td>
<td></td>
<td>May cause chest stiffness</td>
</tr>
<tr>
<td>Morphine</td>
<td>2.5-10 mg</td>
<td>Takes long time to be effective - 10 mins</td>
</tr>
<tr>
<td><strong>Sedative/hypnotic</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Midazolam</td>
<td>0.5-4 mg</td>
<td>Can cause delirium</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Accumulates in liver failure</td>
</tr>
<tr>
<td>Lorazepam</td>
<td>1-4 mg</td>
<td>Can cause delirium</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Accumulates in liver failure</td>
</tr>
<tr>
<td>Diazepam</td>
<td>5-10 mg</td>
<td>Can cause delirium</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Accumulates in renal failure</td>
</tr>
<tr>
<td><strong>Induction agent</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Ketamine</td>
<td>0.5-2 mg/kg</td>
<td>Can cause ↑ intracranial pressure</td>
</tr>
<tr>
<td>Etomidate</td>
<td>0.2-0.3 mg/kg</td>
<td>Can cause adrenal suppression with multiple doses</td>
</tr>
<tr>
<td>Propofol 1%</td>
<td>1-2.5 mg/kg*</td>
<td>Can cause ↓ blood pressure</td>
</tr>
<tr>
<td>Thiopental</td>
<td>3-6 mg/kg</td>
<td></td>
</tr>
<tr>
<td><strong>Neuromuscular blockers</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Suxamethonium</td>
<td>0.6-2 mg/kg</td>
<td>Avoid if hyperkalaemia, neuromuscular patients, recent burn or renal failure</td>
</tr>
<tr>
<td>Rocuronium</td>
<td>0.6-1.2 mg/kg</td>
<td>First-line RSI paralytic</td>
</tr>
<tr>
<td>Vecuronium</td>
<td>0.1 mg/kg</td>
<td></td>
</tr>
<tr>
<td>Atracurium</td>
<td>0.4-0.5 mg/kg</td>
<td></td>
</tr>
<tr>
<td>Pancuronium</td>
<td>0.04-0.1 mg/kg</td>
<td>Prolonged block if renal failure</td>
</tr>
</tbody>
</table>

* Consider dose reduction for critically ill patients or selecting alternate agents if haemodynamically unstable.

Note:
### 9.8 Choice of induction agents in children

**Choice of induction agents in paediatric patients**

<table>
<thead>
<tr>
<th>Category</th>
<th>Agent</th>
<th>Intravenous dose</th>
<th>Notes</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Opiates</strong></td>
<td>Atropine</td>
<td>20 mcg/kg (min. dose 100 mcg); &gt; 12 years 300-600 mcg</td>
<td></td>
</tr>
<tr>
<td></td>
<td>Fentanyl</td>
<td>2.5 mcg/kg</td>
<td>Can cause ↓ blood pressure</td>
</tr>
<tr>
<td></td>
<td>Morphine</td>
<td>0.1-0.2 mg/kg</td>
<td>Relatively prolonged onset: ~10 mins</td>
</tr>
<tr>
<td><strong>Sedative/hypnotic</strong></td>
<td>Midazolam</td>
<td>0.1-0.2 mg/kg</td>
<td>Can cause delirium</td>
</tr>
<tr>
<td></td>
<td></td>
<td></td>
<td>Accumulates in liver failure</td>
</tr>
<tr>
<td></td>
<td>Lorazepam</td>
<td>0.1 mg/kg</td>
<td>Can cause delirium</td>
</tr>
<tr>
<td></td>
<td></td>
<td></td>
<td>Accumulates in liver failure</td>
</tr>
<tr>
<td></td>
<td>Diazepam</td>
<td>0.2-0.3 mg/kg</td>
<td>Can cause delirium</td>
</tr>
<tr>
<td></td>
<td></td>
<td></td>
<td>Accumulates in renal failure</td>
</tr>
<tr>
<td><strong>Induction agent</strong></td>
<td>Ketamine</td>
<td>1-3 mg/kg</td>
<td>Can cause ↑ intracranial pressure</td>
</tr>
<tr>
<td></td>
<td>Etomidate</td>
<td>0.2-0.3 mg/kg</td>
<td>Can cause adrenal suppression with multiple doses</td>
</tr>
<tr>
<td></td>
<td>Propofol 1% (induction only)</td>
<td>2-4 mg/kg (3 years)</td>
<td>Can cause ↓ blood pressure</td>
</tr>
<tr>
<td><strong>Neuromuscular blockers</strong></td>
<td>Suxamethonium</td>
<td>3 mg/kg/dose (neonate); 1-2 mg/kg all other ages</td>
<td>Avoid if hyperkalaemia, neuromuscular patients, recent burn or renal failure</td>
</tr>
<tr>
<td></td>
<td>Rocuronium</td>
<td>0.9-1.2 mg/kg</td>
<td>First-line RSI paralytic</td>
</tr>
<tr>
<td></td>
<td>Vecuronium</td>
<td>0.1 mg/kg</td>
<td></td>
</tr>
<tr>
<td></td>
<td>Atracurium</td>
<td>0.4-0.5 mg/kg</td>
<td></td>
</tr>
<tr>
<td></td>
<td>Pancuronium</td>
<td>0.05-0.15 mg/kg</td>
<td>Vastly prolonged block in renal failure</td>
</tr>
</tbody>
</table>

Note:
- *Consider dose reduction for critically ill patients or selecting alternate agents if haemodynamically unstable.*
9.9 Protocol to deliver lung protective ventilation (LPV)

This protocol to deliver lung protective ventilation (LPV) was used in the low tidal volume (TV) trial published in 2000 (ARDS Network et al., 2000).

**Ventilator set up and adjustment**

- Calculate predicted body weight (PBW):
  - **Males** = 50 + 1.1 [height (cm) - 152]
  - **Females** = 45.5 + 1.1 [height (cm) - 152].
- Select any ventilator mode.
- Set ventilator settings to achieve initial tidal volume (TV) = 6 mL/kg PBW (range 4-8 mL/kg).
- Reduce TV by 1 mL/kg at intervals ≤ 2 hrs until achieve pressure goals.
- Set initial rate to approximate baseline minute ventilation (MV) (not > 35 breaths/min in adults).
  \[ MV = \text{Tidal volume (TV)} \times \text{Respiratory rate (RR)} \]
- Set I:E ratio (ratio of the duration of inspiration and expiration phases):
  - set high inspiratory flow rates
  - monitor for intrinsic PEEP
  - in severe ARDS I:E ratios of 1:1 or inverse ratios may be needed.
- Set inspiratory flow rate above patient demand (commonly > 60 L/min).
- Set FiO₂ at 1.0 and titrate down.
- Set PEEP 5-10 cmH₂O or higher for severe ARDS.
- Adjust TV and RR to achieve pH and Pplat goals below.

**Note:** Principles are the same for children except that children younger than 8 years require a lower maximum PEEP - 15 cmH₂O and the peak Pplat should be < 28 cmH₂O.
9.9.1 ARDS–net PEEP FiO₂ grid to guide PEEP

See (4).

Set PEEP corresponding to severity of oxygen impairment:

**Oxygenation goal:** PaO₂ 55-80 mmHg or SpO₂ 88-95%

- Use a minimum PEEP of 5 cmH₂O
- Consider incremental PEEP/FiO₂ combinations such as shown below to achieve goal
- PEEP levels > 15 should not be used in children < 8 years

There are two PEEP/FiO₂ grids - the second one can be used for more severe hypoxaemia.

<table>
<thead>
<tr>
<th>Lower PEEP/higher FiO₂</th>
<th>FiO₂</th>
<th>0.3</th>
<th>0.4</th>
<th>0.4</th>
<th>0.5</th>
<th>0.6</th>
<th>0.7</th>
<th>0.7</th>
<th>0.8</th>
<th>0.9</th>
<th>0.9</th>
<th>0.9</th>
<th>1.0</th>
</tr>
</thead>
<tbody>
<tr>
<td>PEEP</td>
<td>5</td>
<td>5</td>
<td>8</td>
<td>8</td>
<td>10</td>
<td>10</td>
<td>12</td>
<td>14</td>
<td>14</td>
<td>14</td>
<td>16</td>
<td>16</td>
<td>16</td>
</tr>
<tr>
<td>Higher PEEP/lower FiO₂ for more severe hypoxaemia</td>
<td>FiO₂</td>
<td>0.3</td>
<td>0.3</td>
<td>0.3</td>
<td>0.3</td>
<td>0.3</td>
<td>0.4</td>
<td>0.4</td>
<td>0.5</td>
<td>0.5</td>
<td>0.5-0.8</td>
<td>0.8</td>
<td>0.9</td>
</tr>
<tr>
<td>PEEP</td>
<td>5</td>
<td>8</td>
<td>10</td>
<td>12</td>
<td>14</td>
<td>14</td>
<td>16</td>
<td>16</td>
<td>18</td>
<td>20</td>
<td>22</td>
<td>22</td>
<td>22</td>
</tr>
</tbody>
</table>

9.9.2 Goals of Pplat and pH in lung protective ventilation

**Pplat goal:** ≤ 30 cmH₂O

- Check Pplat using 0.5 second inspiratory pause, at least every 4 hours and after each change in PEEP or TV
  - If Pplat > 30 cmH₂O or > 28 cmH₂O in children: decrease TV by 1 mL/kg steps (minimum = 4 mL/kg).
  - If Pplat < 25 cmH₂O and TV < 6 mL/kg: increase TV by 1 mL/kg until Pplat > 25 cmH₂O or TV = 6 mL/kg.
  - If Pplat < 30 cmH₂O and breath stacking or asynchrony occurs: may increase TV in 1 mL/kg increments to 7-8 mL/kg if Pplat remains ≤ 30 cmH₂O.

**pH goal:** 7.30-7.45

- Acidosis management: (pH < 7.30)
  - If pH 7.15-7.30: increase RR until pH > 7.30 or PaCO₂ < 25 (maximum set RR = 35).
  - If pH < 7.15: increase RR to 35.
  - If pH remains < 7.15, TV may be increased in 1 mL/kg steps until pH > 7.15 (Pplat target of 30 may be exceeded).
  - May give NaHCO₃ to act as a transient buffer.

- Alkalosis management: pH > 7.45
  - Decrease ventilator rate if possible.

- Inspiration to expiration ratio goal
  - Recommend that duration of inspiration be less than duration of expiration.

**Notes:** Paw - airway pressure; PEEP - positive end-expiratory pressure; Pplat - plateau airway pressure.
9.10 Memory aid: comparison of normal waveforms during volume and pressure-limited ventilation

<table>
<thead>
<tr>
<th>Volume control</th>
<th>Pressure control</th>
</tr>
</thead>
<tbody>
<tr>
<td><img src="image1" alt="Volume control waveform diagram" /></td>
<td><img src="image2" alt="Pressure control waveform diagram" /></td>
</tr>
</tbody>
</table>

- Volume cycled ventilation
- Flow is fixed (shown) or decelerating
- Volume is set and the airway pressures are variable
- Inspiration cycles to expiration after fixed time or volume delivered
- To measure Pplat you need to perform an inspiratory pause, which means briefly stopping gas flow at the end of pressure + PEEP inspiration
- Patient can trigger a breath

- Pressure limited ventilation
- Flow is variable
- Pressure is set (see square wave form) and the volumes are variable
- Inspiration cycles to expiration after fixed time
- Pplat is the set inspiratory pressure (Pinsp), equal to driving pressure + PEEP
- Patient can trigger a breath

Notes: Paw - airway pressure; PEEP - positive end-expiratory pressure; Pplat - plateau airway pressure.
9.11 Memory aid: recognizing and interpreting abnormal pressure and flow waveforms during volume control ventilation

### Pressure curves

<table>
<thead>
<tr>
<th>Pressure curves</th>
<th>Characteristics</th>
<th>Interpretation</th>
</tr>
</thead>
<tbody>
<tr>
<td>60 cm H₂O</td>
<td>Normal pressure curve</td>
<td>Normal</td>
</tr>
<tr>
<td>90 cm H₂O</td>
<td>Increased peak airway pressure</td>
<td>Reduced compliance</td>
</tr>
<tr>
<td>60 cm H₂O</td>
<td>Increased peak airway pressure</td>
<td>Increased resistance</td>
</tr>
</tbody>
</table>

### Flow curves

<table>
<thead>
<tr>
<th>Flow curves</th>
<th>Characteristics</th>
<th>Interpretation</th>
</tr>
</thead>
<tbody>
<tr>
<td>2 L/s</td>
<td>Normal flow pattern</td>
<td>Normal</td>
</tr>
<tr>
<td>3 L/s</td>
<td>High expiratory peak flow rate expiratory flow is shorter</td>
<td>Reduced compliance</td>
</tr>
<tr>
<td>2 L/s</td>
<td>Prolonged expiratory flow</td>
<td>Increased resistance</td>
</tr>
</tbody>
</table>

9.12 Guide to distinguishing between the causes of high peak airway pressures: resistance versus compliance

<table>
<thead>
<tr>
<th>Abnormal airway pressure(s)</th>
<th>High peak with high plateau airway pressure</th>
<th>High peak with normal plateau airway pressure</th>
</tr>
</thead>
<tbody>
<tr>
<td>Main physiologic problem</td>
<td>Reduced respiratory system compliance (Crs)</td>
<td>High resistance (R)</td>
</tr>
<tr>
<td>Formula</td>
<td>[ C_{rs} = \frac{\text{Tidal volume}}{\text{P\textsubscript{plat} - P\textsubscript{PEEP}}} ]</td>
<td>[ R = \frac{P\text{peak} - P\text{plat}}{\text{Flow}} ]</td>
</tr>
<tr>
<td>Normal</td>
<td>60-100 mL/cm H\textsubscript{2}O</td>
<td>5-10 cm H\textsubscript{2}O/L/sec for intubated adult</td>
</tr>
</tbody>
</table>
| Problems that can be treated quickly | • mainstem bronchus intubation  
• tension pneumothorax  
• pleural effusion  
• abdominal distension  
• congestive heart failure  
• atelectasis  
• hyperinflation | Patient problems:  
• patient biting, coughing, fighting ventilator  
• secretions  
• bronchospasm  
Ventilator problems:  
• tube kinked  
• circuit filled with water  
• small endotracheal tube |
| Other problems that may improve over the time | • ARDS  
• consolidation  
• fibrosis  
• chest wall oedema  
• thoracic deformity | • asthma  
• chronic obstructive pulmonary disease (COPD) |

Factors influencing peak airway pressure

\[ P\text{airway} = P\text{resistance} + P\text{compliance} \]

<table>
<thead>
<tr>
<th>Airflow resistance</th>
<th>Respiratory system compliance</th>
<th>Chest wall compliance</th>
</tr>
</thead>
<tbody>
<tr>
<td>• size of airway</td>
<td>• chest wall</td>
<td>• chest wall</td>
</tr>
<tr>
<td>• lower airway obstruction</td>
<td>• tidal volume</td>
<td>• patient position</td>
</tr>
<tr>
<td>• mechanical obstruction</td>
<td>• lung elasticity</td>
<td>• external compression of chest from abdomen</td>
</tr>
</tbody>
</table>
9.13 Troubleshooting high peak airway pressures, low tidal volumes, desaturation or haemodynamic instability in ventilated patients

☐ Is the endotracheal tube in the trachea?
  • Large cuff leak or no chest rise with inspiration suggest that ETT is dislodged: assess with direct laryngoscopy and re-intubate.

☐ Is there a problem with the ventilator circuit or oxygen supply?
  • Take the patient off the ventilator and hand ventilate with 100% oxygen while checking equipment.

☐ Can you pass a suction catheter through the endotracheal tube?
  • If no, ETT may be kinked: straighten or insert bite block to prevent patient from biting.
  • If no, ETT may be blocked with secretions: reintubate with new ETT.
  • If yes, suction ETT to remove sputum/mucus plugs.

☐ Are there breath sounds bilaterally?
  • Unilaterally absent breath sounds: evaluate for mainstem intubation/lobar collapse versus pneumothorax by assessing mediastinal shift and by chest X-ray or ultrasound if patient not in extremis.
  • Suspicion of tension pneumothorax mandates immediate needle decompression followed by chest tube placement, without a chest X-ray.
  • Mainstem intubation may be suspected clinically if ETT further in patient than previously. Withdraw to previous position; can confirm with bronchoscopy if available.
  • Lobar collapse or atelectasis may respond to aggressive suctioning and can be confirmed with chest X-ray.
  • Bilateral wheezing: consider bronchospasm; give bronchodilators.
  • Bilateral crackles: consider pulmonary oedema; give diuretic or more PEEP depending on full clinical evaluation of volume status.

☐ Are there other problems causing low compliance?
  • Abdominal distension: drain stomach with NG tube.
  • Auto-PEEP: diagnose by examining ventilator waveforms. Treat with bronchodilators, sedation; may require temporary disconnection from positive pressure.

☐ Is there haemodynamic instability?
  • Restore haemodynamic stability with fluid or vasopressors while determining and treating primary cause.
  • If severe hypotension, evaluate for tension pneumothorax or severe auto-PEEP (often in patients with asthma or COPD).
  • Other causes include high airway pressures reducing venous return, vasodilation due to sedative and analgesic medications or a new problem (sepsis, bleeding, pulmonary embolism, myocardial infarction).

☐ Is the patient agitated and asynchronous with the ventilator?
  • May be secondary to any other problem or may be primary problem and causing asynchrony: treat cautiously with sedation.
9.14 Respiratory care pocket card reference

The following respiratory care pocket card reference is intended to serve as a tool to learn about the fundamentals of oxygen delivery, including mechanical ventilation, ARDS management and ventilator liberation.

The card, which can be printed or saved to your mobile device, is available in:
- English
- French
- Portuguese
- Spanish.

Source: USAID-STAR-UCSF Open Critical Care Project
### Oxygen Sources & Delivery Devices

<table>
<thead>
<tr>
<th>Device Type</th>
<th>Pros</th>
<th>Cons</th>
</tr>
</thead>
<tbody>
<tr>
<td>Nasal Cannula (NC)</td>
<td>Pros: Ubiquitous; commonly used up to 6LPM</td>
<td>Cons: Requires humidification if &gt;4LPM (risk of epistaxis); no work of breathing support</td>
</tr>
<tr>
<td></td>
<td>O₂ works with any pressure source via flow meter; FI O₂ increases 2-4% per LPM; variable FI O₂ delivery based on patient’s minute ventilation &amp; flow rate</td>
<td></td>
</tr>
<tr>
<td>Non-Rebreather/ Face mask (NRB/FM)</td>
<td>Pros: High FI O₂;</td>
<td>Cons: Limited FI O₂ if high respiratory drive; no work of breathing support</td>
</tr>
<tr>
<td></td>
<td>O₂ works with any pressure source via flow meter; simple FM 5-10 LPM (~FI O₂ 35-50%); NRB 10-15 LPM (~FI O₂ 60-95%); enough flow to prevent bag collapse</td>
<td></td>
</tr>
<tr>
<td>High Flow Nasal Cannula (HFNC)</td>
<td>Pros: High FI O₂, even with high minute ventilation; can titrate flow and FI O₂; heated and humidified for comfort; may improve outcomes in acute hypoxic respiratory failure compared to NIPPV or low-flow O₂; small amount positive pressure may help with recruitment; high flow = deadspace washout, may help with work of breathing</td>
<td></td>
</tr>
<tr>
<td></td>
<td>Cons: Requires special device; consumes massive amounts of oxygen</td>
<td></td>
</tr>
<tr>
<td></td>
<td>O₂ requires high pressure/flow source; ~&gt;90% FI O₂ (variable with minute ventilation, entraining room air around cannula) 3 types: 1) With blender to mix compressed air + O₂; 2) With port/Venturi effect to entrain room air and mix with compressed O₂; or 3) Without blender.</td>
<td></td>
</tr>
<tr>
<td></td>
<td><strong>Initial Settings:</strong> infant &lt;1year = 8LPM; child 1-4 years = 10LPM; Child &gt; 4 years = 20LPM; adolescents/adults = 40LPM flow and 100% FI O₂; can titrate flow and/or FI O₂ (max flow depends on cannula size; up to 60 LPM for adults and 100% FI O₂) if tolerated and O₂ source adequate.</td>
<td></td>
</tr>
<tr>
<td>Non-invasive Ventilation (NIV) or Positive Pressure Ventilation (NIPPV)</td>
<td>Pros: May avoid intubation in some patients (COPD, cardiogenic pulmonary edema, upper airway obstruction) by decreasing work of breathing and adding PEEP</td>
<td></td>
</tr>
<tr>
<td></td>
<td>Cons: Risk of infectious aerosol generation (possibly less if helmet NIPPV); risk of aspiration if patient not alert / unable to protect airway or if inspiratory pressures &gt;20cm H₂O; pt must be alert enough to remove mask if uncomfortable; skin breakdown with prolonged use; confusing terminology: IPAP (inspiratory pressure) = P S + PEEP, EPAP (expiratory pressure) = PEEP; P S of “5 over 5” is the same as P S 5 over 5, is the same as IPAP 10/EPAP 5</td>
<td></td>
</tr>
<tr>
<td></td>
<td>O₂ requires high pressure/flow source to achieve high FI O₂</td>
<td></td>
</tr>
<tr>
<td></td>
<td><strong>Initial Settings:</strong> PS ΔS=8/PEEP (EPAP) 5-10; titrate ΔP up to 15 to reduce inspiratory work; use higher initial IPAP with obese patients; higher pressures may require sedation in pediatric patients</td>
<td></td>
</tr>
<tr>
<td>Continuous Positive Airway Pressure (CPAP)</td>
<td>Pros: Delivered via face mask or multiple other potential interfaces to split open the upper airway, increase lung volume &amp; intrathoracic pressure</td>
<td></td>
</tr>
<tr>
<td></td>
<td>Cons: Prolonged use is uncomfortable &amp; causes skin breakdown; limited unloading of inspiratory muscles or provide complete respiratory support</td>
<td></td>
</tr>
<tr>
<td></td>
<td>O₂ requires high flow/pressure source to achieve high FI O₂</td>
<td></td>
</tr>
<tr>
<td></td>
<td><strong>Initial Settings</strong> (adults/peds): CPAP or PEEP 5-10; adul. titrate as needed up to 15; peds ≤12; higher pressures may require sedation in peds</td>
<td></td>
</tr>
</tbody>
</table>

