



Management of COVID-19 in Pregnancy in South Australia

v7.2 03/02/2022

Introduction

South Australia currently has a high level of the Omicron variant of COVID-19 circulating in the general population. The pathways that were initially established across the state in November and December did not anticipate the large volume of pregnant women exposed to the virus, and the management plans discussed in the original document have been heavily modified.

This updated version of the guidance contains the new pathways for Telehealth management, statewide coordination, enoxaparin administration, monoclonal antibody infusions and pre-admission testing.

As it is a statewide guidance, internal logistics have not been included and the protocol from each local health network should also be consulted.

As the treatments for COVID-19 have changed as well, detailed sections within this document on medications have been removed and placed in an appendix at the back for reference if required.

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Special thanks to Monash Medical Centre

Common Acronyms

SALHN	Southern Adelaide Local Health Network
CALHN	Central Adelaide Local Health Network
NALHN	North Adelaide Local Health Network
WCHN	Women's and Children's Health Network
rLHNs	Regional Local Health Networks
LHN	Local Health Network
FMC	Flinders Medical Centre
LMH	Lyell McEwin Hospital
WCH	Women's and Children's Hospital
GPAT	General Practitioner Assessment Team
SAAS	South Australian Ambulance Service
CRCT	COVID Response Care Team
CDCB	Communicable Diseases Control Branch
DOB	Date of birth
ICCU	Intensive and Critical Care Unit
MDT	Multi-disciplinary Team
CCU	COVID Care Unit
PPE	Personal protective equipment
CTG	Cardiotocograph
sCOVID	Suspected COVID

Treatment Streaming and Options

Treatment of COVID-19 in pregnant patients will involve sound teamwork between SALHN, CALHN, NALHN, WCHN, Regional LHN's, the CDCB, GPAT and the COVID Response Care Team (CRCT). The aim of these guidelines is to stratify each model of care and create simple flowcharts that can be utilised from each centre with COVID-19 positive obstetric patients with each degree of severity of illness. There will be individual cases that fall outside of this framework and will need to be managed on a case by case basis with these key stakeholders.

Once a patient is diagnosed, both Communicable Disease Control Branch (CDCB) and COVID Response Care Team (CRCT) are notified simultaneously. The patient will be contacted by their pathology provider with their positive result via text message. The patient will also be sent a link to the online application from CDCB to complete. Part of the initial screening questions for risks for severe disease should include pregnancy for all female patients of appropriate age. Their name, date of birth and booking hospital should be recorded by CDCB, and their local care provider notified. Likewise, if a LHN or private provider were to become aware of a positive patient in the community the clinical staff should contact their local infection control who will ensure the CDCB is notified.

Telehealth is offered to all patients through the LHNs, however at this stage relies on patients informing their obstetric care providers of their positive diagnosis.

Patients considered as Suspected COVID (sCOVID) who need medical review will need to be managed at their designated LHN or their private provider, not by CRCT or FMC. Patients requiring review for obstetric reasons should be asked to attend their LHN or contact their private provider. Those requiring review for medical reason should be asked to attend ED. The relevant area will need to be notified of their pending arrival.

Communication Between LHNs/CRCT/GPAT

The daily huddle takes place at 9:30 am each day of the week, including weekends. The huddle includes a representative from each LHN, and invitations have been extended to CRCT, Country Health, SAAS and MedSTAR to attend when required.

The huddle addresses the current caseload and workflow across the three major LHNS and provides strategies for decanting in the coming few days.

Online Database

There is an online database of all known pregnant women with COVID-19 held within Microsoft Teams. This database is updated daily by medical and midwifery staff at each LHN.

Country and private patients that have been diagnosed with COVID-19 should have their diagnosis in pregnancy form completed by their health care provider and sent to their linked LHN to be added to the database.

The database allows for an overview of the number of active cases, likely upcoming deliveries, and some basic medical data to assist in emergency after hours transfers. It should be recognised that all patients identified in the database have contacted their health provider to self-report their COVID-19 status. SA Health are continuing work on an electronic dashboard that will give up to date and accurate data of all positive pregnant women once completed.

Obstetric COVID Care Teams

The Obstetric COVID Care Team (OCCT) will be decided by each metropolitan maternity unit but initially will be made up of the treating midwives and the consultant on call. This team will function in conjunction with Infectious Diseases, Infection Control, Obstetric Medicine, General Medicine/Respiratory, Anaesthetic, Intensive Care and Neonatology as required.

The Midwife Unit Manager (MUM) or other designated officer of each LHN will keep a hospital record of all patients in the community who are COVID-19 positive or are in lockdown/cannot attend the hospital due to isolation or quarantine. Each patient will have a plan made which will include, frequency of telehealth visits, plan for adjustment to any antenatal care and who is responsible for following up the patient.

Follow up may be medical or midwifery based depending on the model of care the patient has chosen and their sequelae of disease.

The OCCT will review each case and plan for ongoing antenatal care, comprising of deferred care and telehealth review.

A telehealth review should then be scheduled with the patient ahead of time. In the first review, ongoing care should be explained, including whether other parameters of care will be deferred and the booking times of the new care.

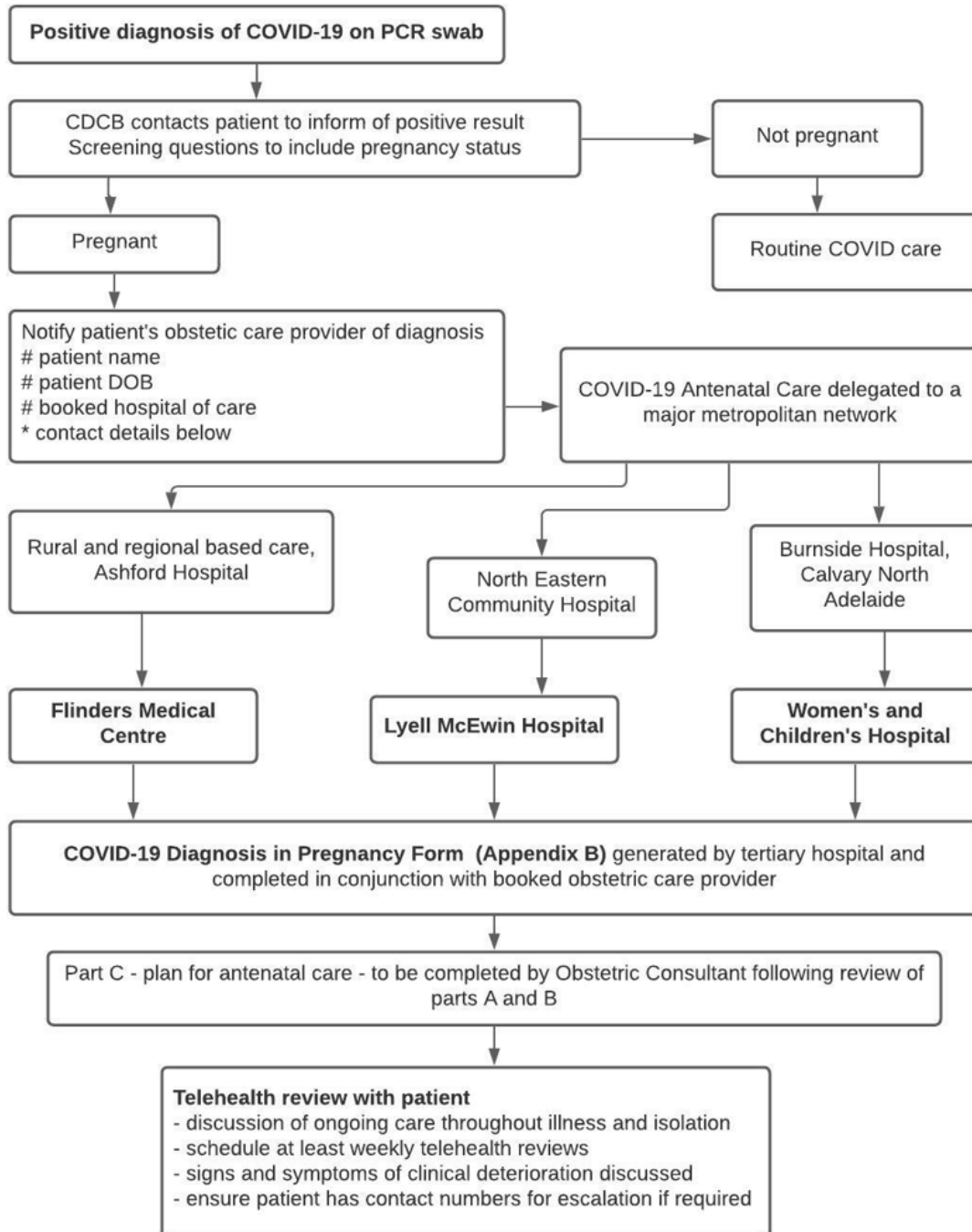
Classification of COVID-19 in pregnancy should be as per the National COVID-19 Evidence Taskforce, with classifications of mild, moderate, severe and critical.

Additional options will need to be made available for smaller services, such as regional hospitals and private care providers, who may not have adequate case numbers or the expertise to coordinate Telehealth care for positive patients. These providers may choose to reallocate care of their patient to an appropriate metropolitan health network (e.g. NALHN, SALHN or WCHN), either for the duration of their infectious period or the remainder of the pregnancy on a case by case basis.

