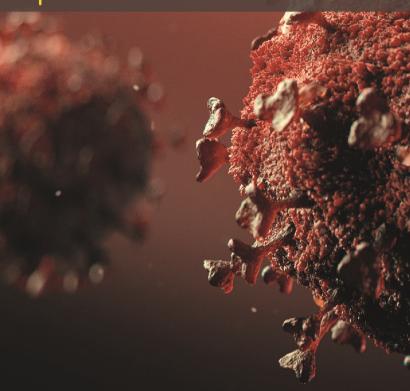


Clinical Management of COVID-19 in Healthcare Setting

Second Edition



Prepared by:



Government of Nepal Ministry of Health and Population Department of Health Services Epidemiology and Disease Control Division

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INTRODUCTION TO THE SECOND EDITION

This is the second edition of the Ministry of Health and Populations, Pocket book for Case Management of COVID-19 in the Healthcare Setting.

The first edition was adapted from the Interim Guidance for Caring of Patients with COVID-19 in Healthcare settings, Nepal Medical Council (NMC) and aligned with Clinical management of COVID-19, Interim Guidance, WHO (May 27, 2020).

The rationale for this updated edition has been to expand the scope of the earlier guidance, bringing together recently updated recommendations from NMCs October 2020 update and COVID -19 Clinical Management: Living Guidance (January 25, 2021) as well as the Clinical Care for Severe Acute Respiratory Infection: Toolkit-COVID-19 Adaptation: WHO 2020 and advice from subject matter experts.

This edition provides updated technical guidance in several case management clinical areas as follows:

- Revision of the release from the COVID-19 clinical pathway criteria
- Addition of Treatment of acute co-infections
- Addition of prevention of complications, immunomodulators and other adjunct therapies
- Revision of recommendations on use of antivirals and corticosteroids in COVID-19
- Assessment of severity by National early Warning Score 2 (NEWS 2)

PURPOSE OF THE POCKET BOOK:

The purpose of this pocketbook is to help physicians, and healthcare workers, to properly manage persons with suspected or confirmed COVID-19 and to standardize case management of COVID-19 cases throughout the country. It is not meant to replace clinical judgement or specialist consultation, but rather to strengthen frontline clinical management.

TARGET GROUPS:

The intended target audience are physicians, nurses, other healthcare personnel, involved in the clinical management of COVID-19 cases.

1. COVID-19 OVERVIEW12:

Coronavirus disease 2019 (COVID-19) is caused by the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2), a single-stranded RNA virus that was first recognized in Wuhan, Hubei province, China, in December 2019. Epidemiology and virologic studies suggest that transmission mainly occurs by close contact through respiratory droplets or by direct contact with infected persons, or by contact with contaminated objects or surfaces, or by aerosols from both symptomatic and asymptomatic people. Clinical and virologic studies have found that shedding of SARS-CoV-2 is highest in the upper respiratory tract (URT) (nose and throat) early in the course of the disease, within the first 3 days from onset of symptoms. The incubation period for COVID-19 is 5-7 days but can be up to 14 days. Some infected persons can be contagious, from 1-3 days before symptom onset (presymptomatic period).

An asymptomatic case is a person infected with SARS-CoV-2 who does not develop symptoms.

Among symptomatic patients, the duration of infectious virus shedding has been estimated at 8 days from the onset of any

¹ COVID-19 Clinical Management, Living Guidance 25 January 2021

² Clinical management of COVID-19 Interim Guidance: May 2020:WHO 2 I Page

symptoms. About 17% (14-20%) patients will be asymptomatic throughout their illness. A living systematic review (4 studies, 462051 women) found that pregnant and recently pregnant women with COVID-19 appear to be less likely to be symptomatic or manifest common symptoms such as fever, dyspnoea and myalgia, compared with nonpregnant women of reproductive age.

Case definition including common symptoms are given in table 1.

Older age, smoking and underlying noncommunicable diseases (NCDs), such as diabetes, hypertension, cardiac disease, chronic lung disease and cancer have been reported as risk factors for severe COVID-19 disease and death. However, most patient with COVID-19 develop only mild (40%) or moderate (40%) disease approximately 15% develop severe disease that requires oxygen support, and 5% have critical disease with complications such as respiratory failure, acute respiratory distress syndrome (ARDS), sepsis and septic shock, thromboembolism, and/or multiorgan failure, including acute kidney injury and cardiac injury. (Table 2)

COVID-19 is also associated with mental and neurological manifestations, including delirium or encephalopathy, agitation, stroke, meningoencephalitis, impaired sense of smell or taste anxiety, depression and sleep problems. In many cases, neurological manifestations have been reported even without respiratory symptoms. Anxiety and depression appear to be common amongst patient hospitalized for COVID-19.

There are few data on the clinical presentation of COVID-19 in specific populations, such as children and pregnant women. Clinical manifestations of COVID-19 are generally mild in children compared with adults. However, most recently, an acute presentation with a hyperinflammatory syndrome leading to multiorgan failure and shock has been described, as 'multisystem inflammatory syndrome temporally associated with COVID-19 in children an adolescents'

While most people with COVID-19 recover and return to normal health, some people can have symptoms that last for weeks or even months after recovery from acute illness named post

COVID condition. Even people who are not hospitalized or have mild symptoms can experience persistent or late symptoms.

2. SCREENING AND TRIAGE^{1,2}:

Screen all persons at the first point of care (POC) at the health facility e.g., OPD, emergency, triage area etc. Ensure IPC precautions are implemented when performing screening of persons (Annex 1). Follow the screening process as follows:

- 1. <u>Screening questionnaire:</u> Screen all persons presenting to the facility by using the simple screening questionnaire (Annex 2)
- 2. <u>Temperature checks</u>: All persons presenting to the facility should have their temperatures taken and recorded
- ! Preferably use a non-touch technique (e.g. infra-red thermometer)
- ! In the absence of a non-touch thermometer, clean the thermometers with 70% alcohol wipe after each use.
- ! Temperatures > 100.4*F or 38* C is considered as fever
- 3. <u>Co-morbidities</u>: Febrile patients should be tested according to facility protocol, irrespective of the presence of respiratory signs and symptoms; screen for high risk of severe COVID-19 disease with priority. Consider other causes of illness (Annex 4 Differential Diagnosis Matrix)
- 4. <u>Diagnosis</u>: After screening, diagnose the patient according to the EDCD case definitions for COVID-19 (Table 1)
- **a.** All suspect cases should be tested to determine if they are a confirmed case. Until proven negative, all suspected cases should remain in the COVID-19 care pathway.
- b. Laboratory diagnostic testing:
- Collection of upper respiratory tract (URT) specimens (nasopharyngeal and oropharyngeal) for testing by RT-PCR (consider using SARS-CoV-2 Antigen RDT when there is no access to RT-PCR) and, where clinical suspicion remains

and URT specimens are negative, collect specimens from the lower respiratory tract (LRT) when readily available (expectorated sputum, or endotracheal aspirate/ bronchoalveolar lavage in ventilated patient).

- II. In addition, testing for other respiratory viruses and bacteria should be considered when clinically indicated.
- III. If testing not available, the person becomes a probable case based on clinical suspicions and should be managed for in the COVID-19 pathway.

Consider, and test for other endemic diseases when clinically indicated e.g.:

Malaria Dengue, Scrub Typhus, TB etc.

Co-infections / morbidity with COVID-19 may be present Recognize high risk groups for severe COVID-19 disease (Table 1.)

- 5. <u>COVID-19 Pregnant and post-partum women:</u> with preexisting or pregnancy related co-morbidities (e.g., pregnancy induced hypertension, gestational diabetes, should also be considered for close monitoring
- 6. <u>Isolation:</u> All suspected or confirmed COVID-19 cases, with or without co-morbidities should be given a medical mask to wear and entered into the COVID-19 pathway

7. <u>Documentation</u>: All information gathered during the screening process, should be documented and recorded on individual patient forms.

PROTECT YOURSELF AND OTHERS

- Follow IPC measures at all times
- Wear PPE according to the activity risk
- Always follow & perform 5 moments of hand hygiene
- Maintain physical distancing of > 2 meters

Jadated in Public health surveillance for COVID-19, aublished 7 August 2020

Suspect COVID-19 case

A person who meets the clinical AND epidemiological criteria: 4

Clinical Criteria:

- Acute onset of fever AND cough;
- 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

AND

Epidemiological Criteria:

- of transmission of virus: closed residential settings, humanitarian settings such as camp and camp-like settings for displaced persons: anytime within Residing or working in an area with high risk
 - 14 days prior to symptom onset;
- transmission anytime within the 14 days prior to Residing or travel to an area with community symptom onset:

Working in any health care setting, including within health facilities or within the community; anytime within the 14 days prior to symptom onset.

(SARI: acute respiratory infection with history of with onset within the last 10 days; and requires A patient with severe acute respiratory illness: fever or measured fever of ≥ 38 C°; and cough; nospitalization). 2

Probable COVID-19 case



A suspect case with chest imaging showing findings suggestive of COVID-19 disease*

m

*Typical chest imaging findings suggestive of COVID-19

include the following:

- Chest radiography: hazy opacities, often rounded in morphology, with peripheral and lower lung distribution.
- Lung ultrasound: thickened pleural lines, B lines Chest CT: multiple bilateral ground glass opacities, often rounded in morphology, with peripheral and ower lung distribution.
 - (multifocal, discrete, or confluent), consolidative patterns with or without air bronchograms.
- (loss of smell) or ageusia (loss of taste) in the A person with recent onset of anosmia absence of any other identified cause.
- with respiratory distress preceding death AND was a contact of a probable or confirmed case or epidemiological linked to a COVID-19 cluster **Death**, not otherwise explained, in an adult with atleast one confirmed case.

