

Novel Coronavirus 2019 (2019-nCoV) Infection: Part I - Preparedness and Management in the Pediatric Intensive Care Unit in Resource-limited Settings

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First reported in China, the 2019 novel coronavirus has been spreading across the globe. Till 26 March, 2020, 416,686 cases have been diagnosed and 18,589 have died the world over. The coronavirus disease mainly starts with a respiratory illness and about 5-16% require intensive care management for acute respiratory distress syndrome (ARDS) and multi-organ dysfunction. Children account for about 1-2% of the total cases, and 6% of these fall under severe or critical category requiring pediatric intensive care unit (PICU) care. Diagnosis involves a combination of clinical and epidemiological features with laboratory confirmation. Preparedness strategies for managing this pandemic are the need of the hour, and involve setting up cohort ICUs with isolation rooms. Re-allocation of resources in managing this crisis involves careful planning, halting elective surgeries and training of healthcare workers. Strict adherence to infection control like personal protective equipment and disinfection is the key to contain the disease transmission. Although many therapies have been tried in various regions, there is a lack of strong evidence to recommend anti-virals or immunomodulatory drugs.

Keywords: COVID-19, Guideline, Pandemic, SARI, Treatment.

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The year 2020 started with the emergence of the 2019 novel corona virus (2019-nCoV) as a threat to the world; shortly afterwards the World Health Organization (WHO) declared it a pandemic. Having begun in China, globalization and travel led its spread all over the globe, overwhelming the healthcare resources and resulting in high mortality and morbidity. About 5% of adults, especially those with comorbidities, were critically ill and required intensive care unit (ICU) care [1]. People of all ages were found to be susceptible but severe illness was rare in children [2]. Most of the experience of critical care management of pediatric patients with coronavirus disease 2019 (COVID-19) is derived from the affected children of present epidemic in China, as well as from the previous coronaviral outbreaks *viz.* Severe acute respiratory syndrome (SARS) and Middle East respiratory syndrome (MERS). We write this review as a guidance statement for preparedness and managing children with suspected or confirmed COVID-19 requiring intensive care in a resource-limited setting like India.

BURDEN

Global: Till March 26, 2020, a total of 416,686 confirmed cases from 197 countries with 18,589 deaths have been reported by WHO. China has reported the maximum cases with a total of 81,869, followed by Italy with 69,176 cases. However, mortality is more in Italy with 6,820 (9.9%) deaths followed by China having 3,287 (4%) deaths. The United States of America has surpassed Spain and Germany over the last few days with 51,914 cases and 673 deaths [3].

Indian scenario: A total of 606 cases with 10 deaths have been reported from India as on March 26, 2020 as reported by the WHO. Among these cases, only one child from Kerala has been tested positive.

EPIDEMIOLOGY

The 2019-nCoV belongs to a group of enveloped positive-sense RNA viruses in the family, Coronaviridae with 4 genera *viz.*, alpha, beta, gamma and delta. Human coronaviruses (HCoV) belong to alpha and beta genus

and are mostly implicated in endemic respiratory infection with mild severity [4]. However, the novel coronaviruses infecting humans namely, SARS-CoV, MERS-CoV and SARS-CoV-2 are believed to have originated from bats with few intermediate hosts like civet cats, camels and pangolins [5]. RNA viruses mutate faster than DNA viruses, single-stranded viruses mutate faster than double-strand virus, and genome size appears to correlate negatively with mutation rate.

Transmission Characteristics

It is speculated that it originated in bat (genetic character matches to bat corona virus) then it got transmitted to pangolins, or scaly anteaters. Humans seem to be accidental host who got this virus from pangolins in Wuhan seafood market. Human to human transmission of COVID-19 started in Wuhan city, Hubei Province of China where it was initially labelled as ‘Pneumonia of unknown etiology’. Epidemiological investigation of early transmission dynamics revealed that 55% of the cases of COVID-19 during December, 2019 were linked to the human seafood wholesale market. The mean incubation period has been reported to be 5.2 days with the 95th centile being 12.5 days. The main modes of transmission include droplet and fomites followed by airborne transmission. Reproduction number of nCoV-19 is between 2.2 to 3.6, which is comparable to SARS-CoV but higher than MERS-CoV[6].