Patients identified as at additional risk may have any of the following comorbidities:

- > Hypertension
- > Asthma
- > Diabetes requiring treatment prior to pregnancy
- > BMI \geq 35
- > Aboriginal or Torres Strait Islander background
- > Other ethnic minority background
- > Chronic respiratory or kidney disease
- > Cardiovascular disease
- > Other immunocompromised
- > Significant psychosocial factors
- > Significant mental health conditions
- > Active cancer
- > Unvaccinated

Coordination of Care



Monitoring of COVID Positive Patients

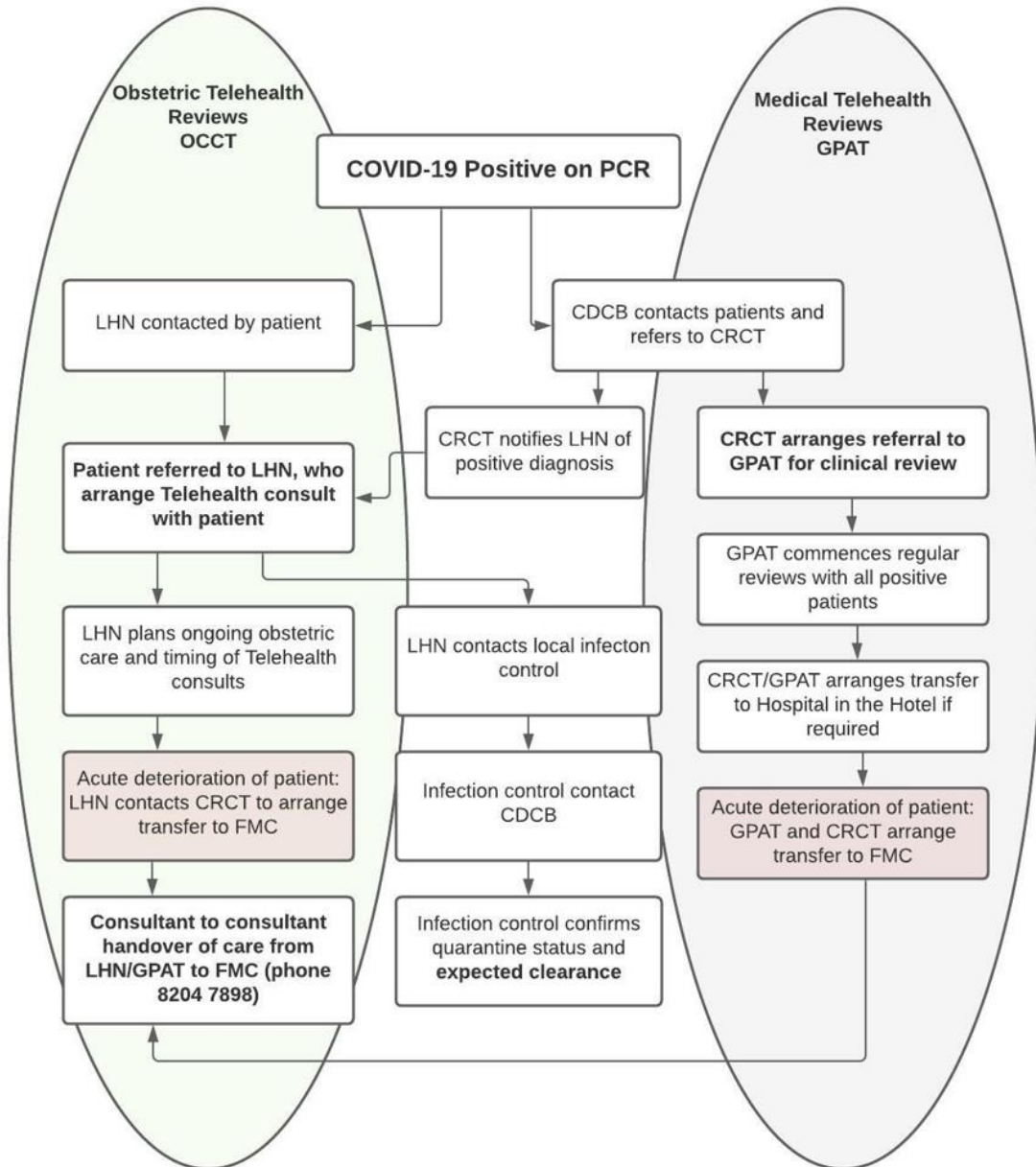
Pregnant patients with COVID-19 are being monitored via Telehealth, both through their obstetric providers and GPAT. Obstetric providers are continuing to monitor patients weekly, or more often if clinically required. GPAT are also providing Telehealth, with escalating responses for patients at higher risk.

Some patients may be supplied with an in-home observation monitoring kit. This will include a Thermometer, O2 probe, Paracetamol, Hydralyte, COVID19 information kit and Rapid Antigen Testing kit x2. The patient will receive instructions on how to take their own heart rate, oxygen saturations, respiratory rate and temperature. They will also have access to the CRCT, GPAT and a live survey for feeding back their observations to GPAT/the CRCT. If available, these observations can be utilised for the obstetric telehealth review.

COVID positive patients in Hospital in the Hotel (dedicated positive sites) requiring obstetric review will be transferred to the FMC for assessment. Patients undergoing supervised quarantine in Hospital in the Hotel for suspected COVID will be transferred to the WCH if they require obstetric assessment.

COVID positive pregnant patients in the community requiring obstetric assessment or admission will need to be reviewed at Flinders Medical Centre.

Model of Care: Telehealth care to remain with booking hospital, inpatient reviews, and admissions to be managed by FMC



Telehealth Reviews

COVID-19 Obstetric Telehealth Review Form (see appendix C)

During obstetric telehealth reviews, patients should be screened for escalating symptoms and fetal concerns. These questions should include:

Maternal:

- > Are you short of breath or having difficulty breathing?
- > Do you need to catch your breath on walking around the room?
- > Are you able to finish your sentences without breathlessness?
- > Have you had any blood on coughing? Amount? (escalate if >1 teaspoon)
- > Do you have any chest pain or pressure, particularly with coughing?
- > Are you able to keep fluids down?
- > Any dizziness with standing?
- > Any drowsiness?
- > Are you making a normal amount of urine?
- > Screening questions for specific obstetric concerns relevant to the patient e.g. pre-eclampsia symptoms, symptoms of chorioamnionitis, symptoms of preterm labour

Fetal:

- > Fetal movement patterns
- > PV loss/bleeding
- > Signs of labour

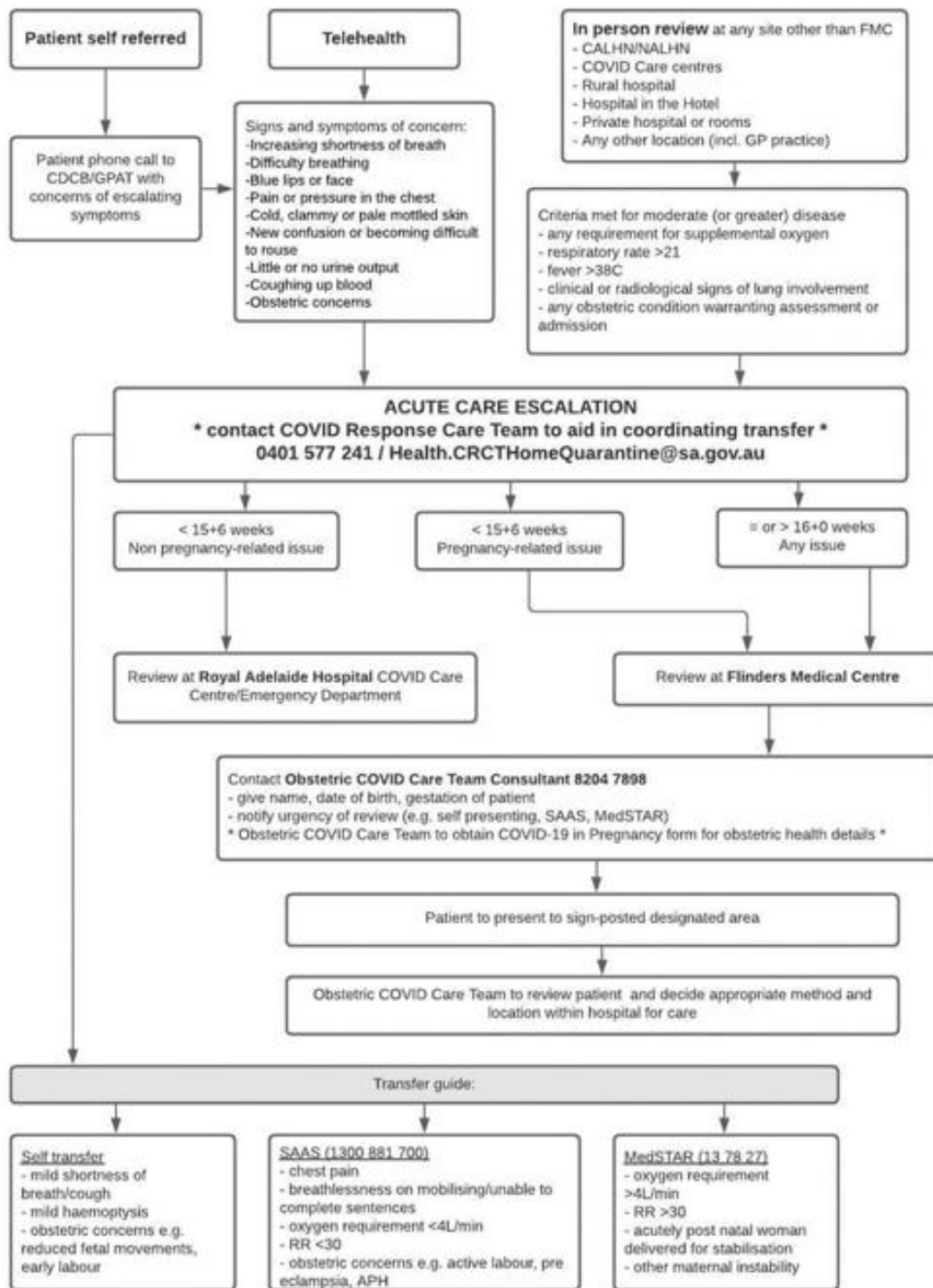
It should be checked that patients have received the appropriate written literature on when to escalate their care and the numbers to call to do this.