Confirmed COVID-19 case

A person with laboratory confirmation of COVID-19 infection, irrespective of clinical igns and symptoms.

⋖

See: Laboratory testing for coronavirus disease (COVID-19) in suspected human cases guidance. For details

NOTE:

Clinical and public health judgment should be investigation in patients who do not strictly meet the clinical or epidemiological criteria. Surveillance case definitions should not be used as the sole basis for guiding clinical used to determine the need for further nanagement. © World Health Organization 2020. Some rights reserved. This work is available under the CC BY-NC-SA 3.0 IGO licence. WHO reference number: WHO/2019-nCoV/Surveillance. Case. Definition/2020.2

SARS-CoV-2- Variant of Interest (VOI)

A SARS-CoV-2 isolate that is phenotypically changed compared to a reference isolate or that has a genome with mutations that lead to amino acid changes associated with established or suspected phenotypic implications; AND has been identified to cause community transmission/multiple COVID-19 case clusters, or has been detected in multiple countries.

OR

is otherwise assessed to be a VOI by WHO in consultation with the WHO SARS-CoV-2 Virus Evolution Working Group (VEWG).

SARS-CoV-2 Variant of concern (VOC):

A VOI (as defined above) that, through a comparative assessment, has been demonstrated to be associated with:

- Increase in transmissibility or change in the epidemiology.
- Increase in virulence or change in disease presentation; or
- Decrease in effectiveness of available diagnostics, vaccines, therapeutics, or public health and social measures.

1. FLOW CHART: TRIAGE AND ISOLATION

Does the person have any of (new-onset) fever, cough, shortness of breath, chills, muscle pain, acute loss of taste or smell, diarrhea or sore throat in the last 14 days? History of contact with a confirmed or Alternative explanation/diagnosis to probable COVID-19 case in the last 14 the person's symptoms/signs? days prior to symptom onset? No No Yes COVID-19 less likely COVID-19 less likely COVID-19 Suspect Moderate / Severe / Critical COVID-19 Infection Asymptomatic/Pre-symptomatic/Mild COVID-19 Infection Pneumonia, ARDS, Sepsis, Shock Admit to ICU Hemodynamically stable Admit to ISOLATION/COVID Hypoxemia requiring Sp02 > 93% in room air W/ARD #Call ICU for review HFNC/NIV/IMV No respiratory distress SpO2 ≤ 93% in room air at rest if 02 requirement Presence of shock/multi-organ RR ≤30 breaths/min >4L {28%} Target SpO2 ≤ 94% on ≤4 L/min O2 SPO2 93-96% {88-PaO2/PaO2 ≤ 200 or SE ratio ≤ Hemodynamically stable 92% if chronic type 235 with worsening respiratory High risk for severe disease** Isolate II respiratory failure) distress** Test-ve Test +ve Test-ve Test +ve Ambulation/Pharmacologic DVT prophylaxis if no Repeat test if suspicion high Repeat sampling if Management of alternative contraindications. strong clinical suspicion diagnosis Supplemental O2/HFNC*/NIV*(aerosolization risk*) Manage alternative diagnosis Awake proning >12-16 hours a day if able to communicate Monitor clinical status for worsening. Dry neubilization (salbutamol MDI 4puffs with spacer) SpO2 and temperature monitoring every 6 hourly) Clinical evaluation if SPO2 drops by >2% Intubate if: i) worsening distress ii) failure to maintain Admission in COVID ward if high-risk for severe disease** SPO2 ≤ 90% @15 lit O2 via NRB or NIV/HFNC iii) Contact and droplet precautions PaO2/FIO2<150 Symptomatic and psychosocial support Antibiotics if sepsis/bacterial pneumonia suspected Ambulation when awake to reduce the risk of VTF Steriods for severe cases (dexa/pred/methylpred/hydrocort X7-10d) **High risk for severe disease Restrictive fluid strategy Age >60 yrs Ventilator management as per the ARDSnet protocol · Cardiovascular disease including HTN Use closed suction catheters & viral/HEPA filters in · DM & other immune compromised conditions Chronic lung/kidney/liver/CNS diseases Consider proning/paralyzing agents early if PaO2/FIO2<150 Nutritional and psychological support Asymptomatic patient Symptomatic patient 10 days after symptoms onset plus 3 days without symptoms 10 days after test is positive (without fever and respiratory symptoms)

3. THE COVID-19 CARE PATHWAY

The patient enters the COVID-19 Care Pathway and follows the continuum of care until they are released from the pathway. All persons enter the COVID-19 pathway after <u>s/he is screened</u> based on a standardized <u>case definition and meets criteria for suspected case.</u>

- ✓ <u>Suspected cases</u> are persons under investigation and should be given and asked to wear a <u>medical mask</u>, and immediately <u>isolated</u> (single isolation rooms preferred) to contain the virus
 - If single isolation rooms are not available, groups the patients with similar diagnosis based on epidemiological risk factors with physical distancing of more than 2 meters apart
- ✓ Suspected cases should <u>not be</u> cohorted with confirmed cases
- ✓ Probable cases are persons for whom testing for COVID-19 (SARS-CoV-2) is inconclusive or not available but clinical suspicion remains high
- Confirmed cases are persons with <u>laboratory confirmation</u> of COVID-19
- Consideration for co-infections and/or chronic diseases must be made within the COVID-19 pathway

COVID-19 care pathways should be coordinated and multidisciplinary to ensure the delivery or safe and quality care in all levels of the health services.

Clinical monitoring of a COVID-19 patients can be done with National Early Warning Score (NEWS) 2. (Annex 5)

Routine tests in moderate, severe and critical COVID-19 disease

- 1. Complete blood count
- 2. Urea, creatinine, sodium and potassium
- 3. Liver function tests
- 4. Where available- CRP, D-dimer, troponin, ferritin, procalcitonin etc.
- Chest X-ray should be done in all hospitalized patients with fever, cough and shortness of breath. Common findings are patchy infiltrates in multiple areas.

3.1 Discontinue transmission-based precautions

(including isolation) and release from the COVID-19 care pathway as follows³:

- ✓ For symptomatic patients: 10 days after symptom onset, plus at least 3 days without symptoms (without fever, and respiratory symptoms)
- ✓ For asymptomatic patients: 10 days after testing positive for SARS-CoV-2
 - ! Release from the COVID-19 pathway is <u>NOT</u> the same as clinical discharge from a facility or from one ward to another, e.g. some patients may require ongoing rehabilitation, or other aspects of care, beyond release for the COVID-19 pathway, based on clinical needs in the COVID-19 pathway
 - ! If discharge from the COVID-19 pathway consider ongoing rehabilitation, medication reconciliation, plan for follow up with clinical provider in place, routine immunization etc.

³Criteria for releasing COVID-19 patients from isolation: Scientific Brief 17 June 2020: WHO

4. DISEASE SEVERITY^{1,2}

The COVID-19 disease is classified into following categories according to severity:

- Mild disease-See section 6.1; Algorithm 1
- Moderate disease- See section 6.2; Algorithm 2
- Severe disease-See section 6.3; Algorithm 3
- Critical disease-See sections 6.4, 6.5; Algorithms 4 & 5
 - ARDS
 - Sepsis
 - Septic shock
 - Or other conditions that would normally require the provision of life sustaining therapies such as mechanical ventilation (invasive or non-invasive) or vasopressor therapy

See Annex 5 for National Early Warning Score 2 to facilitate monitoring of the patient

Table 2. Clinical Symptoms Associated with COVID-19 and Risk Factors²

Clinical Presentation

he spectrum of illness ranges from asymptomatic infection to acute respiratory distress, multiorgan dysfunction. About 40% of patients will be asymptomatic. Presenting signs and symptoms of COVID-19 vary

Most persons experience fever (83–99%) though may be absent in close to half of patients initially, cough (59–82%), fatigue (44–70%), anorexia (40–84%), shortness of breath (31–40%), myalgias (11–35%). Other **non-specific symptoms**, such as sore throat, nasal congestion, headache, diarrhoea, nausea and vomiting, have also been reported. Loss of smell (anosmia) or loss of taste (ageusia) preceding the onset of respiratory symptoms has also been reported

Older people and immunosuppressed patients in particular may present with atypical symptoms such as fatigue, reduced alertness, reduced mobility, diarrhoea, loss of appetite, delirium, and fever may be absent

Symptoms such as dyspnoea, fever, GI symptoms or fatigue due to physiologic adaptations in **pregnant women**, adverse pregnancy events, or other diseases such as malaria, may overlap with symptoms of COVID-19.

Compared with adults, **children** are less likely to present with fever, cough and shortness of breath

Risk factors for severe disease

Age more than 60 years (increasing with age)

Underlying noncommunicable diseases (NCDs): hypertension, cardiac disease, chronic lung disease, cerebrovascular disease, diabetes mellitus (DM), immunocompromising conditions, and cancer have been associated with higher mortality.