Less severe affection in children: Children less than 10 years of age accounted for 1% of the total cases [1]. The median age among pediatric cases was 6.7 years [7]. The lesser proportion of severe cases among children has been attributed to lesser opportunities for exposure and immaturity of angiotensin converting enzyme 2 receptors, which are proposed to be the binding sites for coronaviruses [8,9].

Case Fatality Rate

The overall case fatality rate as per China Centre for Disease Control and Prevention (CDC) is 2.3%, which is much lower compared to SARS (9.6%) and MERS (34%) but significantly higher compared to the latest H1N1 influenza pandemic (0.001 – 0.007%)[1]. However, as per WHO, the global case fatality rate is as high as 4.4% with absolute number of deaths already higher than the total fatality of SARS and MERS combined [10]. The case fatality reported from Italy is 7.2% which has gone up to 9.8% as per WHO (as on March 26, 2020) [11].

CLINICAL MANIFESTATIONS

The common clinical features reported in the critically ill patients include fever (98%), cough (77%), dyspnea

(63%), malaise (35%), myalgia, headache, nausea, vomiting and diarrhea [12]. A prospective study from China involving 171 children with confirmed COVID-19 reported fever (41%) with a median duration of 3 days (1-16), cough (48%), pharyngeal erythema (46%) tachypnea (28%) and diarrhea (8.8%). The cohort had 15% asymptomatic, 19% upper respiratory infection, and 65% pneumonia. Only 3 children (1.7%) required care and mechanical ventilation. All three of them had comorbidities, and one died [7].

ICU Requirements in COVID

The severe and critical categories require admission and management in ICU. Among adults, 7% of patients admitted with SARS-CoV-2 pneumonia required ICU care. The mean age of these ICU patients was 60 years with male: female ratio of 2:1 and 50% had chronic illness. Majority had Multi-organ dysfunction syndrome (MODS) with ARDS (67%), acute kidney injury (29%), liver dysfunction (29%) and cardiac injury (23%). Of the ICU admissions, 71% required mechanical ventilation, 35% vasoactive support, 17% renal replacement therapy and 11% ECMO. Mortality was as high as 61% among the critically ill [12]. As per unpublished data from Italy, 16% of admitted patients with COVID-19 needed ICU care [13]. In the Chinese pediatric cases, 5.9% of all pediatric cases belonged to the severe or critical categories. Based on the experience in managing community-acquired pneumonia, high-risk pediatric population includes children with underlying conditions such as congenital heart disease, broncho-pulmonary hypoplasia, airway/lung anomalies, severe malnutrition, and immunocompromised state; however, more information is needed in the setting of COVID-19 [2].

DIAGNOSIS

Case definitions for suspected, probable and confirmed COVID-19 cases as given by WHO are in **Box I** [16]. The largest series on children analyzing suspected and confirmed COVID cases is from the electronic data base of Chinese CDC [17]. Cases were suspected based on the presence of clinical features and exposure history. They also identified high-risk cases and categorized into groups based on severity (**Box II**).

Laboratory testing of suspected cases is based on clinical and epidemiological factors. Screening protocol should be adapted to local situation and may change with the evolution of the outbreak scenario in the local population. Recent testing strategy in India (as on March 20, 2020) given by ICMR is as per algorithm in **Fig. 1**[18]. Specimen handling for molecular testing would require Biosafety 2 (BSL-2) or equivalent facilities.

BOX I World Health Organization Case Definitions for Coronavirus Disease 19 (COVID-19)**Suspect case**

A. A patient with acute respiratory illness (fever and at least one sign/symptom of respiratory disease (e.g., cough, shortness of breath), AND with no other etiology that fully explains the clinical presentation AND a history of travel to or residence in a country/area or territory reporting local transmission (See situation report) of COVID-19 disease during the 14 days prior to symptom onset.

