ಕರ್ನಾಟಕ ವಿಧಾನ ಪರಿಷತ್ತು

ಚುಕ್ಕೆ ಗುರುತಿಲ್ಲದ ಪ್ರಶ್ನೆ ಸಂಖ್ಯೆ

: (108) 96

ಮಾನ್ಯ ಸದಸ್ಯರ ಹೆಸರು

: ಶ್ರೀ ಕವಟಗಿಮಠ ಮಹಾಂತೇಶ ಮಲ್ಲಿಕಾರ್ಜುನ (ಸ್ಥಳೀಯ

ಸಂಸ್ಥೆಗಳ ಕ್ಷೇತ್ರದಿಂದ ಚುನಾಯಿತರಾದವರು)

ಉತ್ತರಿಸಬೇಕಾದ ದಿನಾಂಕ

: 01.02.2021

ಉತ್ತರಿಸುವ ಸಚಿವರು

: ಮಾನ್ಯ ಆರೋಗ್ಯ ಮತ್ತು ಕುಟುಂಬ ಕಲ್ಯಾಣ ಹಾಗೂ

ವೈದ್ಯಕೀಯ ಶಿಕ್ಷಣ ಸಚಿವರು

ಕ್ರಸಂ.	ಪ್ರಶ್ನೆ	ಉತ್ತರ					
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	ವೈರಸ್ ಅವಾಂತರ ಹೆಚ್ಚಾಗಿದ್ದು,	ಭಯಬೀತರಾಗಿರುವುದು ಸರ್ಕಾರದ ಗಮನಕ್ಕೆ ಬಂದಿರುವುದಿಲ್ಲ					
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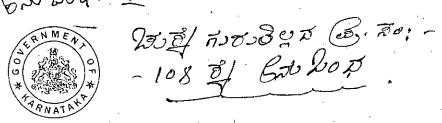
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GOVERNMENT OF KARNATAKA

No: HFW 333 ACS 2020

Karnataka Government Secretariat Vikasa Soudha, Bengaluru, Dated: 03.09.2020

CIRCULAR

SUBJECT - TREATMENT PROTOCOL FOR COVID19 PATIENTS (VERSION 3.0)

<u>DISCLAIMER:</u> This Recommendation is derived from the prevailing trials published and collation of various guidelines. These guidelines would change with the evolving evidences

DEFINITIONS:

- ILl is defined as one with acute respiratory infection
 - with fever $\ge 38^{\circ}$ C (100.4°F) and
 - cough
 - with onset within last 10 days
- SARI is defined as one with acute respiratory infection
 - with fever $\ge 38^{\circ}$ C (100.4°F),
 - cough
 - with onset within the last 10 days and
 - requiring hospitalization
- Respiratory failure
 - Represents the failure of the lung to maintain adequate gas exchange
 - Characterized by ABG abnormalities: PaO2< 60 mmHg with or without hypercarbia
 PaCO2> 46 mmHg (with drop in pH<7.30)

COVID 19 RT-PCR POSITIVE PATIENT

- 1. Management of any COVID 19 patient mandates the Health Care Personnel (HCP) to be in full Personal Protection Equipment(PPE).
- 2. Patient is Categorized in to three groups:

CATEGORY Type of patients who are provided treatment and care					
Group A	Asymptomatic/Patients with mild symptoms				
	RR<24/m & SpO2>94% in room air				
Group B	Symptomatic patient with mild to moderate Pneumonia with no signs of severe disease				
	RR: 24-30/m (or) SPO2: 90%-94% at Room Air				

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Group C	Symptomatic patient with Severe Pneumonia with	
	RR > 30/min (or) SPO2 < 90% at Room Air (or) less than 94% with oxygen, ARDS, Septic Shock	
		İ