At the end of the obstetric Telehealth review, it should be explained that the patient will have a repeat review in 1 week unless otherwise clinically indicated, and this time should be specified.

In Hospital Review and Inpatient Management

If a patient has symptoms during any obstetric Telehealth review that warrant review and escalation of care, the COVID Response Care Team should be notified to assist in coordination of presentation for review at Flinders Medical Centre. The FMC COVID obstetric consultant should be notified.

Acute Care Escalation



COVID-19 Obstetric Escalation Guide

Designed for utilisation within SA Health

Category	Oxygen requirements	Maternal Care	Fetal considerations (>23 weeks)
Green <i>(mild disease)</i>	SpO2 >95% Room air and RR ≤ 20	Exclude other obstetric or medical issues OUTPATIENT CARE May be discharged for in home care Consider monoclonal antibody infusions and thromboprophylaxis	
Yellow <i>(moderate disease)</i>	SpO2 92-98% on < 4L/min And/or RR ≥ 21	INPATIENT CARE - obstetric doctor review Notify - obstetric consultant - obstetric anaesthetist - COVID medical team	Assess fetal well being Discuss timing of birth Consider - steroids for fetal lung maturity - MgSO4 for neuroprotection
Orange <i>(severe disease)</i>	SpO2 92-98% on ≥ 4L/min And/or RR ≥ 25	URGENT Obstetric review Refer for URGENT ICU review	Discuss risks and benefits of emergency caesarean Notify neonatal team
Red <i>(critical disease)</i>	SpO2 <92% on 15L/min via non- rebreather mask	URGENT ICU review Immediately activate MET call URGENT Obstetric attendance Consider awake proning/high flow oxygen	Discuss risks and benefits of emergency caesarean
Peri-arrest	33# OBSTETRIC, NEONATAL MET/MER (CODE BLUE) Multidisciplinary team discussion regarding possible intubation of mother +/- delivery of neonate		

* Adapted from **Coronavirus (COVID-19) in Pregnancy, Information for Health Professionals.**

RCOG and National COVID-19 Clinical Evidence Taskforce Mild Illness

COVID-19 Disease Severity

Disease-modifying agent in pregnancy	Asymptomatic or Mild illness	Moderate illness (requiring supplemental oxygen)	Severe illness	Critical illness
Sotrovimab	Consider in second and third trimester	Not Recommended	Not Recommended	Not Recommended
Casirivimab/Imdevimab (Ronapreve®)	Consider (A)	Consider if seronegative	Consider if seronegative	Not Recommended
Dexamethasone^b	Not Recommended	Start (C)	Start or continue (C)	Start or continue (C)
Remdesivir	Not Recommended	Consider (D)	Continue if commenced (do not start)	Continue if commenced (do not start)
Baricitinib	Not Recommended	Given the limited data on Baricitinib in pregnant and breastfeeding patients, should only be used in clinical trials		
Sarilumab	Not Recommended	Given the limited data on Sarilumab in pregnant and breastfeeding patients, should only be used in clinical trials		
Tocilizumab	Not Recommended	Consider (E)	Consider (E)	Consider (E)

(**A**) Consider using Sotrovimab within five days or Casirivimab/Imdevimab within 7 days of symptom onset in pregnant or breast-feeding patient who do not require oxygen, are not fully vaccinated or immunosuppressed and who have one or more risk factors for disease progression (refer to individual drug monographs below)

(**B**) Seek specialist advice for patients taking long term or high dose corticosteroids prior to admission

(**C**) Obtain guidance from obstetric medicine, prednisolone or hydrocortisone may be preferred in the first trimester.

(**D**) Paucity of evidence of efficacy in COVID-19 infection. Consider using Remdesivir for selected pregnant or breastfeeding patients hospitalised with moderate to severe COVID-19 who do not require ventilation with ID guidance. Pregnant patients were excluded from all clinical trials of Remdesivir in COVID-19.

(**E**) Consider using tocilizumab for the treatment of COVID-19 in pregnant or breastfeeding women who require supplemental oxygen, particularly where there is evidence of systemic inflammation.

VTE prophylaxis

Background

Active COVID-19 is a major risk factor for VTE, with obstetric patients sitting at an elevated baseline risk. Studies in non-obstetric patients have shown up to 22% VTE rate in patients admitted to ICU with early strains of COVID-19. It is not yet known the exact effect the Omicron variant will have on VTE rate, however each patient should be assessed for risk factors for VTE at the time of diagnosis, when their clinical situation changes and for post-partum prophylaxis to ensure our patient come to no harm.

As there is a very large amount of COVID-19 in our community at this point, we will also need to focus on teaching patients to self-administer to limit our usage of community services.

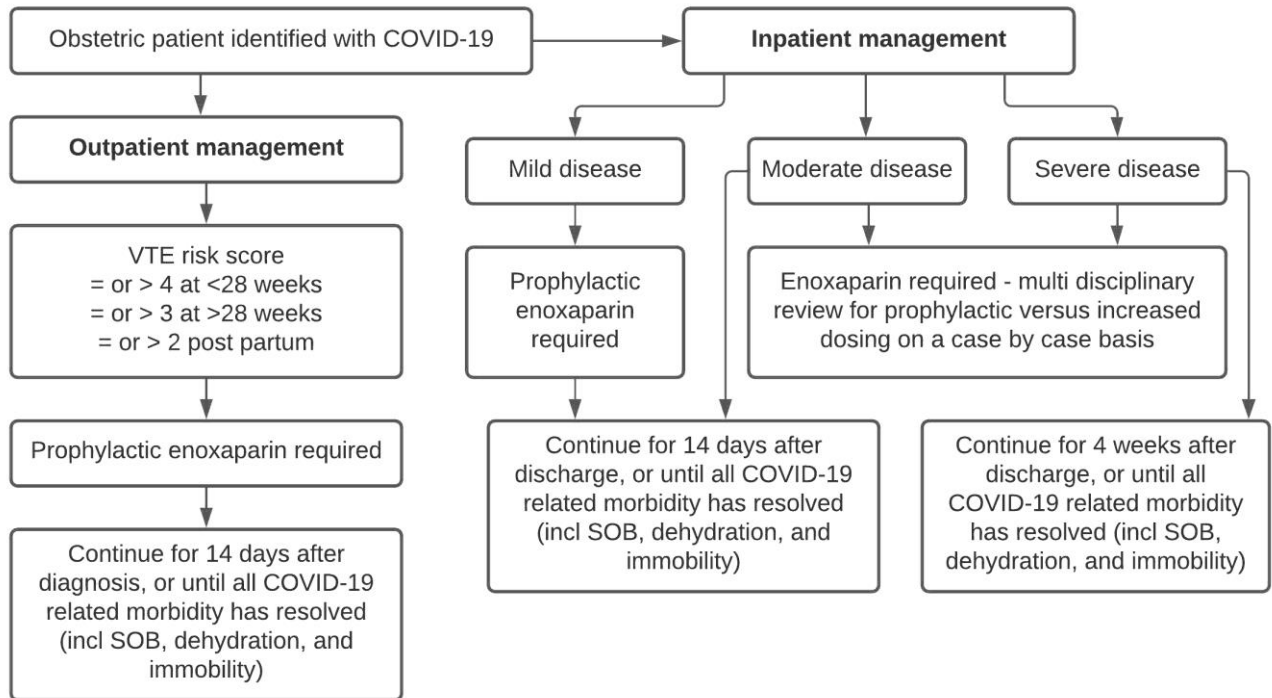
Basic information regarding VTE should be discussed with patients, including the need for hydration and mobilisation, and the signs and symptoms to be aware of. When admitted, there should be consideration for compression stockings and mechanical prophylaxis (calf stimulators) as appropriate for each case.

Who Needs VTE Prophylaxis?

All patients admitted to hospital with COVID-19 will require VTE prophylaxis. The dosing and length of continuance on discharge will depend on their severity of illness.

Patients who are either asymptomatic or mildly symptomatic and managed in the community will require VTE prophylaxis if they are at increased risk.

VTE Prophylaxis



VTE Scoring

Each patient should have a VTE risk score calculated and documented at their first review once they have been diagnosed with COVID-19. The scoring system will follow that suggested in the RCOG Greentop Guideline 37.a and as recommended in the updated Coronavirus (COVID-19) Infection in Pregnancy published Dec 2021.

Thresholds for Treatment

Outpatient management

< 28 weeks	Score \geq 4
\geq 28 weeks	Score \geq 3
Post-partum	Score \geq 2

Inpatient management

Any admitted patient Any score

One point should be added to the score if the patient is dehydrated or immobile.