### Oxygen Supply Calculator

**Oxygen Delivery Device & Supply FAQ**

**Oxygen Supply & Demand Calculator**
Respiratory Mechanics

Positive End Expiratory Pressure (PEEP)
- Pressure within respiratory circuit at end of expiration
- Must be ≥5 cmH₂O in IMV to prevent derecruitment of alveoli
- This value is always set by ventilator operator

Pressure Peak Inspiratory (PIP)
- Reflects pressure generated by airway/ETT resistance and compliance
- Range 10-40 cmH₂O; target <40 cmH₂O

Pressure Plateau (Pplat)
- Reflects pressure in alveoli only
- If in volume control, perform inspiratory pause (when there is no flow, there is no effect of resistance; Pplat = Pressure at alveoli
- Target <30 cmH₂O (adults); <28 (peds); optimal ≤25 cmH₂O

Pressure Driving (Pdr)
- Pdr = Pplat - PEEP
- Tidal stress (lungs injury and mortality risk) if elevated
- Target ≤5 cmH₂O; mortality risk if ≥20 cmH₂O

I:E and Inspiratory Time (T_i)
- I:E = ratio of Inspiration to Expiration
- Normal 1:2 or 1:3, 1:1 is only tolerated when paralyzed (and rarely indicated), 1:4 or 1:5 may be better in asthma or COPD
- Normal Ti = 1-1.5s in non-ARDS; Consider Ti 0.7-1 for ARDS

Minute Ventilation (MV)
- MV = V_in × RR; where V_in is the tidal volume (i.e. volume of each breath) and RR is the respiratory rate (breaths per minute)
- Normal 4-6 LPM; lower if obtunded, hypothermic, deeply sedated;
- Higher 8-14 LPM in hypoxemic respiratory failure
- Adjust for pCO₂ goal (e.g. permissive hypercarbia if ARDS); -6-8 L/min in most intubated adults, may be ≥10-15 L/min in ARDS

Peak Flow
- Highest flow delivered by ventilator during inspiration
- 40-60 LPM common; -50-80 LPM if patient triggered mode
- Sometimes increasing flow can improve patient-ventilator synchrony; caution this may cause elevation in PIP

Compliance (C)
- C = ΔV / ΔP = Tidal volume of breath / Pdr
- Dynamic compliance (VT/PIP-PEEP) or static compliance (VT/Pplat-PEEP) measured at end inspiratory pause
- Range is 60-80 mL/cmH₂O in intubated patients; ARDS ≤40

Inspiratory Resistance (R)
- R = PIP - Pplat/inspiratory flow
- Must be measured during constant flow
- Normal ≤10 cmH₂O/L/sec; concern if ≥15 cmH₂O/L/sec

Choosing a Ventilator Mode
- Assist Control (AC) Volume Mode is default for non-spontaneous breathing patients or ARDS
- AC Pressure Mode & Dual Modes can be used for non-spontaneous breathing patients or ARDS
- PSV if spontaneous breathing and non-ARDS; SIMV and APRV have no data to support regular use
Volume Control

Other Names
AC-VC; Assist Control Volume Control; VCV; -CMV (controlled mandatory ventilation = all modes with RR and fixed T;); (S)CMV

Controlled Variables
RR, Vt, PEEP, FIO2, Trigger level, Flow pattern, I:E (either directly or via peak flow, T, settings)

Initial Settings
Adult & Pediatric
1. Set Vt at 6-8 mL/kg predicted body weight (PBW)
2. Set RR: Adults: set at pt’s most recent RR (do not exceed 35);
   Peds: set at most recent RR (do not exceed 60 bpm in infants, 40 bpm in toddlers/preschoolers, 35 bpm in school-aged children or adolescents)
3. Set T1: Adults 0.70-1 sec; Peds based on RR to maintain a minimum ratio of 1:2
4. Select FIO2 and PEEP (use ARDSnet grid if applicable; see next page)

Flow
Square wave/constant/fixed; or Variable/decreasing ramp (potentially more physiologic); 40LPM healthy, 60LPM ARDS

I:E
- I:E of 1:2 or 1:3 is best for most patients; Normal Ti ~ 1-1.5s in non ARDS patients; Consider Ti 0.7-1 for ARDS
- I:E of 1:1 or >1:1 associated with PEEP, decreased cardiac output (CO) and oxygen delivery
- Process for setting I:E may vary by ventilator make; commonly by changing Ti, inspiratory flow and flow pattern

Pros
Guaranteed MV regardless of changing respiratory system mechanics; precise control of Vt to limit volutrauma

Cons
Will overcome high resistance or compliance to deliver set V (must set pressure limit & alarm); breath stacking (i.e. next breath delivered before exhalation of prior breath); fixed flow and T can increase asynchrony when Vt & flow demand > vent settings

Breath Initiation
Control: Time trigger (60s/set RR): fixed VE
Assist: Pt effort triggers full breath at set T1, Vt, and flow rate

If No Patient Trigger
Delivers full set Vt, at set rate (i.e. guaranteed VE)

Breath Termination
Time cycled = breath ends at set T1; alarms if Vt not achieved; flow is set, breath ends once Vt delivered
Pressure cycled = safety mechanism; breath termination by clinician set high-pressure limit (10-15cmH2O>avg PIP); “pop-off”

Notes
- Inspiratory pause (~0.3s) can be built into each breath, will increase mean airway pressure; can measure Pplat
- Alarms: high pressure 5-10 > PIP, VE 50% above/below actual
- Trigger: 2-5 Lpm for flow; -2 cmH2O for pressure

Decelerating Flow

Constant Flow
### Pressure Control

<table>
<thead>
<tr>
<th>Other Names</th>
<th>AC-PC; Assist Control Pressure Control; -CMV - PC</th>
</tr>
</thead>
<tbody>
<tr>
<td>Controlled Variables</td>
<td>RR, Pinsp (or PC level), PEEP, FiO2, Flow trigger, Rise time, I:E (set directly or by Inspiratory time, T_i)</td>
</tr>
</tbody>
</table>

1. Set inspiratory pressure (Pinsp) at 8-20cmH2O, or set equal to previous Pdr, Pplit or -1/2 of PIP if transitioning from VC (goal 6-8 m:/ Kg PBW)
2. Set RR: Adults: set at pt’s most recent RR (do not exceed 35); Peds: set at most recent RR (do not exceed 60 bpm in infants, 40 bpm in toddlers/preschoolers, 35 bpm in school-aged children or adolescents)
3. Set T_i: Adults 0.70-0.85 sec; Peds based on RR to maintain a minimum ratio of 1:2
4. Select FiO2 & PEEP (use ARDSnet grid if applicable; see next page
   - Variable/decreasing ramp (potentially more physiologic)
   - Peak Flow determined by: 1) Pinsp level, 2) R_i, 3) T_i (shorter = more flow), 4) Pressure rise time (↑ Rise time → ↑ Peak flow), 5. Pt effort (↑ Effort → ↑ Peak flow)
5. Set I:E of 1:2 or 1:3 is best for most patients; Ti 0.7-1s for ARDS
   - I:E 1:1 or >1:1 associated with PEEPi, decreased CO & O2 delivery
   - Determined by set T_i and RR (Volume and flow variable)

| Pros | Avoids high PIPs
- Variable flow (↑ pt effort causes ↑flow to maintain constant airway pressure = potentially better synchrony; ↑ pt effort → ↑ flow and ↓ V_t)
- "Automated/active expiratory valves" (transiently opens expiratory valve to vent off pressure with coughing, asynchrony); ↑ comfort and ↓ barotrauma risk |

| Cons | V_t and MV not guaranteed; V_t determined by C and R (might be bigger or smaller than is optimal) |

| Breath Initiation | Control: Time trigger - (60s/set RR)
Assist: Pt trigger delivers Pinsp for inspiratory time cycle |

| If No Patient Trigger | Delivers Pinsp at set rate and T_i |

| Breath Termination | Time cycled = I:E or T_i set, breath ends at set time |

| Notes | - Pplat is the set inspiratory pressure
- Alarms: high pressure 5-10 > PIP, VE 50% above/below actual
- Trigger: 2-5 Lpm for flow; -2 cmH2O for pressure
- Unlike in VC, in PC the ventilator cannot compensate for volume lost to circuit compliance (i.e. V_t delivered may be less than V_t measured and my be significant especially in pediatrics) |

---

#### Decelerating Flow

- **T_i too short**
- **T_i Appropriate** (flow to zero)
- **T_i too long**

---

9. Acute respiratory distress syndrome (ARDS) 199
### Pressure Support

<table>
<thead>
<tr>
<th>Other Names</th>
<th>PS; PSV; Spontaneous</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Controlled Variables</strong></td>
<td><strong>Pinsp</strong> (PS), PEEP, FiO₂, Flow trigger, Rise time</td>
</tr>
<tr>
<td><strong>Initial Setting</strong></td>
<td><strong>Adult &amp; Pediatric</strong></td>
</tr>
<tr>
<td>(More details on next card)</td>
<td></td>
</tr>
<tr>
<td><strong>Flow</strong></td>
<td>Use for Spontaneous Breathing Trial (SBT): 1. Set Pinsp Δ5-10 cmH₂O accounting for ETT size (3.0/3.5mm = 10 cmH₂O; 4.0/4.5mm = Δ8 cmH₂O; &gt;5mm = Δ5 cmH₂O) 2. Set PEEP 5-8 cmH₂O 3. FiO₂ &lt;0.40 (Peds) or &lt;0.50 (Adults) per SBT initiation criteria</td>
</tr>
<tr>
<td><strong>I:E</strong></td>
<td>Determined by patient effort and flow termination (&quot;Eₚₜ = &quot; - see below &quot;Breath Termination&quot;)</td>
</tr>
<tr>
<td><strong>Pros</strong></td>
<td>Synchrony: allows pt to determine peak flow, Vₜ, and Tₜ</td>
</tr>
<tr>
<td><strong>Cons</strong></td>
<td>• No guaranteed MV; V₁ determined by pt (big or small); high PS and/or low Eₚₜ in COPD can incr air-trapping asynchrony; muscle weakness/fatigue: ↓ effort or ability to sustain effort → hypoventilation, ↑ fatigue</td>
</tr>
<tr>
<td><strong>Breath Initiation</strong></td>
<td>Pt flow or pressure triggered; Flow (3-5LPM) more sensitive than pressure trigger (~2cmH₂O)</td>
</tr>
<tr>
<td><strong>If No Patient Trigger</strong></td>
<td>Apnea (Most vents will have backup rate; all have alarm)</td>
</tr>
<tr>
<td><strong>Breath Termination</strong></td>
<td>Flow cycled: Delivers Pinsp until flow drops to predetermined % of initial peak flow ~Esens (Standard setting ~25%; ~40-50% if obstructive pulmonary disease to prevent air trapping)</td>
</tr>
<tr>
<td><strong>Notes</strong></td>
<td>PS mode is not necessarily equivalent to a spontaneous breathing trial (SBT); must know if PS is relative to PEEP or ambient</td>
</tr>
</tbody>
</table>

![Flow Diagram](image)

### Dual (Control) Mode

| Other Names & Function | • Pressure regulated volume control (PRVC); VC +, AutoFlow  
• -PC with a target Vₜ & variable Pinsp (Δ1-3cmH₂O per breath) to meet goal V₁, despite changing C and R; |
|------------------------|------------------------------------------------------------------------------------------------------------------------|
| **Pros** | • ↓ likelihood of hypo/hyperventilation associated with PC.  
• If R or C changes, Pinsp automatically adjusts to keep target V₁  
• Active expiratory valve (unlike AC-VC) promotes synchrony |
| **Cons** | • C & R can change significantly without notification  
• Vent can’t discern if V₁-target is due to ↑ pt effort or ↑ C; vent response to both = ↓ Pinsp; Can lead to closed-loop “runaway” (↓ Pinsp→↑ Pt Effort→↓ Pinsp)=↑ Pt work; must carefully set alarms |
<table>
<thead>
<tr>
<th>Respiratory Care, Setup, &amp; Monitoring</th>
</tr>
</thead>
</table>
| **Ventilator Setup (prior to connecting patients)** | • Inspect all equipment for cleanliness or damage  
• Review circuit orientation, filters, & heat & humidification system  
• Ensure gas supply connected  
• Perform machine self-test with new patient and per manufacture (ensure leak test included)  
• Confirm initial settings and alarms |
| **Ventilator Performance** | • Perform Full Status Check q4h: (PIP, Pplat, Y, FiO2, auto-PEEP, Alarms, SpO2, ETCO2 in addition to routine ICU monitoring)  
• Evaluate vent & patient within ~1h of ventilator settings changes  
• Wipe down ventilator with approved disinfection qshift |
| **Pulmonary, Endotracheal Tube & Circuit Hygiene** | • Check cuff pressure and auscultate q12h to avoid over-inflation/leak (<25 cmH2O); consider ‘minimal occluding volume’ in peds  
• Check inflation of pilot balloon to ensure it remains inflated  
• Reposition & secure endotracheal tube with skin checks q12h  
• Check ventilator circuit qshift for moisture accumulation (drainage); change circuit only if damaged or gross contamination (Ventilator Associated Pneumonia Prophylaxis -VAP PPx)  
• Head of bed 30 degrees elevated for pneumonia prophylaxis (VAP PPx)  
• Oral hygiene with mouthwash & suctioning TID (VAP PPx)  
• Consider continuous subglottic suctioning or q12h oropharyngeal suctioning (VAP PPx)  
• All external filters should be inspected qdaily (and after nebs)  
• Replace viral filters as frequently as supplies allow in accord with the manufacturer’s recommendations or if damaged/soiled (may last >1 week)  
• For turbine & compressor ventilators, external inlet filters & fan filters must be cleaned at least monthly. For ventilators that allow, bacterial/viral filters should be placed proximal to external intake filters  
• Minimize instrumental/filter deadspace |
| **Filters** | • Active system: must use distilled or sterile water (~500mL daily) to avoid infectious risk and device damage; can be made on site or purchased; check H2O supply q12-24h  
• Passive heat moisture exchanger (HME): Only some HME include pathogen filter capability; Many manufacturers suggest change q24h, but studies show that an unsoiled HME in some circumstances can be used for 3-7 days. Nebs decrease lifespan (and must be given via bypass or with HME removed from circuit). Monitor for signs of an increased resistance (e.g. increase in PIP but no change in Pplat, or a prolonged exp flow). Ensure at least 28-30 mgH2O/L efficiency |
| **Heat & Humidification** | • Continuous pulse oximetry, if unable then spot check as frequently as possible  
• Continuous capnography, if unable then spot check as frequently as possible, especially after major ventilator settings changes  
• Auscultation performed routinely with checks  
• Skin/Mucosal Assessments qshift |
| **Respiratory Specific Monitoring** | • Ensure manual (i.e. bag valve resuscitator) ventilation device is operational and at beside along with a face mask and PEEP valve |

Disclaimer: This card is intended to be educational in nature and is not a substitute for clinical decision making based on the medical condition presented. It is intended to serve as an introduction to terminology. It is the responsibility of the user to ensure all information contained herein is current and accurate by using published references. This card is a collaborative effort by representatives of multiple academic medical centers.
# Lung-Protective Ventilation (LPV)

## When to Use LPV?

All ARDS patients and most intubated non-ARDS patients will benefit from LPV, though there are some instances where departures from LPV are justified.

### ARDS Berlin Definition for Adult ARDS with Kigali Modification
1. Acute (within 1 week of new symptoms or insult)
2. Bilateral opacities on Chest CT or chest US
3. P:F ≤ 200 or S:F ≤ 264 with or without ≥3 cmH2O PEEP
4. Not fully explained by cardiac failure or fluid overload on exam

### Pediatric ARDS (pARDS) Definition
1. Acute (within 1 week of new symptoms or insult)
2. Infiltrate(s) on chest imaging consistent with acute lung disease
3. Non-Invasive Ventilation: P:F ≤ 200 or S:F ≤ 264 with CPAP ≤ 5 cmH2O
4. Not fully explained by cardiac failure or fluid overload on exam; exclude perinatal related lung disease

## Acute Respiratory Distress Syndrome (ARDS)

### Tidal Volume (Vt)

*(Goal: 4-6 mL/Kg PBW)*

- Measure height & calculate predicted body weight (PBW) *(See table)*
- Set initial Vt, 6 mL/kg PBW (AC-VC)
- Check Vt at least every 4h (PC or if weaning PS mode)
- Titrated Vt by pressure goals & pH *(below)*
  - If pH < 7.15 consider increase Vt toward 8 mL/kg regardless of Pplat

### Pressures

*(Adults Goals: Pplat<30cmH2O and Pdr <15 cmH2O)*

*(Pediatric Goal: Pplat<28cmH2O)*

- Check Pplat (0.5s inspiratory pause) & Pdr *(deltaP=Vt/Cv = Pplat-PEEP)* ~q4-h after each change in PEEP or Vt
  - If adult Pplat >30 cmH2O (-28 Pediatrics), optimize sedation *(paralysis)* and decreasing Vt by 0.5-1 cc/kg toward ~4 mL/kg
  - If Pplat >30 cmH2O and severe patient-ventilator dysynchrony that cannot be addressed pharmacologically, consider increase Vt in 1 mL/kg steps up to 8 mL/kg
  - If Pplat >25 cm H2O and Vt <6 mL/kg, increase Vt to 6 mL/kg
  - If PEEP >20 then use Pdr goal <15 (rather than Pplat goal)

### Respiratory Rate (RR) & Inspiratory Time (Ti)

*(Goal based on pH)*

- Set RR at pre-intubation RR don’t exceed ~35 breaths/minute *(Adults)*
- Set Ti 0.70-0.85 sec *(may be longer if low RR)* *(avoid Ti <0.70 sec)*
  - When changing Vt, adjust RR to keep target VE by goal pH *(~8-12 L/min in acute ARDS)*
  - Consider lower RR if evidence of obstructive ventilatory defect
  - Increase RR if pH <7.30 and decrease RR if pH >7.45
  - Keep duration of inspiration ≤ expiration

### PEEP & FiO2

*(Goal to minimize)*

- Start at 5 cmH2O PEEP for 2min, if stable hemodynamics, then
- Select one of the following PEEP / FiO2 titration strategies for goal PaO2: 55-80 mmHg or SpO2: 88-95% *(In ARDS, PEEP usually ~10-14 cmH2O)*
  - When + PEEP, if Pplat ↑ more than ∆ PEEP, think over-distension

#### Lower PEEP/higher FiO2 Strategy *(Default - May consider if low Pdr or pediatrics)*

<table>
<thead>
<tr>
<th>FiO2</th>
<th>0.3</th>
<th>0.4</th>
<th>0.4</th>
<th>0.5</th>
<th>0.5</th>
<th>0.6</th>
<th>0.7</th>
<th>0.7</th>
<th>0.8</th>
<th>0.9</th>
<th>0.9</th>
<th>0.9</th>
<th>1.0</th>
</tr>
</thead>
<tbody>
<tr>
<td>PEEP</td>
<td>5</td>
<td>5</td>
<td>8</td>
<td>8</td>
<td>10</td>
<td>10</td>
<td>10</td>
<td>12</td>
<td>14</td>
<td>14</td>
<td>16</td>
<td>16</td>
<td>18-24</td>
</tr>
</tbody>
</table>

#### Higher PEEP/lower FiO2 Strategy *(May consider if PaO2/FiO2 is <100, high Pdr, or BMI>40)*

<table>
<thead>
<tr>
<th>FiO2</th>
<th>0.3</th>
<th>0.3</th>
<th>0.3</th>
<th>0.3</th>
<th>0.3</th>
<th>0.4</th>
<th>0.4</th>
<th>0.5</th>
<th>0.5-0.8</th>
<th>0.8</th>
<th>0.9</th>
<th>1.0</th>
<th>1.0</th>
</tr>
</thead>
<tbody>
<tr>
<td>PEEP</td>
<td>5</td>
<td>8</td>
<td>10</td>
<td>12</td>
<td>14</td>
<td>16</td>
<td>16</td>
<td>18</td>
<td>20</td>
<td>20</td>
<td>22</td>
<td>22</td>
<td>24</td>
</tr>
</tbody>
</table>
Adjunctive Therapies for ARDS Hypoxemia

Fluid Management
- Concentrate IV medications and consider diuresis once hemodynamically tolerated with goal of euvoeemia
- FACTT Trial of conservative vs. liberal fluid strategy showed conservative fluid strategy improved oxygenation, more ventilator-free & ICU-free days, no increased shock. However, no mortality benefit.