CLINICAL CATEGORIES

Clinical category	. Description	Parameters
Asymptomatic	No Symptoms	SpO2: ≥94% in room air RR: ≤ 24/m No evidence of hypoxemia or breathlessness
Mild	Patients with uncomplicated upper respiratory tract infection.	SpO2: ≥94% in room air RR: ≤ 24/m No evidence of hypoxemia or breathlessness
Moderate	Pneumonia with no signs of severe disease	Sp02: 94%-90% in room air RR: 24-30/m
Severe	Severe Pneumonia	SpO2: < 90% room air RR: >30/m
Critical	Acute Respiratory Distress Syndrome (ARDS)	Onset: new or worsening respiratory symptoms within one week of known clinical insult. Chest imaging (Chest X ray and portable bed side lung ultrasound): bilateral opacities, not fully explained by effusions, lobar or lung collapse, or nodules. Origin of Pulmonary infiltrates: respiratory failure not fully explained by cardiac failure or fluid overload. Need objective assessment (e.g. echocardiography) to exclude hydrostatic cause of infiltrates/ oedema if no risk factor present. Oxygenation impairment in adults: Mild ARDS: 200 mmHg < PaO2/FiO2 ≤ 300 mmHg (with PEEP or CPAP ≥5 cm H2O) Moderate ARDS: 100 mmHg < PaO2/FiO2 ≤200 mmHg with PEEP ≥5 cm H2O) Severe ARDS: PaO2/FiO2 ≤ 100 mmHg with PEEP ≥5 cm H2O) When PaO2 is not available, SpO2/FiO2 ≤315 suggests ARDS (including in non- ventilated patients).
Critical	Septic Shock	Adults: persisting hypotension despite volume resuscitation, requiring vasopressors to maintain MAP ≥65 mmHg and serum lactate level > 2 mmol/L



IDENTIFICATION OF HIGH-RISK PATIENT

CO MORBIDITIES	CLINICALLY	LABORATORY VALUE
Age>50 yrs	Hypoxia- SPO2<94%	Lymphopenia (<20) with Neutrophil/Lymphocyte ratio >17
Ischecmic Heart Disease Diabetes	Tachycardia>100/min Respiratory Distress RR>30/min	CRP>100 mg/L Serum Ferritin >300 microg/L
Hypertension	Hypotension Systolic BP < 90mmHg	LDH >450
Lung Disease (COPD/Asthma/Post TB Sequele)	Altered Sensorium	D-Dimer > 1000ng/ml
Chronic Kidney Disease/ Chronic Liver Disease Immunosuppression / HIV / Malignancy Obesity		

Note: Calculation tool for predicting critically ill COVID-19 at admission can be used as reference tool. (Development of Validation of Clinical risk score to predict the occurrence of critical illness in hospitalised patient with COVID19. JAMA internal Medicine -published online, May 12/05/2020)

GENERAL MEASURES AND GUIDELINES

- 1) Categorize in to A, B, C based on Symptoms, SpO2 & Respiratory Rate
- 2) Supportive Care:
- Finger Pulse Oximeter for continuous monitoring of Heart rate and Oxygen saturation
- Start oxygen with Mask at saturation of 94% or lower
- HFNC to be used if there is failed oxygen therapy and Non-invasive ventilation (NIV) to be used appropriately with two limb circuit expiratory filters
- Counselling of COVID19 patients (By Counsellor/psychologist/psychiatrist)
- Normal feeding, no dietary restrictions, good oral hydration
- Maintenance IV fluids (If indicated)
- Maintain blood glucose levels <180 mg/dl.
- If Patient is on ACE inhibitors/ARBs, should be continued
- Avoid using NSAIDs other than Paracetamol Unless Absolutely Necessary
- Avoid using Nebulized drugs to avoid aerosolization of virus. PREFER MDI with SPACER
- Antibiotic selection in case of superadded bacterial pneumonia should be according to institution antibiogram.

<u>INVESTIGATIONS</u>

Timing	Mild	Moderate	Severe/Critical
At	CBC	Complete Blood Count	•Complete Blood Count
admission	RBS	(with N/L RATIO)	(with N/L RATIO)
, .	ECG	• LFT, RFT, RBS	•LFT, RFT, RBS
	HbA1C (if Diabetic)	S.Electrolytes	•S. Electrolytes
	D-Dimer	• 12 lead ECG	•12 lead ECG
		CHEST X Ray -PA view	•CHEST X Ray –PA view
	(If starting on Tab	CRP, D-DIMER	•CRP, D-DIMER
	Favipiravir)	S. FERRITIN, S.LDH	•S. FERRITIN, S.LDH
. •	RFT	PROCALCITONIN	•PROCALCITONIN
•	S.Electrolyte	• TROP-1&T	•TROP – I & T
	S. Uric Acid	• PT/INR	•PT/INR
		• ABG	•ABG
		CT Thorax (if Available)	•CT Thorax (if Available)
		Blood culture (if total	·Blood culture (if total
		count is high)	count is high)
		• IL-6	•IL - 6
		• S. Cortisol	•S. Cortisol
	•	· 2D	•S.Mg ²⁺ , S.Ca ²⁺
		ECHOCARDIOGRAP	•2D
		HY	ECHOCARDIOGRA
	1.	COVID Antibody	РНУ
	· ·	_	·NTproBNP
ı	· ·	IgM/IgGT ests	•HsCRP
		\	•S. Lactate
			•COVID Antibody
·			
		O III DI IO III	IgM/IgGTests
Repeat		Complete Blood Count, LFT,	Complete Blood Count, LFT,
Daily	_ :	RFT	RFT
		ABG ·	ABG
Repeat		CRP, D-DIMER	CRP, D-DIMER
Every 72hrs	If initial D-Dimer is	S. FERRITIN, S.LDH	S. FERRITIN, S.LDH
	high	Chest X ray	Chest X ray
At the time		CRP, D-DIMER	CRP, D-DIMER
1		S. FERRITIN, S.LDH	S. FERRITIN, S.LDH
of discharge	-	· 1	Chest X ray
	ļ	Chest X ray	RT-PCR – Nasal & Throat
		1	
1 .		<u> </u>	swab