Risk factors for VTE		
Pre-existing risk factors	Tick	Score
Previous VTE (except a single event related to major surgery)		4
Previous VTE provoked by major surgery		3
Known high-risk thrombophilia		3
Medical comorbidities e.g. cancer, heart failure; active systemic lupus erythematosus, inflammatory polyarthropathy or inflammatory bowel disease; nephrotic syndrome; type I diabetes mellitus with nephropathy; sickle cell disease; current intravenous drug user		3
Family history of unprovoked or estrogen-related VTE in first-degree relative		1
Known low-risk thrombophilia (no VTE)		1 ^a
Age (> 35 years)		1
Obesity		1 or 2 ^b
Parity ≥ 3		1
Smoker		1
Gross varicose veins		1
Obstetric risk factors		
Pre-eclampsia in current pregnancy		1
ART/IVF (antenatal only)		1
Multiple pregnancy		1
Caesarean section in labour		2
Elective caesarean section		1
Mid-cavity or rotational operative delivery		1
Prolonged labour (> 24 hours)		1
PPH (> 1 litre or transfusion)		1
Preterm birth < 37 ^{wo} weeks in current pregnancy		1
Stillbirth in current pregnancy		1
Transient risk factors		
Any surgical procedure in pregnancy or puerperium except immediate repair of the perineum, e.g. appendicectomy, postpartum sterilisation		3
Hyperemesis		3
OHSS (first trimester only)		4
Current systemic infection		1
Immobility, dehydration		1
TOTAL		

Abbreviations: ART assisted reproductive technology; IVF in vitro fertilisation; OHSS ovarian hyperstimulation syndrome; VTE venous thromboembolism.

^a If the known low-risk thrombophilia is in a woman with a family history of VTE in a first-degree relative postpartum thromboprophylaxis should be continued for 6 weeks.

^b BMI ≥ 30 = 1; BMI ≥ 40 = 2

Dosing of Enoxaparin

Therapeutic anticoagulation should be given in the form of enoxaparin as a first line.

Prophylaxis	Dosing
Creatinine Clearance (CrCl) <30mL/min or body weight < 50kg	Enoxaparin 20mg daily or Unfractionated Heparin 5000 units BD*
Weight 50-90kg + CrCl >30mL/min	Enoxaparin 40mg subcut daily
Weight 91-130kg + CrCl >30mL/min	Enoxaparin 60mg subcut daily
Weight 131-170kg + CrCl >30mL/min	Enoxaparin 80mg subcut daily
Weight >170kg + CrCl >30mL/min	Consult Obstetrics Medicine or Haematology

Cautions and contraindications for VTE Prophylaxis

- > Imminent delivery
- > Known bleeding disorder (e.g. haemophilia, von Willebrand's disease or acquired coagulopathy)
- > Active antenatal or post-partum bleeding
- > Increased risk of major haemorrhage (e.g. placenta praevia)
- > Thrombocytopenia (Plt <75 x 10⁹/L)
- > Acute stroke in previous 4 weeks (haemorrhagic or ischaemic)
- > Severe renal disease (GFR <30ml/minute/1.73m²)
- > Severe liver disease (prothrombin time above normal range or known varices)
- > Uncontrolled hypertension (blood pressure >200mmHg systolic or >120 mmHg diastolic)

Post-Partum

Active COVID-19 within the first 6 weeks after birth

If a patient develops COVID-19 in the first 6 weeks after birth and is suitable for outpatient management, they should be assessed and offered prophylactic enoxaparin if their VTE risk score is ≥ 2 . This should continue for 14 days post diagnosis or until their symptoms have completely resolved.

If a patient requires admission post-partum for COVID-19, they should be offered prophylactic enoxaparin as per the mild, moderate and severe guidance above, with continued dosing on discharge as per antenatal patients.

Women who have had COVID-19 in pregnancy but are negative at the time of birth

If a patient was managed as an outpatient during her COVID-19 infection but has recovered at the point of birth, she should be offered 14 days of prophylaxis if her VTE risk score is ≥ 2 .

Women who have had COVID-19 during pregnancy requiring hospitalisation for their illness will all require post-partum prophylaxis.

Mild-moderate illness 14 days prophylaxis

Severe-critical illness 6 weeks prophylaxis

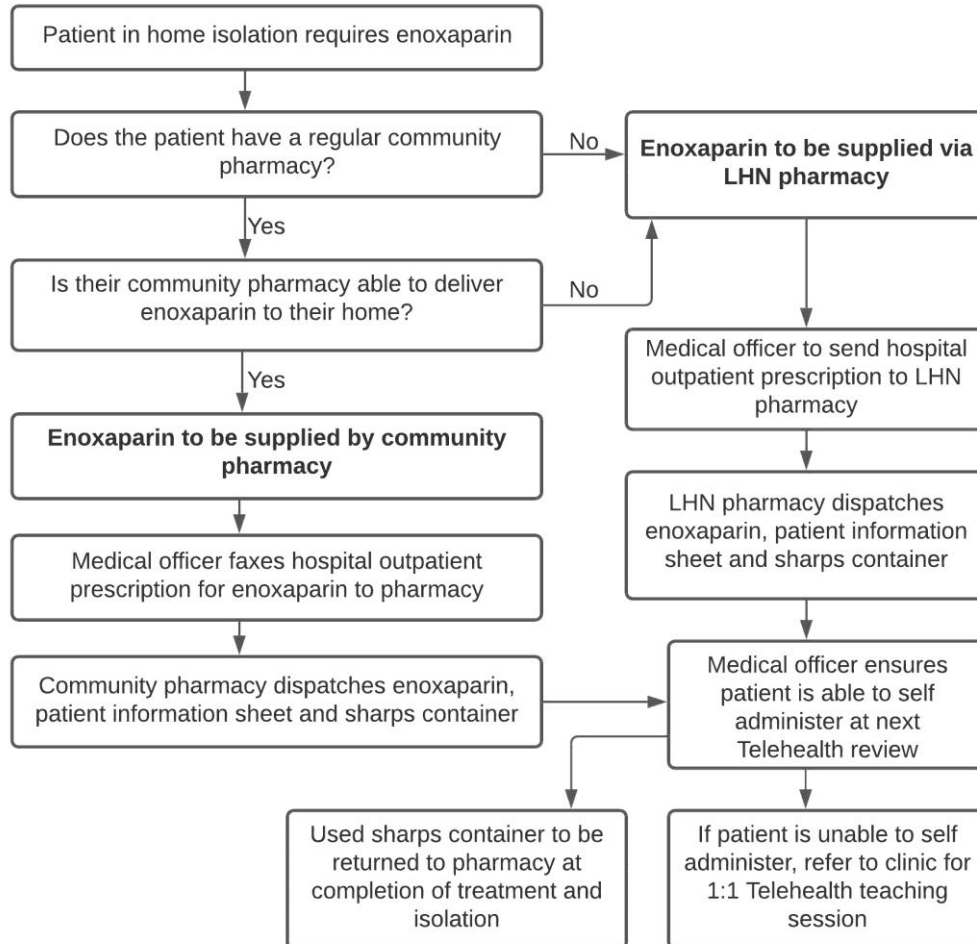
Administration Logistics

Antenatal Outpatients

Patients who are identified as requiring VTE prophylaxis whilst in home isolation will require the medication to be delivered to their house and will need written and/or virtual education for administration.

If the patient has a community pharmacy they are linked in with for their normal medications, the enoxaparin can be ordered through this pharmacy. The contact details of the pharmacy should be obtained and a hospital outpatient prescription faxed through. The pharmacy can then deliver the enoxaparin, product brand instruction sheet and sharps container to the patient.

If the patient does NOT have a regular community pharmacy, or their regular community pharmacy is unable to deliver medication to their home, the enoxaparin should be obtained through the LHN pharmacy.



Once a script for enoxaparin is sent to the Hospital pharmacy, the pharmacy team will dispense the medication with the patient information and video link together with a small sharps container to the patients home address via courier. Instructions to the patient will include advice on return the sharps bin to their local pharmacy following isolation.

Antenatal Inpatients

Patients admitted with COVID-19 should be taught to self-administer their own enoxaparin so that they are as comfortable and confident as possible at the point they are discharged back to the community.

The medical officer responsible for discharge should order their enoxaparin and medical waste container from the hospital pharmacy that can be sent home with

the patient. The patient will have ongoing Telehealth reviews through their LHN, through which the timing to cease can be decided (i.e. 14 days post discharge, or longer if the patient remains symptomatic).

Post-partum patients

Patients identified as requiring prophylactic enoxaparin post-partum should be taught to self-administer during their inpatient stay for birth. The medical officer responsible for discharge should order their enoxaparin and medical waste container that can be sent home with the patient.

Communication with GPAT

The GP Assessment Team is involved with overseeing all COVID-19 positive cases within the community and wish to be informed when patients are started on enoxaparin.

If enoxaparin is started on a patient in the community, please send an email to:

DL.HealthMediHotelGPAT@sa.gov.au

Please state the patients name, date of birth, dosage and duration of enoxaparin therapy.

This email will not require any further follow up or action on behalf of the medical officer.

Monoclonal Antibody Infusions

Monoclonal antibody infusions should be considered in early disease to reduce the likelihood of development of moderate-severe COVID-19.

Sotrovimab appears to have greater efficacy against the Omicron variant than Casirivimab/Imdevimab and as such, is the current monoclonal antibody infusion being offered through the COVID Care Centres.

To be eligible for a MAB infusion, pregnant women need to be in their second or third trimester and:

- > with symptom onset of no more than 7 days
AND
- > who do not require oxygen
AND
- > Immunosuppressed irrespective of vaccine status

OR

- > who have reduced immunity to COVID-19 e.g. not vaccinated, not fully vaccinated (including patients who received a second dose of vaccine < 2 weeks prior) AND who have one or more risk factors for severe or critical illness.