OR

B. A patient with any acute respiratory illness AND having been in contact with a confirmed or probable COVID-19 case (see definition of contact) in the last 14 days prior to onset of symptoms

OR

C. A patient with severe acute respiratory infection (fever and at least one sign/symptom of respiratory disease (e.g., cough, shortness breath) AND requiring hospitalization AND with no other etiology that fully explains the clinical presentation.

Probable case

A suspect case for whom testing for COVID-19 is inconclusive. Inconclusive being the result of the test reported by the laboratory

Confirmed case

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

Source: World Health Organization [16].

BOX II Risk Stratification and Severity Categorization for Coronavirus Disease-19 (COVID-19)**High risk cases**

Clinical features	Fever, respiratory/ digestive symptoms, fatigue
Laboratory tests	Leukopenia, lymphopenia, high C-reactive protein
Radiology	Abnormal chest ray

Severity categorization

Asymptomatic infection	No clinical or radiological features but tested positive
Mild	Upper respiratory or gastrointestinal symptoms and signs
Moderate	Clinical/radiological features of lower respiratory involvement
Severe	Presence of dyspnea or hypoxemia requiring oxygen, refusal to feed, altered sensorium
Critical	Organ dysfunction including Acute respiratory distress syndrome (ARDS), shock, encephalopathy, myocardial dysfunction, coagulation dysfunction and acute kidney injury

Modified From Dong, et al. [17].

Attempts to culture the virus require minimum of BSL-3 facilities [19].

Type of Sample

Upper respiratory specimens: nasopharyngeal and oropharyngeal swabs; both swabs are placed together in a viral transport medium and transported to the laboratory in ice.

Lower respiratory specimens: sputum and/or endotracheal aspirate or bronchoalveolar lavage in patients with more severe respiratory disease (obtained with aerosol precautions)

Confirmatory Tests

- Respiratory tract or blood samples tested positive for 2019-nCoV nucleic acid using Real-time Reverse Transcriptase – Polymerase Chain Reaction (RT-PCR)
- Genetic sequencing of respiratory tract or blood samples is highly homologous with the known 2019-nCoV, but this is not done routinely.

Serological tests may help in epidemiological investigation but there could be cross reactivity with other coronaviruses. Viral isolation is not done routinely for diagnosis. Rapid diagnostic test kits like Xpert Xpress

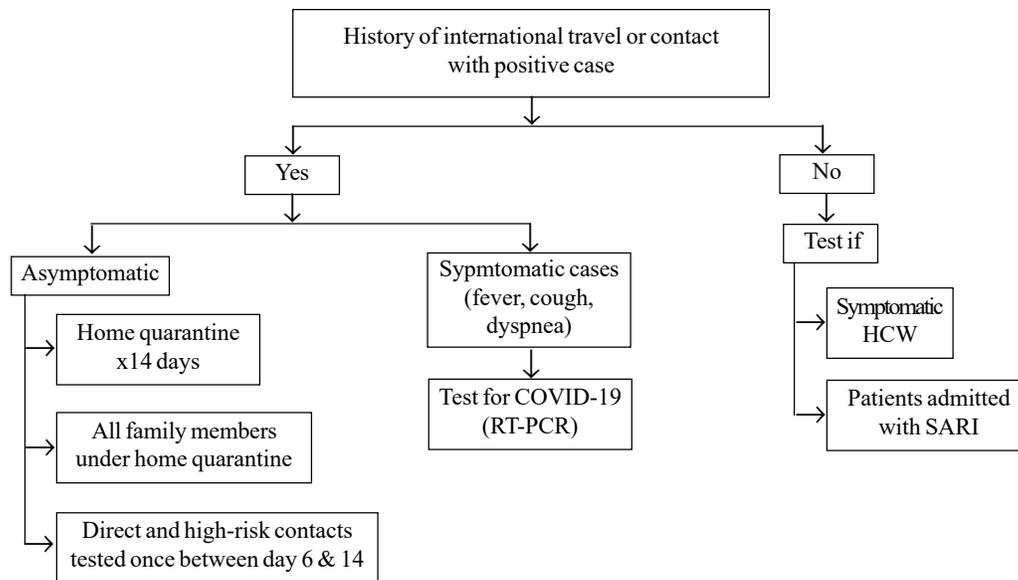


Fig. 1 Testing strategy for suspected cases as per Indian Council of Medical Research.