Other Investigations should be done based on patient's Co-morbid status

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GROUP A - MILD CASES

TREATMENT

ANTIVIRAL THERAPY

TAB HYDROXYCHLOROQUININE(HCQ) 400MG BD FOR 1 DAY Followed by 200MG 1-0-1 X 4 DAY for patients in COVID CARE CENTER/HOME ISOLATION (OR)

Tab FAVIPIRAVIR 1800mg 1-0-1 on Day 1 f/b 800mg 1-0-1 for 6 days (total 7 days) for PATIENTS IN DCHC

(OR)

If Tab HCO/Tab FAVIPIRAVIR is contraindicated, then combination of Cap DOXYCYCLIN 100mg 1-0-1 for 5 days

Tab IVERMECTIN 12mg 1-0-0 for 3 days

"Cap Oseltamavir 75mg 1-0-1 for 5 days

ANTICOAGULATION

Inj ENOXAPARIN 40mg S/C 1-0-0 X 7 DAYS (IF D-DIMER IS MORE THAN 1000NG/ML (OR) X-RAY/CT THORAX SHOWING **GROUND GLASS OPACITIES)**

SUPPORTIVE THERAPY-

- TAB ZINC 50 MG 0-1-0 X 7 DAYS
- TAB VITAMIN C 500 MG 1-1-1 X 7 DAYS
- Tab N Acetylcysteine 600mg 1-1-1 If Patients Having Cough

PRECAUTIONS

- CATEGORIZATION SHOULD BE REASSESSED REGULARLY
- CONTRAINDICATION FOR HCQ-
 - OT INTERVAL > 480ms
 - Pre-existing cardiomyopathy and cardiac rhythm disorders
 - History of Unexplained Syncope
 - Retinopathy,
 - Hypersensitivity to HCQ or 4aminoquinoline compounds
 - G6PD deficiency
 - Epilepsy
 - Hypokalemia ($K^+ < 3$ Meq)
- Contraindications for Tab FAVIPIRAVIR: Hyperuricaemia, severe hepatic & renal impairment, Pregnant women and lactating mothers
- PREGNANCY IS NOT A CONTRAINDICATION FOR HCQ
- # Cap OSELTAMAVIR is advised due to possibility of H1N1 co infection along with COVID19 disease with present weather condition. Its usage will be reviewed at a later date.

GROUP B - MODERATE CASES

TREATMENT

PRECAUTIONS

ANTIVIRAL THERAPY

• Inj REMDESIVIR 200 mg IV on day I followed by 100 mg IV daily for 4 days (total 5 days)

IF REMEDESVIR IS NOT AVAILABLE TO START TAB HYDROXYCHLOROQUININE(HCQ) 400MG BD FOR 1 DAY followed by 200MG 1-0-1 X 4 DAY

Co-administration of Inj REMDESIVIR with HCQ or chloroquine should be avoided

• Cap Oseltamavir 75mg 1-0-1 for 5 days

STEROIDS

 Inj. Methyl Prednisolone 0.5 -1 mg/kg (or)

Inj. Dexamethasone 0.1 - 0.2 mg/kg for 3-5 Days

ANTICOAGULATION

Inj ENOXAPARIN 40MG S/C 1-0-0 x 7 DAYS

IV ANTIBIOTICS ACCORDING TO LOCAL ANTIBIOGRAM

AWAKE PRONING

 CONVALASCENT PLASMA THERAPY: 4 to 13 ml/kg (usually 200 ml single dose given slowly over not less than 2 hours)

SUPPORTIVE THERAPY

- TAB ZINC 50 MG 0-1-0 X 7 DAYS
- TAB VITAMIN C 500 MG 1-1-1 X 7 DAYS
- TAB N-ACETYL CYSTEINE 1-1-1 IN PATIENTS WITH COUGH