Smoking

5. MANAGEMENT OF COVID-19 CASES ACCORDING TO DISEASE SEVERITY

First, <u>protect yourself and others</u> by implementing appropriate IPC measures at all times:

- ! Instruct the patient to wear a medical mask
- ! Maintain physical distancing of 2 meters where possible
- ! Instruct patients to adopt; good hand hygiene, respiratory hygiene, and cough etiquette (Annex 3)
- ! Depending on the activity risk:
- ! Apply standard precautions
- ! Apply contact and droplet precautions
- ! Apply airborne precautions

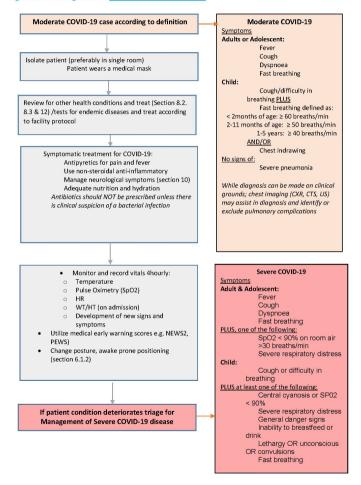
5.1 Case Management of Mild COVID-19 cases

5.1 Case Management of MILD COVID-19 cases Algorithm 1: Management of Mild COVID-19 disease Disease severity: Mild COVID-19 Mild COVID-19 diagnosed according to definition Symptoms (most people): √ Fever Cough ✓ Fatigue Isolate patient (preferably in single room, do not ✓ Anorexia cohort suspected cases with confirmed COVID-1 19 ✓ Shortness of breath patients) No evidence of: Patients wears a medical mask ✓ Viral Pneumonia ✓ Hypoxia Older & Immunocompromised people may present with atypical Review for other health conditions and treatment / symptoms such as fatigue, reduced tests for endemic diseases and treat according to alertness, reduced mobility. the facility protocol (section: 8.2, 8.3 & 12) diarrhea, appetite loss etc. Dyspnoea, fever, GI symptoms, or fatigue due to physiological Prescribe symptomatic treatment for COVID-19: adaptation in pregnant women. ✓ Antipyretics for pain and fever adverse pregnancy events or other ✓ Use non-steroidal anti-inflammatory endemic diseases may overlap with ✓ Manage neurological symptoms ✓ Adequate nutrition COVID-19 symptoms ✓ Adequate hydration × Antibiotics should NOT be prescribed unless Children (compared to adults) are there is clinical suspicion of a bacterial less likely to present with fever, infection cough and shortness of breath Monitor and record patient vitals at least twice Moderate COVID-19 daily: Symptoms Adults or Adolescent: ✓ Temperature ✓ Fever ✓ Pulse Oximetry (SpO2) Cough Dyspnoea ✓ WT/HT (on admission) √ Fast breathing ✓ Development of new signs and Child: ✓ Cough/ difficulty in breathing √ Fast breathing AND/OR If patient condition deteriorates triage for ✓ Chest indrawing Management of Moderate COVID-19 disease No signs of: × Severe pneumonia Refer to Moderate COVID-19 algorithm for additional information

5.1.1 Case Management of MODERATE COVID-19 cases

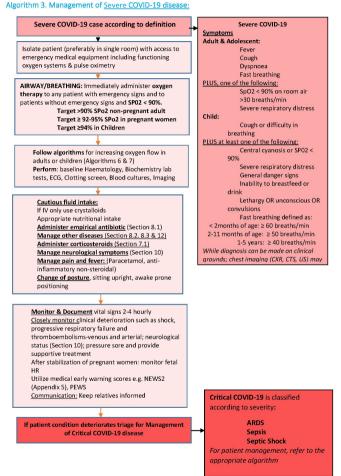
5.1.1 Case Management of MODERATE COVID-19 cases

Algorithm 2. Management of Moderate COVID-19 disease



5.1.2 Case Management of SEVERE COVID-19 cases

5.1.2 Case Management of SEVERE COVID-19 cases



5.1.3 Case Management of CRITICAL COVID-19

Algorithm 4. Acute Respiratory Distress Syndrome (ARDS)

4 hourly oral hygiene and eye care 2 hourly pressure points o Catheter/ drain/ wound care Change ETT tape when necessary EWS: Utilize medical early warning scores e.g.

Changes in medical treatment Keep relatives informed

NEWS2, PEWS) Communicate:

Critical COVID-19 disease: ARDS Critical COVID-19: ARDS Symptoms: Onset: within 1 week of pneumonia, or new worsening Isolate in single ICU room (preferable) respiratory symptoms Airborne precautions should be adopted CHEST: Bilateral opacities, (not fully explained by effusions), lobar/ lung collapse, nodules AIRWAY-Oxygen Management & Targets Oedema origins: Respiratory failure not fully explained by cardiac failure or fluid overload: Need objective Mild-Moderate ARDS: Trial of CPAP or RIPAP or HENC for 1 hour (provided no hypoxemic assessment (ECHO) to exclude hydrostatic oedema if no hemodynamic instability; multi-organ failure or abnormal risk factor mental status) ADULT: Oxygenation impairment: If failure to respond to increasing oxygenation - proceed to ✓ Mild ARDS: 200 mmHg <PaO2/ FiO2a ≤ 300</p> Severe ARDS: Proceed to intubation and IMV: mmHg (with PEEP or CPAP ≥ 5cmH2O) b oung Children, pregnant women; obese patients may desaturate quickly: Pre oxygenate with 100% FIO2 for 5 ✓ Moderate ARDS: 100 mmHg < PaO2/FiO2 ≤ 200</p> minutes, using face mask with reservoir mmHg (with PEEP ≥ 5 cmH2O) b Assess airway -> rapid intubation ! Avoid bag-valve mask ventilation to reduce exposure to Severe ARDS: PaO2/FiO2 ≤100 mmHg 9with PEEP ≥ 5 cmH2O) b IMV: Use low tidal volumes (TV): Lower plateau pressures: ✓ ADULT: Initial target TV is 6ml/kg predicted body weight (PBW); up to 8ml/kg PBW is allowed if pH <7.5); Lower **CHILDREN: Oxygenation impairment:** Note OI and OSIc. Use OI when available. If PaO2 not ✓ Prone positioning for 12-16 hours available, wean FiO2 to maintain SpO2 < 97% to ✓ CHILD: Adapt TV to disease severity: 3-6mL/kg PBW (in...) poor respiratory compliance) and 5-8mL/kg PBW (in calculate OSI or SpO2/FiO2 ratio: ✓ Bilevel (NIV or CPAP) ≥ 5cmH2O via full face (<28cmH2O) is targeted, lower target of pH is permitted mask: paO2/FiO2 ≤ 300 mmHg or SpO2/FiO2 ≤ (7.15-7.30) Positioning: Adults: prone ventilation for 12-16 hours (expertise needed). <u>Pregnant women</u> (3rd Trimester) may benefit from lateral decubitus positioning ✓ Mild ARDS (invasively ventilated): 4 ≤ OI < 8 or</p> 5 < OSI < 7.5 Avoid disconnecting the patient from ventilator, which results in loss of PEEP, atelectasis and increased risk of airborne ✓ Moderate ARDS (invasively ventilated): 8 ≤ OI <</p> transmission to HCWs 16 or 7.5 ≤ OSI <12.3 Sedate patients, use neuromuscular blockades intermittently ✓ Severe ARDS (invasively ventilated): OI ≥ 16 or OSI ≥12.3 Cardiovascular-Nursing management ✓ Administer corticosteroids: (Section:8.1) √ <u>Administer</u> vasopressors to maintain MAP: (Section:8.2.5) **MAP Targets** ✓ Administer drugs to prevent complications; (Section 8.4) ADULT: > 65mmHg ✓ Use conservative fluid to prevent fluid overload √ Manage neurological symptoms (Section 10) ADULT: >65 years: 60-65mmHg is equivalent ✓ Monitor and Document; vital signs according to ICU to ≥ 65mmHg protocol; neurological status; ABGs; laboratory results, fluid balance, thromboembolisms-venous/ arterial CHILD: Any hypotension (systolic BP) <5th ✓ Quality nursing care:

centile or >2SD below normal for age)

5.1.4 ARDS protocol and PEEP titration



Mechanical Ventilation Protocol Summary NIH NHLBI ARDS Clinical Network

INCLUSION CRITERIA: Acute onset of

PaO₂/FiO₂ ≤ 300 (corrected for altitude)

Bilateral (patchy, diffuse, or homogeneous) infiltrates consistent with pulmonary edema

PART I: VENTI LATOR SETUP AND ADJUSTMENT

No clinical evidence of left atrial hypertension

e

Males = 50 + 2.3 [height (inches) - 60] Calculate predicted body weight (PBW)

Females = 45.5 + 2.3 [height (inches) -60] Select any ventilator mode

Set initial rate to approximate baseline minute ventilation (not > 35 Reduce V_T by 1 ml/kg at intervals ≤ 2 hours until V_T = 6ml/kg PBW. Set ventilator settings to achieve initial V_T = 8 ml/kg PBW

Adjust V_T and RR to achieve pH and plateau pressure goals below.

bpm).