SARS-CoV-2 by Cepheid has been approved by the US-FDA (United States Food and Drug Administration) for Emergency Use Authorization (EUA) and RealStar SARS-CoV-2 RT-PCR kit 1.0 by Altona Diagnostics and Patho Detect by MY LAB have been approved by ICMR[20,21].

Ancillary Investigations

Complete blood count: Lymphopenia was seen in 85% of critically ill adults, suggesting it a marker of severe disease while among the overall pediatric cases, it was seen in 3.5% [7,12].

Infection markers: Elevation of C-reactive protein (CRP) was reported in 20% and procalcitonin in 64% of cases [7].

Radiological findings: Chest radiography (CXR) or computed tomography (CT) are not recommended as a routine for children but only in specific cases presenting with pneumonia and/or acute respiratory distress syndrome (ARDS). Parenchymal abnormalities with peripheral consolidations on CXR have been reported in a small case series from Korea [14]. Ground glass opacities (32%), local patchy shadows (18%) and bilateral patchy shadows (12%) on CT chest were the common findings in children [7]. Bilateral pneumonia (75%), unilateral pneumonia (25%) and multiple mottling and ground-glass opacity (14%) were reported based on CXR and CT findings from adult patients in Wuhan, China [15].

Laboratory markers of organ dysfunction: Elevation of transaminases is seen in 12-14% and d-Dimer in 14% cases [7].

PREPAREDNESS AND ADMINISTRATIVE CONCERNS FOR ICU

A phased and tiered plan for ICU during the pandemic needs to be made based on the assessment of healthcare burden and resource utilization [13,22,23].

Intensive care units: Create cohort intensive care units where critically ill confirmed COVID-19 patients will be managed. This would be a different area from where other PICU patients are being managed in order to reduce transmission within the hospital. In addition, a separate area should be developed where suspected COVID-19 patients will be managed. With increasing burden of patients, general beds may have to be converted to ICU beds and provided with suitable infrastructure. Predictive models based on local epidemic need to be developed for expected number of patients as well as need of equipment.

Setting up of isolation rooms : Negative pressure isolation is the standard recommendation for management of a suspected or proven COVID-19 patient. However, in case of non-availability of these rooms, use single rooms with separate air outlet/exhaust, preferably on the higher floor of the building. These rooms should be equipped with resuscitation trolley, essential drugs, multipara monitor and ventilator. Positive pressure rooms

like operation theatres are not suitable for airway management as aerosol generation is higher.

Reducing the ICU burden: All elective non-urgent admissions and surgeries need to be halted during the outbreak in order to rationalize resource-utilization, and ensure adequate back-up to handle the crisis.

Re-allocation of staff: During the crisis, there may be acute shortage of critical care specialists and nursing staff. It is essential to identify staff from respiratory medicine, infectious disease and other units who may be trained in infection control, personal protective equipment (PPE) use and management of critically ill patients.

Rotation of staff and reserve for back-up: Adequate reserve of healthcare providers needs to be ensured as a back-up in case of emergencies or healthcare professionals falling sick. The team members should be working on rotation (in a shift of 4-7 days) with adequate rest in between.

Training of all staff: All those who are likely to come in close contact with the patient or are handling equipment, surroundings, and waste management should receive training regarding infection control including correct technique of donning and doffing of PPE and disinfection of surfaces and equipments. Proper training and a written plan (Standard Operating Procedure) should be there for waste disposal.