Contraindications for Inj REMDESIVIR:

- AST/ALT > 5 times Upper limit of normal (ULN)
- Severe renal impairment (i.e., eGFR < 30ml/min/m2 or need for hemodialysis)
- Pregnancy or lactating females
- Children (< 12 years of age)
- No dose adjustment for Inj REMDESIVIR if eGFR >30ml/min
- Formula to calculate eGFR in Adults
- eGFR, Male: (140 age in years) × (weight in kg)
 / 72 × (serum creatinine in mg/dL);
- eGFR, Female: (140 age in years) × (weight in kg) × 0.85 / 72 × (serum creatinine in mg/dL)

STEROIDS

- to be started preferably within 48 hours of admission (or) if oxygen requirement is increasing and if inflammatory markers are increased.
- PATIENT SHOULD BE REASSESSED EVERY 12 HRLY AND CONTINOUS MONITORING OF SATURATION.
- START ON OXYGEN-NASAL PRONGS 2-5 L/MIN or FACE MASK 5L/MIN

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GROUP C - SEVERE/CRITICAL CASES

TREATMENT

ANTIVIRAL THERAPY

- If the patient has not received Inj
 REMDESIVIR, such patients can be started on
 Inj REMDESIVIR.
 Inj REMDESIVIR 200 mg IV on day 1 followed
 by 100 mg IV daily for 4 days (total 5 days)
- Inj. TOCILUZUMAB 8mg/kg (maximum 800 mg at one time) given slowly in 100 ml NS over 1 hour; dose can be repeated once after 12 to 24 hours if needed

(Or)

Inj ITOLIZUMAB: 1st dose – 1.6mg/kg dose iv infusion. Subsequent dose: weekly ... 0.8mg/kg dose infusion over 4hours if required

• Cap Oseltamavir 75mg 1-0-1 for 5 days

STEROIDS

Inj. Methyl Prednisolone 1-2 mg/kg (or)
 Inj. Dexamethasone 0.2 - 0.4 mg/kg for 5-7
 Days

ANTICOAGULATION

 Inj ENOXAPARIN 1mg/kg body wt s/c 1-0-1 X 7 DAYS

PRONE VENTILLATION

Inj CEFTRIAXONE 1gm IV 1-0-1 AND CAN BE ESCALATED ACCORDING TO LOCAL ANTIBIOGRAM OR TREATING PHYSICIAN

CONSIDER SEPSIVAC (IF AVAILABLE) 0.3ml INTRADERMAL ONCE A DAY FOR 3 DAYS IN CASE OF SEPTIC SHOCK

IV Diuretics in case of evidence of Heart Failure secondary to Myocarditis
SUPPORTIVE THERAPY

- TAB ZINC 50 MG 0-1-0X 7 DAYS
- INJ. VITAMIN C 1.5GM IV 6 HOURLY X 5DAYS
- TAB N-ACETYL CYSTEINE 1-1-1

PRECAUTIONS

Indication for TOCILUZUMAB/ITOLIZUMAB;-

- 1. IL-6 levels 50-100 fold higher than normal (Normal range 0 9.5pg/ml
- 2. Worsening trend of the inflammatory markers (Ferritin, LDH, CRP)
- 3. Deteriorating clinical condition with worsening of PaO2/Fio2 ratio (more than 25% deterioration from the immediate previous value).

Contraindications for Ini TOCILIZUMAB/ITOLIZUMAB

PLHIV, those with active infections (systemic bacterial/fungal), High Serum. Procalcitonin, Tuberculosis, active hepatitis, Absolute Neutrophil Count < 2000/mm3 and Platelet count < 1,00,000/mm3, hepatic and renal impairment; patients on chronic steroid therapy, Paediatric patients <18 years old; Pregnancy and, Nursing mothers.