Paralysis
- May be considered in severe ARDS if high PEEP and FiO2, especially if asynchrony present; requires adequate sedation and train of four monitoring
- Choice of agent (each with pros & cons, may vary by setting): cisatracurium, atracurium, rocuronium, pancuronium, or vecuronium
- ACURASYS Trial showed mortality benefit; PETAL Trial did not
- Short term paralysis eliminates work of breathing and can be helpful to accurately assess respiratory mechanics & asynchronies associated w/ ARDS

Prone Positioning
- Prone patient for ~12-16h at a time, continue proning until P:F >150 with PEEP remaining <10 cmH2O while patient is supine for >4h
- Alternate with supine positioning which allows for patient care
- Do not need special bed: manually proning requires a team
- If unable to prone, could put less diseased lung down to improve V/Q match
- PROSEVA Trial showed mortality benefit

Additional LPV Reference Calculations

Predicted Body Weight (PBW) (kg)
- Males = 50 + 2.3 [height (inches) - 60]
- Females = 45.5 + 2.3 [height (inches) - 60]

<table>
<thead>
<tr>
<th>Height (inches)</th>
<th>PBW (lb)</th>
<th>4mL/Kg l/m</th>
<th>5mL/Kg l/m</th>
<th>6mL/Kg l/m</th>
<th>7mL/Kg l/m</th>
<th>8mL/Kg l/m</th>
</tr>
</thead>
<tbody>
<tr>
<td>58&quot; (147cm)</td>
<td>40.9/45.4 kg</td>
<td>164/182</td>
<td>205/227</td>
<td>245/272</td>
<td>286/318</td>
<td>327/363</td>
</tr>
<tr>
<td>60&quot; (152cm)</td>
<td>45.5/50 kg</td>
<td>182/200</td>
<td>228/250</td>
<td>273/300</td>
<td>319/350</td>
<td>364/400</td>
</tr>
<tr>
<td>62&quot; (157cm)</td>
<td>50.1/54.6 kg</td>
<td>200/218</td>
<td>251/273</td>
<td>301/328</td>
<td>351/382</td>
<td>401/437</td>
</tr>
<tr>
<td>64&quot; (163cm)</td>
<td>54.7/59.2 kg</td>
<td>219/237</td>
<td>274/296</td>
<td>328/355</td>
<td>383/414</td>
<td>438/474</td>
</tr>
<tr>
<td>66&quot; (168cm)</td>
<td>59.3/63.8 kg</td>
<td>237/255</td>
<td>297/319</td>
<td>356/383</td>
<td>415/447</td>
<td>474/510</td>
</tr>
<tr>
<td>68&quot; (173cm)</td>
<td>63.9/68.4 kg</td>
<td>256/274</td>
<td>320/342</td>
<td>383/410</td>
<td>447/479</td>
<td>511/547</td>
</tr>
<tr>
<td>70&quot; (178cm)</td>
<td>68.5/73 kg</td>
<td>274/292</td>
<td>343/365</td>
<td>411/438</td>
<td>480/511</td>
<td>548/584</td>
</tr>
</tbody>
</table>

Imputed Values for P:F Ratio
- Use when blood gas analysis unavailable (Link to source data)

SpO2 Values Corresponding to P:F ≤150:

<table>
<thead>
<tr>
<th>Measured SpO2</th>
<th>Imputed PaO2</th>
<th>FiO2</th>
<th>Imputed P:F</th>
</tr>
</thead>
<tbody>
<tr>
<td>96%</td>
<td>82 mmHg</td>
<td>≥0.6</td>
<td>≤137</td>
</tr>
<tr>
<td>95%</td>
<td>76 mmHg</td>
<td>≥0.5</td>
<td>≤152</td>
</tr>
<tr>
<td>94%</td>
<td>71 mmHg</td>
<td>≥0.5</td>
<td>≤142</td>
</tr>
<tr>
<td>92%</td>
<td>67 mmHg</td>
<td>≥0.5</td>
<td>≤134</td>
</tr>
<tr>
<td>91%</td>
<td>64 mmHg</td>
<td>≥0.5</td>
<td>≤128</td>
</tr>
<tr>
<td>90%</td>
<td>61 mmHg</td>
<td>≥0.4</td>
<td>≤153</td>
</tr>
<tr>
<td>&lt;89%</td>
<td>57 mmHg</td>
<td>≥0.4</td>
<td>≤148</td>
</tr>
</tbody>
</table>

Scan for PBW Calculator
Scan for Imputed P:F Calculator
# 9.15 Adult ventilation order set (ARDS)

**Clinical Care for Severe Acute Respiratory Infection—Toolkit—Update 2022**

## Adult Ventilator (ARDS) Order Set

**Mode:**  
- [ ] AC-VC  
- [ ] AC-PC  
- [ ] PRVC  
- [ ] ___  

### Tidal Volume:

1. Initiate at 6-8 mL/kg predicted body weight (PBW). Titrates down in 0.5-1 mL/kg steps 0-1 h if needed to prevent acute acidosis.
2. If EIP > 30 cm H2O, F V T ≤21 as low as 4 mL/kg.
3. pH < 7.35: F VT ≤3 mL/kg steps to 8 mL/kg regardless of EIP until pH ≥ 7.20.
4. When severe asynchronous occurs and sedation cannot be increased, then VT 1 mL/kg to upper limit of 8mL/kg.
5. If EIP ≤ 25 cm H2O and VT ≤ 6 mL/kg, ↑ VT to 6 mL/kg.
6. When using AC-PC adjust PIP for VT≤6mL/kg

### Plateau Pressure:

- 1. 25-30 cm H2O.
- 2. EIP > 30 cm H2O allowed if PEEP > 20 cm H2O or pH < 7.20

### Respiratory Rate:

- 1. Set Rate 6-35 (Typical starting rate 28, consider ≥25 if profound acidosis or markedly increased preintubation RR; consider lower if signs of obstructive pulmonary physiology)
- 2. Adjust RR to target Minute Ventilation (prior to intubation), though be cautious for possible air trapping if not too fast and/or too large of tidal volumes. (Typical minute ventilation requirements for adult ARDS ~6-12 LPM, 130-190 mL/kg PBW)
- 3. RR if pH < 7.30 (Max ~35)
- 4. RR when pH ≥ 7.45.

### Inspiratory Time:

- 0.70-0.85 sec. Avoid Tinsp < 0.70. Upper EE Limit of 1:1 (Tinsp of 0.85 @ RR of 35)

### Peak Flow Rate:

- 60-75 L/min for comfort. Avoid peak flow rate < 50 L/min

### Arterial pH:

- 1. Target: 7.30-7.45
- 2. pH < 7.25 and PaCO2 > 25: Consider Buffer Rx
- 3. pH < 7.15: Modify VT/EIP goals to achieve pH ≥ 7.20. Buffer Rx indicated to maintain lung-protective ventilation.

### PEEP FIO2 Titration:

- **ARDS Net ARMA Trial Low PEEP Table** (Default for most patients)
  - FIO2: 0.3 0.4 0.5 0.6 0.7 0.8 0.9 1.0
  - PEEP 5 5-8 7-10 10 10-14 14 14-18 18-26

- **ARDS Net Modified from ALVEOLI Trial High PEEP Table** (May consider if PaO2/FIO2 is <100, or BMI >40)
  - FIO2: 0.3 0.4 0.5 0.60-0.70 0.8 0.9 1.0
  - PEEP 5-14 14-16 16-18 18-20 20-22 22 22-24

### Free Form PEEP:

- Maintain PEEP at ___ cm H2O and Titrate FIO2 for Protocol PaO2/SpO2 range.

### Alternative Arterial:

For patients concurrently managed on the traumatic brain injury protocols, or patients with/at risk for myocardial or gastrointestinal ischemia:

- PaO2 range: __________
- PaCO2 Range: __________
- pH Range: __________

### Arterial Blood Gas:

- 1. Protocol initiation and/or immediately after intubation
- 2. Every day, first 7 days of protocol
- 3. Severe acidosis (pH < 7.35) after VT ↑ to 7 and 8 mL/kg
- 4. Every PEEP ↑ > 10 cm H2O
- 5. Check SpO2 at least Q6h and < 30 min after adjustments in PEEP/FIO2

### VD/VT (deadspace):

- When volumetric capnography available (or by crude estimation with time-based capnography - i.e., PaCO2-end tidal CO2/PaCO2), check on protocol initiation, after PEEP titrations and daily.

### Routine Care:

- Head of bed 30 degrees
- Mouthwash oral solution (15 mL) swish and suction orally q12h white intubated
- Oropharyngeal suctioning q 12h
- Perform full status check q48h (PIP, Pplat, VT, FIO2, RR)

### Source:

Adapted with permission from San Francisco General Hospital, San Francisco (CA) and USAID-STAR-UCSF Open Critical Care Project.
9.16 Checklist for proning in severe ARDS

This checklist is adapted from Messerole et al. (2002) and the most recent randomized control trial by Guérin et al. (2013). These studies found an improved mortality in patients treated with LPV plus prone position.

Prone ventilation should be carried out by four to five team members using a protocol rehearsed in advance. It is easier to perform in children. See the following article and video ( ).

Timing and duration of prone position
The most recent clinical trial (Guérin et al., 2013) observed mortality benefit in patients with severe ARDS. Patients were turned prone within 24 hours of recognition and kept prone for at least 12-16 consecutive hours a day.

Contraindications
- Elevated intracranial pressure > 30 mmHg or cerebral perfusion pressure < 60 mmHg
- Massive haemoptysis
- Recent tracheal surgery or sternotomy
- Serious facial trauma or facial surgery
- Deep venous thrombosis treated for less than 2 days
- Cardiac pacemaker inserted in the last 2 days
- Unstable spine, femur or pelvic fractures
- MAP < 65 mmHg
- Pregnancy
- Single anterior chest tube with air leaks.

Preparation of the patient for proning while IMV

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>Check for contraindications.</td>
</tr>
<tr>
<td>2</td>
<td>Consider possible adverse effects of prone positioning (e.g. if on chest tube drainage).</td>
</tr>
<tr>
<td>3</td>
<td>Whenever possible, explain the manoeuvre to the patient or their family.</td>
</tr>
<tr>
<td>4</td>
<td>Confirm from a recent chest X-ray that the tip of the endotracheal tube is located 2-4 cm above the main carina.</td>
</tr>
<tr>
<td>5</td>
<td>Inspect and confirm that the endotracheal tube and all central and large bore peripheral catheters are firmly secured.</td>
</tr>
<tr>
<td>6</td>
<td>Consider exactly how the patient’s head, neck and shoulder girdle will be supported after they are turned prone. Assemble all needed pillows, foam pads or other supports that might be needed.</td>
</tr>
<tr>
<td>7</td>
<td>Stop tube feeding, check for residual, fully evacuate the stomach, and cap or clamp the feeding and gastric tubes.</td>
</tr>
<tr>
<td>8</td>
<td>Prepare endotracheal suctioning equipment, and review what the process will be if copious airway secretions abruptly interfere with ventilation.</td>
</tr>
<tr>
<td>9</td>
<td>Decide whether the turn will be rightward or leftward.</td>
</tr>
<tr>
<td>10</td>
<td>Prepare all IV tubing and other catheters and tubing for connection when the patient is prone:</td>
</tr>
<tr>
<td></td>
<td>• Assure sufficient tubing length.</td>
</tr>
<tr>
<td></td>
<td>• If chest drainage: relocate all drainage bags on the opposite side of the bed, move chest tube drains between the legs.</td>
</tr>
<tr>
<td></td>
<td>• Reposition IV tubing toward the patient’s head, on the opposite side of the bed.</td>
</tr>
</tbody>
</table>
### Turning procedure

1. Place one (or more) people on both sides of the bed (to be responsible for the turning processes) and another at the head of the bed (to assure the central lines and the endotracheal tube do not become dislodged or kinked).

2. Increase the FiO2 to 1.0 and note the mode of ventilation, the tidal volume, the minute ventilation, and the peak and plateau airway pressures.

3. Pull the patient to the edge of the bed furthest from whichever lateral decubitus position will be used while turning.

4. Place a new draw sheet on the side of the bed that the patient will face when in this lateral decubitus position. Leave most of the sheet hanging.

5. Turn the patient to the lateral decubitus position with the dependent arm tucked slightly under the thorax. As the turning progresses the nondependent arm can be raised in a cocked position over the patient’s head. Alternatively, the turn can progress using a log-rolling procedure.

6. Remove ECG leads and patches. Suction the airway, mouth and nasal passages if necessary.

7. Continue turning to the prone position.

8. Reposition in the centre of the bed using the new draw sheet.

9. If the patient is on a standard hospital bed, turn their face toward the ventilator. Assure that the airway is not kinked and has not migrated during the turning process. Suction the airway if necessary.

10. Support the face and shoulders appropriately avoiding any contact of the supporting padding with the orbits or the eyes.

11. Position the arms for patient comfort. If the patient cannot communicate avoid any type of arm extension that might result in a brachial plexus injury.

12. Auscultate the chest to check for right mainstem intubation. Reassess the tidal volume and minute ventilation.

13. Adjust all tubing and reassess connections and functions.

14. Reattach ECG patches and leads to the back.

15. Tilt the patient into reverse Trendelenburg. Slight, intermittent lateral repositioning (20-30°) should also be used, changing sides at least every 2 hours.


### Criteria for stopping proning in severe ARDS:

- Oxygenation improvement defined as PaO2/FiO2 ≥ 150 mmHg with PEEP ≤ 10 cmH2O and FiO2 ≤ 0.6; in the prone group, these criteria had to be met in supine at least 4 hours after the end of the last prone session.

- PaO2/FiO2 ratio deterioration by more than 20% relative to supine before two consecutive prone sessions.

- Complications occurring during a prone session such as:
  - non-scheduled extubation
  - mainstem bronchus intubation
  - endotracheal tube obstruction
  - haemoptysis, SpO2 < 85%
  - PaO2 < 55 mmHg for more than 5 minutes under FiO2 1.0
  - cardiac arrest
  - HR < 30 beats per minute for more than 1 minute
  - SBP < 60 mmHg for more than 5 minutes
  - or any other life-threatening reason for which the clinician decided to stop.
9.17 Ventilator circuit types, filter and humidifier locations for SARI

When managing a patient with infectious SARI, careful consideration must be given to the placement of adequate bacterial-viral (BV) filters to protect health care workers and patients from potential exposure to infectious particles.

Filters may be placed at the air intake, inspiratory limb, patient wye, expiratory limb and/or exhaust port; however, placement at each of these sites does not provide equivalent function. Ideally a two-filter setup should be used:

- **Inspiratory/patient filter:** The first BV filter should be placed between the patient and the inspiratory limb take off from the ventilator. This filter is often placed between the inspiratory limb and the ventilator or between the circuit wye connector and the patient. The inspiratory/patient filter has two purposes: 1) to protect the ventilator from exhaled gases from an infected patient; and 2) to protect a non-infected patient from a possibly contaminated ventilator.

- **Expiratory filter:** A second BV filter should be placed between the patient and the exhalation valve on the expiratory limb of the circuit. This filter is intended to protect the room environment and health care staff from aerosolized particles, and to protect the device in a dual limb circuit setup.

If using an active heat and humidification system, then the inspiratory filter should be a BV filter, without heat-moisture exchange (HME) function. If not using an active heat and humidification system, then the circuit must include either: 1) an HME with BV filter function (also known as an HMEF) between the wye and patient; or 2) an HME without BV filter function placed at the wye, in addition to an inspiratory and expiratory BV filter.

Placement of HME, BV or HMEF filters between the circuit wye and the patient’s endotracheal tube can add significant dead space to the circuit, especially for paediatric patients, and must always be accounted for when using this setup.
Invasive mechanical ventilation devices

Ventilator Circuit Setup
Filter Placement & Humidification Types

**SINGLE LIMB** CIRCUITS*

Heated Wire
(Active heat & humidification)

**Air Intake BV Filter** may be recommended for some devices and may require menu settings changes. When invasive BV filter is in use, BV filter below the expiratory and inspiratory limb may not be needed.

Heat Moisture Exchanger Filter (HMEF)
(BV filter by Exhalation Valve)

HMEFs must always be placed between the circuit and the endotracheal tube. They should be used at the ventilator or exhalation valve.

Heat Moisture Exchanger (HME)
(BV filter by Exhalation Valve)

When using an HME, the expiratory BV filter should ideally be placed either at the exhalation valve or at the start of the expiratory limb from the HME (not shown). This illustration shows a slightly different type of expiratory valve than the other on this page. There is no implication for the location of HME or filters.

---

Source: USAID-STAR-UCSF Open Critical Care Project (  )
DUAL LIMB CIRCUITS

Heated Wire
(Active heat & humidification)

Heat Moisture Exchanger (HME)

Heat Moisture Exchange Filter (HMEF)

END TIDAL CO₂ PLACEMENT

Sidestream CO₂
Sampling HMEF with integrated sampling port

Sidestream CO₂
Sampling HME and Bacterial Viral Filter

Mainstream CO₂
Monitoring HME

Source: USAID-STAR-UCSF Open Critical Care Project (©)
**Bag valve resuscitator setup**

HMEF may be considered over a bacterial/viral filter if duration of use on the bag valve resuscitator is anticipated to be prolonged.

**In-line suction catheter setup**

This illustration shows the location of the in-line suction catheter in relation to the patient circuit. There are other potential configurations not shown here, but in all, the in-line suction catheter is placed on the endotracheal tube.

Source: USAID-STAR-UCSF Open Critical Care Project (Download Icon)
Non-invasive mechanical ventilation

CPAP/NIPPV Circuit Setup
Mask types, filter and humidification placement

Dual Limb Circuit + Non-Vented Mask
(Exhalation valve in the circuit)

- Because this setup has an expiratory valve at or in the ventilator (not shown), a non-vented mask must be used for CPAP/NIPPV.
- See Figure "Ventilator Circuit Setup" for alternative humidification & filter setups. An HMEF is shown here but may not be the optimal setup depending on local resources.
- The addition of filters between the circuit, Wye and the patient introduce dead space.

"Single Limb" Circuit + Non-Vented Mask
(Exhalation valve in the circuit)

- The term "single limb" circuit is a commonly used moniker when referring to the type of circuit shown here. This setup shown has a short expiratory limb and expiratory valve and functions similarly to a dual limb circuit.
- Because this setup has an expiratory valve, a non-vented mask must be used for CPAP/NIPPV.
- See Figure "Ventilator Circuit Setup" for alternative humidification & filter setups.

True Single Limb Circuit + Vented Mask
(No exhalation valve)

- The true single limb circuit is commonly found in home CPAP/NIPPV machines and has no expiratory valve.
- Exhaled breath must leave through intentional leak ports - in this setup a vented mask is used.
- The BV filter in this scenario protects the patient from a potentially contaminated device/circuit but does not protect personnel from potentially infectious aerosolized particles in the exhaled gas.

True Single Limb Circuit + Leak Adapter + Non-Vented Mask
(No exhalation valve)

- This true single limb circuit is commonly found in home CPAP/NIPPV machines and has no expiratory valve.
- Exhaled breath must leave through intentional leak ports - in this setup a non-vented mask is used and the leak occurs via a specialized fixed leak adapter in the circuit.
- Either a bacterial/viral filter if active heated humidification is being used, or HMEF if no active heated humidification should be placed between the leak adapter & the patient.

References and resources


Hess DR. Ventilator waveforms and the physiology of pressure support ventilation. Respir Care. 2005;50(2):166-186.


10 Manage pain, sedation and delirium
Summary

In critically ill patients, comfort and safety should be the goals when managing pain control, anxiolysis and prevention and treatment of delirium.