Rational use of PPE: In view of current global shortage, WHO has formulated guidelines for the rational use of PPE. This includes co-ordination of PPE supply chain management mechanism, appropriate PPE use based on indication, minimizing the need of PPE by bundling activities, using physical barriers and telemedicine where appropriate, and restricting visitors [24].

MANAGEMENT IN RESOURCE-LIMITED SETTINGS

Triage and Transport

A dedicated area for screening and triaging of patients with suspected COVID-19 is essential. Once the patient fits to the case definition and requires admission, unnecessary movement must be avoided and minimum staff should accompany the patient. Ensure that the patient (if self-breathing) and the accompanying persons should be on a 3-ply surgical mask.

ICU Management

Severe and critical cases need ICU care for monitoring, ventilation and organ support therapy.

Severe acute respiratory illness (SARI): SARI is defined by the presence of cough and fast breathing plus at least one of the following [25]: (i) Oxygen saturation (SpO_2) <90%, (ii) severe chest indrawing and grunting, and (iii) altered mental status.

SARI is the most common indication for ICU transfer and most guidelines are similar to management of any viral pneumonia with ARDS with an emphasis on minimizing risk of transmission to others, especially healthcare workers [26,27]. The details on the management of SARI are given in Part II of this write-up and **Table I**.

Septic shock: Management of septic shock in COVID is not very different from the routine. However, the Surviving Sepsis Campaign (SSC) guidelines for COVID-19 recommend conservative fluid strategy, avoiding colloids as resuscitation fluid, and to use low dose steroids in catecholamine refractory shock [28]. In children, epinephrine is the first vasoactive of choice for septic shock.

Co-infections: Co-infections like secondary bacterial pneumonia are common, especially in children (50%) and addition of broad spectrum antibiotic to cover gram positive, gram negative, and staphylococcal infection is recommended [29].

Myocarditis: Cardiogenic shock with elevations in hypersensitive Troponin-I have been seen in 12% of patients. Management includes inodilators like milrinone, diuretics, immunomodulators (methylprednisolone and IVIG) and circulatory support with ECMO (extracorporeal membrane oxygenation) have also been used in a few cases [30,31].

Acute kidney injury : This has been reported in 7% and renal replacement therapy may be necessary [32].

Supportive care: This includes conservative fluid management, nutrition, appropriate sedo-analgesia, and prevention and treatment of healthcare associated infections.

Specific Therapy

Although no definitive therapy till date has proven benefit for SARS-CoV2, antiviral drugs like Remdesivir, Lopinavir/Ritonavir are being used in over 50% of the critically ill adults based on *in vitro* viral inhibition and recovery in SARS and MERS but there is no strong evidence [33–36]. Chloroquine has been found to increase endosomal pH and hinder virus cell fusion and also interfere with ACE2, a receptor for binding of SARS-CoV2 [37]. A combination of hydroxychloroquine and azithromycin showed reduction in viral load [38].

Table I Treatment Based on Severity of Disease in Proven Coronavirus Disease-19 (COVID-19)

Symptomatic proven case	Admit in	Treatment	Discharge
Mild	Designated COVID isolation room	Symptomatic treatment	Discharge if 72 h afebrile or 7d after symptom onset and two samples negative 24 h apart followed by home quarantine for total 14 d
Moderate	Designated COVID isolation room	Supportive care, oxygen Oseltamivir	Clinical improvement and two negative nCoV PCR tests 24 h apart
Severe	COVID ICU	Provide nasal prong oxygen Escalate to invasive ventilation if worsening Avoid HFNC/NIV Oseltamivir Ritonavir/Lopinavir OR Hydroxychloroquine Supportive care	Clinical improvement and two negative nCoV PCR tests 24 h apart
Critical	COVID ICU	In addition to the above: Intubate based on clinical/blood gas/radiological features Use all airborne precautions Ventilation ARDS protocol Other organ support Once improving, wean from ventilator and extubate as per protocol	Clinical improvement and two negative nCoV PCR tests 24 h apart

HFNC: High-flow nasal cannula, NIV: Non-invasive ventilation, ICU: intensive care unit, ARDS: Acute respiratory distress syndrome.