Risk factors for severe or critical illness include:

- i. Vaccination status
 - > No vaccination
 - > Received 1 vaccine
 - > < 2 weeks since second vaccine
 - > > 6 months since second vaccine
 - > Fully vaccinated but immunocompromised
- i. Risk factors for progressing to severe illness
 - > Age \geq 55 years or \geq 35 for Aboriginal and Torres Strait Islander
 - > Diabetes or pregestational diabetes AND requiring medication
 - > Chronic kidney disease (eGFR < 60 mL/min)
 - > Chronic liver disease (cirrhosis)
 - > Obesity (BMI > 30kg/m²)
 - > Moderate to severe asthma (on inhaled corticosteroid or prescribed course of oral steroid in previous 12 months)
 - > Chronic lung disease (chronic bronchitis, COPD, emphysema with dyspnoea on exertion)
 - > Congestive heart failure (NHYA Class II or above)
 - > Cardiovascular disease
- i. Immunocompromising conditions
 - > Haematological neoplasm (leukaemia, lymphoma or myelodysplastic syndrome)
 - > Haematopoietic stem cell transplant within 24 months
 - > Solid organ transplant on immunosuppressive therapy
 - > Primary or acquired (HIV/AIDS) immunodeficiency
 - > Current or recent immunosuppressive therapy
 - > Chemotherapy or radiotherapy
 - > High dose corticosteroids (\geq 20mg prednisolone per day or equivalent) for \geq 14 days
 - > Biological therapy or disease-modifying anti-rheumatic drugs

If a patient is a potential candidate for a MAB infusion, the centralised online form should be completed:

<https://forms.office.com/Pages/ResponsePage.aspx?id=9yilvan8L0O8mL1-kNQJBkHkTVb2yj1EgO3yQqVPjONUNTVTWjIGTDROU1pNUTIQSVcxWTJRQkY1Qi4u>

The application will be reviewed by SA Health and Infectious Diseases, the patient will then be contacted by the CCC to arrange infusion timing and location if suitable.

Mild Illness

Adults not presenting any clinical features suggestive of moderate or severe disease or a complicated course of illness

Mild Illness

Characteristics

- > No symptoms
- > Or mild upper respiratory tract symptoms
- > Or cough, new myalgia or asthenia without new shortness of breath a reduction in oxygen saturation

Monitoring of Pregnant COVID Patients

Pregnant patients who have mild disease only and are asymptomatic will be managed at home with Telehealth support via GPAT and their designated LHN Obstetric COVID Care Team.

Investigations: no routine investigations are required at this point.

When to Escalate Care

Patients should be referred to Flinders Medical Centre for review if they report:

- > Increasing shortness of breath
- > Difficulty breathing
- > Blue lips or face
- > Pain or pressure in the chest
- > Cold, clammy or pale mottled skin
- > New confusion or becoming difficult to rouse
- > Little or no urine output
- > Coughing up blood

Treatments

Disease modifying treatments are available for mild disease and are appropriate for use in pregnancy. Providing all appropriate criteria are fulfilled, pregnant patients may be considered for monoclonal antibody infusions. VTE prophylaxis should be discussed for all patients.

Moderate/Severe Illness

Moderate Illness

Stable adult patient presenting with respiratory and/ or systemic symptoms or signs. Able to maintain oxygen Saturation above 92% (or above 90% for patients with chronic lung disease) with up to 4L/min oxygen via nasal prongs

Characteristics:

- > Prostration, severe asthenia, fever $>38^{\circ}\text{C}$ or persistent cough
- > Clinical or radiological signs of lung involvement
- > No clinical or laboratory indicators or clinical severity or respiratory impairment

Severe Illness

Adult patients meeting any of the following criteria

- > Respiratory rate >30 breaths/min
- > Oxygen saturation $<92\%$ at rest state
- > Arterial partial pressure of oxygen (PaO_2) inspired oxygen fraction (FiO_2) <300

In pregnancy, patients with moderate or severe disease will be managed as inpatients at Flinders Medical Centre.

Investigations: baseline CBC, EUC, LFT, CXR and ECG

Observations: all patients with moderate illness will begin on 4 hourly observations. If there is any clinical deterioration or progression to severe illness, observations will be hourly. Clinical escalation may be based on any observations, however guidance by oxygen requirements and respiratory rate is attached as an addendum to this document.

Fetal monitoring: should be decided on a case-by-case basis by the Obstetric COVID Care team taking into account the gestation and clinical status of the mother. Any acute deterioration >28 weeks should prompt a baseline CTG, with the decision for continuous monitoring to be decided by the clinical scenario.

Treatment

Once a patient is diagnosed with moderate disease, there are increased options for pharmacotherapies.

Corticosteroids are recommended as part of the supportive care once oxygen administration has begun. Disease modifying treatments include Remdesivir for patients who are not ventilated and Casirivimab plus Imdevimab for patients who are seronegative. Once there are biochemical signs of systemic inflammation (CRP >75, Ferritin >500) it is recommended to treat with either Tocilizumab. Given the worldwide shortage of Tocilizumab, any immunomodulators in this group should be started after multidisciplinary input including infectious diseases and pharmacy.

Thromboprophylaxis

Should be considered for all patients with moderate to severe disease. Consideration of 40mg enoxaparin SC BD in patients with severe disease.

Corticosteroid Treatment

Corticosteroid treatment is recommended for the treatment of \geq moderate disease, however the choice of steroid will be guided by Obstetric Medicine, Infectious Disease and ICU at the time of treatment. Patients may be offered dexamethasone or prednisolone depending on their gestation, pregnancy details, comorbidities and other illness factors.

Other Considerations

Bacterial pneumonia:

Treat with antibiotics as appropriate in consultation with respiratory

Antenatal corticosteroids:

Should be given if there is concern for preterm delivery <34+6. Does not need to be given if patient on dexamethasone

Magnesium sulphate

Should be given for eclampsia prophylaxis and neuroprotection of the preterm neonate as routinely indicated.

Delivery

This is a multidisciplinary team decision within the Obstetric COVID Care Team and potentially in conjunction with ICCU. Decision for delivery will be decided on a case by case basis and encompass maternal health status, gestation and disease progression. All patients with \geq moderate disease will need to have a delivery plan discussed and documented in case of acute deterioration.

Critical Illness

Critical Illness

Adult patient meeting any of the following criteria:

Respiratory failure

- > Occurrence of severe respiratory failure ($paO_2/FiO_2 < 200$) respiratory distress or acute respiratory distress syndrome (ARDS).
This includes patients deteriorating despite advanced forms of respiratory support (non-invasive ventilation (NIV), high flow nasal oxygen (HFNO) OR patients requiring mechanical ventilation

OR other signs or significant deterioration

- > Hypotension or shock
- > Impairment of consciousness
- > Other organ failure

Patients with critical illness in pregnancy will be managed in the Intensive and Critical Care Unit at Flinders Medical Centre. The patient will need to be managed in a multi-disciplinary context, with liaison between ICCU, obstetrics, obstetrics physicians, anaesthetics, infectious diseases and neonatology.

Medications:

These patients may be offered dexamethasone, Tocilizumab / Baricitinib however will not be suitable for treatment with Remdesivir if mechanical ventilation is required. Each medication recommendation is as per the COVID-19 National Evidence Taskforce.

All patients in ICCU should have thromboprophylaxis unless there is a major contraindication or delivery is imminent

- > Enoxaparin SC 40mg twice daily or
- > Dalteparin 5000IU twice daily

Mechanical calf stimulators should also be considered for this group.

If delivery is planned, betamethasone should be considered for fetal lung maturity. Dexamethasone will cross the placenta, and if the patient has already been commenced on steroids betamethasone may not need to be administered.

Consider a loading dose of magnesium sulphate (4g IV over 20 minutes) for neuroprotection for all deliveries <30 weeks.

Positioning:

Pregnant patients >24 weeks gestation will need to be managed at 30° left lateral tilt whilst supine to prevent aortocaval compression.

Proning is recommended for patients with critical illness and patients on mechanical ventilation should be managed in the prone position for up to 12 hours per day, noting prone position may be less feasible later in pregnancy as per national guidelines due to the risk of hypofusion, compartment syndrome, pressure ulcers, airway swelling and peripheral arterial compression. A pictorial and video tutorial for appropriate positioning is available through the Green Journal at the following link:

https://journals.lww.com/greenjournal/fulltext/2020/08000/prone_positioning_for_pregnant_women_with.7.aspx

Delivery:

Any patient admitted to ICCU should have a delivery plan discussed and documented. If the patient requires intubation there should be a robust discussion regarding the timing of her delivery plans in consultation with their partner / family.

Gestation at admission	Pregnancy planning
<28 weeks	Expectant management
≥ 28-34 weeks	Individualised care balancing maternal and fetal health
>34 weeks	Low threshold for delivery of fetus in deteriorating mother

Patients with Critical Disease outside of FMC:

In the event that a patient presents with critical disease to a location outside of Flinders Medical Centre, care should be escalated as per the Acute Care Escalation plan detailed earlier in this document.

Acute management will need to be decided on a case by case basis involving (but not limited to) the treating team at the remote location, the COVID Care Response Team and MedSTAR. If the patient is deemed too unstable to transfer and requires delivery and ventilation at a remote location, MedSTAR Kids may also need to be involved to attend for neonatal retrieval.