Implementation of a protocolized management approach to pain, agitation and delirium (PAD protocol) improves patient outcomes

<table>
<thead>
<tr>
<th>Short-term outcomes</th>
<th>Long-term outcomes</th>
</tr>
</thead>
<tbody>
<tr>
<td>• Fewer days of delirium</td>
<td>• Better cognitive function</td>
</tr>
<tr>
<td>• Fewer days of IMV (quicker time to extubation)</td>
<td>• Better physical mobility</td>
</tr>
<tr>
<td>• Fewer days in ICU</td>
<td>• Fewer psychiatric conditions (e.g. PTSD, depression, anxiety)</td>
</tr>
<tr>
<td>• Lower hospital costs</td>
<td>• Increased survival</td>
</tr>
<tr>
<td>• More mobilization during ICU stay</td>
<td></td>
</tr>
<tr>
<td>• Increased survival</td>
<td></td>
</tr>
</tbody>
</table>

Steps to develop a PAD protocol adapted to the health facility

Step 1  Assess and recognize pain, agitation and delirium using standardized and reproducible scales (i.e. VAS, RASS, CAM-ICU).

Step 2  Manage pain first: prevention and treatment (treat pain with a multimodal approach using opioids and non-opioids to minimize the harmful effects of sedatives).

Step 3  Manage agitation: choice of targeted sedation (treat anxiety using non-benzodiazepines sedatives and target light sedation). Evaluate patients daily for potential reductions of sedation, anxiolytics, and analgesics as well as for sedation interruptions in intubated patients when criteria are met.

Step 4  Manage delirium: prevention and treatment (prioritize non-pharmacologic interventions to prevent or treat delirium, over pharmacologic treatments).

Step 5  Special situations that may need deep sedation and neuromuscular blockade (i.e. severe ARDS).

Step 6  Monitor-record-interpret-respond.

Step 7  Deliver quality care: implementation as part of ABCDEF bundle.

Short-term outcomes

• Fewer days of delirium
• Fewer days of IMV (quicker time to extubation)
• Fewer days in ICU
• Lower hospital costs
• More mobilization during ICU stay
• Increased survival

Long-term outcomes

• Better cognitive function
• Better physical mobility
• Fewer psychiatric conditions (e.g. PTSD, depression, anxiety)
• Increased survival

Short-term outcomes

• Fewer days of delirium
• Fewer days of IMV (quicker time to extubation)
• Fewer days in ICU
• Lower hospital costs
• More mobilization during ICU stay
• Increased survival

Long-term outcomes

• Better cognitive function
• Better physical mobility
• Fewer psychiatric conditions (e.g. PTSD, depression, anxiety)
• Increased survival
Tools

10.1 Numerical pain assessment scales
10.2 Behavioural pain assessment scales
   10.2.1 Behavioural Pain Scale (BPS)
   10.2.2 Critical-Care Pain Observation Tool (CPOT)
   10.2.3 COMFORT-B Scale
   10.2.4 FLACC Behavioural Pain Assessment Scale
10.3 Richmond Agitation-Sedation Scale (RASS)
10.4 Flowchart and worksheet for the Confusion Assessment Method of the ICU for adults (CAM-ICU)
10.5 Flowchart and worksheet for the Confusion Assessment Method of the ICU for children (pCAM-ICU)
10.6 Procedure for assessing attention: attention screening exam (ASE) for adults
10.7 Guide to commonly used sedatives in adults
10.8 Guide to commonly used opioid analgesics in adults
10.9 Guide to using neuromuscular blockers in adults
10.10 Guide to commonly used antipsychotics (haloperidol) in adults
10.11 Guide to paediatric analgesics, sedatives and neuromuscular blockers
10.1 Numerical pain assessment scales

Visual analogue scale
The visual analogue scale (VAS) for pain assessment in adults and adolescents is a validated and widely used method of monitoring the subjective level of pain experienced by patients. It is a 10 cm long scale, which ranges from 0 (no pain) to 10 (the worst pain that one can imagine). It is flexible, in that patients can make verbal or visual responses (i.e. if verbal communication is not possible, the patient can be shown a 10 cm scale and can point to the region which corresponds to their pain).

A major limitation of the VAS is that it requires an awake patient who grasps the concept of a scale. These conditions are frequently not satisfied in ICU patients.

A major limitation of the VAS is that it requires an awake patient who grasps the concept of a scale. These conditions are frequently not satisfied in ICU patients.

The lower the VAS score, the higher the quality of the analgesia. However, a low VAS score with excessive sedation must be avoided, if possible. The level of sedation must be also closely monitored (see the Richmond Agitation-Sedation Scale tool).

Wong-Baker Faces Scale
The Wong-Baker Faces Scale can be used in younger children - they are asked to point to the face that reflects their pain level.

10.2 Behavioural pain assessment scales

There are two validated behavioural pain assessment scales that can be used to assess pain in non-communicative or sedated patients (e.g. mechanical ventilation); these are recommended to use instead of physiological indicators alone. The sedation and pain levels of children in intensive care should be assessed at least 4 hourly in intensive care.

- **BPS**: Behavioural Pain Scale
- **CPOT**: Critical-Care Pain Observation Tool
- In children: **Comfort-B Scale** and **FLACC** Behavioural Pain Assessment Scale

### 10.2.1 Behavioural Pain Scale (BPS)

BPS score ranges from 3 (no pain) to 12 (maximum pain).

<table>
<thead>
<tr>
<th>Item</th>
<th>Description</th>
<th>Score</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Facial expression</strong></td>
<td>Relaxed</td>
<td>1</td>
</tr>
<tr>
<td></td>
<td>Partially tightened (e.g. brow lowering)</td>
<td>2</td>
</tr>
<tr>
<td></td>
<td>Fully tightened (e.g. eyelid closing)</td>
<td>3</td>
</tr>
<tr>
<td></td>
<td>Grimacing</td>
<td>4</td>
</tr>
<tr>
<td><strong>Upper limb movements</strong></td>
<td>No movement</td>
<td>1</td>
</tr>
<tr>
<td></td>
<td>Partially bent</td>
<td>2</td>
</tr>
<tr>
<td></td>
<td>Fully bent with finger flexion</td>
<td>3</td>
</tr>
<tr>
<td></td>
<td>Permanently retracted</td>
<td>4</td>
</tr>
<tr>
<td><strong>Compliance with mechanical ventilation</strong></td>
<td>Tolerating movement</td>
<td>1</td>
</tr>
<tr>
<td></td>
<td>Coughing but tolerating ventilation most of the time</td>
<td>2</td>
</tr>
<tr>
<td></td>
<td>Fighting ventilator</td>
<td>3</td>
</tr>
<tr>
<td></td>
<td>Unable to control ventilation</td>
<td>4</td>
</tr>
</tbody>
</table>
### Critical-Care Pain Observation Tool (CPOT)

<table>
<thead>
<tr>
<th>Indicator</th>
<th>Score</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>Facial expressions</td>
<td>Relaxed, neutral</td>
<td>0  No muscle tension</td>
</tr>
<tr>
<td>Tense</td>
<td>1</td>
<td>Presence of frowning, brow lowering, orbit tightening and levator contraction or any other change (e.g. opening eyes or tearing during nociceptive procedures)</td>
</tr>
<tr>
<td>Grimacing</td>
<td>2</td>
<td>All previous facial movements plus eyelid tightly closed (the patient may present with mouth open or biting endotracheal tube)</td>
</tr>
<tr>
<td>Body movements</td>
<td>Absence of movements or normal position</td>
<td>0  Does not move at all (doesn’t necessarily mean absence of pain) or normal position (movements not aimed toward the pain site or not made for the purpose of protection)</td>
</tr>
<tr>
<td>Protection</td>
<td>1</td>
<td>Slow, cautious movements, touching or rubbing the pain site, seeking attention through movements</td>
</tr>
<tr>
<td>Restlessness/agitation</td>
<td>2</td>
<td>Pulling tube, attempting to sit up, moving limbs/thrashing, not following commands, striking at staff, trying to climb out of bed</td>
</tr>
<tr>
<td>Compliance with the ventilator (intubated patients) or Vocalization (extubated patient)</td>
<td>Tolerating ventilator or movement</td>
<td>0  Alarms not activated, easy ventilation</td>
</tr>
<tr>
<td>Coughing but tolerating</td>
<td>1</td>
<td>Coughing, alarms may be activated but stop spontaneously</td>
</tr>
<tr>
<td>Fighting ventilator</td>
<td>2</td>
<td>Asynchrony; blocking ventilation, alarms frequently activated</td>
</tr>
<tr>
<td>Talking in normal tone or no sound</td>
<td>0</td>
<td>Talking in normal tone or no sound</td>
</tr>
<tr>
<td>Sighing, moaning</td>
<td>1</td>
<td>Sighing, moaning</td>
</tr>
<tr>
<td>Crying out, sobbing</td>
<td>2</td>
<td>Crying out, sobbing</td>
</tr>
<tr>
<td>Muscle tension Evaluation by passive flexion and extension of upper limbs when patient is at rest or evaluation when patient is being turned</td>
<td>Relaxed</td>
<td>0  No resistance to passive movements</td>
</tr>
<tr>
<td>Tense, rigid</td>
<td>1</td>
<td>Resistance to passive movements</td>
</tr>
<tr>
<td>Very tense or rigid</td>
<td>2</td>
<td>Strong resistance to passive movements or incapacity to complete them</td>
</tr>
<tr>
<td><strong>Total</strong></td>
<td>_</td>
<td>_ /8</td>
</tr>
</tbody>
</table>

Source: Adapted from Gélinas et al (2006).
Facial expressions

0  Relaxed, neutral
   (no muscle tension)

1  Tense
   (frowning, brow lowering, orbit tightening, little levator contraction)

2  Grimacing
   (contraction of the whole face: frowning, brow lowering, eyes tightly closed, levator contraction - mouth may be opened or the patient may be biting the endotracheal tube)

Source: Adapted from Payen et al (2001).
Note: A score of 1 may be attributed when a change in the patient’s facial expression is observed compared with rest (e.g. opening or weeping).

How to use the Critical-Care Pain Observation Tool

1. The patient must be observed at rest for 1 minute to obtain a baseline value of the CPOT.
2. Then, the patient should be observed during nociceptive procedures known to be painful (e.g. turning, wound care) to detect any changes in the patient’s behaviours to pain.
3. The patient should be evaluated before and at the peak effect of an analgesic agent to assess whether the treatment was effective or not in relieving pain.
4. For the rating of the CPOT, the patient should be attributed the highest score observed for each item during the observation period.
5. The patient should be attributed a score for each behaviour included in the CPOT and muscle tension should be evaluated last, especially when the patient is at rest because the stimulation of touch alone (when performing passive flexion and extension of the arm) may lead to behavioural reactions.

Free teaching CPOT video available from the Society of Critical Care Medicine.
**Observation of patient at rest (baseline)**
The nurse looks at the patient’s face and body to note any visible reaction for an observation period of 1 minute. She/he gives a score for all items except for muscle tension. At the end of the 1-minute period, the nurse holds the patient’s arm in both hands – one at the elbow, and one to hold the patient’s hand. Then she/he performs and passive flexion and extension of the upper limb, and feels any resistance the patient may exhibit. If the movements are performed easily, the patient is found to be relaxed with no resistance (score 0). If the movements can still be performed but with more strength, then it is concluded that the patient is showing resistance to movement (score 1). Finally, if the nurse cannot perform the movement, strong resistance is felt (score 2). This can be observed in patients who are spastic.

**Observation of patient during turning**
Even during the turning procedure, the nurse can still assess the patient’s pain. While she/he is turning the patient on one side, she/he looks at the patient’s face to note any reactions such as frowning or grimacing. These reactions may be brief or can last longer. The nurse also looks out for body movements. For instance, she/he looks for protective movements like the patient trying to reach or touching the pain site (e.g. surgical incision, injury site). In the mechanically ventilated patient the nurse pays attention to alarms and if they stop spontaneously or require that she/he intervenes (reassurance, administering medication). According to muscle tension, the nurse can feel if the patient is resisting to the movement or not. A score of 2 is given when the patient is resisting against the movement and attempts to get on his/her back.

### 10.2.3 COMFORT-B Scale

It cannot be used in children who are receiving muscle relaxant drugs or children with severe neurological impairment. The child should be observed for 2 minutes and six behaviours are scored as below (score either respiratory response or crying, depending on the child’s intubation status).

Children scoring 11–22 are in the optimal range of sedation; children scoring < 10 may be oversedated (consider weaning); and children > 23 are undersedated.

<table>
<thead>
<tr>
<th>Item</th>
<th>Description</th>
<th>Score</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Alertness</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td>1. Deeply asleep</td>
<td></td>
<td></td>
</tr>
<tr>
<td>2. Lightly asleep</td>
<td></td>
<td></td>
</tr>
<tr>
<td>3. Drowsy</td>
<td></td>
<td></td>
</tr>
<tr>
<td>4. Fully awake and alert</td>
<td></td>
<td></td>
</tr>
<tr>
<td>5. Hyperalert</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>Calmness/agitation</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td>1. Calm</td>
<td></td>
<td></td>
</tr>
<tr>
<td>2. Slightly anxious</td>
<td></td>
<td></td>
</tr>
<tr>
<td>3. Anxious</td>
<td></td>
<td></td>
</tr>
<tr>
<td>4. Very anxious</td>
<td></td>
<td></td>
</tr>
<tr>
<td>5. Panicky</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Item</td>
<td>Description</td>
<td>Score</td>
</tr>
<tr>
<td>----------------------------------</td>
<td>--------------------------------------------------------------------------------------------------</td>
<td>-------</td>
</tr>
<tr>
<td><strong>Respiratory response</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td>(ventilated children)</td>
<td>1. No coughing and no spontaneous respiration</td>
<td></td>
</tr>
<tr>
<td></td>
<td>2. Spontaneous respiration with little or no response to ventilation</td>
<td></td>
</tr>
<tr>
<td></td>
<td>3. Occasional cough or resistance to ventilator</td>
<td></td>
</tr>
<tr>
<td></td>
<td>4. Actively breathes against ventilator or coughs regularly</td>
<td></td>
</tr>
<tr>
<td></td>
<td>5. Fights ventilator, cough or choking</td>
<td></td>
</tr>
<tr>
<td><strong>Cry</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td>(non-ventilated children)</td>
<td>1. Quiet breathing, no crying</td>
<td></td>
</tr>
<tr>
<td></td>
<td>2. Sobbing or gasping</td>
<td></td>
</tr>
<tr>
<td></td>
<td>3. Moaning</td>
<td></td>
</tr>
<tr>
<td></td>
<td>4. Crying</td>
<td></td>
</tr>
<tr>
<td></td>
<td>5. Screaming</td>
<td></td>
</tr>
<tr>
<td><strong>Physical movement</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>1. No movement</td>
<td></td>
</tr>
<tr>
<td></td>
<td>2. Occasional, slight movements</td>
<td></td>
</tr>
<tr>
<td></td>
<td>3. Frequent, slight movements</td>
<td></td>
</tr>
<tr>
<td></td>
<td>4. Vigorous movement limited to extremities</td>
<td></td>
</tr>
<tr>
<td></td>
<td>5. Vigorous movements including torso and head</td>
<td></td>
</tr>
<tr>
<td><strong>Muscle tone</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>1. Muscles totally relaxed, no muscle tone</td>
<td></td>
</tr>
<tr>
<td></td>
<td>2. Reduced muscle tone</td>
<td></td>
</tr>
<tr>
<td></td>
<td>3. Normal muscle tone</td>
<td></td>
</tr>
<tr>
<td></td>
<td>4. Increased muscle tone and flexion of fingers and toes</td>
<td></td>
</tr>
<tr>
<td></td>
<td>5. Extreme muscle rigidity and flexion of fingers and toes</td>
<td></td>
</tr>
<tr>
<td><strong>Facial tension</strong></td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>1. Facial muscle totally relaxed</td>
<td></td>
</tr>
<tr>
<td></td>
<td>2. Facial muscle tone normal; no facial muscle tension evident</td>
<td></td>
</tr>
<tr>
<td></td>
<td>3. Tension evident in some facial muscles</td>
<td></td>
</tr>
<tr>
<td></td>
<td>4. Tension evident throughout facial muscles</td>
<td></td>
</tr>
<tr>
<td></td>
<td>5. Facial muscles contorted and grimacing</td>
<td></td>
</tr>
</tbody>
</table>

**Total score**

Source: Adapted from Ambuel et al (1992).
10.2.4 FLACC Behavioural Pain Assessment Scale

The FLACC scale is a measurement used to assess pain for children between 2 months and 7 years or for individuals who are unable to communicate their pain.

<table>
<thead>
<tr>
<th>FLACC Behavioural Pain Assessment Scale</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Categories</strong></td>
</tr>
<tr>
<td>Face</td>
</tr>
<tr>
<td>Legs</td>
</tr>
<tr>
<td>Activity</td>
</tr>
<tr>
<td>Cry</td>
</tr>
<tr>
<td>Consolability</td>
</tr>
</tbody>
</table>


How to use the FLACC

In patients who are awake: observe for 1 to 5 minutes or longer. Observe legs and body uncovered. Reposition patient or observe activity. Observe body for tenseness and tone. Initiate consoling interventions if needed.

In patients who are asleep: observe for 5 minutes or longer. Observe legs and body uncovered. If possible, reposition the patient. Touch the body and observe for tenseness and tone.

**Face**
- Score 0 if the patient has a relaxed face, makes eye contact, shows interest in surroundings.
- Score 1 if the patient has a worried facial expression, with eyebrows lowered, eyes partially closed, cheeks raised, mouth pursed.
- Score 2 if the patient has deep furrows in the forehead, closed eyes, an open mouth, deep lines around nose and lips.

**Legs**
- Score 0 if the muscle tone and motion in the limbs are normal.
- Score 1 if the patient has increased tone, rigidity, or tension; if there is intermittent flexion or extension of the limbs.
- Score 2 if the patient has hypertonicity, the legs are pulled tight, there is exaggerated flexion or extension of the limbs, tremors.

**Activity**
- Score 0 if the patient moves easily and freely, normal activity or restrictions.
- Score 1 if the patient shifts positions, appears hesitant to move, demonstrates guarding, a tense torso, pressure on a body part.
- Score 2 if the patient is in a fixed position, rocking; demonstrates side-to-side head movement or rubbing of body part.

**Cry**
- Score 0 if the patient has no cry or moan, awake or asleep.
- Score 1 if the patient has occasional moans, awake or asleep.
- Score 2 if the patient has frequent or continuous moans, cries, grunts.

**Consolability**
- Score 0 if the patient is clam and does not require consoling.
- Score 1 if the patient responds to comfort by touching or talking in 30 seconds to 1 minute.
- Score 0 if the patient requires constant comforting or is inconsolable.

Interpreting the Behavioural Score

Each category is scored on the 0-2 scale, which results in a total score of 0-10; 0 = Relaxed and comfortable; 1-3 = Mild discomfort; 4-6 = Moderate pain; 7-10 = Severe discomfort or pain or both.
10.3 Richmond Agitation-Sedation Scale (RASS)

Assess agitation, anxiety and sedation levels on a regular basis using a standardized scale and set a daily sedation target based on clinical condition and management plans for the day. Consider the use of the Richmond Agitation-Sedation Scale (RASS). This has been validated in many clinical trials and can be easily taught to staff.

<table>
<thead>
<tr>
<th>Score</th>
<th>Term</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>+4</td>
<td>Combative</td>
<td>Overtly combative, violent, immediate danger to staff</td>
</tr>
<tr>
<td>+3</td>
<td>Very agitated</td>
<td>Pulls or removes tube(s) or catheter(s); aggressive</td>
</tr>
<tr>
<td>+2</td>
<td>Agitated</td>
<td>Frequent non-purposeful movement, fights ventilator</td>
</tr>
<tr>
<td>+1</td>
<td>Restless</td>
<td>Anxious but movements not aggressive vigorous</td>
</tr>
<tr>
<td>0</td>
<td>Alert and calm</td>
<td></td>
</tr>
<tr>
<td>-1</td>
<td>Drowsy</td>
<td>Not fully alert, but has sustained awakening (eye-opening/eye contact) to voice (&gt; 10 seconds)</td>
</tr>
<tr>
<td>-2</td>
<td>Light sedation</td>
<td>Briefly awakens with eye contact to voice (&lt; 10 seconds)</td>
</tr>
<tr>
<td>-3</td>
<td>Moderate sedation</td>
<td>Movement or eye opening to voice (but no eye contact)</td>
</tr>
<tr>
<td>-4</td>
<td>Deep sedation</td>
<td>No response to voice, but movement or eye opening to physical stimulation</td>
</tr>
<tr>
<td>-5</td>
<td>Unarousable</td>
<td>No response to voice or physical stimulation</td>
</tr>
</tbody>
</table>

Algorithm for RASS assessment

In most patients, this assessment is very quick and takes only 30 seconds (only 10% take a few minutes).