Use a minimum PEEP of 5 cm H₂O. Consider use of incremental FO₂/PEEP combinations such as shown below (not required) to achieve goal. **DXYGENATION GOAL: PaO₂ 55-80 mmHg or SpO₂ 88-95%**

Lower	PEEP/	Lower PEEP/ higher FIO2	F102					
FiO ₂	0.3	0.4	0.4	0.5	0.5	9.0	0.7	0
PEEP	5	2	8	8	10	10	10	12
							1	į
FiO ₂	0.7	8.0	6.0	6.0	6.0	1.0	П	
PEEP	14	14	14	16	18	18-24		

Higher PEEP/ lower FiO2	PEEP/ IA	ower Fi	102					
FiO ₂	0.3	0.3	0.3	0.3	0.3	0.4	0.4	0.5
PEEP	5	8	10	12	14	14	16	16
FiO ₂	0.5	0.5-0.8	г	8.0	6.0	1.0	1.0	_
PEEP	18	20	Г	22	22	22	24	

Check Pplat (0.5 second inspiratory pause), at least q 4h and after each PLATEAU PRESSURE GOAL: ≤ 30 cm H₂O change in PEEP or V_T.

increase V_T in 1ml/kg increments to 7 or 8 ml/kg if Pplat remains ≤ 30 cm If Pplat < 25 cm H₂O and V₇< 6 ml/ kg, increase V₇ by 1 ml/kg until If Pplat < 30 and breath stacking or dys-synchrony occurs: may If Pplat > 30 cm H₂O: decrease V₇ by 1ml/kg steps (minimum = 4 Polat > 25 cm H₂O or V_T = 6 ml/kg. ml/kg)

Source: ards.net

Algorithm 5: Management of Critical COVID-19

Management of Critical COVID-19 Sepsis and Septic Shock

Critical COVID-19 disease

Resuscitation for septic shock:

ADULT See page 18:

NOTE: Pregnant women: Compression of the vena cava (VC) can cause a decrease in venous return-place in lateral decubitus position to off load the VC

CHILD (see Algorithm 6):

Vasopressors (Section 10):

ADULT:

When shock persists during or after fluid resuscitation administer vasopressors to reach MAP targets

CHILD (see Algorithm 6):

Inotropes (Section 11):

If signs of poor perfusion and cardiac dysfunction persist despite achieving MAP targets with fluid and vasopressors, consider inotrope such as Dobutamine

Other treatments/ management

- Administer corticosteroids: (Section 7.1) ✓ Use measures to prevent complications: (Table 3)
- ✓ Monitor and Document:
- o vital signs according to ICU
 - protocol
 - neurological status o IMV & ABGs 4 hourly & PRN (if available)
 - Laboratory results: Lactate. Biochemistry, Hematology,
 - Clotting, LFTs, Cardiac enzyme o Fluid balance-in-put & out-put
 - Thromboembolisms s-venous/ arterial: perfusion -
 - Capillary refill

Quality nursing care:

- o 4 hourly eye care & oral hygiene
- 2 hourly pressure points
- o Catheter/ drain/ wound care
- Change ETT tape when necessary
- ✓ EWS: Utilize medical early warning scores e.g. NEWS2, PEWS)
- Communicate:
 - o Changes in medical treatment
 - Keep relatives informed

Critical COVID-19: Sepsis Definition

Symptoms

ADULTS: Acute life-threatening organ dysfunction caused by a dysregulated host response to suspected or proven infection.

Signs of organ dysfunction include:

- ✓ Alerted mental status
- ✓ Difficult or fast breathing
- √ Low oxygen saturation ✓ Reduced urine output
- ✓ Fast heart rate
- √ Weak pulse
- ✓ Cold extremities or low BP
- √ Skin mottling √ Laboratory evidence of
- coagulopathy ✓ Thrombocytopenia
- √ Acidosis
- ✓ High lactate or

hyperbilirubinemia CHILD:

✓ Suspected or proven

infection and ≥ 2 age based systematic inflammatory response syndrome (SIRS), of which one must be

Critical COVID-19: Septic Shock Definition

Symptoms:

ADULTS: persistent hypotension despite volume resuscitation, requiring vasopressors to maintain MAP ≥ 65mmHg and serum lactate level > 2mmol/L in

the absence of hypovolemia CHILD: any hypotension (systolic BP) <5th centile or > 2SD below normal for age)

OR 2 or more of the following:

- ✓ Altered mental state
- ✓ Bradvcardia or Tachycardia (HR <90bpm or > 160 bpm in infants; and HR
- ✓ Prolonged capillary refill (>2 sec) or feeble nulsas.
- Tachypnoea
- ✓ Mottled or cold skin OR petechial OR purpuric
- ✓ Increased lactate
- ✓ Oliguria

Targets

MAP: ADULT: ≥ 65mm Hg

ADULT >65 years: 60-65mm Hg is equivalent to ≥

CHILD: Any hypotension (systolic BP) < 5th centile or >2SD below normal for age)

URINE OUTPUT:

ADULT: > 0.5 mL/kg/hr.

Child: 1ml/kg/hr.

Lactate: <2mmol/L

Recommended resuscitation strategies for adult and pediatric patients with septic shock 1

- In resuscitation for septic shock in adults, give 250–500 mL crystalloid fluid (Normal saline or Ringer lactate) as rapid bolus in first 15–30 minutes and for septic shock in children, give 10–20 mL/kg crystalloid fluid as a bolus in the first 30–60 minutes. Fluid resuscitation may lead to volume overload, including respiratory failure, particularly with ARDS.
- If there is no response to fluid loading or signs of volume overload appear (e.g. jugular venous distension, crackles on lung auscultation, pulmonary oedema on imaging, or hepatomegaly), then reduce or discontinue fluid administration.
- Determine the need for additional fluid boluses (250–500 mL in adults; 10–20 mL/kg in children) based on clinical response and improvement of perfusion targets and reassess for signs of fluid overload after each bolus.
- Perfusion targets include MAP (> 65 mmHg or ageappropriate targets in children), urine output (> 0.5 mL/kg/ hr in adults; 1 mL/kg/hr in children), and improvement of skin mottling and extremity perfusion, capillary refill, heart rate, level of consciousness, and lactate.
- Consider dynamic indices of volume responsiveness:
 passive leg raises, fluid challenges with serial stroke
 volume measurements, or variations in systolic pressure,
 pulse pressure etc. to guide volume administration beyond
 initial resuscitation based on local resources and
 experience.

6. PRONE POSITIONING

6.1.1 Prone positioning in intubated severe COVID-19 patients⁵⁻⁷

Early prone positioning for 16-18 hours should be considered in COVID-19 patients with ARDS when PaO2/FiO2 is <150 mmHg on FiO2 >0.6 and PEEP >10 cm H2O). It helps to reduce ventilation perfusion mismatching, hypoxemia and shunting. It requires expertise and patient need to be monitored closely for possible complications- accidental extubation, tube dislodgement, worsening of hemodynamics and hypoxemia. Spinal injury, open chest wound, and 3rd trimester of pregnancy are contraindications of prone positioning.

6.1.2 Awake prone positioning in moderate to severe disease

UK Intensive care society advocates awake prone positioning to become standard of care for suspected or confirmed COVID-19 in patients requiring oxygen supplementation >28%. This will lead to better ventilation perfusion matching believed due to more homogenous lung aeration and strain distribution with recruitment of dorsal lung. Awake prone positioning appears to be safe and may slow or avoid the respiratory deterioration in COVID-19 patients requiring oxygen supplementation. The optimal frequency and duration of awake prone positioning in non-intubated patient is not known. Patients are required to adopt the prone position for 1 hour each, five sessions per day during waking hours.

PHOTOS BELOW TO DEMONSTRATE THIS

- 1. 30 minutes-2 hours: lying on your belly
- ३० मिनेट देखि २ घन्टाः घोप्टो परेर सुत्ने वा घोप्टो परेर कुइनो, घुँडा र टाउकोले टेकेर बस्ने





- 2. 30 minutes-2 hours: lying on your right side
- २. ३० मिनेट देखि २ घन्टाः दाहिने कोल्टो फर्कने



Then back to position #1. Lying on your belly अनि फेरि शुरुको (घोप्टो) आसनमा फर्कने







Self Positioning Guide. Elmhurst Hopsital_SB आफँ आसन बदल्ने निर्देशिका (एल्महर्स्ट अस्पताल, अमेरिका)

The awake prone positioning is advised in patients who are alert, able to communicate and has stable hemodynamics. Close monitoring of mental status, oxygen saturation and blood pressure is essential to find out deterioration⁸

⁵ Koeckerling D, Barker J, Mudalige NL, *et al.* Awake prone positioning in COVID-19. *Thorax* 2020;**75**:833–834

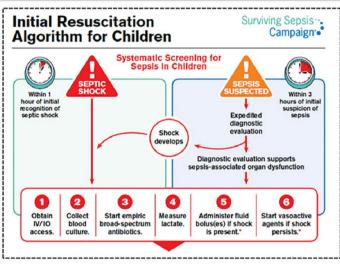
⁶ McNicholas, Bairbre et al. "Prone positioning in COVID-19 acute respiratory failure: just do it?." *British journal of anaesthesia* vol. 125,4 (2020): 440-443. doi:10.1016/j.bja.2020.06.003