Birthing Management and Considerations

The birthing plan of a patient affected by COVID-19 may be altered by their disease, however in patients with mild disease or who have recovered it is important to aim to normalise care as much as possible. In patients with severe and critical disease, the birthing plan may be dictated by maternal instability with the decision to deliver made by the multidisciplinary team involved in patient care.

Patients who are birthing whilst positive for COVID-19 will need to be managed with appropriate personal protective equipment as per the SA Health matrix. It is important for the midwifery and obstetric team to remember that standard interventions may take longer due to the need for donning and doffing, which may complicate care in an acute emergency. Caution should be taken with abnormal CTGs, and delivery interventions such as caesarean section or instrumental deliveries offered in a timely manner to limit the need for high category emergency cases or rapid neonatal resuscitation. Whilst fetal scalp sampling is not contraindicated in a patient with COVID-19, the practitioner should be aware of the complexities of management of potential delays in resuscitation if an emergency situation arises at the end of labour.

Mode of delivery:

In patients with mild disease and recovered disease, the mode of delivery should be based on obstetric indications alone. If the patient had previously planned a vaginal delivery, then this plan can remain.

Delayed cord clamping and skin to skin contact may remain as per standard procedure, as can breastfeeding plans and rooming in on well infants. With each of these items, it is important to ensure appropriate hand hygiene and masks are used during the mother's infectious period at times of close contact, to reduce the risk of transmission to the neonate.

CTG Monitoring:

Patients that are positive with mild COVID-19 and asymptomatic do not need continuous electronic fetal monitoring, they will require CTG evaluation for the same reasons as per standard guidelines.

Any patient that is symptomatic regardless of severity will require continuous CTG monitoring throughout labour.

Support persons:

A nominated person will be able to attend as a support person. It must be assumed that they are COVID-19 positive or likely to become positive, so they will be required to wear PPE as per the SA Health matrix, they should remain within the room with the patient and are not to walk through other areas of the hospital. They can attend when the patient is in active labour and will have to leave within a specified timeframe based on current restrictions.

The support person must be someone in the same household who is also in isolation for COVID-19

Observations:

Observations should be taken as per routine care throughout labour and include standard obstetric observations (heart rate, blood pressure, contractions, fetal heart rate) respiratory rate, oxygen saturations and oxygen requirements should also be documented.

Increasing oxygen requirements should be escalated as per the escalation chart in this document. Obstetric and ICU review sought at appropriate intervals.

Observations may be increased to 15 minutely in unwell patients as dictated by the obstetric and intensive care treating teams.

Analgesia

Patients should be offered analgesia throughout the induction and labour process:

- > water immersion in first stage
 - patients who are positive for COVID-19 will not be able to utilise water immersion for first stage of labour
 - water immersion in a shower may be utilised provided the patient is clinically well with mild disease and no medical contraindications
 - water immersion for second stage is not recommended due to the presence of COVID-19 within faecal particulate matter and the increased risk of exposure to staff and the neonate
- > inhaled nitrous oxide
 - inhaled nitrous oxide may be utilised providing a single-patient filter is available on the handpiece

- > opiate medication
 - oral paracetamol + codeine, oxycodone, IM morphine and IV/SC fentanyl can all be used as per normal management

- > epidural anaesthesia
 - epidural should be offered early in the labour process
 - early placement will allow time for the anaesthetist to don PPE and set up appropriately
 - placement of an epidural may reduce the need for a general anaesthetic in the event of fetal or maternal compromise. A general anaesthetic in these circumstances may incur more risks for a mother with COVID-19, and potentially expose staff at the time of intubation

Fluid management:

Patients with COVID-19 will need cautious fluid monitoring to avoid fluid overload and pulmonary oedema. The aim should be for euvolaemia throughout the labour process with hourly fluid input matching output, measured on a strict hourly fluid balance chart.

Third Stage:

Active management of the third stage is recommended to reduce the risk of post-partum haemorrhage and need for additional resuscitation for the patient.

Medications for management of third stage may be as per local guidelines and may include IM or IV stat oxytocin, IM Syntometrine or IV Carbetocin.

- > Medications for post-partum haemorrhage may be given on obstetric indications, with standard considerations for asthma and underlying hypertensive disorders
- > Oxytocin infusion should be given in a low volume regimen as described for Cardiac Disease in Pregnancy in the South Australian Perinatal Practice Guidelines
 - 40 units' oxytocin diluted in a 100mL bag of Normal Saline
 - Infusion run at 25mL/hour for 4 hours

Placental Histology:

All patients who have had COVID-19 at any stage of pregnancy should be offered placental histology. The severity of illness and any sequelae should be noted on the pathology form.

If the patient has active disease at the time of delivery, the placenta will need to be appropriately labelled and double bagged as per local guidelines for sending COVID-19 specimens to the laboratory.

Neonatal Management

Babies born to COVID positive patients at FMC, that require care within the first 14 days post-partum will be treated at SALHN, noting that if clinically indicated the baby be transferred to the WCH. Neonates greater than 14 days post-partum requiring care will receive treatment at the WCH. COVID positive children and babies not needing hospitalisation will be linked to COVID Kids for surveillance and follow up.

Please refer to the state neonatal and paediatric guidance for further information.

COVID-19 Admission Testing in Pregnancy

Patients and their partners are to have Rapid Antigen Testing upon arrival to hospital. The location and timing of the RAT test will be conducted as per local protocol.

If a patient returns a positive RAT result, there will need to be a discussion as to whether it is safe to delay care, arrange transfer to Flinders Medical Centre or continue with the care in the current location.

This decision should be based on the reason for the current presentation and risk to mother and baby, gestation, active labour, bed status at Flinders Medical Centre and availability for transfer through SAAS/MedSTAR if required.

Clearance and Discharge to Treating Hospital

All patients are cleared of their infection at day 10 following their positive diagnostic test, regardless of timing of symptom onset or cessation.

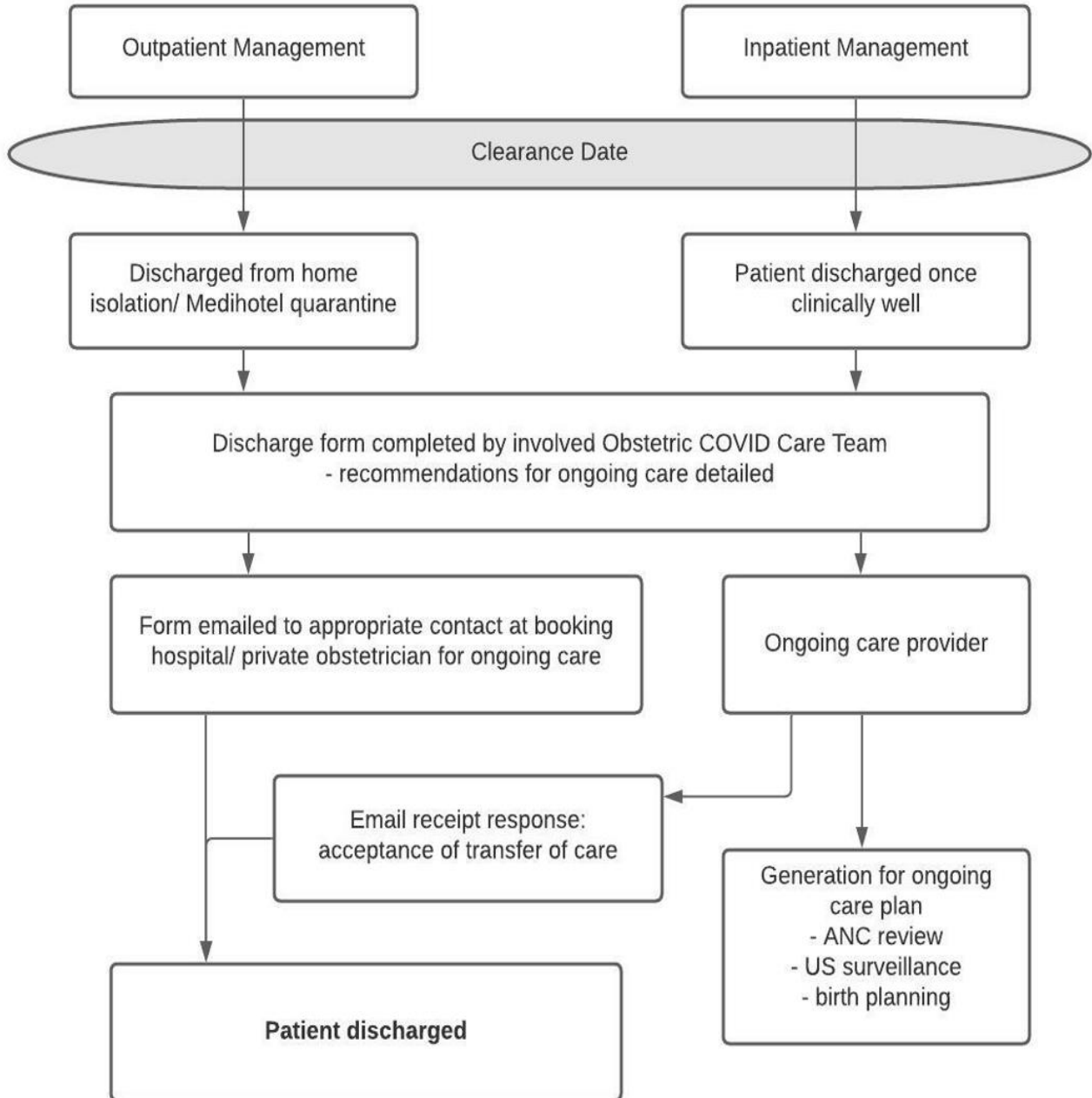
Patients do not require a nasopharyngeal PCR test for clearance, as this may remain positive beyond the time for which they are infectious. Patients should not undergo repeat RAT or PCR testing for one month following diagnosis and are not deemed as close contacts within this time period.