1. Patient is alert, restless or agitated
   2a. Patient awakens with sustained eye opening and eye contact
   2b. Patient awakens with eye opening and eye contact, but not sustained
   2c. Patient has any movement in response to voice but no eye contact

2. Patient is not alert
   State patient’s name and say to open eyes and look at speaker
   2a. Patient awakens with sustained eye opening and eye contact
   2b. Patient awakens with eye opening and eye contact, but not sustained
   2c. Patient has any movement in response to voice but no eye contact

3. Patient has no response to verbal stimulation
   Physically stimulate patient by shaking shoulder and/or rubbing sternum
   3a. Patient has any movement to physical stimulation
   3b. Patient has no response to any stimulation

Score: 0 to +4
   ➔ If RASS is +4 to +3, then proceed to the delirium assessment
   ➔ If RASS is +2 to 0, then proceed to the delirium assessment

Score: -1 to -5
   ➔ If RASS is -1 or -2, then proceed to the delirium assessment
   ➔ If RASS is -3 or -4, then stop and reassess patient at later time

Source: Adapted from Sessler et al (2002).
10.4 Flowchart and worksheet for the Confusion Assessment Method of the ICU for adults (CAM-ICU)

Use the CAM-ICU flowsheets and worksheet reproduced below, to assess delirium in conjunction with the RASS scale. Additional training materials on how to do the CAM-ICU and train staff can also be found at the same link.

**CAM-ICU flowchart**

The flowchart can be used as a pocket card or wall poster to easily reference the procedure to assess for the presence of delirium.

**Feature 1. Acute change or fluctuating course of mental status**
- Is there an acute change from mental status baseline? or
- Has the patient’s mental status fluctuated during the past 24 hours?

**Feature 2. Inattention**
- “Squeeze my hand when I say the letter ‘A’”.
  - Read the following sequence of letters:
    - S A V E A H A R T or C A S A B L A N C A or A B A D B A D A A Y
  - Errors: No squeeze with ‘A’ and squeeze with letter other than ‘A’.
  - If unable to complete letters → pictures.

**Feature 3. Altered level of consciousness**
- Current RASS level.

**Feature 4. Disorganized thinking**
- Will a stone float on water?
- Are there fish in the sea?
- Does one pound weigh more than two?
- Can you use a hammer to pound a nail?
  - Command: “Hold up this many fingers.”
    - (Hold up two fingers)
      - “Now do the same thing with the other hand.”
        - (Do not demonstrate)
    - "Add one more finger.”
      - (If patient unable to move both arms)

**CAM-ICU negative**
- NO DELIRIUM

**CAM-ICU positive**
- DELIRIUM present

**CAM-ICU worksheet**

<table>
<thead>
<tr>
<th>Feature 1: Acute onset or fluctuating course</th>
<th>Score</th>
<th>Check here if present</th>
</tr>
</thead>
<tbody>
<tr>
<td>Is the patient different than his/her baseline mental status? or Has the patient had any fluctuation in mental status in the past 24 hours as evidenced by fluctuation on a sedation/level of consciousness scale (i.e. RASS/SAS, GCS or previous delirium assessment)?</td>
<td>Either question Yes →</td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Feature 2: Inattention</th>
<th>Score</th>
<th>Check here if present</th>
</tr>
</thead>
<tbody>
<tr>
<td>Letters attention test:</td>
<td>Number of errors &gt; 2 →</td>
<td></td>
</tr>
<tr>
<td>Directions: Say to the patient, “I am going to read you a series of 10 letters. Whenever you hear the letter ‘A’, indicate by squeezing my hand.” Read the letters from the following list in a normal tone 3 seconds apart.</td>
<td></td>
<td></td>
</tr>
<tr>
<td>save a h a r t or c a s a b l a n c a or a b a b a d a a y</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Errors are counted when patient fails to squeeze on the letter “A” and when the patient squeezes on any letter other than “A”.</td>
<td></td>
<td></td>
</tr>
<tr>
<td>If unable to complete letters attention test → use pictures (see Tool 10.6)</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Feature 3: Altered level of consciousness</th>
<th>Score</th>
<th>Check here if present</th>
</tr>
</thead>
<tbody>
<tr>
<td>Present if actual RASS score is anything other than alert and calm (zero)</td>
<td>RASS anything other than zero →</td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Feature 4: Disorganized thinking</th>
<th>Score</th>
<th>Check here if present</th>
</tr>
</thead>
<tbody>
<tr>
<td>Yes/No questions:</td>
<td>Combined number of errors &gt; 1 →</td>
<td></td>
</tr>
<tr>
<td>Will a stone float on water?</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Are there fish in the sea?</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Does one pound weigh more than two?</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Can you use a hammer to pound a nail?</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Errors are counted when the patient incorrectly answers a question. Command:</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Say to the patient, “Hold up this many fingers.” (Hold up two fingers in front of the patient)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>“Now do the same thing with the other hand.” (Do not repeat the number of fingers)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Note: If patient is unable to move both arms, for second part of command ask patient to “Add one more finger”</td>
<td></td>
<td></td>
</tr>
<tr>
<td>An error is counted if patient unable to complete entire command.</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Overall CAM-ICU</th>
<th>Score</th>
<th>Check here if present</th>
</tr>
</thead>
<tbody>
<tr>
<td>CAM-ICU positive = Feature 1 + Feature 2 + either Feature 3 or Feature 4</td>
<td>Criteria met →</td>
<td>CAM-ICU positive (DELIRIUM present)</td>
</tr>
<tr>
<td></td>
<td>Criteria not met →</td>
<td>CAM-ICU negative (NO DELIRIUM)</td>
</tr>
</tbody>
</table>

10.5 Flowchart and worksheet for the Confusion Assessment Method of the ICU for children (pCAM-ICU)

This tool is adapted from Smith et al (2011) (see References and resources).

pCAM-ICU flowchart

- **Feature 1. Acute change or fluctuating course of mental status**
  - Is there an acute change from mental status baseline? or
  - Has the patient’s mental status fluctuated?

- **Feature 2. Inattention**
  - “Squeeze my hand when I say ‘A’.”
  - Read the following sequence of letters: A B A D B A D A A Y.
  - Errors: No squeeze with ‘A’ and squeeze with letter other than ‘A’.
  - If unable to complete ASE Letters → ASE pictures.

- **Feature 3. Altered level of consciousness**
  - Refer to current RASS (sedation assessment) score.

- **Feature 4. Disorganized thinking**
  1. Is sugar sweet? (Alternate: Is a rock hard?)
  2. Is ice cream hot? (Do rabbits fly?)
  3. Do birds fly? (Is ice cream cold?)
  4. Is an ant bigger than an elephant? (Is a giraffe smaller than a mouse?)

  **Command:**
  - “Hold up this many fingers.” (Hold up two fingers)
  - “Now do that with the other hand.” (Do not demonstrate)
  - or
  - “Add one more finger.” (If patient is unable to move both arms)
### pCAM-ICU worksheet

<table>
<thead>
<tr>
<th>Feature 1: Acute change or fluctuating course of mental status</th>
</tr>
</thead>
<tbody>
<tr>
<td>A. Is there an acute change from mental status baseline? Yes or No</td>
</tr>
<tr>
<td>B. Has my patient’s mental status fluctuated during the past 24 hours? Yes or No</td>
</tr>
<tr>
<td>Evidenced by fluctuation on a sedation scale (RASS), SAS, GCS or previous delirium assessment.</td>
</tr>
<tr>
<td>If either answer YES then circle +</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Feature 2: Inattention ➔ FEATURE POSITIVE if SCORE 0-7 on Vigilance “A” test or ASE picture test</th>
</tr>
</thead>
<tbody>
<tr>
<td>Vigilance “A” test:</td>
</tr>
</tbody>
</table>
| I want my patient to squeeze my hand when I say ONLY the letter “A”.
I will read the 10-letter sequence in the same order every day, with my normal voice, saying each letter once every second. |
| Directions to patient: “Squeeze my hand when I say the letter ‘A’. Let’s practise, ‘A’”. |
| To score: |
| When I say the letter “A” and the patient does not squeeze my hand, I subtract 1 point. |
| When I say the other letters and the patient squeezes my hand, I subtract 1 point. |
| A____B____A____D____B____A____D____A____A____Y____ |
| or |
| ASE picture test: |
| I will show the patient “5 memory pictures”. I want the patient to remember the 5 “memory pictures” when shown a larger “deck” of 10 pictures. |
| Directions to patient: “I am going to show you 5 pictures that I want you to remember”.(Show 1 picture every 3 seconds and state object’s name.) |
| Directions if patient can verbalize: “Say yes when you see 1 of those 5 pictures again”. (Show all pictures from deck and state object’s name.) |
| Directions to intubated patient: “Nod your head yes when you see 1 of those 5 pictures again”. |
| To score: |
| If the patient nods or says “yes” to ONLY the 5 memory pictures they have completed the task successfully - SCORE 10/10. |
| If patient does not nod or say “yes” to 1 of the 5 memory pictures, I will subtract 1 point. |
| If the patient nods or says “yes” to the other pictures in the deck, I will subtract 1 point. |
| Memory picture: _____ / 5 Deck pictures: _____ / 5 |
| If the SCORE is 0-7 then circle + | + / – |

<table>
<thead>
<tr>
<th>Feature 3: Altered level of consciousness ➔ FEATURE POSITIVE if the current RASS score is anything other than 0</th>
</tr>
</thead>
<tbody>
<tr>
<td>At the time of sedation assessment the RASS score was ________</td>
</tr>
</tbody>
</table>
**Feature 4: Disorganized thinking**

Directions if patient can verbalize: “I am going to ask you 4 questions, say ‘yes’ or ‘no’ to answer”.

Directions to intubated patient: “I am going to ask you 4 questions, nod your head yes or no to answer”.

**Set A:**
1. Is sugar sweet?
2. Is ice cream hot?
3. Do birds fly?
4. Is an ant bigger than an elephant?

**Set B:**
1. Is a rock hard?
2. Do rabbits fly?
3. Is ice cream cold?
4. Is a giraffe smaller than a mouse?
5. Directions to patient: “Hold up this many fingers.” (Examiner hold up two fingers for patient to see)
   - Directions to patient: “Now do the same thing with the other hand.” (Do not show fingers again to patient)
   - Directions to patient if unable to move both arms: “Now, add one more finger.” (Do not show fingers again to patient)

**To score:**
- If the patient answers a question incorrectly, I will subtract 1 point.
- If the patient is not able to complete the command no. 5, I will subtract 1 point.

<table>
<thead>
<tr>
<th>Score</th>
<th>Paediatric delirium = Feature 1 + Feature 2 + either Feature 3 or Feature 4</th>
</tr>
</thead>
<tbody>
<tr>
<td>0-3</td>
<td>+ / -</td>
</tr>
</tbody>
</table>

If the SCORE is 0-3 then circle +
10.6 Procedure for assessing attention: attention screening exam (ASE) for adults

This procedure is to be used to assess for feature 2 (inattention - a cardinal feature of delirium), when the patient is unable to complete the letters attention test (SAVEHAART). This happens in only about 10% of patients.

**Step 1**
- Say to the patient: “Mr or Mrs ..., I am going to show you pictures of some common objects. Watch carefully and try to remember each picture because I will ask what pictures you have seen.”

- Present five pictures: naming them and showing them each for 3 seconds.

**Step 2**
- Say to the patient: “Now I am going to show you some more pictures. Some of these you have already seen and some are new. Let me know whether or not you saw the picture before by nodding your head yes (demonstrate) or no (demonstrate).”

- Present ten pictures (five new, five repeated): naming them and showing them each for 3 seconds.

**Scoring**
This test is scored by the number of correct “yes” or “no” answers during Step 2 (out of a possible 10).

⚠️ **Important:** Alternate daily between Forms A and B (see next tool) if repeat measures are taken. If a patient wears glasses make sure they have them on when attempting the ASE.

Source: Adapted from Ely and Vanderbilt University (2002).
Form A

![Form A with icons]

Source: Adapted from Ely and Vanderbilt University (2002).

Form B

![Form B with icons]

Source: Adapted from Ely and Vanderbilt University (2002).
10.7 Guide to commonly used sedatives in adults

There are many sedative medications available to treat agitation and anxiety. You will need to see which medications your hospital currently has and consider which medications you may want to use in the future. It is important to familiarize yourself with the basic pharmacokinetics and side-effects of any drug you use. The goal is to reach the sedation target with the lowest possible sedative medication to minimize toxicity. The doses provided below are intended to be used for adult patients who are intubated and receiving mechanical ventilation. Continuous infusions of benzodiazepines should be avoided in adult patients if safe alternatives are to reduce the risks of oversedation, prolonged days of IMV and delirium.

<table>
<thead>
<tr>
<th>Medication</th>
<th>Onset</th>
<th>Infusion</th>
<th>Time to arousal</th>
<th>Risks</th>
</tr>
</thead>
<tbody>
<tr>
<td>Propofol</td>
<td>&lt; 1 minute</td>
<td>10-80 µg/kg/min</td>
<td>10-15 minutes</td>
<td>Respiratory depression, Hypotension, Idiosyncratic rhabdomyolysis, cardiovascular collapse (Propofol Infusion Syndrome, PRIS), Raised triglycerides</td>
</tr>
<tr>
<td>Midazolam</td>
<td>1-5 minutes</td>
<td>0.04-0.2 mg/kg/hr</td>
<td>1-2 hours</td>
<td>Respiratory depression, Hypotension, Prolonged sedation with infusions due to active metabolite, Reduce dose in renal and liver failure, Delirium</td>
</tr>
<tr>
<td>Lorazepam</td>
<td>5-20 minutes</td>
<td>0.01-0.1 mg/kg/hr</td>
<td>2-6 hours</td>
<td>Respiratory depression, Hypotension, Propylene glycol carrier may irritate veins and cause metabolic acidosis with prolonged administration, Delirium</td>
</tr>
<tr>
<td>Diazepam</td>
<td>2-5 minutes</td>
<td>Not used</td>
<td>2-4 hours</td>
<td>Respiratory depression, Hypotension, Oversedation with repeated boluses with accumulation of drug and active metabolite, Delirium</td>
</tr>
<tr>
<td>Dexmedetomidine</td>
<td>1-3 minutes</td>
<td>0.2-1.5 µg/kg/hr</td>
<td>6-10 minutes</td>
<td>Hypotension, Bradycardia, More pronounced in elderly, Laryngospasm, Emergence delirium, Respiratory depression (especially in neonates or when used concurrently with opiates), Porphyria, Genitourinary symptoms</td>
</tr>
<tr>
<td>Ketamine</td>
<td>&lt; 1 minute</td>
<td>0.2-0.5 mg/kg/hr</td>
<td>1-2 hours</td>
<td></td>
</tr>
</tbody>
</table>

Note: 
* Less commonly available.

Early in severe ARDS, however, deep sedation targets may be needed to safely achieve LPV targets and reduce asynchrony. In cases when NMB are administered, remember to also give a continuous sedative for amnesia and analgesic for pain.
10.8 Guide to commonly used opioid analgesics in adults

There are several opioids available to treat pain. You will need to see which medications your hospital currently has and consider which medications you may want to use in the future. Familiarize yourself with the basic pharmacokinetics and side-effects of any drug you use. Be sure to set a therapeutic analgesia plan and communicate to all caregivers for a consistent approach.

These considerations are adapted from the Clinical practice guidelines for the sustained use of sedatives and analgesics in the critically ill adult (Jacobi et al., 2002) (see References and resources).

The doses provided below are suggestions and will need adjustment based on the amount of pain and whether the patient is receiving mechanical ventilation.

<table>
<thead>
<tr>
<th></th>
<th>Morphine</th>
<th>Hydromorphone</th>
<th>Fentanyl</th>
</tr>
</thead>
<tbody>
<tr>
<td>Intermittent dose IV</td>
<td>0.01-0.15 mg/kg every 1-2 hr</td>
<td>10-30 µg/kg every 1-2 hr</td>
<td>0.35-1.5 µg/kg every 0.5-1 hr</td>
</tr>
<tr>
<td>Infusion</td>
<td>0.07-0.5 mg/kg/hr</td>
<td>7-15 µg/kg/hr</td>
<td>0.7-10 µg/kg/hr</td>
</tr>
<tr>
<td>Half-life</td>
<td>3-7 hr</td>
<td>2-3 hr</td>
<td>1.5-6 hr</td>
</tr>
<tr>
<td>Equianalgesic IV dose⁴</td>
<td>10 mg</td>
<td>1.5 mg</td>
<td>200 µg</td>
</tr>
<tr>
<td>Situations where drug is preferred</td>
<td>Intermittent dosing</td>
<td>Intermittent dosing</td>
<td>Rapid onset in acutely distressed patients</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Haemodynamic instability</td>
<td>Haemodynamic instability</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Renal failure</td>
<td>Renal failure</td>
</tr>
<tr>
<td>Risks&lt;sup&gt;5&lt;/sup&gt;</td>
<td>Histamine release causing hypotension</td>
<td>Prolonged effect in renal failure due to metabolite</td>
<td>Rigidity with high doses</td>
</tr>
<tr>
<td></td>
<td></td>
<td></td>
<td>Repeated dosing may cause accumulation and prolonged effects</td>
</tr>
</tbody>
</table>

Notes:
⁴ These doses produce approximately the same analgesic effects;
<sup>5</sup> Side-effects common to all agents include respiratory depression, coma and delirium, hypotension (especially with morphine) and ileus.

![Warning]

Meperidine (pethidine), tramadol and codeine may be available at many hospitals; however, each drug has noteworthy limitations and thus are not optimal for critically ill patients. Meperidine has variable efficacy for analgesia, has an active metabolite that causes neuroexcitation (aprehension, tremors, delirium and seizures) and may interact with antidepressants (contraindicated with monoamine oxidase inhibitors and best avoided with selective serotonin-re-uptake inhibitors), so it is not recommended for repetitive use. Tramadol is generally considered inferior to many other analgesics and can cause adverse reactions when used with other serotonin re-uptake inhibiting medications. Codeine has markedly different pharmacodynamics across different patients due to variability in drug metabolizing rates.
10.9 Guide to using neuromuscular blockers in adults

WHO suggests that in moderate-severe ARDS (PaO₂/FiO₂ < 150), neuromuscular blockade by continuous infusion should not be routinely used.

A trial found that neuromuscular blockade improved survival in adult patients with severe ARDS without causing significant weakness (Papazian et al., 2010), but results of a recent larger trial found that use of neuromuscular blockade with high PEEP strategy was not associated with a survival benefit when compared with a light sedation strategy without neuromuscular blockade (NHLBI PCTN et al., 2019). Continuous neuromuscular blockade may still be considered in patients with ARDS, both adults and children, in certain situations: ventilator dyssynchrony despite sedation, such that tidal volume limitation cannot be reliably achieved; or refractory hypoxaemia or hypercapnia. Neuromuscular blockade may also be useful to ascertain accurate pulmonary mechanics (e.g. plateau pressure measurements), but this alone is not an indication for neuromuscular blockade infusion.

Suxamethonium (ie succinylcholine) is a depolarizing muscle relaxant used most commonly for rapid sequence induction for intubation. This medication can cause bradycardia (especially in infants or children < 5 years of age) and life-threatening hyperkalaemia. Because of this latter concern, suxamethonium is contraindicated in chronic skeletal muscle disease, denervating events or neuromuscular disease, recent burns, rhabdomyolysis, history of malignant hyperthermia or concurrent hyperkalaemia.

<table>
<thead>
<tr>
<th></th>
<th>Pancuronium</th>
<th>Vecuronium</th>
<th>Cisatracurium</th>
<th>Rocuronium</th>
</tr>
</thead>
<tbody>
<tr>
<td>IV dose</td>
<td>Intermittent: 0.08-0.1 mg/kg</td>
<td>Intermittent: 0.08-0.1 mg/kg</td>
<td>Intermittent: 0.15-0.20 mg/kg</td>
<td>Intermittent: 0.6-1.2 mg/kg</td>
</tr>
<tr>
<td></td>
<td>Infusion: 0.2-0.6 µg/kg/min (usually 1-2.5 mg/hr)</td>
<td>Infusion: 0.2-0.8 µg/kg/min (usually 1-4 mg/hr)</td>
<td>Infusion: 3 mcg/kg/min for first 20 minutes then reduce to 1-2 mcg/kg/min (range: 0.5-10 mcg/kg/min)</td>
<td>Infusion: 0.5-0.7 mg/kg/hr</td>
</tr>
<tr>
<td>Common points on dosing</td>
<td>Tailor intermittent dose to patient response. Titrate infusion dose clinically or to achieve one or two twitches with train of four stimulation on peripheral nerve stimulator, if available</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Onset</td>
<td>&lt; 4 minutes</td>
<td>2-3 minutes</td>
<td>2-3 minutes</td>
<td>2-3 minutes</td>
</tr>
<tr>
<td>Specific risks</td>
<td>Long duration of activity: ~90-160 minutes. Accumulation in hepatic and renal dysfunction. Dose-dependent increased HR and blood pressure (due to vagolytic and weak sympathomimetic effects)</td>
<td>Intermediate duration of activity: ~30-45 minutes. Accumulation in hepatic and renal dysfunction</td>
<td>Duration of action: ~45-75 minutes. Slight accumulation in hepatic and renal dysfunction</td>
<td>Intermediate duration of activity: ~30-45 minutes. Accumulation in hepatic and renal dysfunction. Can be reversed completely by sugammadex when available</td>
</tr>
<tr>
<td>Common risks</td>
<td>Appropriate sedation and analgesia should be administered concurrently. HR and blood pressure should be routinely monitored; increases may indicate inadequate sedation or analgesia. ICU-acquired weakness if used for prolonged period</td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>
10.10 Guide to commonly used antipsychotics (haloperidol) in adults

Antipsychotic agents can be used to keep actively delirious patients safer. Haloperidol is a typical antipsychotic that has been available for many years. Atypical antipsychotics can also be used (e.g. quetiapine, olanzapine and risperidone). Dexmedetomidine is a newer agent that has both sedative and anti-delirium effects.