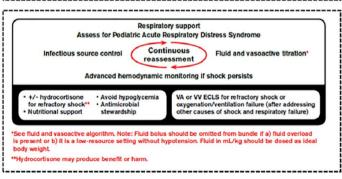
⁷ Ng Z, Tay WC, Ho CHB. Awake prone positioning for non-intubated oxygen dependent COVID-19 pneumonia patients. Eur Respir J 2020;56:2001198

⁸ Interim Clinical Guidance for Care of Patients with COVID-19 in Healthcare Settings. Nepal Medical Council, October 20202

Surviving Sepsis ... Fluid and Vasoactive-Inotrope Campaign: Management Algorithm For Children Healthcare Systems Healthcare Systems WITH Intensive Care WITHOUT Intensive Care SEPTIC SHOCK Abnormal Perfusion with Abnormal perfusion Abnormal perfusion WITHOUT WITH hypotension or without Hypotension hypotension · If signs of fluid overload If signs of fluid overload are absent, administer Do NOT give fluid are absent, administer fluid bolus, 10-20 mL/kg. fluid bolus, 10-20 mL/kg. bolus unless there are signs of · Repeat assessment of Assess hemodynamic dehydration with hemodynamic response response to fluid and ongoing fluid losses to fluid and consider fluid repeat fluid boluses, 10-20 (eg. diarrhea). boluses, 10-20 mL/kg, until mL/kg, until hypotension shock resolves or signs of resolves or signs of Start maintenance fluid overload develop. fluid overload develop. fluids. Assess cardiac function. Monitor Assess cardiac function (if available) hemodynamics Consider epinephrine closely. if there is myocardial Consider epinephrine/ dysfunction or epinephrine/ norepinephrine if Consider vasoactivenorepinephrine if shock inotropic support hypotension persists persists after 40-60 mL/ (if available). after 40 mL/kg or sooner if signs of fluid kg (or sooner if signs of fluid overload develop). overload develop. Fluid in mL/kg should be dosed as ideal body weight. Shock resolved, perfusion improved Do not give more Consider Monitor for signs/symptoms fluid boluses. of recurrent shock. maintenance fluids. *Hypotension Presence of all 3 World < 50 mm Hg < 60 mm Hg < 70 mm Hg Health Organization criteria: in healthcare systems WITHOUT in children in children in children OR cold extremities, prolonged intensive care is aged < 12 aged 1 to 5 aged > 5 capillary refill > 3 seconds, defined as either: months years years weak/fast pulse

This algorithm, 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 (see References and resources).





Algorithm 7. Delivering increasing oxygen in ADULT

How to deliver increasing oxygen



Place prongs inside the nostril Hook tubing behind ears. Flow rates higher than 5 L will dry mucous membranes.



Secure mask firmly on face over nose and mouth. Pull strap over head.



Make sure bag is full to deliver highest oxygen concentration. An empty bag is dangerous.

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



- → Use face mask
- → Increase oxygen to 6–10 L/min
- → Assess response



- → Use face mask with reservoir
- Increase oxygen to 10-15 L/min
- Make sure bag inflates
- → Call for help from district dinician
- → Assess response

If increasing respiratory distress or SpO₅ < 90%^a, transfer to a hospital with available invasive mechanical ventilator possible

- → Call for help from district dinician for possible tracheal intubation
- Start manual ventilation (bagging) with high-oxygen flow

Estimating FiO, when delivering oxygen

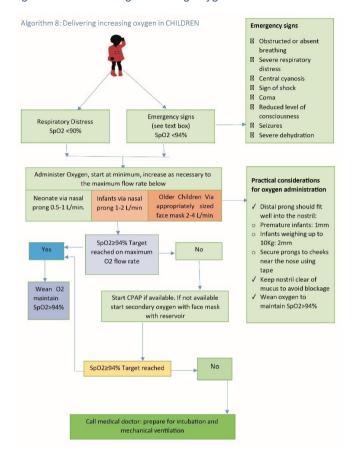
Adults

- 2-4 L/min ~ Fi0, 0.28-0.36
- 5 L/min ~ Fi0, 0.40
- 6-10 L/min ~ Fi0, 0.44-0.60 • 10-15 L/min ~ Fi0. 0.60-0.95

- * Patients presenting with emergency signs should receive oxygen therapy if SpO₂ is < 94%.
- Emergency signs:

 Obstructed or absent breathing
- Severe respiratory distress
- Central cyanosis
 Signs of shock, defined as cold extremities with capillary refill time > 3 sec and weak and fast pulse
- Coma (or seriously reduced level of consciousness)
- Seizures
 Seizures
 Signs of severe dehydration:
 lethargy or unconscious, sunken
 eyes, very slow return after
 pinching the skin.

Algorithm 8: Delivering increasing oxygen in CHILDREN



Footnotes for algorithm boxes 1-6

- **a** If altitude is higher than 1000 m, then the correction factor should be calculated as follows: PaO2/FiO2 x barometric pressure/760.
- **b** When PaO2 is not available, SpO2/FiO2 ≤ 315 suggests ARDS (including in non-ventilated patients).
- c Oxygenation Index (OI) is an invasive measurement of the severity of hypoxaemic respiratory failure and may be used to predict outcomes in paediatric patients. It is calculated as follows: percentage of fraction of inhaled oxygen multiplied by the mean airway pressure (in mmHg), divided by the partial pressure of arterial oxygen (in mmHg). Oxygen saturation index (OSI) is a non-invasive measurement and has been shown to be a reliable surrogate marker of OI in children and adults with respiratory failure. OSI replaces PaO2 with oxygen saturation as measured by pulse oximetry (SpO2) in the OI equation.
- d The SOFA score ranges from 0 to 24 and includes points related to six organ systems: respiratory (hypoxaemia defined by low PaO2/FiO2); coagulation (low platelets); liver (high bilirubin); cardiovascular (hypotension); central nervous system (low level of consciousness defined by Glasgow Coma Scale); and 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 (62).
- **e** SIRS criteria: abnormal temperature (> 38.5 °C or < 36 °C); tachycardia for age or bradycardia for age if < 1 year; tachypnoea for age or need for mechanical ventilation; abnormal white blood cell count for age or > 10% bands.

Abbreviations: BP blood pressure; bpm beats per minute; CPAP continuous positive airway pressure; FiO2 fraction of inspired oxygen; MAP mean arterial pressure; NIV non-invasive ventilation; OI Oxygenation Index; OSI Oxygenation Index using SpO2; 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

Indications of intubation

- Worsening respiratory distress with SpO2 < 90% despite oxygen supplementation with non-rebreathing face mask and or failure of HFNO/NIV
- PaO₂/FiO₂ <150 mmHg
- Full airborne precautions need to be taken during bag and mask ventilation and intubation.

High flow Nasal Oxygen (HFNO) and Non-invasive ventilation (NIV) 9,10

For adult patients with acute hypoxic respiratory failure despite supplementation of oxygen by non-rebreather mask, and no urgent indication for intubation, high flow nasal oxygen is feasible. Almost half of those who receive it can be successfully weaned without the need for mechanical ventilation. It appears to be safe for health care workers and could well liberate critical ICU resources. If HFNO is not available, non-invasive ventilation can be tried. Awake prone positioning can be considered in alert and hemodynamic patient on NIV or HFNO.

Precautions:

 It should not be used in patients who needs urgent intubation such as hypoxaemic respiratory failure and haemodynamic instability, multiorgan failure or abnormal mental status.

⁹ G.L. Calligaro et al., The utility of high-flow nasal oxygen for severe COVID-19 pneumonia in a resource-constrained setting: A multi-centre prospective observational study, EClinicalMedicine (2020), https://doi.org/10.1016/j.eclinm.2020.100570

¹⁰ Guy, Tiphaine et al. "High-flow nasal oxygen: a safe, efficient treatment for COVID-19 patients not in an ICU." *The European respiratory journal*, 2001154. 9 Sep. 2020, doi:10.1183/13993003.01154-2020

- 2. Patients receiving a trial of HFNO or NIV should be in a monitored by personnel experienced with HFNO and/or NIV and capable of performing endotracheal intubation in case the patient acutely deteriorates or does not improve after a short trial (about 1 hour). Intubation should not be delayed if the patient acutely deteriorates or does not improve after a short trial. Adult HFNO systems can deliver 60 L/min of gas flow and FiO2 up to 1.0. Paediatric circuits generally only handle up to 25 L/min, and many children will require an adult circuit to deliver adequate flow.
- 3. Aerosols generation is a concern and precautions should be taken including putting surgical mask over HFNO.
- Viral filters or high efficiency particulate air (HEPA) filter should be used in NIV.

7.

7.1 CORTICOSTEROID THERAPY AND COVID-19

In patients with severe or critical COVID-19 disease dexamethasone is strongly recommended. Hydrocortisone or prednisolone can be used when dexamethasone is not available. Other indication of corticosteroids is patients requiring supplemental oxygen to meet their desired oxygen level. Continue corticosteroids for up to 10 days unless there is a clear indication to stop early which includes discharge from hospital or a hospital supervised virtual COVID ward.