- PATIENT SHOULD BE CONTINOUSLY MONITORED
- TO START ON OXYGEN WITH FACE MASK WITH NON REBREATHING BAG @ 8-10 lt/m
- BASED ON PaO2/FiO2 ratio, HIGH FLOW NASAL OXYGEN (HFNC)/NIV SHOULD BE GIVEN AND IF PATIENT DETERIORATES INTUBATION SHOULD BE CONSIDERED AND LUNG PROTECTIVE VENTILATION TO BE FOLLOWED AS PER ARDSnet PROTOCOL
- ABG TO BE DONE REGULARLY FOR MONITORING OF ACIDOSIS AND HYPOXEMIA
- INOTROPHIC SUPPORT (NORADRENALINE
 - TITRATE ACCORDING TO THE MEAN
 ARTERIAL PRESSURE)
- CORRECTION OF ACIDOSIS
- MAINTAIN Hb% GREATER THAN 8gm%

(and

SUMMARY OF TREATMENT OF COVID-19 PATIENTS

BASED on CLINICAL CATEGORIES

Antiviral Therapy

Fraction of the property of

- 1. Continous monitoring of oxygen saturation by pulse eximeter and early diagnosis of hypoxemia is essential in all group of patients
- 2. Indications and contraindications of the drugs are to be considered before use which is mentioned in detail below
- 3. Transition of patients between the clinical categories is based on SpO2, RR & PaO2/FiO2 ratio
- 4:. Treatment of all co morbid illness to continue

Antiviral Therapy aini REMDESVIR 200 mg IV on d Nifollowed by 100 ing Nydaily for days (Or)

(O))

FFSIF REMEDES/IRISNOT

AVAILABLE TO START

Tab. Hydroxychlorogwinine(HCQ)
(400mg BD.For 11 Bay: F/B 200mg

1-0-1 X 4 Day (\$24) K o-administration of Inj

Co administration of initial REMDES VIR with HEQ or significant of the

ANTICOAGULATION:
Initenoxaparin 40mglS/C 1-0-0x7
days(if.Wt. 65kg 50md 1-0 Lion of the continuous section of the conti

Start on oxygen -Nasal Prongs 2 \$51/min or face mask 51/min CONVALASCENT PLASMA

THERAPY: 410-13 ml/kg (usually 200 ml/singletdose/given/slowly overing these than 2 hours a Supportive Therapy Tab Zinc 50 Mg 0-1-0x 7/Days

Tab.Vitamin C 500 Mg 1-1-1-X Days Tab:N/Acetylcysteine 1-1-1-1 B/2 Patients Having Cough

Special Note:

- *Cap Oseltamavir 75mg 1-0-1 for 5 days to be added to patients of all categories
- All the investigational therapies and drugs approved recently by DGCI should be used with caution. and after informed consent from the patient

Antiviralinherapy PhilanochuzuMA88mg

catter Z to Z4hb urshinged (On) \$ In | FOUZUMAB FIN dose = 1.6 o Adoseli Vintusion Subsequento weeki vo 8mg/kg doselintusion 44hours i drequired STEROIDS Init sMethyl Prednisoloned 2 of Tores 7 Days (Or)

ANTICOAGULATION

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Injlenoxapar in Bilyle/Agebody Agent

for 10 ft/7 days

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mask+NRM and change over to general the NC/NIV (based jon Pa02/Fi02)

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IE PATIENTIDETERIORATES, with:
HENC/NIV trial/(repear ABG arter)
Ghrs suggests worsening or convergention) it hence and the convergence of the convergence

4. Hydroxychloroquine (HCQ)

<u>Dose:</u> Tab HCQ 400MG BD FOR 1 DAY Followed by 200MG 1-0-1 X 4 Days

CONTRAINDICATION FOR HCQ

- QT INTERVAL > 480ms
- Pre-existing cardiomyopathy and cardiac rhythm disorders
- History of Unexplained Syncope
- Retinópathy,
- Hypersensitivity to HCQ or 4-aminoquinoline compounds
- G6PD deficiency
- Epilepsy
- Hypokalemia (K⁺ < 3 Meq)

5. Anticoagulant Agents

Pro Coagulant factors are increased in COVID-19 infection and associated with increased risk of thrombosis

Pneumonia and sepsis are complicated by DIC, but although COVID-19 patients do have abnormalities of coagulation and are not atypical of DIC.

The most marked abnormality is an elevation of D-Dimer (if D-dimer is more than 1000ng/ml) but without a parallel fall in platelet or prolongation of clotting time, this suggests that local rather disseminated thrombin generation and fibrinolysis is taking place

Dose:

Inj ENOXAPARIN 40MG S/C Once daily for mild and moderate. Twice daily in severe cases.