### Haloperidol

| Loading dose | Begin with 2-5 mg IV  
|              | Double dose every 15 minutes until desired effect is achieved  
|              | Do not exceed total of 20 mg/day |
| Onset        | 10-20 minutes |
| Risks        | Torsade de pointes arrhythmia, do not use if the QTc interval on ECG is prolonged to > 460 milliseconds  
|              | Suspect neuroleptic malignant syndrome if patient develops hyperthermia, muscle rigidity and rhabdomyolysis |

### Dosing recommendations

| Quetiapine   | Begin with 25-50 mg po twice or three times daily  
|              | Increase up to 200 mg po total daily dose (halve dose in elderly) |
| Olanzapine   | Begin with 5-10 mg IV/IM/po  
|              | Repeat dose in 2 hours to maximum of 30 mg/day |
| Risperidone  | Begin with 1-2 mg po daily  
|              | Increase to maximum of 6 mg po daily |

⚠️ Side-effects of atypical antipsychotics are prolonged QTc interval and extrapyramidal effects (less common than with typical antipsychotic agents).
10.11 Guide to paediatric analgesics, sedatives and neuromuscular blockers

There are several agents available for analgesia, sedation and neuromuscular blockade. You will need to see which medications your hospital currently has and consider which medications you may want to use in the future. Familiarize yourself with the basic pharmacokinetics and side-effects of any drugs you use. The doses provided below are suggestions and will need titration in individual patients based on the amount of pain and whether the patient is receiving mechanical ventilation. Appropriate sedation and analgesia should be administered concurrently with neuromuscular blockade, which has no sedative or analgesic properties.

Propofol is contraindicated for sedation in children < 16 years old in the ICU because of the risk of propofol infusion syndrome (acidosis and rhabdomyolysis).

<table>
<thead>
<tr>
<th>Drug</th>
<th>Enteral dose</th>
<th>Bolus IV dose</th>
<th>IV infusion</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Analgesia</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Paracetamol</td>
<td>10-15 mg/kg/6hr</td>
<td>10mg/kg q6-8hr</td>
<td>N/A</td>
</tr>
<tr>
<td>Oxycodone</td>
<td>0.05-0.2 mg/kg/4-6hr</td>
<td>N/A</td>
<td>N/A</td>
</tr>
<tr>
<td>Ibuprofen</td>
<td>5-10 mg/kg/6-8hr</td>
<td>N/A</td>
<td>N/A</td>
</tr>
<tr>
<td>Morphine</td>
<td>0.2-0.4 mg/kg/6hr</td>
<td>0.1-0.2 mg/kg</td>
<td>0-40 µg/kg/hr</td>
</tr>
<tr>
<td>Fentanyl</td>
<td>N/A</td>
<td>1-2 µg/kg</td>
<td>0-8 µg/kg/hr</td>
</tr>
<tr>
<td><strong>Sedation</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Hydromorphone</td>
<td>0.03-0.06 mg/kg/4hr</td>
<td>0.01-0.015 mg/kg</td>
<td>0.003-0.005 mg/kg/hr</td>
</tr>
<tr>
<td>Midazolam</td>
<td>N/A</td>
<td>0.05-0.2 mg/kg</td>
<td>0-4 µg/kg/min</td>
</tr>
<tr>
<td>Diazepam</td>
<td>N/A</td>
<td>0.1-0.2 mg/kg</td>
<td>N/A</td>
</tr>
<tr>
<td>Chloral hydrate</td>
<td>30-50 µg/kg/6hr</td>
<td>N/A</td>
<td>N/A</td>
</tr>
<tr>
<td>Triciofos</td>
<td>30-50 µg/kg/6hr</td>
<td>N/A</td>
<td>N/A</td>
</tr>
<tr>
<td>Allmazene</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
</tr>
<tr>
<td>Ketamine</td>
<td>6-8 mg/kg</td>
<td>0.5-2 mg/kg</td>
<td>0.3-1.22 mg/kg/hr</td>
</tr>
<tr>
<td>Dexmedetomidine</td>
<td>N/A</td>
<td>1 mcg/kg over 10 minutes</td>
<td>0.2-0.7 mcg/kg/hr</td>
</tr>
<tr>
<td>Clonidine</td>
<td>4-5 mcg/kg</td>
<td>1-2 mcg/kg</td>
<td>0.18-3.16 mcg/kg/hr</td>
</tr>
<tr>
<td><strong>Neuromuscular blockade</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Vecuronium</td>
<td>N/A</td>
<td>0.1 mg/kg as required</td>
<td>0-4 µg/kg/min</td>
</tr>
<tr>
<td>Cisatracurium</td>
<td>N/A</td>
<td>0.1-0.15 mg/kg</td>
<td>0.5-10 µg/kg/min</td>
</tr>
<tr>
<td>Rocuronium</td>
<td>N/A</td>
<td>0.6 mg/kg</td>
<td>0.12 µg/kg/min</td>
</tr>
<tr>
<td>Pancuronium</td>
<td>N/A</td>
<td>0.04-0.1 mg/kg</td>
<td>0.1 mg/kg/h</td>
</tr>
<tr>
<td>Suxamethonium</td>
<td>N/A</td>
<td>1-2 mg/kg</td>
<td>0.3-0.6 mg/kg q5-10min PRN</td>
</tr>
</tbody>
</table>
References and resources


Liberation from invasive mechanical ventilation
Liberation from invasive mechanical ventilation

Summary

Use a daily coordinated spontaneous awakening trial (SAT) with spontaneous breathing trial (SBT) protocol to liberate patients from mechanical ventilation as soon as possible as this improves patient outcomes. In patients who fail SBT, recognize and treat the reason for failure, and try again the next day. In patients who pass SBT, consider extubation after evaluating their ability to protect the airway and clear secretions. After extubation, monitor the patient closely for signs of respiratory failure and the need for prompt re-intubation.

Steps for liberation from invasive mechanical ventilation (IMV)

<table>
<thead>
<tr>
<th>Step</th>
<th>Task</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>Daily assessment for patient readiness to breath spontaneously</td>
</tr>
<tr>
<td>2</td>
<td>Conduct the SBT safely</td>
</tr>
<tr>
<td>3</td>
<td>Evaluate patient’s performance on the SBT</td>
</tr>
<tr>
<td>4</td>
<td>Assess safety for extubating</td>
</tr>
<tr>
<td>5</td>
<td>Extubate</td>
</tr>
<tr>
<td>6</td>
<td>Monitor-record-interpret-respond</td>
</tr>
<tr>
<td>7</td>
<td>Deliver quality care: implement as part of ABCDEF protocol</td>
</tr>
</tbody>
</table>

There are limited data to guide the timing of tracheostomy, though routinely this is considered at 10-14 days if the need for prolonged mechanical ventilation persists. During the COVID-19 pandemic, the optimal timing of tracheostomy has been debated. These debates have included arguments to perform tracheostomy as early as possible to assist with early mobilization, reduced sedation requirements, easier pulmonary hygiene and less potential risk for tube malposition. The debates have also included arguments to perform tracheostomy considerably later than routine in order to decrease the chance of health care worker exposure to aerosolized particles during the procedure.

Tools

11.1 Algorithm for daily sedation interruption and daily spontaneous breathing trial (SBT)
11.2 How to perform a cuff leak test
11.3 Respiratory care pocket card reference
11.4 Spontaneous breathing trial (SBT) order set
11.1 Algorithm for daily sedation interruption and daily spontaneous breathing trial (SBT)

This protocol outlines the steps for daily consideration of sedation interruption, spontaneous breathing trial, and extubation - modified by the OpenCriticalCare.org Project from Girard et al. (2008) and MacIntyre (2007).

Source: USAID-STAR-UCSF Open Critical Care Project (Tool 11.2)
11.2 How to perform a cuff leak test

Performing a cuff leak test may help to determine that the airway is patent, without swelling.

*When an adequate cuff leak is present*, it signifies that the patient’s airway is unlikely to be profoundly edematous, and the patient is less likely to have post-extubation stridor (PES). Of note, studies of the predictive value of cuff leak tests vary, and generally the cuff leak test is considered to have higher specificity than sensitivity. The presence of a cuff leak does not ensure that the patient’s airway will remain open after extubation.

*When an adequate cuff leak is not present, consider steroids and elevating the head of the bed to reduce swelling; check again in 12–24 hours.*

The lack of a cuff leak makes it more likely that the patient could have post-extubation stridor if steroids are not given prior (Kuriyama et al., 2020; Miller and Cole, 1996).


<table>
<thead>
<tr>
<th>Risk factors for post-extubation stridor</th>
</tr>
</thead>
<tbody>
<tr>
<td>neck trauma</td>
</tr>
<tr>
<td>prone positioning</td>
</tr>
<tr>
<td>anasarca</td>
</tr>
<tr>
<td>multiple recent intubations</td>
</tr>
<tr>
<td>intubation for &gt; 6 days</td>
</tr>
</tbody>
</table>

**Protocol to perform a cuff leak test**

1. Ensure proper PPE and IPC for providers as the leak test is an aerosol-generating procedure.
2. Patient should be sedated, or the test done when the patient is asleep (to prevent measurement artifacts).
3. Standard vent settings; volume-controlled ventilation with:
   - TV: 8–10 mL/kg
   - RR: 10–12
   - flow rate: 50–60 L/min.
4. Suction oropharynx (to prevent secretions from passing into the trachea when the ETT cuff is deflated).
5. Measure the expired TV.
6. Deflate the ETT cuff.
7. Re-measure the expired TV over six breaths:
   - cuff leak is the difference in TV with cuff inflated and deflated;
   - > 110 mL difference in expiratory TV before and after cuff down has negative predictive value (NPV) 98% for post-extubation stridor.
8. Reinflation of the cuff.
Cuff leak protocol

Cuff leak test necessity screen
Patient meets criteria for SBT* and ≥ 1 of the following:
- Traumatic intubation
- Intubation more than 6 days
- Large endotracheal tube (ETT)
- Female sex
- Multiple recent intubations
- Other clinical concern for airway swelling (e.g. prone positioning or anasarca)

Prepare patient for cuff leak test
- Ensure patient is synchronous with ventilator and adequately sedated (a transient RASS goal of -4 should be considered)
- Set ventilator to volume control mode with tidal volume (TV) 8–10 mL/kg predicted body weight
- Suction oropharynx to prevent oral secretions from passing into the trachea with cuff deflation

Perform the cuff leak test
- Observe and record the measured expired TV
- Deflate the ETT cuff by attaching syringe to the pilot balloon and pulling back on syringe until the balloon is flat
- Observe the measured expired TV on the vent over six breaths

Difference between expired TV before and after cuff deflation is > 110 mL

Cuff leak test has been PASSED
- Proceed to extubation if all other parameters in place as in Tool 11.1

Difference between expired TV before and after cuff deflation is < 110 mL

Cuff leak test has been FAILED
- Address potential causes of failure
- Consider dexamethasone 5–10 mg IV q6–12h x 2 doses and then reassess patient for extubation in 12–24h
- Elevate head of bed

*SBT – spontaneous breathing trial criteria (see Tool 11.1).

Source: USAID-STAR-UCSF Open Critical Care Project ( 
# 11.3 Respiratory care pocket card reference

The card, which can be printed or saved to your mobile device, is available in:
- English
- French
- Portuguese
- Spanish.

## High Pressures, Desaturations & Dyssynchrony

### General Considerations
- Is the ventilator set to FiO₂ 100% while you troubleshoot?
- Is the patient hemodynamically unstable? If so, consider pressets & urgent evaluation for pneumothorax or severe auto-PEEP.
- Is the ventilator circuit connected & set correctly?
- Are breath sounds bilateral? If unilateral, consider mainstem ETT, lobar collapse, atelectasis, and pneumothorax.

### High Pressures
- Is the patient agitated or asynchronous?
- Can a suction catheter pass easily through the endotracheal tube? If no, consider kinked tube, biting/bite block, secretions/suction, or replace ETT.
- What is the Pplat?

### Troubleshooting

**Low Compliance:**
- Decrease V̇ & evaluate for breath stacking (auto-PEEP).
- Consider ↓ PEEP or adjunct therapies for hypercapnia if decrease in PEEP causes desaturations.

**High Resistance:**
- Work outside (machine) to inside (alveoli): circuit problem, ETT kink/occlusion/biting, ETT obstructed/mainstem, large airway obstruction (mucus plug), small/medium airway obstruction (bronchospasm); auscultation & passing a suction catheter can quickly eliminate many of these.

### Desaturations
- Is the endotracheal tube in good position? (consider CXR)
- Is the pulse oximetry waveform good quality?
- Is there acute hypotension? Evaluate for tension pneumothorax, air trapping and pulmonary embolism.
- Is the source of oxygen faulty or is there an air leak? Check each connection/element sequentially from source of oxygen to the patient.
- Are there concomitant pressure elevations? If so, see “High Pressures” (next column)
- Is P:F <150 in setting of worsening ARDS? If so, consider adjunct therapies (above).
- Are there signs of infection? Consider ventilator-associated pneumonia.

### Patient-Ventilator Dyssynchrony
- Detect signs of dyssynchrony: coughing, paradoxical breathing, ventilator alarms (low tidal volumes or high pressures), breath stacking.
- Treat underlying causes: 1) Ineffective triggering (patient or ventilator); 2) Inappropriate triggering (patient inspires while ventilator expires); 3) Auto-triggering (non-respiratory muscle movement triggers ventilator); 4) Flow dyssynchrony (too fast or too slow)

### General Approach
- Treat pain or anxiety if present
- If ineffective ventilator triggering change trigger sensitivity, decrease VT or pressure
- Increase V̇, to 8 mL/kg and increase flow rate if pressures allow. Consider change to decelerating flow delivery if setting available.
- If still dysynchronous, paralyze patient (and sedate patient to RASS goal -5)
**Discomfort (pain, agitation, anxiety) & Delirium**

- Ensure appropriate analgesia & sedation to minimize ICU/IMV duration & risk of long-term neuropsychiatric impact.
- Reassess every 24 hours using a standardized scale
  - Goal RASS: -4 to 0 in intubated patients
  - If RASS > -4 for >5 consider lightening sedation unless patient is paralyzed or dysynchronous with ventilator.
  - Target RASS: -5 for paralyzed patients. Never paralyze without sedation.
- Perform Daily Sedation Interruptions (DSIs) in non-paralyzed patients to reassess sedation & analgesia needs, which can guide weaning these medications.

**Delirium:** Prevention & treatment of delirium reduces mortality and ICU/IMV duration.

- Screen every 12h using standardized tool (i.e. CAM-ICU)
- Treat delirium by addressing underlying causes (pain, agitation, anxiety, or physiologic derangements)

**Ventilator Weaning & Extubation**

**Spontaneous Breathing Trial (SBT) Initiation Criteria & Considerations**

- Patient likely to trigger ventilator, stable FiO₂ ≤ 0.50 adults (≤ 0.40 pediatrics) and PEEP ≥ 8
- pH = 7.30, VE < 15 L/min
- MAP ≥ 60 mmHg (minimal pressors)
- ICP: non-labile and < 20 mmHg w/ CPP > 60 mmHg
- No MI in previous 48hr

**Weaning Strategies**

- Once daily SBT PS ∆≤ 7 cmH₂O if ETT size ≥ 5 (ø 8 cmH₂O) if ETT 4-4.5; ∆> 10 cmH₂O if ETT 3-3.5 in addition to PEEP ≥ 8 cmH₂O (2nd daily trial ok if failure sedation-related or due to transient issue); Consider additional Δ 3-5 cmH₂O of PS if HNE used
- SBT ≥ 30min - probably as good as SBT ≥ 2hr or longer for most adult patients
- If adult with cardiogenic pulmonary edema risk: Consider 15min T-piece (i.e. no PS or PEEP)
- RSI (rapid Shallow Breathing Index) = f/V; RSI is unreliable; <0 goal for extubation; sensitive, not specific (if >10; good predictor of failure)
- Coupling daily sedation interruption, early mobility, & SBT is associated with faster extubation

**Exubation Criteria**

- Have you fixed the original problem and no upcoming procedures?
- Adequate oxygenation? (PaO₂ > 60 on PEEP ≥ 8 cmH₂O, FiO₂ ≤ 0.50)
- Adequate ventilation without excessive work of breathing? (PaCO₂ Δ of < 10 mmHg with remaining pH ≥ 7.30 during SBT)
- Secrecions? (assess cough strength, suction frequency & secretion volume)
- Airway protection? (assess gag, spontaneous cough and GCS)
  - Assess risk of post-extubation airway obstruction:
    - Consider cuff leak test if: intubation >6d, trauma, multiple intubations, prolonged prone, flat, volume overload, head/neck trauma, among others
- Cuff Leak Test: 1. pt must be sedated (interaction with vent = incr PIP, incr leak = false reassurance); 2. Suction oropharynx; 3. Initiate AC-VC V, 8 mL/kg, RR 12, Ti: 1.5sec, Flow 50 LPM; 4. Measure expired V; 5. Deflate cuff and wait 6 breaths & Measure expired V, (should be +110mL to predict low risk of stridor post extubation); 7. Reinflate cuff
  - Decrease aspiration risk by holding tube feeds for safe interval (~4-8h)
  - Extubation criteria/goals for neuro patients may be different (e.g. visual tracking, swallowing, GCS>10, <40yo)
- Hemodynamics - re-intubation of an unstable patient can be lethal

Source: USAID-STAR-UCSF Open Critical Care Project.
11.4 Spontaneous breathing trial (SBT) order set

Spontaneous Breathing Trial (SBT) Order Set

- SBT initiation
  - FIO2 ≤ 0.50 and PEEP ≥ 5 or CI ≤ 4.0 in pediatrics patients
  - No No. 1 in PS or FIO2 requirements over past 24 hrs.
  - >7.30 and no significant INR required

Tube Feeds During SBT: [ ] Continue Tube Feeds

- Standard SBT Strategy:
  - Daily SBT until patient is extubated, or until at least 3 consecutive failed trials (see below).
  - If failure is sedation related can repeat SBT later on same day
  - When possible, coordinate with Daily Sedation Interruption/Spontaneous Awakening Trial (adults only)
  - Suspend SBT on days when weaning readiness criteria are not met (see protocol for details)
  - May withhold SBT for pending special procedures/surgery

Mode:
- Default if ETT size > 5mm, Pressure Support (PS) = delta 5-7 cmH2O above PEEP of 5-8 cmH2O; FIO2 ≥ 0.50
- Default if ETT size 4-4.5mm, Pressure Support (PS) = delta 8 cmH2O above PEEP of 5 cmH2O; FIO2 ≥ 0.50
- Default if ETT size 3-3.5mm, Pressure Support (PS) = delta 10 cmH2O above PEEP of 5 cmH2O; FIO2 ≥ 0.50
- T-piece Trial (Consider if resolving cardiac pulmonary edema - Adults only)

PS = [ ]
PEEP = [ ]
FIO2 = [ ]

Duration: [ ] 30 minutes [ ] 2 hours

- Initial intolerance of SBT: PS can be titrated down from PS of 15-20 to SBT settings over 30 min.
- Evaluation: RT at bedside for initial 5 min until a 30 min SBT is passed. Full Vent check @ 30 min or 2 hrs.
- Failure Criteria: MTT: Return to previous ventilator settings (or new written orders for post-SBT). When ARDS Net protocol is in place resume previous ARDS Net settings.

After SBT Passed:
- [ ] Notify ICU team to evaluate for extubation
- [ ] Extended SBT as tolerated
- [ ] Return to prior settings

When Tracheostomy tube is present:

Follow orders as written above. When patient passes a 2 hr SBT, then:
- [ ] Extend SBT as tolerated if failure criteria met, then return to prior settings
- [ ] Tracheostomy Mask Trials at FIO2 ≤ 0.50 as tolerated. If failure criteria met, then return to SBT PS settings. If these PS settings are also not tolerated then return to 0.50 or newly ordered vent settings.