Systemic corticosteroid therapy of:

6 mg Dexamethasone orally or intravenously daily for 10 days (or until hospital discharge whichever comes earlier) for adult and 150 micrograms/kg (as a base) orally, nasogastrically or intravenously once a day for 10 days (max 6 ma) in children

OR

50 mg of Hydrocortisone intravenously every 8 hours for 10 davs

OR

Prednisolone 40 mg once daily for 10 days for adults and 1 mg/kg orally, nasogastrically or intravenously once a day for 10 days (max 40 mg; doses can be rounded as per routine clinical practice)

Do not routinely use corticosteroids to treat COVID-19 in people who do not need supplemental oxygen, unless there is another medical indication to do so.

Pregnant Women: WHO recommends antenatal corticosteroid therapy for women at risk of preterm birth from 24 to 34 weeks of gestation when there is no clinical evidence of maternal infection, and adequate childbirth and newborn care is available. However, in cases where the woman presents with mild COVID-19, the

clinical benefits of antenatal corticosteroid might outweigh the risks of potential harm to the mother. In this situation, the balance of benefits and harms for the woman and the preterm newborn should be discussed with the woman to ensure an informed decision. This assessment may vary depending on the woman's clinical condition, her wishes and that of her family, and available health care resources⁵.

7.2 Remdesivir

The conditional recommendation against the use of REMDESIVIR for patients with COVID-19 in Therapeutcs and COVID-19: living quideline - World Health Organizaton is based on the evidence of possibly no effect on mortality, need for mechanical ventilation. recovery from symptoms and other patient-important outcomes. though of albeit of low certainty.

NICE guidelines put conditional recommendation for COVID-19 pneumonia in adults, and young people 12 years and over weighing 40 kg or more, who are in hospital and on supplemental oxygen but not on invasive mechanical ventilation for 5 days with a dose of 200 mg on day 1 followed by 100 mg daily for 4 days.

7.3 Tocilizumab

Tocilizumab is recommended (strong recommendation) to adults in hospital with COVID-19 if all of the following apply:

- they are having or have completed a course of corticosteroids such as dexamethasone, unless they cannot have corticosteroids
- they have not had another interleukin-6 inhibitor during this admission
- there is no evidence of a bacterial or viral infection (other than SARS-CoV-2) that might be worsened by tocilizumab.
- And they either:

¹¹ Corticosteroids for COVID-19: Living guidance: 2 September 2020: WHO ¹² Therapeutics and COVID-19: living guideline - World Health Organization (WHO), 31 March 2021

3 COVID-19 rapid guideline: managing COVID-19 NICE guideline Pub-

lished: 23 March 2021 www.nice.org.uk/guidance/ng191

- need supplemental oxygen and have a CRP level of 75 mg/L or more, or
- are within 48 hours of starting high-flow nasal oxygen, continuous positive airway pressure, noninvasive ventilation or invasive mechanical ventilation

Common side effects are URT infections, nasopharyngitis, headache, hypertension, increased alanine aminotransferase (adverse effects are ALT), injection site reactions)

Dosage: A single dose of 8 mg/Kg by intravenous infusion (total dose not exceeding 800 mg)

Convalescent plasma therapy:

Only a few clinical trials of convalescent plasma have produced results, and the evidence, at least so far, has not been convincing enough to endorse it beyond use as an experimental therapy. WHO: There are a number of side effects ranging from mild fevers to severe lung injuries or circulatory overload.

Not recommended agents

WHO recommends AGAINST administering HYDROXYCHLOROQUINE or CHLOROQUINE OR IVERMECTIN or LOPINAVIR/RITONAVIR for treatment of COVID-19 to patents with any disease severity and any duration of symptoms.

8. TREATMENT OF OTHER ACUTE AND CHRONIC INFECTIONS IN PATIENTS WITH COVID-19

8.1 <u>Use of antimicrobials:</u> The prevalence of acute coinfections or secondary infections coinciding with COVID-19 has been not adequately described but appears to be low and will be based on local factors and endemic or other emerging infections.

The following recommendations for patients with:

- suspected or confirmed mild COVID-19,
- o against the use of antibiotic therapy or prophylaxis
- ! suspected or confirmed moderate COVID-19,
- antibiotics should not be prescribed unless there is clinical suspicion of a bacterial infection.

Suspected or confirmed severe COVID-19

- ✓ Administer empiric antimicrobials to treat all likely pathogens, based on clinical judgment, patient host factors and local epidemiology, and this should be done as soon as possible (within 1 hour of initial assessment if possible), ideally with blood cultures obtained first. Antimicrobial therapy should be assessed daily for de-escalation and reviewed to treat based on microbiology results.
 - ADULT: Ceftriaxone 1-2g, once daily <u>PLUS</u>
 Azithromycin 500mg, once daily <u>OR</u> Ampicillin 2g IV, 4 times day <u>PLUS</u> Azithromycin 500mg, once daily
 - CHILD: Ampicillin 50mg/kg or Benzylpenicillin 50,000U/kg IM or IV every 6 hours for 5 days PLUS Gentamycin 7.5 mg/kg IM or IV once daily for at least 5 days
- ✓ If bacterial pneumonia or sepsis is suspected use Ceftriaxone or Amoxicillin-Clavulanic Acid
 - Add Azithromycin for atypical coverage of pneumonia substitute with Doxycycline if allergic to macrolides

- 8.2 Chronic infections: It is currently unknown whether immunosuppression caused by chronic co-infections such as HIV put persons at greater risk for severe COVID-19 disease. However, people living with HIV with advanced disease have an increased risk of opportunistic infections (notably TB) and related complications in general. Facility-based HIV testing services should be continued and those newly diagnosed should start antiretroviral therapy as soon as possible. For people living with HIV already on treatment, continuity of antiretroviral therapy and prophylaxis for co-infections is essential, with multimonth prescribing.
- 8.3 Endemic diseases (co-morbidities) e.g. malaria, dengue, typhoid etc. should be treated in accordance with the appropriate treatment protocols at the facility level. Many of the endemic diseases have similar clinical presentation as COVID-19 disease, use the differential diagnostic matrix (Annex 4) to facilitate diagnosis.

9 USE OF OSELTAMIVIR

Oseltamivir is currently not proven to be effective for COVID-19. Oseltamivir can be used when influenza is suspected or known to be circulating. If testing for influenza is not possible, empirical treatment is indicated.

Oseltamivir has not been evaluated in severely ill patients, pregnancy, or pediatric populations. Oseltamivir should be used with caution.

10 USE OF VASOPRESSORS

In adults, administer vasopressors when shock persists during or after fluid resuscitation. The initial blood pressure target is MAP ≥ 65 mmHg in adults and improvement of markers of perfusion. Norepinephrine is considered the first-line treatment in adult patients; epinephrine or vasopressin can be added to achieve the MAP target. Because of the risk of tachyarrhythmia, reserve dopamine for selected patients with low risk of tachyarrhythmia or those with bradycardia. For children, refer to algorithm 6. Vasopressors (i.e. norepinephrine, epinephrine, vasopressin and dopamine) are most safely given through a central venous catheter at a strictly controlled rate, but it is also possible to safely administer them via peripheral vein and intraosseous,

11 USE OF INOTROPES

Side-effects of inotropes, such as dobutamine, include tachyarrhythmias and hypotension due to peripheral vasodilation. 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.

12 NON-COMMUNICABLE DISEASES (NCD) and COVID-19

When caring for patients with suspected and confirmed COVID-19 with underlying NCDs, we may need to continue or modify previous medical therapy according to the patient's clinical condition.

12.1 Antihypertensive patients: antihypertensive drugs should not routinely be stopped in patients with COVID-19, but therapy may need to be adjusted based on general considerations of patients with acute illness, with reference to maintaining normal blood pressure and renal function

- 12.2 Asthmatic patients: Respiratory viral infections, such as COVID-19, can trigger and worsen asthma symptoms. However, at present, there is no evidence suggesting that people with asthma are any more likely to contract COVID-19 than anyone else. Continue to administer regular treatment for asthmatic patients, monitor peak flow and respiratory function tests according to facility protocol.
- 12.3 Diabetic patients: are at a higher risk of severe COVID 19 disease and associated mortality due to COVID-19. However, controlled diabetic patients have more favorable outcomes than non-controlled. Monitor blood sugar levels regularly and manage according to facility protocol.
- 12.4 Patients with myocarditis: refer to cardiologist consultant for appropriate management.

13 PREVENTION OF COMPLICATIONS IN HOSPITALIZED AND CRITICALLY ILL PATIENTS WITH COVID-19

13.1 Thromboembolism

Adults and adolescents:

- Use low molecular weight heparin, if contraindicated use mechanical prophylaxis including intermittent pneumatic devices. Enoxaparin 40 mg by subcutaneous injection every 24h or UFH subcutaneously 12 hourly.
- Monitor patients for signs and symptoms suggestive of thromboembolism.
- If thromboembolism is suspected proceed immediately to the area/center with diagnostic and management facility.

13.2 Other complications

These interventions are based on surviving sepsis or other guidelines and are based on limited feasible recommendations based on high-quality evidence.