Interferons, IVIG, and convalescent plasma from recovered SARS patients are other tested treatment options [39]. Vaccination for RNA viruses (measles, influenza, polio) has shown higher titers of neutralizing antibodies against SARS-CoV [40] (**Table II**). Based on the current experience, we may use broad spectrum antibiotics, oseltamivir, protease inhibitors, hydroxychloroquine and azithromycin. Lopinavir/Ritonavir along with Chloroquine should be avoided in combination.

Course and Recovery

In adult patients with COVID-19 pneumonia, onset of symptoms to respiratory failure takes an average of 7 days with peak severity at 10 days. Signs of improvement starts occurring by day 14. However, at the time of reporting of most studies, many patients were still admitted and their course needs to be followed to know the exact prognosis [40].

INFECTION PREVENTION AND CONTROL

In the intensive care setting, disinfection of high-touch surfaces like monitors, ventilator screen, other equipment, resuscitation trolleys etc are essential and need to be carried out every 4 hours.

Surface decontamination: Alcohol (e.g. isopropyl 70%

or ethyl alcohol 70%) can be used to wipe down surfaces where the use of bleach is not suitable for e.g. Mobiles, laptops, keys, pens etc.

Disinfection: Freshly prepared 1% sodium hypochlorite should be used as a disinfectant for cleaning and disinfection with at least 10 minute contact period.

Aerosol: Ensure room disinfection within 20 minutes of any procedure generating aerosol.

Social distancing: Maintain at least 1 meter distance unless required for examination or procedure.

Contact and droplet precautions: minimize direct contact, ensure hand hygiene, and cough etiquette.

Healthcare Worker (HCW) Risks

Apart from risks related to droplet spread and from contaminated surfaces, ICU professionals face the challenge of acquiring infection during aerosol generating procedures (*see table in Part II*). HCW should wear a medical mask and gown when entering a room where patients with suspected or confirmed COVID-19 are admitted and use full personal protective equipment (PPE), which includes N95 mask, goggles or face shield, cap, full sleeve gown and shoe cover, when performing aerosol-generating procedures [41]. The entire PPE is

TABLE II Pharmacotherapy in COVID-19

Drug class	Drug name; stage	Mechanism	Dose	Additional points	Evidence
<i>Antiviral drugs</i>					
Nucleoside analogue	Ribavirin; Pneumonia	Inhibits RNA synthesis and viral replication	IV 8 mg/kg 8 hourly × 14 d	Side effects: Hemolytic anemia, Hypocalcemia, Hypomagnesemia May increase viral load in combination with steroid	In vitro studies SARS data Not recommended
Neuraminidase inhibitor	Oseltamivir; Pneumonia	Reduces viral replication	<12 mon: 6 mg/kg/dose BD >12 mon: <15 kg: 60 mg/d 15-23 kg: 90 mg/d 23-40 Kg: 120 mg/d >40 kg: 150 mg/d Given PO /BD for 5 d (max dose 150 mg)	If co-infection with influenza suspected	MERS-CoV data
Protease inhibitor	Lopinavir/ Ritonavir; Early ARDS	Inhibit CoV main protease required in replication	Low dose: 200/100 mg BD High dose: BD for 6-15 d 14 d-12 mon: 16 mg/kg/dose < 15 kg: 12 mg/kg/dose 15-40 kg: 10 mg/kg/dose (Based on Lopinavir) >40 kg: 400/100 mg Given PO/BD for 5-14 d (Max dose Lopinavir 400 mg / ritonavir 100 mg)		In-vitro studies SARS data [33] Weak recommendation [44]
Adenosine analogue	Remdesivir; Pneumonia	Incorporates into viral RNA and leads to premature chain	Adult dose: 200 mg IV on d 1 followed by 100 mg daily ×5-10 d	Avoid in children, pregnant, renal and hepatic impairment	In vitro studies [35] Case report in US [36] On-going trials termination
Aminoquinoline	Chloroquine Hydroxy-chloroquine; Pneumonia	Increases endosomal pH and hinder virus cell fusion Inhibits viral binding to ACE-2 Immunomodulatory effect	CQ: 10 mg/kg base stat followed by 5 mg/kg base BD HCQ: 8 mg /kg/ loading dose, then 4 mg/kg /dose PO /BD/ 5 d (max dose 400 mg) Prophylaxis: 400 mg BD on d 1 then 400 mg weekly	Inhibits pneumonia exacerbation Negative conversion Shortens disease	Unpublished data [45] Ongoing phase III trial for prophylaxis and reducing transmission ICMR recommendation for prophylaxis