Difficult to Wean:

PS Range: delta 5-5.5 cmH2O above PEEP 5-8 cm H2O; FIO2 ≤ 0.50

Titrate PS as needed to maintain spontaneous RR 25-35 (1.5x age adjusted normal) and VT 5-8 ml/kg

Daily Routine: After 2 hrs of stable breathing pattern at a stable PS level, then:
- [ ] PS Decrement challenge: reduce by 3.5 cmH2O every 2-4 hrs
- [ ] Trial Duration: not to exceed [ ] hrs/day
- [ ] Notify ICU team when patient passes 2 hr SBT at standard PS settings to evaluate for extubation

Cuff Leak Check:

Consider if: intubation > 6 days, neck trauma, multiple intubations, prolonged prone positioning, volume overload
- [ ] Perform cuff leak PRIOR TO SBT
- [ ] No cuff leak necessary

Alternative approach (pediatrics):
- 1. PS must be sedated (interaction with vent can incr PIP which can incr breath - false reassurance)
- 2. Suction oropharynx
- 3. Initiate AC-VC VT 8 ml/kg, RR 12, TI 1.5sec, Flow 50 LPM
- 4. Measure expired VT
- 5. Deflate cuff and wait 8 breaths
- 6. Measure expired VT expiratory VT (goal is [ ] by >110 mL in measure expired VT for adults)
- 7. Re-inflate cuff

Date (time): [ ] Name: [ ] Signature: [ ] Contact #: [ ]

Source: USAID-STAR-UCSF Open Critical Care Project (148)
References and resources


11. Liberation from invasive mechanical ventilation 247
12 Best practices to prevent complications
12 Best practices to prevent complications

Summary

ICU patients are at high risk for complications. This chapter provides tools for implementation of key interventions to reduce the risk of complications in the ICU which include:

Tools

12.1 Interventions to prevent complications in hospitalized and critically ill patients with COVID-19
12.2 Checklist for central venous catheter (CVC) insertion
12.3 Checklist for preventing ventilator-associated pneumonia (VAP)
12.4 Checklist for preventing urinary tract infections (UTI)
12.5 Procedure for providing enteral nutrition (EN) for adults
12.6 Procedure for providing enteral nutrition (EN): paediatric considerations
12.7 Algorithm for early mobility in the ICU
12.8 Thromboembolic prophylaxis in COVID-19
12.9 ABCDE bundle

Notes: CRBSI - catheter-related bloodstream infection; UTI - urinary tract infection; VAP - ventilator-associated pneumonia; VTE - venous thromboembolism.
### 12.1 Interventions to prevent complications in hospitalized and critically ill patients with COVID-19

See WHO COVID-19 Clinical management: living guidance (2021)

<table>
<thead>
<tr>
<th>Anticipated outcome</th>
<th>Interventions</th>
</tr>
</thead>
</table>
| Reduce days of invasive mechanical ventilation          | • Use weaning protocols that include daily assessment for readiness to breathe spontaneously  
• Minimize continuous or intermittent sedation, targeting specific titration endpoints (light sedation unless contraindicated) or with daily interruption of continuous sedative infusions  
• Early mobilization  
• Implementation as a bundle of care |
| Reduce incidence of ventilator-associated pneumonia (VAP) | • Oral intubation is preferable to nasal intubation in adolescents and adults  
• Keep patient in semi-recumbent position (head of bed elevation 30-45°)  
• Use a closed suctioning system; periodically drain and discard condensate in tubing  
• Use a new ventilator circuit for each patient; once patient is ventilated, change circuit if it is soiled or damaged, but not routinely  
• Change heat moisture exchanger when it malfunctions, when soiled, or every 5-7 days |
| Reduce incidence of catheter-related bloodstream infection (CRBSI) | • Use a checklist with completion verified by a real-time observer as a reminder of each step needed for sterile insertion and as a daily reminder to remove catheter if no longer needed |
| Reduce incidence of pressure ulcers                      | • Turn patient every 2 hours |
| Reduce incidence of stress ulcers and GI bleeding       | • Give early enteral nutrition (within 24-48 hours of admission)  
• Administer histamine-2 receptor blockers or proton-pump inhibitors in patients with risk factors for GI bleeding. Risk factors for GI bleeding include:  
  – mechanical ventilation for ≥ 48 hours  
  – coagulopathy  
  – renal replacement therapy  
  – liver disease  
  – multiple comorbidities  
  – higher organ failure score |
| Reduce the development of antimicrobial resistance       | Utilize de-escalation protocols as soon as patient is clinically stable and there is no evidence of bacterial infection |
| Reduce the development of adverse drug effects           | Expose patient to empiric antimicrobial therapy for the shortest time possible, to prevent nephrotoxicity, cardiac and other side-effects from unnecessary antimicrobial use |
| Promote appropriate antimicrobial prescribing and use during the COVID-19 pandemic | Do not prescribe antibiotics to suspected or confirmed COVID-19 patients with low suspicion of a bacterial infection, to avoid more short-term side-effects of antibiotics in patients and negative long-term consequences of increased antimicrobial resistance |
12.2 Checklist for central venous catheter (CVC) insertion

☑ In the literature, a research collaborative found that using a central line checklist as a reminder for the inserter significantly reduced the incidence of central venous catheter-related bloodstream infections. This checklist is adapted from An intervention to decrease catheter-related bloodstream infections in the ICU (Provonost et al., 2006).

☐ Patient verification, allergies and informed consent.

☐ Hand hygiene before the procedure.

☐ Wear maximal barrier precautions on insertion: full
  ☐ sterile gown
  ☐ face mask
  ☐ face shields
  ☐ sterile gloves
  ☐ hair cover
  ☐ cover the patient in a full sterile sheet from head to toe.

☐ Use chlorhexidine 2% in 70% isopropyl alcohol for skin preparation and apply in a back and forth friction rub motion for 30 seconds.

☐ Let dry completely before puncturing site. It should not be blotted dry.

☐ Choose the optimal site: subclavian or internal jugular vein preferred in adults; internal jugular or femoral vein preferred in children depending on age.

☐ Once in place, evaluate the continuing need for the central line on a daily basis.

☐ Remove line immediately when no longer needed or when non-functional.
12.3 Checklist for preventing ventilator-associated pneumonia (VAP)

In order to prevent VAP, a complication of endotracheal intubation and invasive mechanical ventilation, consider the following procedures, when possible:

☐ Oral intubation instead of nasal intubation.

☐ Keep the patient in a semi-recumbent position (head of bed elevated up to ≥ 30–45°).

☐ Use a closed suctioning system.

☐ Periodically drain and remove condensation in tubing.

☐ Use a new ventilator circuit for each patient. Change if soiled or damaged.

☐ Do not routinely change ETT or ventilator circuit, only if they malfunction.

☐ Change heat and moisture exchanger when malfunctions, soiled, or every 5-7 days.

☐ Perform regular antiseptic oral care with chlorhexidine gel or mouthwash.

☐ Discontinue invasive ventilation in a safe and prompt manner: Daily
  ☐ sedation interruption of continuous sedative infusions. Daily evaluation for SBT readiness (see Chapter 11).
  ☐ Extubation to non-invasive ventilation when appropriate (i.e. primarily for patients ventilated because of a COPD exacerbation, and only in centres with sufficient expertise in non-invasive ventilation).
  ☐ ABCDE bundle.

Note: Heat and moisture exchangers (HME) are not routinely used in infants and small children as they significantly increase dead space. Use heated humidifiers instead.

When HMEs are used, ensure the size of the selected filter is appropriate for the size of the patient to minimize dead space and calculate/account for increased dead space in the circuit when choosing ventilator settings (and target minute ventilation).
12.4 Checklist for preventing urinary tract infections (UTI)

☑ Prevention of UTI requires an appropriate technique for catheter insertion as well as appropriate management of indwelling catheters. Consider the following procedures when possible:

**Catheter insertion**
- Insert catheter only when necessary.
- Hand hygiene before procedure.
- Use aseptic technique and sterile equipment.
- Use as small a catheter as possible, consistent with proper drainage.

**Catheter management**
- Maintain unobstructed urine flow.
- Empty collection bag regularly:
  - Separate collecting container for each patient.
  - Do not allow draining spigot to touch collecting container.
- Keep collecting bag below level of bladder at all times.
- Cleaning urethral meatus with antiseptic is unnecessary. Routine cleaning is adequate.
- Secure catheter to prevent movement and urethral traction.
- Sterile, continuously closed drainage system.
  - Do not disconnect catheter and drainage tube unless catheter must be irrigated.
  - Replace collecting system aseptically and after disinfecting catheter-tubing junction if the following occur:
    - break in aseptic technique
    - disconnection
    - leakage.
- Daily check and remove as soon as there is no indication.
12.5 Procedure for providing enteral nutrition (EN) for adults

The goal is to start enteral nutrition, even in small volumes, as soon as the patient is stable.

This tool can be used to start enteral nutrition.
1. Place a feeding tube.
2. Confirm placement with radiograph (gastric [NG] or small bowel [NJ] feeding are acceptable).
3. Once the feeding tube has been confirmed, start with an infusion of up to 30 mL/hr of clear fluid or feed.
4. Aspirate the NG tube every 4 hours.
5. Gradually increase the volume of feed with the aim of building up to full feeding within 48 hours.

Feeding intolerance
Intolerance of feeding may result from poor gastric emptying and lead to high residual gastric volumes.

The absolute value that is too high and should prompt cessation of tube feeds is not clear.

Stop feeding when:
- volumes high (between 250-500 mL)
- clinical signs of intolerance (abdominal pain, abdominal distension and diarrhoea).

None of the features are specific for feed intolerance.

Possible treatments include advancing the feeding tube into the small bowel (can be done at the bedside) or adding prokinetic medications (e.g. metoclopramide intravenously).

Note: With an NJ tube, only continuous feeds can be delivered (no bolus) and residuals cannot be checked.

Set calorific target and aim to reach this within a few days

Estimate the patient’s daily caloric needs, or basal energy expenditure (BEE). Adjust for fever and stress:
- BEE (kcal/day) = 25 × body weight (kg)
- fever: BEE × 1.1 (for each degree above the normal body temperature)
- mild to moderate stress: BEE × 1.2-1.4
- moderate to severe stress: BEE × 1.4-1.6.

Estimate your patient’s daily protein requirements:
- normal 1.2-2.0 g/kg
- hypercatabolism: 2-3 g/kg
- ratio of non-protein calories to nitrogen (70:1-100:1).

Note: Hypocaloric feeding (40-60% of non-protein caloric needs) may be as beneficial as full caloric feeds (> 70%).

Note:
- With an NJ tube, only continuous feeds can be delivered (no bolus) and residuals cannot be checked.
12.6 Procedure for providing enteral nutrition (EN): paediatric considerations

Enteral feeding via NG tube is the preferred method of providing maintenance fluid.


Initial fitting
1. Measure the distance from the nose to the ear and then to the epigastrium.
2. Insert NG tube to the measured distance.
3. Check correct placement of tube:
   - check the pH of aspirate using pH indicator strips
   - position can be seen on chest X-ray
   - if in doubt remove and replace.
4. Secure the NG tube by taping to the cheek avoiding upwards pressure on the nares.
5. Once correct placement has been confirmed, flush the tube with water. It is now safe to use the tube for administration of feed and medication.
6. Flush the NG tube with sterile water after administration of NG drugs otherwise it will block.

Ongoing checks
Check the position of the NG tube:
- before each use
- every 6 hours if continuous feeds
- after episodes of vomiting or retching, increased respiratory distress or excessive coughing
- if the tube looks dislodged (i.e. with more tubing visible).

NG tube sizes
This is only a rough guide; the bore of tube must fit easily in the child’s nostril.

<table>
<thead>
<tr>
<th>Description of patient</th>
<th>Tubes sizes</th>
</tr>
</thead>
<tbody>
<tr>
<td>&lt; 2 kg, preterm</td>
<td>4 Fr</td>
</tr>
<tr>
<td>2–4 kg</td>
<td>6 Fr</td>
</tr>
<tr>
<td>Term to 1 year</td>
<td>8 Fr</td>
</tr>
<tr>
<td>Younger children</td>
<td>10 Fr</td>
</tr>
<tr>
<td>Older children and adolescents</td>
<td>12 Fr</td>
</tr>
<tr>
<td>Small adult</td>
<td>14 Fr</td>
</tr>
<tr>
<td>Large adult</td>
<td>16 Fr</td>
</tr>
</tbody>
</table>
12. Best practices to prevent complications

- **Check NG tube position**

- **Hourly bolus feeds** - consider continuous pump for large hourly volumes

**Aspirate every 4 hours**

- **No aspirate**
  - **Continue**
  - **No aspirate**
  - **Abdominal distension**
    - **Stop feeds**
    - **Medical review**
  - **Aspirate**
    - **Discard aspirate**

- **Aspirate**
  - **Replace aspirate (max. 4 hours)**, continue at same rate and reassess after 4 hours
  - **No abdominal distension**
    - **Reduce rate by 50%** and reassess at 4 hours
12.7 Algorithm for early mobility in the ICU

An adapted early mobility algorithm is presented below. It is adapted from Balas et al. (2014).

The patient’s level of consciousness will be determined prior to the daily physical rehabilitation session using the Richmond Agitation-Sedation Scale.

- **A patient who is only arousable to physical stimulation** (RASS -4/-5) will undergo passive range of motion (ROM) exercises.

- **Once a patient can open their eyes to voice** (RASS -2/-3), passive ROM exercises will be performed, and the patient will be placed in the chair position in bed.

- Finally, **once a patient is alert and calm** (RASS -1/+1), they will progress from active ROM up through ambulation as they are able.

Sessions will continue until hospital discharge or a patient meets certain functional milestones. Early mobility in select intubated patients is feasible when adequate staff are available and safety assessment has been completed.

**Physical rehabilitation protocol**

![Algorithm diagram](image)
12.8 Thromboembolic prophylaxis in COVID-19

Coagulopathy is common in patients with severe COVID-19, and both venous and arterial thromboembolism have been reported.

Monitor patients with COVID-19 for signs or symptoms suggestive of thromboembolism, such as stroke, deep vein thrombosis, pulmonary embolism or acute coronary syndrome. If these are clinically suspected, proceed immediately with appropriate diagnosis and management pathways.

In hospitalized patients with COVID-19 without an established indication for higher dose anticoagulation, WHO recommends to administer standard thromboprophylaxis dosing of anticoagulation (rather than therapeutic or intermediate dosing).

<table>
<thead>
<tr>
<th>Type of heparin</th>
<th>Dose</th>
<th>Comments</th>
</tr>
</thead>
<tbody>
<tr>
<td>Enoxaparin</td>
<td>40 mg/day SC</td>
<td>Clinical observation if &lt; 45 kg (women) or &lt; 57 kg (men); risk of bleeding</td>
</tr>
<tr>
<td></td>
<td></td>
<td>BMI &gt; 40 mg/m² or weight &gt;120 kg; 40 mg/12h SC</td>
</tr>
<tr>
<td>Unfractionated heparin (UFH)</td>
<td>5000 units/12h or 8h SC</td>
<td>BMI &gt; 40 mg/m² or weight &gt;120 kg; 7500 units/12h or 5000 units/8h</td>
</tr>
<tr>
<td>Tinzaparin</td>
<td>4500 units/day SC</td>
<td>BMI &gt; 40 mg/m² or weight &gt;120 kg; 9000 units/day</td>
</tr>
<tr>
<td>Dalteparin</td>
<td>5000 units/day SC</td>
<td>BMI &gt; 40 mg/m² or weight &gt;120 kg; 5000 units/12h</td>
</tr>
<tr>
<td>Fondaparinux</td>
<td>2.5 mg/day SC</td>
<td></td>
</tr>
</tbody>
</table>

Notes:
- SC - subcutaneous injection.
- In renal failure, reduce dose of low molecular weight heparin (LMWH) (except dalteparin).
- Patients on thromboprophylaxis dosing of anticoagulation do not require monitoring, except for platelet count monitoring after 5-7 days if UFH is used.
- Dosing should be adjusted according to body weight/BMI and renal function according to local protocols.

Use mechanical prophylaxis if moderate VTE risk, in patient at risk of bleeding:
- intermittent pneumatic compression devices;
- graduated compression stockings.

Switch to pharmacologic agent once risk of bleeding has decreased.
12.9 ABCDE bundle

This algorithm is adapted from Balas et al. (2014). Implementing this bundle reduced the time patients spent on invasive mechanical ventilation by 3 days. Additionally, patients experienced less delirium and were more likely to be mobilized.

Adapt this bundle to fit your ICU and implement using a quality improvement mechanism.

**ABCDE bundle algorithm**

**AWAKENING:** Applied daily to patient receiving continuous sedative medications and mechanical ventilation - RN driven

- Every 24 hours
  - SAT safety screen
  - Perform SAT
  - If SAT fails, perform SAT again
  - COORDINATION: Prompt performance of SBT safety

**BREATHING:** Applied daily to patients receiving mechanical ventilation - RN driven

- Every 24 hours
  - SAT safety screen
  - Perform SBT
  - COORDINATION: If SBT tolerated for > 2 hours, prompt extubation

**DELIRIUM MONITORING/MANAGEMENT:** Applied daily to all patients receiving mechanical ventilation - RN driven

- Administer RASS every 2 hours
- Administer CAM-ICU every 8 hours
- COORDINATION: Discuss RASS, CAM-ICU, and treatment plans

**EARLY EXERCISE/MOBILITY:** Applied daily to all patients - RN/PT driven

- Exercise/mobility safety screen
- Perform exercise/mobility
- COORDINATION: Discuss mobility progression on daily rounds

**Notes:**

- * Continuous sedative medications maintained at previous rate if spontaneous awakening trial (SAT) safety screen failure. Mechanical ventilation continued, and continuous sedative medications restarted at half the previous dose only if needed due to SAT safety screen failure.
- Continuous sedative infusions stopped, and sedative boluses held. Bolus doses of opioid medications allowed for pain. Continuous opioid infusions maintained only if needed for active pain.
- Continuous sedative medications restarted at half the previous dose, and then titrated to sedation target if SAT failed. Interdisciplinary team determines possible causes of SAT/SBT failure during rounds. Mechanical ventilation restarted at previous settings, and continuous sedative medications restarted at half the previous dose only if needed if SBT failed.
- SAT pass if the patient can open their eyes to verbal stimulation without failure criteria (regardless of trial length) or does not display any of the failure criteria after 4 hours of shutting off sedation.
- Each day on interdisciplinary rounds, the RN will inform the team of the patient’s target RASS score, actual RASS score, CAM-ICU status, and sedative and analgesic medications the patients is receiving. If delirium is detected, team will discuss possible causes, eliminate risk factors, and employ non-pharmacologic management strategies.
- Each eligible patient is encouraged to be mobile at least once a day, with the specific level of activity geared to their readiness.