Other options:

- Inj Fondaparinux 2.5mg OD SC
- Unfractioned Heparin 5000 Units BD SC

Contraindications:

ESRD, active bleeding, emergency surgery, platelets < 20,000/mm3, BP > 200/120 mmHg)

INVESTIGATIONAL THERAPIES (as per MOHFW)

- 1. Remdesivir (under Emergency Use Authorization) may be considered in patients with moderate disease (those on oxygen) with none of the following contraindications:
 - AST/ALT > 5 times Upper limit of normal (ULN)
 - Severe renal impairment (i.e., eGFR < 30ml/min/m2 or need for hemodialysis)
 - · Pregnancy or lactating females
 - Children (< 12 years of age)

Dose: 200 mg IV on day 1 followed by 100 mg IV daily for 4 days (total 5 days)

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- 2. Convalescent plasma (Off Label) may be considered in patients with moderate disease who are not improving (oxygen requirement is progressively increasing) despite use of steroids.

 Special prerequisites while considering convalescent plasma include:
 - ABO compatibility and cross matching of the donor plasma
 - Neutralizing titer of donor plasma should be above the specific threshold (if the latter is not available, plasma IgG titer (against S-protein RBD) above 1:640 should be used)
 - Recipient should be closely monitored for several hours post transfusion for any transfusion related adverse events
 - Use should be avoided in patients with IgA deficiency or immunoglobulin allergy

Dose: Dose is variable ranging from 4 to 13 ml/kg (usually 200 ml single dose given slowly over not less than 2 hours

- 3. <u>Tocilizumab (Off Label)</u> may be considered in patients with severe disease with progressively increasing oxygen requirements and in mechanically ventilated patients not improving despite use of steroids. Long term safety data in COVID 19 remains largely unknown. Special considerations before its use include:
 - o IL-6 levels 50-100 fold higher than normal (Normal range 0 9.5pg/ml
 - o Worsening trend of the inflammatory markers (Ferritin, LDH, CRP)
 - o Deteriorating clinical condition with worsening of PaO2/Fio2 ratio (more than 25% deterioration from the immediate previous value)

The drug is contraindicated in

PLHIV, those with active infections (systemic bacterial/fungal), High Serum. Procalcitonin, Tuberculosis, active hepatitis, Absolute Neutrophil Count < 2000/mm3 and Platelet count < 1,00,000/mm3, hepatic and renal impairment; patients on chronic steroid therapy, Paediatric patients <18 years old; Pregnancy and, Nursing mothers

Dose: 8mg/kg (maximum 800 mg at one time) given slowly in 100 ml NS over 1 hour; dose can be repeated once after 12 to 24 hours if needed

Drugs Recently approved by DGCI

- 1. ITOLIZUMAB (An anti-CD6 IgG1 monoclonal antibody) Indication:
 - 1. IL-6 levels 50-100 fold higher than normal (Normal range 0 9.5pg/ml
 - 2. Worsening trend of the inflammatory markers (Ferritin, LDH, CRP)
 - 3. Deteriorating clinical condition with worsening of PaO2/Fio2 ratio (more than 25% deterioration from the immediate previous value).

Dose: 1st dose - 1.6mg/kg dose iv infusion

• Subsequent dose: weekly 0.8mg/kg dose infusion over 4hours if required based on lung function parameters

Jarl

Contraindication:

PLHIV, those with active infections (systemic bacterial/fungal), High Serum. Procalcitonin, Tuberculosis, active hepatitis, Absolute Neutrophil Count < 2000/mm3 and Platelet count < 1,00,000/mm3, hepatic and renal impairment; patients on chronic steroid therapy, Paediatric patients <18 years old; Pregnancy and, Nursing mothers

Side effects:

- In trial Infusion reactions have been reported in 15% of the patients
 - In clinical practice also infusion reaction ranged from 12% to 15%
 - Other adverse events include Diahorea, Pruritus in 7 12 % of cases

2. Tab. FAVIPIRAVIR

Mechanism of action: It is considered that favipiravir is metabolized in cells to a ribosyl triphosphate form (favipiravir RTP) and that favipiravir RTP selectively inhibits RNA polymerase involved in influenza viral replication

Indications: mild to moderate cases of COVID19 in adults >18yrs old

Dose: 1800mg bid followed by 800mg bid upto maximum of 14days

Contraindications: Hyperuricaemia, severe hepatic & renal impairment, Pregnant women and lactating mothers

Side Effects: increased Uric Acid levels, diarrhea, decreased neutrophil counts, increase in AST/ALT levels

<u>Drug Interactions:</u> metabolised partly by Aldehyde Oxidase(AO) and partly by Xanthine Oxidase(XO). Precauitons for co-administration with Pyrazinamide, Repaglinide, Theophyline, Famciclovir

PRONE VENTILATION

Early self-proning in awake, non-intubated patients - Moderate cases

- Any COVID-19 patient with respiratory embarrassment severe enough to be admitted to the hospital may be considered for rotation and early self-proning.
- Care must be taken to not disrupt the flow of oxygen during patient rotation