Contd....

Table 1 Contd....

Drug class	Drug name; stage	Mechanism	Dose	Additional points	Evidence
<i>Immuno-modulators</i>					
Corticosteroids	Methylprednisolone; Pneumonia ARDS	To suppress cytokines storm, HLH	1-2 mg/kg/day × 5-7 d	Delays clearance of viral RNA	Reduced duration of supplemental oxygen and radiological improvement [46] SSC guidelines recommend use in ARDS but meta-analysis in viral pneumonia- harm > benefit [28]
Immunoglobulin	IVIg/Convalescent plasma; Critical stage	Immunomodulator	1-2 g/kg over 2-5 d	After all therapies failed	Critically ill SARS [47]
<i>Immuno-modulator and antiviral</i>					
Immunomodulator & antiviral	Interferon-α; Early phase of URTI, Pneumonia	Reduces viral load	Nebulization of 200,000 -400,000 IU/kg (2-4 µg/kg) in 2 mL sterile water BD for 5-7 d		Weak recommendation [48]
	Interferon-α2b spray; Close contacts URTI	Reduces viral load	1-2 sprays (8000 IU/spray) on each side of the nasal cavity, 8-10 sprays on the oropharynx, once every 1-2 hrs for 5-7 d		
<i>Immunotherapy</i>					
Interleukin -6 inhibitor	Tocilizumab; Cytokine release syndrome	Immunosuppression	<30 kg - 12 mg/kg/dose >30 kg - 8 mg/kg/dose IV BD as infusion 1-2 d (max dose 800 mg)	For HLH and cytokine storm	On-going clinical trials

recommended to be used for 4-6 hours and changed earlier if there is any soiling. Team should not include staff vulnerable to infection like immunocompromised person, pregnant ladies, age >60 years or those with comorbidities. In the event of exposure and manifestation of infection, management as per guidelines as well as psychosocial support needs to be ensured. Adequate communication, education and adherence to strict personal protection can minimize the risk of transmission to HCW [26]. ICMR recommends prophylactic use of hydroxychloroquine 400 mg twice a day on day 1, followed by 400 mg once weekly for next 7 weeks for HCW managing suspected or confirmed COVID-19 patients [42].

Special Considerations for Resuscitation

It is important to minimize the number of people inside the room during high aerosol generating events like cardiopulmonary resuscitation. One airway specialist, one nurse/doctor for chest compression and one nurse for medication are essential. Other assistants may remain outside the room and may enter only if necessary after donning full PPE. Hand bagging needs to be avoided. During any disconnection from ventilator, endotracheal (ET) tube needs to be clamped and/or viral filter attached to the ET tube. In case re-intubation is required, follow the standard procedure described (*see Part II in this issue*).

CONCLUSION

The COVID-19 pandemic caused by 2019-nCoV has become a serious concern for mankind all over the world. It has challenged and overwhelmed the existing intensive care facilities globally. SARI is the most common indication for intensive care management and is associated with high mortality. The disease so far appears to be less common in children and seems to have a milder course. Preparation for handling crisis during this outbreak is essential for early identification, stratification and management of cases. Prevention by ensuring strict infection control practices minimizes transmission to other patients and healthcare workers, especially in intensive care units.

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