Patients who have been admitted with \geq moderate disease may have a lengthened stay in hospital and should remain under the care of Flinders Medical Centre until they are discharged back into the community.

Upon discharge, the Obstetric COVID Care Team should complete a discharge form that will details (Appendix D):

- > Clearance status
- > Hospital that the patient will be returning to for care/delivery
- > Next planned care contact
- > Any additional care changes as a result of the patients COVID infection
 - Ultrasound at 14 days after symptom onset
(consideration for women admitted with symptomatic disease)
 - Continuance of acute thromboprophylaxis
 - Plan for post-natal thromboprophylaxis
 - Delivery planning if affected by the disease

Discharge Planning



References and Guidelines

The authors of this guidelines thank Monash Health for access and reference to their protocols for COVID-19 in pregnancy.

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The Northern Hospital COVID VTE Prophylaxis Guidelines. Updated 21.12.2021

Appendix A

Elective LSCS “Outsourcing” During the COVID Pandemic

The following list is a guide to patients who may be suitable for outsourcing of their elective LSCS to decant public hospitals during the COVID pandemic. It would be expected that the patient is seen at a mutually convenient time for the patient and surgeon prior to the LSCS but in some circumstances this might be deferred to the day of surgery.

Suitable

- > BMI <40 (level 4 facility can take BMI up to 45 if no other contraindications)
- > Gestation >37 weeks
- > Primary CS – maternal request, previous 3rd / 4th degree tear
 - LGA
 - Breech
 - Uncomplicated DC/DA twins >+37wks
 - Primary genital herpes
- > Repeat CS (= <3) - maternal request or medical recommendation for repeat
 - Other conditions listed for primary CS

Relative Exclusions

- > BMI 40-45
- > CS for maternal medical condition
 - type 1 DM
 - inflammatory bowel conditions
 - Musculoskeletal & neurological conditions
 - Stable cardiac conditions
- > CS for fetal/placental condition not requiring admission
 - Minor anterior placenta previa
 - Posterior major previa
 - No evidence of accrete
- > Previous history of postnatal obstetric, medical or mental health issues
- > Patients who required inpatient management during the pregnancy
- > Complex DC/DA twins >37+ weeks gestation (although if complicated would usually be delivered before 37 weeks)

Absolute Exclusions

- > BMI >45
- > Unstable lie
- > Suspected placenta accrete
- > Placenta previa not managed in the community
- > Vasa previa
- > Significant cardiac condition (any condition needing cardiac opinion during pregnancy)
- > Significant anemia (Hb <100)
- > Bleeding disorder
- > MC/MA twins or triplets or higher order pregnancy (*would most likely be delivered before 37 weeks GA)

VTE prophylaxis

Patients with mild disease in home isolation with no other risk factors for VTE do not require thromboprophylaxis and should be counselled regarding hydration and mobility.

VTE prophylaxis should be considered for patients who are isolating at home/ Hospital in the Hotel (dedicated positive site) with any of the following risk factors:

- > Prior VTE
- > Age >35
- > BMI >30 Plus another risk factor
- > BMI >40
- > Blood dyscrasias
- > Smoking
- > Multiple pregnancy
- > Pre-eclampsia
- > Immobility

VTE prophylaxis should consist of 40mg enoxaparin daily unless delivery imminent and continue for at least 14 days or until all COVID-19 morbidity has resolved. Consider increased dosage to 60mg daily if maternal weight >100kg. Refer to National COVID-19 Evidence Taskforce if enoxaparin is not appropriate for any specific patient

Thromboprophylaxis required:	<input type="checkbox"/> Yes	<input type="checkbox"/> No
Medication:	Dosage:	
Route:	Timing:	
<input type="checkbox"/> GPAT notified to arrange administration		
Signed:	Name/Designation:	

Appendix D

Discharge to Obstetric Care Provider Form (1 page)

NALHN / SALHN / WCHN / Other <u>COVID-19 Discharge to Obstetric Care Provider Form</u>	PATIENT LABEL UR Number: Surname: Given Name: DOB: Sex:
Discharge Details	
Date of discharge:	
Service/ provider discharged to:	
Details of Illness	
Severity of Illness:	
<input type="checkbox"/> Mild	<input type="checkbox"/> Outpatient management <input type="checkbox"/> Inpatient admission <input type="checkbox"/> Moderate <input type="checkbox"/> Severe/Critical
Treatments required:	
<input type="checkbox"/> Casirivmab/imdevimab	<input type="checkbox"/> Dexamethasone <input type="checkbox"/> Prednisolone/Hydrocortisone <input type="checkbox"/> Remdesivir <input type="checkbox"/> Tocilizumab
<input type="checkbox"/> Respiratory support	<input type="checkbox"/> High flow oxygen <input type="checkbox"/> Intubation
<input type="checkbox"/> Thromboprophylaxis	<input type="checkbox"/> Ceased <input type="checkbox"/> Date to cease:
Disease complications:	
.....	
.....	
Ongoing Antenatal Management	
<input type="checkbox"/> Routine antenatal care – no change to management	
<input type="checkbox"/> Ultrasound growth surveillance. Details and timing:	
<input type="checkbox"/> Other interventions:	
.....	
.....	
Birth Planning	
<input type="checkbox"/> Routine planning – no change to management	
<input type="checkbox"/> Recommended alterations to birth planning or timing:	
.....	
.....	
.....	
Post Partum Thromboprophylaxis	
<input type="checkbox"/> Thromboprophylaxis on obstetric indications only	
<input type="checkbox"/> Thromboprophylaxis recommended	<input type="checkbox"/> Medication:
<input type="checkbox"/> Dosage:	<input type="checkbox"/> Duration:
Signed: Date:	

Appendix E: Medications

Sotrovimab

Mechanism of Action	Monoclonal antibody targeting the spike protein of SARS-CoV-2, designed to block virus attachment and entry into human cells
Dose	500mg IV single dose Give over 30 minutes (no dosing adjustment required for hepatic or renal impairment)
Indications	<p>Treatment of COVID-19 within 5 days of symptom onset in adults weighing >40kg who do not require oxygen and who have one or more risk factors for disease progression. These include</p> <ul style="list-style-type: none"> • Diabetes prior to pregnancy requiring medication • BMI >30 • Chronic kidney disease (eGFR < 60) • Congestive heart failure (NYHA ≥II) • Moderate to severe asthma (requiring inhaled corticosteroid, or has been prescribed oral steroids in the last 12 months) • Age ≥ 55 <p>Only for use in</p> <ul style="list-style-type: none"> • Unvaccinated or partially vaccinated patients who meet the above criteria • Immunocompromised: consider regardless of vaccination status • Women in their second or third trimester <p>Do NOT use in fully vaccinated patients unless immunocompromised, such as</p> <ul style="list-style-type: none"> • Primary or acquired immunodeficiency <ul style="list-style-type: none"> ○ Hematologic neoplasms: leukaemia's, lymphomas, myelodysplastic syndromes ○ Post-transplant: solid organ (on immunosuppressive therapy), haematopoietic stem cell transplant (within 24 months) ○ Immunocompromised due to primary or acquired (HIV/AIDS) immunodeficiency ○ Other significantly immunocompromising conditions • Immunosuppressive therapy (current or recent) <ul style="list-style-type: none"> ○ Chemotherapy or radiotherapy ○ High-dose corticosteroids (≥ 20mg of prednisolone per day, or equivalent) for ≥ 14 days ○ All biologics and most disease-modifying anti-rheumatic drugs (DMARDs)
Pregnancy	<p>There is no data on use of Sotrovimab in pregnant or breastfeeding patients. However, its use should be considered in pregnant or breastfeeding patients, particularly for patients in their second and third trimesters of pregnancy, with additional risk factors for severe COVID-19.</p> <p>Sotrovimab is a monoclonal antibody directed specifically against the SARS-CoV-2 virus and therefore is not expected to have significant off target effects. Because Sotrovimab is a large protein molecule, the amount in breast milk is likely to be very low. It is also likely to be partially destroyed in the infant's gastrointestinal tract and absorption both infant is probably minimal.</p> <p>There are no available data on the excretion of Sotrovimab in human milk, and the potential benefits and risks to a breastfed baby are not known. The median elimination half-life of Sotrovimab is 49 days, and human IgGs are known to be excreted in breast milk. A decision whether to discontinue breastfeeding or to abstain from Sotrovimab therapy should consider the benefit of breastfeeding for the baby and the benefit of therapy for the woman.</p>
Contraindications	For discussion in first trimester
Adverse effects	Allergy, diarrhoea, transfusion reaction, fever, rash
Monitoring	Observe for 1-hour post infusion