Criteria to be fulfilled	Avoid proning
 Patients with oxygen requirement of >4L Normal mental status Able to self-prone or change position with minimal assistance 	 Hemodynamic instability Close monitoring not possible

Typical protocols include 30–120 minutes in prone position, followed by 30–120 minutes in left lateral decubitus, right lateral decubitus, and upright sitting position (Caputo ND, Strayer RJ, Levitan R. Academic Emergency Medicine 2020;27:375–378)

Jap

Requirements for safe prone positioning in ARDS

- Pre-oxygenate the patient with FiO2 1.0
- Secure the endotracheal tube and arterial and central venous catheters
- Adequate number of staff to assist in the turn and to monitor the turn
- Supplies to turn (pads for bed, sheet, protection for the patient)
- Knowledge of how to perform the turn as well as how to supine the patient in case of an emergency

Contraindications to prone ventilation

- Spinal instability requires special care
- Intra cranial pressure may increase on turning
- Rapidly return to supine in case of CPR or defibrillation

When to start proning in SEVERE CASES?

- P/F ratio <150 while being ventilated with FiO2 >0.6 and PEEP >5 cm, H2O When to stop proning?
- When P/F exceeds 150 on FiO2 > 0.6 and > 6 PEEP

What portion of the day should patients be kept prone?

- As much as possible (16-18 hours a day)
- Adult patients with severe ARDS receive prone positioning for more than 12 hours per day (strong recommendation, moderate-high confidence in effect estimates)

(ATS-ERS Guideline. Am J.RespirCrit Care Med; 2017; 195(9): 1253-1263)

Oxygen delivery protocol

SpO2 < 94% ~ Supplement with nasal prongs or simple face mask at 2-5L/min



- Monitor continuous SpO2 with finger pulse oximetry
- If SpO2 < 94% on simple face mask or nasal prongs, change to non-rebreather mask oxygen (NRB) at 10-15L/min



Oxygen Delivery Devices & approximate FiO2%

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Nasal Cannula	
1	24
	28
- 191 3 - 19 15 - 1915 - 1915 - 1915 - 1915 - 1915 - 1915 - 1915 - 1915 - 1915 - 1915 - 1915 - 1915 - 1915 - 1915	. 32
	36
	40
	44
Oxygen Mask 5-6	
6-7	40
- 1977年 - 1874年 7日本	50
Mask with Reservoir Bag	60
6	60
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6 (68)	23
9 (50) 12 (50)	
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*Number in parentheses indicates total flow of entrained room air with Venturi mixture.

HFNO (High Frequency Nasal Oxygen) and NIV (Non-invasive Ventilation)

- When oxygen requirement increases to needing NRB, options of High Frequency Nasal Oxygen (HFNO) or NIV should be considered.
- HFNC flow rates to be set from 30 -60 L/min titrating to maintain SpO2 \geq 92%
- HFNC provides PEEP up to 5-6 cm H₂0 and can deliver FiO2 up to 100%



• If HFNC non-available or patient not maintaining SpO2 on flow rates up to 60L/min, initiate on non-invasive ventilation (NIV) only with an ICU ventilator with two limbed circuit and expiratory HME filter with a NIV mode available. Caution is to be exercised to not use potable home BiPAP or CPAP machines with single circuit for these patients.

Jan



N I V settings:

FiO2 to be titrated to maintain SpO2 \geq 92%.

Pressure Support (IPAP): 12-15cm

H₂O (to target

tidal volume of 6ml/kg)

PEEP (EPAP): 5-15 cmH20 as

tolerated to achieve

 $SpO2 \ge 90-92\%$

Backup rate: 15 breaths/min

Backup I:E ratio 1:3

Trigger: maximum sensitivity

- Appropriate mask with good seal to be ensured when initiated on NIV. Helmet masks/hoods if available, to be preferred to minimize aerosol contamination.
- Once initiated on NIV, close monitoring of respiratory variables hourly is important.
- Reassess clinical condition hourly, monitor and observe ABG's 4-6hrly
 - Look for signs of clinical improvement in the form of settling tachycardia, improving SpO2, reduced tachypnea and reduced work of breathing.
 - On NIV when there are signs of clinical deterioration in the form of worsening sensorium, increased accessory muscles of breathing, raising Pco2, worsening pH on ABG ~ failure of NIV has to be considered and patient has to be planned for intubation and mechanical ventilation after consent from the family.