**CAM-ICU** = confusion assessment method for the intensive care unit; **PT** = physical therapist; **RASS** = Richmond Agitation-Sedation Scale; **RN** = registered nurse; **RT** = respiratory therapist; **SAT** = spontaneous awakening trial; **SBT** = spontaneous breathing trial.
## ABCDE bundle safety screen questions and success/fail criteria

<table>
<thead>
<tr>
<th>ABCDE bundle component</th>
<th>Safety screen criteria: conditions for exclusion</th>
<th>Pass/fail criteria: conditions denoting failure</th>
</tr>
</thead>
</table>
| **Spontaneous awakening trial** | 1. Active seizures  
2. Alcohol withdrawal  
3. Neuromuscular blockade  
4. Control of increased ICP  
5. ICP > 20 mmHg  
6. Receiving ECMO  
7. Documentation of MI in past 24 hours  
8. Current RASS > 2 | 1. RASS score > 2 for ≥ 5 minutes  
2. Pulse oximetry < 88% for ≥ 5 minutes  
3. Respiration > 35 BPM for ≥ 5 minutes  
4. Acute cardiac arrhythmia  
5. ICP > 20 mmHg  
6. Two or more of the following: (heart rate increase ≥ 20 BPM, heart rate < 55 BPM, use of accessory muscles, abdominal paradox, diaphoresis or dyspnoea) |
| **Spontaneous breathing trial** | 1. Chronic ventilator dependence  
2. Pulse oximetry reading < 88%  
3. FiO₂ > 50%  
4. Set PEEP > 7  
5. ICP > 20 mmHg  
6. Receiving mechanical ventilation in attempt to control ICP  
7. Documentation of MI in past 24 hours  
8. Increasing doses of vasopressor medications  
9. Lack of inspiratory effort | 1. RR > 35 BPM for ≥ 5 minutes  
2. RR < 8  
3. Pulse oximetry < 88% > 5 minutes  
4. ICP > 20 mmHg  
5. Mental status changes  
6. Acute cardiac arrhythmia  
7. Two or more of the following:  
   - use of accessory muscles  
   - abdominal paradox diaphoresis  
   - dyspnoea |
| **Early exercise/mobility** | 1. RASS -3  
2. FiO₂ > 0.6  
3. Set PEEP > 10 cmH₂O  
4. Increasing doses of vasopressor infusions in the last 2 hours  
5. Evidence of active MI  
6. Administration of a new antiarrhythmic agent  
7. Receiving therapies that restricted mobility (e.g. ECMO, open-abdomen, etc.)  
8. Injuries in which mobility is contraindicated (e.g. unstable fractures, etc.) | 1. Symptomatic drop in mean arterial pressure  
2. Heart rate < 50 or > 130 BPM ≥ 5 minutes  
3. RR < 5 or > 40 BPM ≥ 5 minutes  
4. Systolic blood pressure > 180 mmHg > 5 minutes  
5. Pulse oximetry < 88% ≥ 5 minutes  
6. Marked ventilator dyssynchrony  
7. Patient distress  
8. New arrhythmia or evidence of active MI  
9. Concern for airway device integrity or endotracheal removal  
10. Fail to knees |

---

**Notes:** ABCDE - Awakening and Breathing Coordination, Delirium Monitoring/Management and Early Mobility Bundle; BPM - beats per minute; ECMO - extracorporeal membrane oxygenation; FiO₂ - fraction of inspired oxygen; ICP - intracranial pressure; MI - myocardial ischaemia; PEEP - positive end-expiratory pressure; RASS - Richmond Agitation-Sedation Scale; RR - respiratory rate.
Bedside checklist for ABCDE protocol

Date: ________/_______/_______

**Awakening and Breathing Coordination**

<table>
<thead>
<tr>
<th></th>
<th>Check if yes or indicate reasons</th>
</tr>
</thead>
<tbody>
<tr>
<td>SAT screen passed?</td>
<td>If not, why?</td>
</tr>
<tr>
<td>SAT done?</td>
<td>If not, why?</td>
</tr>
<tr>
<td>SBT screened passed?</td>
<td>If not, why?</td>
</tr>
<tr>
<td>SBT done?</td>
<td>If not, why?</td>
</tr>
<tr>
<td>SAT and SBT coordinated/paired?</td>
<td></td>
</tr>
</tbody>
</table>

**D**

**Delirium nonpharmacologic interventions**

<table>
<thead>
<tr>
<th>Intervention</th>
<th>Check if done</th>
</tr>
</thead>
<tbody>
<tr>
<td>Pain assessment/management</td>
<td></td>
</tr>
<tr>
<td>Orientation</td>
<td></td>
</tr>
<tr>
<td>Sensory (eyes/ears)</td>
<td></td>
</tr>
<tr>
<td>Sleep (nonpharm)</td>
<td></td>
</tr>
<tr>
<td>Check any intervention that was performed during your shift (including night shift)</td>
<td></td>
</tr>
</tbody>
</table>

**E**

**Early Exercise and mobility**

<table>
<thead>
<tr>
<th>Intervention</th>
<th>Check if done</th>
</tr>
</thead>
<tbody>
<tr>
<td>Active ROM</td>
<td></td>
</tr>
<tr>
<td>Sitting up on side of bed</td>
<td></td>
</tr>
<tr>
<td>Standing</td>
<td></td>
</tr>
<tr>
<td>Walking</td>
<td></td>
</tr>
<tr>
<td>Check any level of activity the patient performed during your shift (including night shift)</td>
<td></td>
</tr>
</tbody>
</table>

Notes: ROM - range of motion; SAT - spontaneous awakening trial; SBT - spontaneous breathing trial.
References and resources


Quality in critical care

Summary

<table>
<thead>
<tr>
<th>Quality elements</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>Safety</td>
<td>Avoiding harm to people for whom the care is intended.</td>
</tr>
<tr>
<td>Timeliness</td>
<td>Reducing waiting times and sometimes harmful delays for both those who receive and those who give care.</td>
</tr>
<tr>
<td>Effectiveness</td>
<td>Providing evidence-based health care services to those who need them.</td>
</tr>
<tr>
<td>Efficiency</td>
<td>Maximizing the benefit of available resources and avoiding waste.</td>
</tr>
<tr>
<td>Equity</td>
<td>Providing care that does not vary because of age, sex, gender, race, ethnicity, geography, religion, socio-economic status, linguistic or political affiliation.</td>
</tr>
<tr>
<td>Patient-centredness</td>
<td>Providing care that responds to individual preferences, needs and values in health services that are organized around the needs of people.</td>
</tr>
</tbody>
</table>

Systematic and continuous quality improvement work is essential because health care delivery is complex and imperfect, even with the best efforts. Quality measures are related to ICU resources/structure, processes of care and patient outcomes. The focus should be on processes of care, instead of hard-to-measure outcomes.
Use the iterative, real-time, **plan-do-act-check** cycle to test changes/improvement.

Create an inclusive team and culture of change for a successful and sustainable quality improvement programme.

See further information on quality interventions see the WHO *Handbook for national quality policy and strategy*.

**Tools**

13.1 Checklist for ICU daily best practices  
13.2 Surviving Sepsis Campaign bundles  
13.3 Checklist for high-quality use of invasive mechanical ventilation for ARDS  
13.4 Process for selecting problem to focus on in the ICU and quality improvement process  
13.5 Checklist for initiating, improving, evaluating and sustaining a quality improvement programme
13.1 Checklist for ICU daily best practices

<table>
<thead>
<tr>
<th>ICU Daily Rounding Checklist</th>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Can sedation be reduced?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Can analgesia be reduced?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Is delirium being assessed, addressed and prevented?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Spontaneous awakening trial and/or Spontaneous breathing trial candidate?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Total fluid balance goal reviewed? And targeting: ____________ [12h] [24h]</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Is the patient at goals for lung protective ventilation?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Head of bed elevation (30 degrees)</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Skin breakdown assessment (prevention) completed</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Is enteral nutrition at goal?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Is blood glucose at goal?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Deep venous thrombosis prophylaxis?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Gastric ulcer prophylaxis?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Can Antibiotics be narrowed or discontinued?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Early mobility candidate and physical therapy consulted?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Is foley catheter needed?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Is central venous/arterial catheter needed?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Has the family been updated?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
<tr>
<td>Are the patient's goals of care and code status current?</td>
<td>☐ Yes</td>
<td>☐ NA</td>
</tr>
</tbody>
</table>

Source: Open Critical Care Project (OpenCriticalCare.org).

DISCLAIMER: This tool is intended to serve as a framework for local modification in accordance with local resources, standards and guidelines. This is intended to be educational in nature and is not a substitution for clinical decision making based on the medical condition presented. It is the responsibility of the user to ensure all information contained herein is current and accurate by using published references. Tools based in part on ACCP Guidelines: Critical Care Medicine, 2013.
13.2 Surviving Sepsis Campaign bundles

Hour-1 bundle: initiate bundle upon recognition of sepsis/septic shock

May not complete all elements within first hour.

1. Measure lactate level.
   Remeasure lactate if initial lactate elevated (> 2 mmol/L).

2. Obtain blood cultures before administering antibiotics.

3. Administer broad-spectrum antibiotics.

4. Begin rapid administration of 250-500 mL crystalloid for hypotension or lactate ≥ 4 mmol/L.

5. Apply vasopressors if hypotensive during or after fluid resuscitation to maintain a mean arterial pressure ≥ 65 mm Hg.

Remember
1. Act quickly upon sepsis and septic shock recognition;
2. Minimize time to treatment - sepsis and septic shock are medical emergencies;
3. Monitor closely for response to interventions;

Consider using an adapted version of this tool to monitor performance for sepsis care. This bundle was recently revised based on the most recent version of these tools. See the Surviving Sepsis Campaign - adult patients website for full details.

The Paediatric Surviving Sepsis Campaign Bundle can be found in Chapter 8 (Tool 8.6) or on the Surviving Sepsis - paediatric patients website.
13.3 Checklist for high-quality use of invasive mechanical ventilation for ARDS

☑ Consider using this tool if you are using IMV to deliver quality care to your patients with ARDS.

Technical competence
☐ Type of mechanical ventilator available.
☐ Able to deliver internal PEEP 5-20 cmH₂O.
☐ Able to measure plateau airway pressure.
☐ Able to deliver FiO₂ 100%.
☐ Able to titrate FiO₂.
☐ Able to titrate tidal volumes from 4-10 mL/kg.
☐ Intubation equipment readily available. See full WHO specifications for mechanical ventilators.
☐ Infection prevention materials readily available (airborne precautions). Skilled person to intubate available.
☐ Skilled personnel to use and troubleshoot IMV. Arterial blood gas analyser available and working. Pulse oximeter available and working.
☐ Supply chain for ventilator consumables (circuits, filters).
☐ Skilled personnel and equipment for biomedical/ventilator troubleshooting and preventive maintenance.

Safety
☐ Protocols and equipment for difficult airway (e.g. backup personnel, equipment and plan - e.g. cricothyrotomy).
☐ Plan for IMV complications (e.g. chest tube for pneumothorax, sedation for agitation).
☐ Protocols for prevention while on IMV (e.g. daily SBT evaluation, daily sedation interruption, VAP prevention).
☐ Protocols for lung protective ventilation. See OpenCritical Care Respiratory care order set (protocol) templates.

Process measures
☐ Process measures (e.g. lung protective targets met).

Outcome measures
☐ Complications (e.g. VAP, pneumothorax).
13.4 Process for selecting problem to focus on in the ICU and quality improvement process

This flowchart provides a framework for selecting a problem to focus on for quality improvement among the many that might be considered.

![Flowchart diagram]

Prioritize
1. Is the problem measurable? How are we currently doing?
2. Is there evidence linking this problem to quality of care/patient safety?
3. Is it a hospital requirement? What is the deadline?
4. Interferes or has synergy with other ongoing projects?
5. How easy is it to implement?
   • Human resources
   • Other resources (equipment, etc.)
   • Does it require changing individual clinical behaviour vs health care system?

Before proceeding:
• audit
• review literature
• discussion with stakeholders

Essential steps in the **plan-do-study-act** cycle:

- Objective
- Questions and predictions, why?
- Plan to carry out the cycle (who, what, where, when)
- Plan for data collections

- What changes are to be made
- Next cycle

- Complete the analysis of the data
- Compare data to predictions
- Summarize what was learned

- Carry out the plan
- Document problems and unexpected observations
- Begin analysis of the data

Source: Adapted from Dr Andre Amaral, Sunnybrook Health Sciences Centre and University of Toronto, Canada.
13.5 Checklist for initiating, improving, evaluating and sustaining a quality improvement programme

This checklist provides steps for initiating, improving, evaluating and sustaining a quality improvement programme in the ICU (adapted from Curtis et al., 2006).

Initiating or improving a quality improvement programme

☐ Do background work: identify motivation, support team and develop strong leadership. Prioritize potential projects and choose the projects to begin.

☐ Prepare for the project by operationalizing the measures, building support for the project and developing a business plan.

☐ Do an environmental scan to understand the current situation (structure, process or outcome), the potential barriers, opportunities and resources for the project.

☐ Create a data collection system to provide accurate baseline data and document improvement.

☐ Create a data reporting system that will allow clinicians and other stakeholders to see and understand the problem and the improvement.

☐ Introduce strategies to change clinician behaviour and create the change that will produce improvement.

Evaluating and sustaining a quality improvement programme

☐ Determine whether the target is changing with ongoing observation, periodic data collection and interpretation.

☐ Modify behaviour change strategies to improve, regain or sustain improvements.

☐ Focus on sustaining interdisciplinary leadership and collaboration for the quality improvement programme.

☐ Develop and sustain support from the hospital leadership.

Common ICU quality indicators

☐ Deep venous thrombosis prophylaxis - number of patients receiving prophylaxis per eligible day.

☐ Stress ulcer prophylaxis - percentage of patients receiving prophylaxis per eligible day.

☐ Ventilator-associated pneumonia prevention strategies - percentage of patients receiving VAP bundle per eligible day.

☐ Central venous catheter blood stream infection prevention strategies - percentage of patients receiving checklist per eligible central venous catheter insertion.
References and resources


14 Ethical considerations
Summary

During a pandemic, the need for critical care services can exceed available resources. Triage decisions may need to be made on how to allocate scarce resources and prioritize patients.

Before patients are prioritized, allocation planners need to prioritize among often mutually exclusive substantive values and principles.

Key values and principles include: maximizing the number of lives or life-years saved; first come, first served; allocating in ways that reduces, rather than maintains or increases inequities; however, there is neither international nor, typically, national consensus on which values and principles to include and/or how to rank them.

Public engagement in pandemic preparedness is essential to develop a prioritization strategy that is fair, transparent and builds trust.

Tools

14.1 Ethical values and principles
14.2 Triage decision process flow
14.3 Hospital scarce resource decision-making
14.1 Ethical values and principles

Ethical analysis involves identifying relevant principles, applying them to a particular situation, and making judgements about how to weigh competing principles when it is not possible to satisfy them all.

The fundamental ethical challenge in triage is how to justify who to initially allocate a ventilator to when there are insufficient resources for all patients who require one, and under what circumstances patients might be removed from ventilators.

Ethical triage requires:
• first, identifying relevant substantive and procedural principles;
• second, processes to apply these principles in a transparent way to particular situations; and
• third, monitoring that triage accomplishes its goals and that there are no unintended consequences.

While there is international agreement on the need for transparent processes, consensus on substantive principles is elusive at the international level and typically also at national level.

Historically, the dominant trend underlying triage frameworks has been to focus on maximizing benefits; and allocation frameworks comprised clinical measures such as prognosis that help ensure that triage saves the most lives, or the most life-years. However, drawing on such frameworks in the COVID-19 pandemic made clear that maximizing benefits has a tendency to exacerbate existing inequities due to the social and political determinants of health. For example, considering prior comorbidities, remaining life expectancy, or even seemingly objective measures such as the Sequential Organ Failure Assessment (SOFA) score, tend to favour economically and otherwise privileged population groups, while simultaneously disadadvantaging less privileged groups. A major challenge is, therefore, how to weigh competing principles when it is not possible to satisfy them all.
### Key ethical principles and descriptions (in alphabetical, not lexical, order)

<table>
<thead>
<tr>
<th>Ethical principle</th>
<th>Description</th>
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<tbody>
<tr>
<td>Autonomy</td>
<td>Enabling, and letting individuals make their own choices based on their values and preferences.</td>
</tr>
<tr>
<td>Equity</td>
<td>• Treating the same those who have the same needs, and treating differently those with different needs, which usually requires reducing, rather than maintaining, or worse, exacerbating, avoidable unfair differences in opportunities or health outcomes.</td>
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<tr>
<td></td>
<td>• Equity important if policies appear to systematically disadvantage groups already experiencing disproportionate discrimination, injustice or other forms of disadvantage.</td>
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<tr>
<td>First come, first served</td>
<td>Allocates resources by respecting the temporal sequence of requests. Typically this assumes that all those requesting resources are reasonably equal in their ability to access services.</td>
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<tr>
<td>Informed consent</td>
<td>Process in which a competent individual authorizes a course of action based on sufficient relevant information, without coercion or undue inducement.</td>
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<tr>
<td>Instrumental value</td>
<td>Giving priority to those who have made, or are likely to make, relevant contributions, for example, essential key health care staff.</td>
</tr>
<tr>
<td>Justice</td>
<td>Encompasses equity - fairness in the distribution of resources, opportunities and outcomes - and procedural justice - a fair process for making important decisions.</td>
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<tr>
<td>Liberty</td>
<td>Includes a broad range of social, religious and political freedoms (e.g. freedom of movement, peaceful assembly, speech), many of which are protected as fundamental human rights.</td>
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<tr>
<td>Life cycle</td>
<td>Captures the notion that it is desirable that everyone should be able to live a life of normal length, and that, while all deaths are tragic, deaths of individuals who are younger are more tragic than those of people near the end of normal life expectancy.</td>
</tr>
<tr>
<td>Maximizing benefits</td>
<td>Actions or policies are right insofar as they maximize benefits. Most commonly this is understood to require policies that save the most lives or life-years.</td>
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<tr>
<td>Procedural justice</td>
<td>Includes:</td>
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<tr>
<td></td>
<td>• Due process - acknowledgement of people and give opportunity to be heard.</td>
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<tr>
<td></td>
<td>• Transparency - clear, accurate information about the basis for decisions and the decision-making process.</td>
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<td></td>
<td>• Inclusiveness/community engagement - ensuring all relevant stakeholders participate.</td>
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<tr>
<td></td>
<td>• Accountability - allocating and enforcing responsibility for decisions.</td>
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<tr>
<td></td>
<td>• Oversight - ensuring appropriate mechanisms for monitoring and review.</td>
</tr>
<tr>
<td>Respect for persons</td>
<td>Treating individuals in recognition of our common humanity, dignity and inherent rights. Key aspects include: autonomy; informed consent; privacy; confidentiality; social, religious and cultural beliefs; important relationships (e.g. family); and transparency and truth telling in public health and research.</td>
</tr>
<tr>
<td>Random selection</td>
<td>Assumes that everyone is equal irrespective of factors such as age, comorbidities, prognosis or other factors and should have an equal chance that can be implemented through a lottery, coin-toss or similar methods. Sometimes proposed as a secondary principle, for example, for patients with similar prognosis.</td>
</tr>
<tr>
<td>Solidarity</td>
<td>Social relations in which a group, community, nation or global community stands together. Justifies collective action in the face of common threats and supports efforts to overcome inequalities that undermine the welfare of minorities and groups that suffer discrimination.</td>
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</tbody>
</table>

Source: Adapted from Guidance for managing ethical issues in infectious disease outbreaks (WHO, 2016); WHO SAGE values framework for the allocation and prioritization of COVID-19 vaccination, 14 September 2020 (WHO, 2020).
14.2 Triage decision process flow

This is adapted from the 2020 *Triage of scarce critical care resources in COVID-19 an implementation guide for regional allocation* (Expert Panel Report of the Task Force for Mass Critical Care and the American College of Chest Physicians) (Maves et al., 2020). It is presented as a framework only, and has not been validated in any population.

**Triage decision algorithm**

1. **Step 1:** Does this patient meet criteria for critical care services (i.e. need for mechanical ventilation or circulatory support)?
   - No: Admit to medical ward with supportive care.

2. **Step 2:** Does the patient (or designated surrogate) agree to receive intensive care?
   - No: Admit to medical ward with supportive care. Palliative care consultation. Consider comfort measures.

3. **Step 3:** Does the patient’s expected risk of mortality, based on comorbid conditions and acute illness using standardized assessments, exceed the threshold set by the regional/local triage plan?
   - No: Admit with intensive care services, with periodic reassessment and possible reallocation by triage team.
   - Yes: Reassess. Periodically consider reallocation of critical care resources based on changes in availability, demand and patient status.

Source: Maves et al. (2020).
14.3 Hospital scarce resource decision-making

Clinical resources inadequate or will be inadequate to meet demands AND:
• HCS implemented.
• Regional transfers not possible at off-load.
• Resources not available in a timely manner.
• Event of sufficient duration/magnitude to require proactive decision-making.

Incident commander directs Clinical Care Committee (CCC) to examine situation, timeframe, available guidance, recommend changes to usual practice and use of resources.

Are case-specific triage resources required?

Each operational period:
• Maintain situational awareness/anticipate issues.
• Communicate clinical and operational changes to patients/families, staff and public as required.
• Reassess response and decisions during prior cycle.

CCC consults with Office of Medical Director (OMD), appoints Triage Team (TT) composed of:
• OMD on call.
• Two specialty staff with expertise in resource area (example critical care for ventilators, CV for ECMO, infectious disease for antivirals, burn surgeon for mass burn event - most situations will involve at least one, if not two critical care staff.

Triage team reviews situation, patients, and available triage guidelines taking into account:
• Clinical/prognostic data (including relevant scoring systems).
• Duration of use.
• Likelihood of benefit.

Team should not consider subjective factors in decision-making.

Team is consulted whenever a demand for specific resource occurs - may be in-person or virtual. Triage team confers and decides which patient(s) will receive the resources available.

Triage team documents decision-making in affected patient charts according to incident-specific template.

Decision communicated to patient’s attending MD and inpatient supervisor.

Attending MD may make clinical appeal within 15 minutes if new clinical information suggest improvement (TT re-evaluates in light of new information and communications decision).

Inpatient supervisor and attending physician determine:
• Family/patient notification - may request OMD assistance.
• Level of care required for ongoing support (floor, intermediate, ICU).
• Transition plan for patient and family (location, support - clinical and emotional, etc.).

Notes: Algorithm and plan does not apply to immediate, reactive triage decisions in the early phase of a disaster (e.g. ED, trauma surgery) or to non-emergency circumstances (specialty providers will engage colleagues in decision-making). This algorithm is a summary of select actions in the HCMM Crisis Care Annex which should be referred to for further details.

CCC = clinical care committee; CV = cardiovascular specialist; ECMO = extracorporeal membrane oxygenation; HICS = hospital incident command system; MD = medical doctor; OMD = office of medical director; TT = triage team.

Source: Maves et al. (2020).
References and resources


For more information, please contact:

Emerging Diseases Clinical Assessment and Response Network
World Health Organization
Avenue Appia 20
CH-1211 Geneva 27
Switzerland

Email: COVID_ClinPlaftorm@who.int
Website: www.who.int/groups/edcarn