Casirivimab/Imdevimab

Mechanism of Action	<p>Combination of 2 monoclonal antibodies targeting different sites on the receptor binding domain of the SARS-CoV-2 spike protein.</p>
Dose	<p>Post-exposure prophylaxis: 1200mg (600mg Casirivimab + 600mg Imdevimab) as a single dose via subcutaneous injection or intravenous infusion as soon as possible following exposure to COVID-19</p> <p>Treatment: 1200mg (600mg Casirivimab + 600mg Imdevimab) as a single dose via intravenous infusion.</p> <ul style="list-style-type: none"> ○ Larger doses can be used in severe infections in seronegative patients. Advice on dosing will be provided by Infectious Diseases in these cases. ○ Subcutaneous injection is an alternative route if IV infusion is not feasible or would lead to a delay in treatment however IV infusion is preferred.
Indications	<p>Post-exposure prophylaxis: Prevention of COVID-19 infection in adults and adolescents (aged ≥ 12 years and weighing ≥ 40kg) who have been exposed to COVID-19 and who have: Been a close contact of a confirmed COVID-19 case within the previous 96 hours</p> <p>AND A medical condition making them unlikely to respond to or be protected by vaccination (i.e. immunosuppressed)</p> <p>OR Are considered at high risk of developing severe illness and are not vaccinated or only partially vaccinated against COVID-19</p> <p>Treatment: Treatment of mild to moderate COVID-19 infection in adults and adolescents (aged ≥ 12 years and weighing ≥ 40kg) who:</p> <ul style="list-style-type: none"> ○ do not require supplemental oxygen <p>AND are at an increased risk of progressing to severe COVID-19 and are not vaccinated or only partially vaccinated against COVID-19 OR are immunosuppressed</p> <p>AND are between day 5 and 7 of symptom onset (if < 5 days since symptom onset use Sotrovimab) OR if pregnant and within 7 days of symptom onset</p>
Pregnancy	<p>Consider using Casirivimab plus Imdevimab within 7 days of symptom onset in pregnant or breastfeeding women who are outpatients with mild COVID-19 and who have one or more risk factors for disease progression:</p> <ul style="list-style-type: none"> ● Age ≥ 50 years ● Obesity (≥ 30 kg/m²) ● Cardiovascular disease (including hypertension) ● Chronic lung disease (including asthma) ● Type 1 or 2 diabetes mellitus ● Chronic kidney disease, including those that are on dialysis ● Chronic liver disease <p>Immunocompromised patients (including individuals with rheumatoid arthritis, HIV/AIDS and systemic lupus erythematosus)</p>

	<p>There are no available data on the excretion of Casirivimab plus Imdevimab in human milk, and the potential benefits and risks to a breastfed baby are not known. Human IgGs are known to be excreted in breast milk. A decision whether to discontinue breastfeeding or to abstain from Casirivimab plus Imdevimab therapy should consider the benefit of breastfeeding for the baby and the benefit of therapy for the woman.</p> <p>Consider using Casirivimab plus Imdevimab in pregnant or breastfeeding women who are seronegative patients hospitalised with moderate to critical COVID-19 with ID guidance.</p>
Contraindications	Hypersensitivity to Casirivimab / Imdevimab including previous anaphylactic reactions
Adverse effects	<p>It may be difficult to distinguish between adverse effects of Casirivimab/Imdevimab and signs and symptoms of COVID-19.</p> <p>As a new medication, adverse reactions to Casirivimab / Imdevimab continue to be investigated. Refer to the product information for a complete list of possible adverse effects. To date reactions include:</p> <p>Common/uncommon: injection site reactions, nausea, dizziness, rash and lymphadenopathy</p> <p>Rare: urticaria, flushing, anaphylaxis</p> <p>Suspected or confirmed adverse reactions should be reported via Safety Learning System and also via the Therapeutic Goods Administrations adverse effects online form: TGA adverse event reporting</p>
Monitoring	Observe the patient for 60 minutes after the infusion is completed in case of infusion reaction or anaphylaxis

Dexamethasone

Mechanism of Action	Immunosuppressant and anti-inflammatory, including suppression of cytokine release
Dose	6mg once daily (oral or intravenously) for up to a total of 10 days (can be discontinued if patient is well enough for discharge) Alternative steroids may be used at equivalent doses
Indications	Adults with moderate, severe or critical COVID-19 including those on mechanical ventilation who are requiring oxygen supplementation
Contraindications	Adults who do not require oxygen supplementation, other than for other non-COVID-19 based indications
Adverse effects	Infection, oedema, hypertension, hyperglycaemia, dyspepsia/peptic ulceration, mood and sleep disturbance
Monitoring	Serology for Hepatitis B (surface antigen, surface antibody, core antibody), Hepatitis C, HIV, Strongyloidiasis (serology) and tuberculosis (QuantiFERON gold), however this should not delay use of the medication QID blood glucose monitoring for at least 72 hours after the first dose of dexamethasone For patients in ICU or those with persistent, severe hyperglycaemia, Endocrinology review +/- insulin infusion may be required <i>Blood glucose monitoring can be ceased in patients without diabetes if all blood glucose levels are <7.8mmol/L after 72 hours without the need for glucose lowering therapy and there is no plan to increase the glucocorticoid dose</i>

Prednisolone/Hydrocortisone

Mechanism of Action	Immunosuppressant and anti-inflammatory, including suppression of cytokine release
Dose	Prednisolone: 50mg oral daily for up to 10 days Hydrocortisone: 50mg intravenously 6-hourly for up to 10 days (can be discontinued earlier if patient is well enough for discharge)
Indications	<u>Pregnant patients in the first trimester</u> with moderate, severe or critical COVID-19 including those on mechanical ventilation who are requiring oxygen supplementation.
Contraindications	Adults who do not require oxygen supplementation, other than for other non-COVID-19 based indications
Adverse effects	Infection, oedema, hypertension, hyperglycaemia, dyspepsia/peptic ulceration, mood and sleep disturbance
Monitoring	Serology for Hepatitis B (surface antigen, surface antibody, core antibody), Hepatitis C, HIV, Strongyloidiasis (serology) and tuberculosis (QuantiFERON gold), however this should not delay use of the medication <u>QID blood glucose monitoring for at least 72 hours after the first dose of prednisolone/hydrocortisone.</u> For patients in ICU or those with persistent, severe hyperglycaemia, Endocrinology review +/- insulin infusion may be required <i>Blood glucose monitoring can be ceased in patients without diabetes if all blood glucose levels are <7.8mmol/L after 72 hours without the need for glucose lowering therapy and there is no plan to increase the glucocorticoid dose</i>

Remdesivir

Mechanism of Action	Prodrug metabolised to a C-adenosine nucleotide triphosphate analogue inhibits RNA-dependent RNA polymerase
Dose	200mg IV loading dose, then 100mg daily (from day 2) Total course 5 days (can be discontinued if patient well enough for discharge)
Indications	Adult patients who require supplemental oxygen, but NOT invasive or non-invasive ventilation, ALT < 5x upper limits of normal (ULN) and/or ALT <3x ULN and Bilirubin <2 ULN
Pregnancy	Paucity of evidence of efficacy in COVID-19 infection. Consider using Remdesivir for selected pregnant or breastfeeding patients hospitalised with moderate to severe COVID-19 who do not require ventilation with ID guidance. Pregnant patients were excluded from all clinical trials of Remdesivir in COVID-19. Animal studies do not suggest reproductive toxicity, and use for COVID-19 in pregnant or breastfeeding patients overseas has not shown safety concerns
Contraindications	Ventilated patients Evidence of multi-organ failure including coagulopathy, hepatic failure or renal failure (low urine output or eGFR <30 mL/min or dialysis) or significant cardiomyopathy with low cardiac output. Known hypersensitivity to Remdesivir, the metabolite, or formulation excipient
Adverse effects	Bradycardia, hypotension, gastrointestinal disturbance, rash, hypalbuminaemia, hypokalaemia, anaemia, thrombocytopenia, abnormal LFTs, AKI, respiratory distress
Monitoring	Monitor CBC/EUC/LFTs regularly

Tocilizumab

Mechanism of Action	Humanised anti-IL-6 receptor monoclonal antibody which antagonises IL-6 binding and thus inhibiting its pro-inflammatory effects, reducing inflammation
Dose	For intravenous administration over 60 minutes 800mg if weight >90kg 600mg if weight between 65 – 90kg 400mg if weight 40 – 65kg 8 mg/kg if weight <40kg A second dose should be considered 12-24 hours if no clinical improvement is noted, or the CRP or ferritin do not start to fall
Indications	Adults with hypoxia requiring oxygen supplementation (with oxygen saturations below 92-94% on room air) and signs of inflammation
Pregnancy	Tocilizumab safety information is largely derived from pregnant patients with non-COVID indications such as rheumatoid arthritis. There is no Embryopathy at doses used to treat COVID-19. There is insufficient data to estimate other effects on the pregnancy, but they are likely to be less significant than the effect of COVID. For the babies of patients who received Tocilizumab during pregnancy (after 20 weeks' gestation), live vaccines (rotavirus and BCG) should be avoided in the first 6 months of life. All non-live vaccinations are safe and should be undertaken. Only small amounts of Tocilizumab are detected in breastmilk in patients who receive Tocilizumab whilst breastfeeding only, live vaccines (rotavirus and BCG) can be given to the baby.
Contraindications	<ul style="list-style-type: none"> • Presence of serious bacterial, fungal or serious viral infection (non-COVID) • Active tuberculosis infection • Bowel perforation/diverticulitis • Abnormal ALT/AST >5 times limit of normal • Platelet <50, neutrophil count <0.5 • Prior hypersensitivity to Tocilizumab
Adverse effects	Infections, gastritis, mouth ulcers, hypertension, allergic reactions, gastrointestinal perforation, Cytopenia and hepatotoxicity
Monitoring	Before commencing treatment measure CBC and LFTs; screen for Hepatitis B (surface antigen, surface antibody, core antibody), Hepatitis C, HIV, Strongyloidiasis (serology) and tuberculosis (QuantiFERON gold), however this does not delay use of the medication. Observe for hypersensitivity reactions for 30 minutes Monitor inflammatory markers 12 hours after dose