Table 3. Anticipated outcome*	Interventions
Reduce days if 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 of the above as a bundle of care (may also reduce delirium); such as the Awakening and Breathing Coordination, Delirium assessment/management,
Reduce incidence of ventilator-associated pneumonia	and Early mobility (ABCDE) 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 blood-stream infection	Use a checklist with completion verified by a real-time observer as 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 gastro- intestinal (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, and higher organ failure score
Reduce the development of antimicrobial resistance Reduce the development of adverse drug effects Promote appropriate antimicrobial prescribing and use during the COVID19 pandemic	Utilize de-escalation protocols as soon as patient is clinically stable and there is no evidence of bacterial infection Expose patient to empiric antimicrobial therapy for the shortest time possible, to prevent nephrotoxicity, cardiac and other side-effects from unnecessary antimicrobial use 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

*Refer to: Clinical Management of COVID-19 Interim Guidance; May 2020; Page 27: WHO

13.3 Nutrition

- ✓ Start enteral feeding early.
- ✓ Nasogastric or orogastric tube feeding in intubated patients
- Consider parenteral nutrition if enteral feeding is not tolerated despite prokinetics use or if enteral feeding is contraindicated.

14 MANAGEMENT OF PREGNANT AND LACTATING WOMEN AND COVID-19¹⁴⁻¹⁵

There is limited data on the clinical presentation and maternal and perinatal outcomes of COVID-19 disease during or after pregnancy. Existing evidence has not identified major risks or complications in babies born to mothers with COVID-19. However, COVID-19 Pregnant and post-partum women: with pre-existing or pregnancy related co-morbidities (e.g. pregnancy induced hypertension, gestational diabetes), should also be considered for close monitoring.

All pregnant women with history of contact with a person with confirmed COVID-19 be carefully monitored considering asymptomatic transmission of COVID-19. Pregnant or recently pregnant women with suspected or confirmed mild COVID-19 may not require acute care in hospital, unless there is concern for rapid deterioration or an inability to promptly return to hospital; but isolation to contain virus transmission is recommended, and can be done at a health facility, community facility or at home, according to established COVID-19 care pathways but women with moderate or severe COVID-19 require acute care in the hospital, as there is concern for rapid deterioration.

WHO recommends that induction of labour and caesarian section should only be undertaken when medically justified and based on maternal and fetal conditions. COVID-19 status alone is not an indication for caesarian sectio.

- Delayed umbilical cord clamping (not earlier than 1 minute) is recommended for improved maternal and infant health and nutritional outcomes. The risk of transmission of COVID-19 through blood is likely to be minimal
- Mothers with suspected or confirmed COVID-19 should be encouraged to initiate or continue to breastfeed. Mothers should be counselled that the benefits of breastfeeding substantially outweigh the potential risk for transmission
- ✓ Symptomatic mothers who are breastfeeding or practicing skin-to-skin contact or kangaroo mother care should practice respiratory hygiene (wear a cloth mask), including during feeding perform hand hygiene before and after contact with

- the child, and routinely clean and disinfect surfaces which the symptomatic mother has been in contact with.
- ✓ See algorithms 1, 2, 3, 4, 5, 7 for management of pregnant women according to disease severity.

15 MANAGEMENT OF NEUROLOGICAL AND MENTAL MANIFESTATIONS ASSOCIATED WITH COVID-19¹

In patients with COVID-19, measures to prevent delirium, an acute neuropsychiatric emergency, should be implemented; and patients be evaluated using standardized protocols, for the development of delirium. If detected, then immediate evaluation by a clinician is recommended to address any underlying cause of delirium and treat appropriately.

- Manage any underlying cause of delirium by monitoring oxygenation and fluid status, correcting metabolic or endocrine abnormalities, addressing co-infections, minimizing the use of medications that may cause or worsen delirium, treating withdrawal from substances, understanding and minimizing the effects of any harmful drug-drug interactions and maintaining normal sleep cycle as much as possible.
- ✓ In patients receiving invasive ventilation, minimize continuous or intermittent sedation, targeting specific titration endpoints (light sedation unless contraindicated) or with daily interruption of continuous sedative infusions, to reduce delirium
- ✓ In patients experiencing agitation (defined as marked restlessness or excessive motor activity, often accompanied by anxiety), use calming communication strategies and attempt to reorient the person. Acute pain due to physical illness or air hunger should be considered as triggers for agitation and need to be addressed immediately. If the person continues to be agitated despite the strategies

WHO recommendations for induction of labour. Geneva: World Health Organization; 2011 (https://www.who.int/reproductivehealth/publications/maternal_perinatal_health/9789241501156/en/, accessed 14 May 2020).
 https://www.cdc.gov/breastfeeding/breastfeeding-special-circumstances/maternal-or-infant-illnesses/covid-19-and-breastfeeding.html

- described above and is experiencing severe distress, it may be necessary to use psychotropic medications.
- Provide basic mental health and psychosocial support (MHPSS) for all persons with suspected or confirmed COVID-19.
- Prompt identification and assessment for anxiety and depressive symptoms in the context of COVID-19 is important, for the management of new anxiety and depressive symptoms.
- ✓ If a person's anxiety or depressive symptoms persist beyond recovery from COVID19 and/or discharge from the hospital, then an underlying anxiety or depressive disorder may be suspected, and a mental health professional should be consulted, and these conditions should be managed appropriately.

16. REHABILITATION FOR PATIENTS WITH COVID-191

Patients with COVID-19, regardless of the disease severity, might present with persistent symptoms and a functional decline which may not be obviously apparent. Few common symptoms are new illness-related fatigue, breathlessness, PTSD symptoms, pain, voice change, cough, dysphagia, anxiety, depression, and problems with concentration, memory and continence are increasingly reported in patients who have recovered from acute COVID-19 even up to months. Prior to hospital discharge, COVID-19 patients should be screened for rehabilitation needs in order to facilitate onward referral. A multidisciplinary approach is recommended for better outcome. Nevertheless, the following interventions are useful

- Patients with COVID-19, should be provided with education and support for the self-management of breathlessness and resumption of activities.
- All rehabilitating patients should be educated about resuming everyday activities conservatively at an appropriate pace that is safe and manageable for energy levels within the limits of current symptoms and should not be pushed for post-

- exertional fatigue. A gradual increase in exercise should be based on symptoms.
- ✓ For patients having difficulties with memory, concentration and problem solving, education should be provided, and advice on strategies to help establish expectations (including from family members) and to alleviate stress and anxiety.
- ✓ For patients with anxiety, depression and PTSD, basic mental health and psychosocial support by appropriately trained health or non-health workers should be provided.

ANNEX 1: APPROPRIATE USE OF PPE ACCORDING TO TASK

Guidelines for use of personal protective equipment

(Developed by the Expert Team of NMC

and Government of Nepal with reference from WHO, published on March 26, 2020)

A. For Aerosol Generating procedures: Dental procedures, bronchoscopy, Upper GI Endoscopy, ENT procedures, Nebulization, Intubation of a patient, CPR, Non-invasive ventilation, endotracheal suctioning, when obtaining nasopharyngeal or oropharyngeal

suctioning, when obtaining hasopharyngeal or oropharyngeal swab, etc. **in Covid-19 suspected or**

confirmed cases health personnel need to use the following protective equipment: Category I PPE:

- a. N-95 mask
- b. Goggles or visor
- c. Gloves (loose gloves acceptable)
- d. Water resistant OR standard disposable gowns
- e. Cap: Regular disposable

B. For Non aerosol generating covid-19 suspected or confirmed patients: Health personnel need to use the following protective equipment: Category II PPE:

- a. Surgical mask (seal the top edge with tape)*
- b. Goggles or visor
- c. Gloves (loose gloves acceptable)
- d. Water resistant or standard disposable gowns
- e. Cap: Regular disposable

C. For Physician/Staff running the fever/screening clinics the following PPE is recommended: Category II PPE

- a. Surgical mask, (seal the top edge with a tape)*
- b. Goggles or visor
- c. Water resistant or standard disposable gowns
- d. Regular disposable Cap
- e. Gloves (loose gloves acceptable)

D. For escorts or drivers, the following PPE is recommended: Category III PPE:

- a. Surgical masks
- b. Gloves
- c. If physical contact is expected, depending on circumstances, a gown PLUS goggles or face shield are also recommended, otherwise need to maintain minimum 2-meter distance from the

patient.

- d. The patient should be given surgical mask and instructed to perform hand-hygiene.
- E. For Laboratory staff: category II or III PPE; depending upon the chance of splash:
- a. surgical mask
- b. Gown
- c. Loose Gloves
- d. Eye protection (if risk of splash)
- F. For all staff, including health care workers involved in any activity that does not involve contact with
- COVIĎ-19 patients and working in other areas of patient transit (e.g. wards, corridors). **No PPE**

required.

For Everyone:

- o Maintain 3-6 feet distance while visiting patients, if no need to touch the patient.
- o Mandatory hand-hygiene after each use of PPE and between patients.

Mandatory surface cleaning of bed or furniture with 0.5% Chlorine disinfectant (Virex* or similar) between each patient in OPD or in an inpatient setting.

*Use N-95 masks if close contact with COVID-19 suspect or confirmed case expected.