watch for accessory muscles of breathing

Intubation and Mechanical Ventilation

- Indication for intubation: ARDS with PaO2/FiO2 < 200
- Worsening respiratory distress even on NIV
- Patient in Shock

Initial Settings: Controlled Mode ventilation: VCV (volume-controlled ventilation) or PCV (pressure-controlled ventilation)

- Tidal Volume (Vt) 6-8ml/PBW (predicted body weight)
- PEEP 8 18 cmH2O (follow FiO2-PEEP table) to titrate to target SpO2 90-92%
- FiO2 ~ target SpO2 90-92% with lowest FiO2 possible
- Respiratory rate 14-18/min (maximum up to 35/min)
- Plateau pressure < 30 cmH2O and driving pressure < 16cmH2O
- ABG targets: PaO2:55-80 mmHg, pH > 7.3
- Measure compliance 6hrly ~ Vt in ml /Pplat PEEP

Notes:

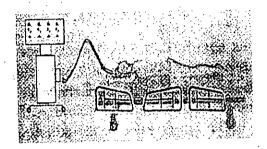


Incremental PEEP FiO2 table

Fi02 0.3	0.4 0.4	0.5 0.5	0.6	0.7	0.7	0.7	0.8	0.9	0.9	0.9	1.0
	5 8	8 10	10	10	12	14	14	14	16	18	18 - 24

Additional steps: If Pplat > 30cmH2O, reduce Vt upto 4ml/PBW

- If SpO2 < 88% despite ARDSnet protocol: increase depth of sedation
- Optimize secretions clearance/bronchodilation
- Initiate early muscle relaxant infusion (cis-atracurium or vecuronium)
- Early prone ventilation



Indication for prone ventilation:

- Intubation and mechanical ventilation < 36hrs
- PaO2/FiO2 < 150, FiO2 > 60%, PEEP > 5, Vt 6ml/PBW
- Duration of proning: 12-16 hrs.
- Multiple sessions until favorable trends are achieved.

Adjunctive measures when intubated and mechanically ventilated:

- Antibiotics guided by protocols
- > Steps to reduce VAP (ventilator associated pneumonia) by following VAP bundles
- > Head-end elevation
- Thrombo-prophylaxis
- Adequate analgesia and sedation

In absence of ABG facility at the hospitals, use SpO2/FiO2 ratio as described in the below table

Derivation of SpO2/FiO2 values corresponding to PaO2/FiO2 ratios in the combined anesthesia and ARMA database*

SOFA Respiratory score	PaO ₂ /FiO ₂	SpO ₂ /FiO ₂
1	<400	<512
2	≺300	<357
3 .	<200	<214
4	<100	<89

Data derived from 4728 matched SpO2/FiO2 and PaO2/FiO2 measurements from the combined anesthesia and ARMA database

	of SpO ₂ /FiO ₂ v		volvonding (O FAOMETC	2 strantied b	y PEEP in	the ARM	À database
SOFA Respiratory score	PaO ₂ /FIO ₂ ratio	SpO2/FiO2 ratio						
		PEEP <8	PEEP 8-12	PEEP>12	•			
Ţ	≪ 100	<502	<515	< 425				
2	<300	<370	<387	<332	•			
3	<200	<240	<259	<234		-		
4	<100	<115	<130	<129	,		•	

The above Treatment protocol for COVID 19 is approved by the Joint Expert Committees, Rajiv Gandhi University of Health Sciences (RGUHS) & The Technical Advisory Committee of Dept. of Health & Family Welfare, Govt. of Karnataka. Hence, all the Government & Private Health Establishments treating COVID 19 persons should strictly adhere to the above protocol, in the larger interest of reducing mortality due to COVID 19.

> Additional Chief Secretary to Government Health & Family Welfare Department

To,

- 1. The Commissioner, BBMP, Bengaluru
- 2. The Deputy Commissioners, All Districts
- 3. The Chief Executive Officers, Zilla Panchayath, All Districts
- 4. The District Health Officer & District Surgeons All Districts
- 5. The Dean cum Directors, All Medical Colleges
- 6. The President IMA, IAP, PHANA, for circulation to all the members & Heads of all Private Medical Establishments.

Copy for information to:

- 1. The Chief Secretary, Government of Karnataka, Bengaluru
- 2. The Principal Secretary to Government, Medical Education Dept., Bengaluru
- 3. The Commissioner, Health & Family Welfare Services, Bengaluru
- 4. The Mission Director, NHM, Bengaluru
- 5. The Director, Health & Family Welfare Services, Bengaluru
- 6. The Director, Medical Education Department, Bengaluru

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