ANNEX 2: SCREENING QUESTIONNAIRE

SCREENING QUESTIONNAIRE:

All individuals presenting to the OPD or ER entrance should be screened with the following questions:

- a. Symptoms: Do you have any of the following symptoms?
 - Cough? Fever? Shortness of breath? (common)
 - Sore throat, headache or body ache, new loss of smell, or taste? (less common)
- **b**. Travel history or contact with traveler:

Have you

- · Recently returned from travel in, or been living in, an affected area in the past 2 weeks?
- · Been in close contact in the past 2 weeks with someone returning from travel in an affected area?

c. Exposures:

Did you have any exposures to any of the following?

- · Close contact with anyone with fever or respiratory illness of unknown cause
- · Known or suspected COVID-positive contact

How to Handwash?

WASH HANDS WHEN VISIBLY SOILED! OTHERWISE, USE HANDRUB

Duration of the entire procedure: 40-60 seconds



Wet hands with water:



Apply enough soap to cover all hand surfaces;



Rub hands palm to palm:



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



Palm to palm with fingers interlaced;



Backs of fingers to opposing palms with fingers interlocked;



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



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



Rinse hands with water;



Dry hands thoroughly with a single use towel;



Use towel to turn off faucet;



Your hands are now safe.

How to Handrub?

RUB HANDS FOR HAND HYGIENE! WASH HANDS WHEN VISIBLY SOILED

Duration of the entire procedure: 20-30 seconds







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

Rub hands palm to palm;



Right palm over left dorsum with interlaced fingers and vice versa:



Palm to palm with fingers interlaced;



Backs of fingers to opposing palms with fingers interlocked:



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



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



Once dry, your hands are safe.

Protect others from getting sick

When coughing and sneezing cover mouth and nose with flexed elbow or tissue





Throw tissue into closed bin immediately after use

Clean hands with alcohol-based hand rub or soap and water after coughing or sneezing and when caring for the sick



HOW TO WEAR A MEDICAL MASK SAFELY

who.int/epi-win

Do's





Wash your hands before touching the mask

Inspect the mask for tears or holes



Find the top side, where the metal piece or stiff edge is



Ensure the colored-side faces outwards



Place the metal piece or stiff edge over your nose



Cover your mouth, nose, and chin



Adjust the mask to your face without leaving gaps on the sides



Avoid touching the mask



Remove the mask from behind the ears or head



Keep the mask away from you and surfaces while removing it



Discard the mask immediately after use preferably into a closed bin



Wash your hands after discarding the mask

Don'ts -



Do not wear a loose mask



Do not touch the front of the mask



Do not remove the mask to talk to someone or do other things that would require touching the mask



Do not Use a ripped or damp mask



Do not wear the mask only over mouth or nose



Do not leave your used mask within the reach of others



Do not re-use the mask

Remember that masks alone cannot protect you from COVID-19. Maintain at least 1 metre distance from others and wash your hands frequently and thoroughly, even while wearing a mask.





ANNEX 4: DIFFERENTIAL DIAGNOSIS MATRIX

Symptoms	COVID-19	Flu	Common Cold	Malaria	Dengue	Scrub Typhus	Allergies
Fever	Common	Common	Rare	Соттоп	Соттоп	Common often with chills	Rare
Dry Cough	Common	Less Common	Less Common	Rare	Rare	Less Common	Less Common
Tiredness	Common	Common	Less Common	Common	Common	Less Common	Less Common
Shortness of Breath	Less Common	Less Common	Rare	Rare	Less Common	Common	Less Common
Aches and Pains	Less Common	Common	Rare	Less Common	Common	Less Common	Rare
Headache	Less Common	Common	Rare	Соттоп	Common	Соттоп	Less Common
Sore throat	Less Common	Less Common	Common	Rare	Rare	Rare	Rare
Diarrhea	Less Common	Less Common	Rare	Common	Rare	Rare	Rare
Stuffy Nose	Rare	Less Common	Common	Rare	Rare	Rare	Common
Running Nose	Rare	Common	Common	Rare	Less Common	Rare	Common
Sneezing	Rare	Rare	Common	Rare	Rare	Rare	Common
Rash	Less Common	Rare	Rare	Rare	Common, Common, exanthematous maculopapular rash, petechiae	Common, maculopapular	Common

ANNEX 5: NATIONAL EARLY WARNING SCORE 2

The National Early Warning Score (NEWS) 2 is based on a simple aggregate scoring system allocated to following physiological measurements which are routinely recorded in hospitalized patients.

- 1. Respiration rate
- 2. Oxygen saturation
- 3. Systolic blood pressure
- 4. Pulse rate
- Level of consciousness or new confusion*
- Temperature

*The patient may respond to questions coherently, but there may be some new-onset confusion, disorientation and/or agitation which score 3 on NEWS system (3 or 4 in GCS).

The use of NEWS 2 will ensure that patients who are deteriorating, or at risk of deteriorating, will have a timely initial assessment by a competent clinical decision maker. NEWS2 should supplement clinical judgement in assessing the patient's condition.

A score is allocated to each parameter as they are measured. The score is then aggregated and frequency of observations and review by appropriate team is decided. If patient is receiving oxygen supplementation, 2 points will be added to score.

In the context of **COVID-19 infection**, oxygen requirement of a patient might increase rapidly but this may not result in additional significant increase (>3) in NEWS 2 score. It is recommended that in patients with COVID-19, all staffs should be aware that any increase in oxygen requirement should trigger an escalation call to a competent clinical decision maker along with increase in observations to at least hourly until a clinical review.

Chart 1: The NEWS scoring system

Physiological				Score			
parameter	3	2	1	0	1	2	3
Respiration rate (per minute)	≤8		9–11	12–20		21–24	≥25
SpO ₂ Scale 1 (%)	≤91	92–93	94–95	≥96			
SpO ₂ Scale 2 (%)	≤83	84–85	86–87	88–92	93–94 on	95–96 on	≥97 on
	303	04-03	00-07	≥93 on air	oxygen	oxygen	oxygen
Air or oxygen?		Oxygen		Air			
Systolic blood pressure (mmHg)	≤90	91–100	101–110	111–219			≥220
Pulse (per minute)	≤40		41–50	51–90	91–110	111–130	≥131
Consciousness				Alert			CVPU
Temperature (*C)	≤35.0		35.1–36.0	36.1–38.0	38.1–39.0	≥39.1	

Chart 2: NEWS thresholds and triggers

NEW score	Clinical risk	Response
Aggregate score 0–4	Low	Ward-based response
Red score Score of 3 in any individual parameter	Low-medium	Urgent ward-based response*
Aggregate score 5–6	Medium	Key threshold for urgent response*
Aggregate score 7 or more	High	Urgent or emergency response**

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

^{**}The response team must also include staff with critical care skills, including airway management.

Chart 3: NEWS observation chart

NEWS key		FULL NAME																					
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Use Scale 2 if target narge is 85-92% ag in hypercapnic respiratory fallure	≥93 on air																						≥93 on
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	Device						\perp		\perp				_				_				Ш		Device
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Chart 4: Clinical response to the NEWS trigger thresholds

NEW score	Frequency of monitoring	Clinical response
0	Minimum 12 hourly	Continue routine NEWS monitoring
Total 1–4	Minimum 4–6 hourly	Inform registered nurse, who must assess the patient Registered nurse decides whether increased frequency of monitoring and/or escalation of care is required
3 in single parameter	Minimum 1 hourly	Registered nurse to inform medical team caring for the patient, who will review and decide whether escalation of care is necessary
Total 5 or more Urgent response threshold	Minimum 1 hourly	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
Total 7 or more Emergency response threshold	Continuous monitoring of vital signs	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 facility, ie higher-dependency unit or ICU Clinical care in an environment with monitoring facilities

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ANNEX 6: MULTISYSTEM INFLAMMATORY SYNDROME IN CHILDREN AND ADOLESCENTS

WITH COVID-19^{16,17}

Multisystem Inflammatory syndrome in children (MIS-C) is a new potentially lethal disease temporally associated with COVID-19 disease. WHO has developed a preliminary case definition.

Children and adolescents 0–19 years of age with fever ³ 3 days **AND** two of the following:

- Rash or bilateral non-purulent conjunctivitis or mucocutaneous inflammation signs (oral, hands or feet)
- b. Hypotension or shock.
- Features of myocardial dysfunction, pericarditis, valvulitis, or coronary abnormalities (including ECHO findings or elevated Troponin/NT-proBNP),
- d. Evidence of coagulopathy (by 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 inflammation, including bacterial sepsis, staphylococcal or streptococcal shock syndromes.

AND

Evidence of COVID-19 (RT-PCR, antigen test or serology positive), or likely contact with patients with COVID-19.

The common clinical symptoms at presentation are fever (100%), abdominal pain or diarrhea (73%), and vomiting (68%). Signs and

symptoms of MIS-C typically appears after three to four weeks after COVID-19 infection and may progress rapidly to shock and cardiorespiratory failure in many cases. Various immunomodulatory treatments (intravenous immunoglobulin, steroids) have been used. Patients may need mechanical ventilation apart from supportive therapy. Early recognition and referral to a tertiary care center is the key for improving survival.

¹⁶ Ahmed M, Advani S, Moreira A et al. Multisystem inflammatory syndrome in children: A systematic review. E Clinical Medicine 26 (2020) 100527. https://doi.org/10.1016/j.eclinm.2020.10052

¹⁷ Multisystem inflammatory syndrome in children and adolescents with COVID-19. Scientific brief. 15 May 2020

