Severe ARDS in COVID-19-infected pregnancy: obstetric and intensive care considerations

William T. Schnettler, MD, Yousef Al Ahwel, MD, Anju Suhag, MD

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2	obstetric and intensive care considerations
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5	William T. Schnettler, MD <sup>1</sup> ; Yousef Al Ahwel, MD <sup>2</sup> ; Anju Suhag, MD <sup>1</sup>
6	
7	1. Division of Maternal-Fetal Medicine, TriHealth-Good Samaritan Hospital, Cincinnati, Ohio
8 9	2. Division of Pulmonology and Critical Care Medicine, TriHealth-Good Samaritan Hospital, Cincinnati, Ohio
10	The authors report no conflicts of interest.
11	
12	Correspondence should be addressed to: William T. Schnettler, MD. Seton Center for
13	Advanced Obstetrics, Good Samaritan Hospital. 375 Dixmyth Ave. Cincinnati, OH,
14	45220. Telephone: (513) 862-6200, Fax: (513) 862-4358, Email:
15	William Schnettler@trihealth.com
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20	considerations for the management of COVID-19 related ARDS in pregnancy
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23	acute respiratory distress syndrome
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Since the emergence of a novel coronavirus (SARS-CoV-2) in Wuhan, China, at the end of December 2019, its infection – COVID-19 – has been associated with severe morbidity and mortality and has left world governments, healthcare systems and providers caring for vulnerable populations, such as pregnant women, wrestling with the optimal management strategy. Unique physiologic and ethical considerations negate a one-size-fits-all approach to the care of critically ill pregnant women with COVID-19, and few resources exist to guide the multi-disciplinary team through decisions regarding optimal maternal-fetal surveillance, intensive care procedures, and delivery timing. We present a case of rapid clinical decompensation and development of severe Acute Respiratory Distress Syndrome (ARDS) in a woman at 31 weeks' gestation to highlight these unique considerations and present an algorithmic approach to the disease's diagnosis and management.

48	Introduction
49	As of April 3, 2020, the COVID-19 global pandemic totaled 972,303 cases with 50,322
50	deaths (5.2% mortality rate) worldwide, and it is spreading rapidly with a basic
51	reproduction number (R0) of 2-2.5 suggesting that 2-3 people will become infected from
52	an index patient [1,2]. Although the United States now leads the world in total cases
53	(239,279), the U.S. mortality rate is less than half that seen worldwide at 2.3% (5,443
54	confirmed deaths), and the hospitalization rate remains low at 24.1 per 100,000
55	population [3]. Such statistics may embolden skeptics eager to challenge the severity of
56	this public health crisis. However, the rates of critical illness and mortality associated
57	with COVID-19 infection among pregnant women - a potentially highly vulnerable
58	population, remain unclear [4-7].
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60	We present the clinical challenges and potential strategies for optimal maternal-fetal
61	surveillance, intensive care procedures, and delivery timing posed by a case of a
62	pregnant woman at 31 weeks' gestation who presented to a tertiary care hospital in
63	Cincinnati, Ohio, with COVID-19 symptoms, laboratory abnormalities, and chest-
64	imaging findings immediately prior to the development of rapid clinical decompensation
65	and severe Acute Respiratory Distress Syndrome (ARDS) requiring prolonged
66	mechanical ventilation and ultimately indicated preterm delivery.
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68	Case:
69	A 39-year-old Caucasian G6 P2031 with a 31.0-week live singleton intrauterine
70	gestation conceived via in-vitro fertilization required admission on 3/24/2020 from the

71	emergency department of a tertiary care center in Cincinnati, Ohio, due to complaints of
72	5 days of worsening non-productive cough, shortness of breath, fever, and malaise.
73	Four days prior (3/20/2020 - COVID disease day #1 in Figure 1), she had been
74	discharged from the emergency department following work-up for a milder presentation
75	of these symptoms that included normal findings on chest x-ray, normal vital sign
76	assessment, and normal results of a respiratory viral pathogen laboratory analysis
77	including influenza A and B rapid screening. Of note, a nasopharyngeal swab for
78	COVID-19 reverse-transcriptase polymerase chain reaction (RT-PCR) was sent to the
79	Ohio Department of Health at that time, but results did not return for 8 days and were
80	still pending when she re-presented on 3/24/2020 (COVID disease day #5). She had
81	previously established early and complete prenatal care through the Maternal-Fetal
82	Medicine service due to underlying mild myotonic dystrophy (without cardiomyopathy),
83	bicuspid aortic valve (without aortic dilation, stenosis or regurgitation), history of two
84	prior low-transverse cesarean deliveries, and history of a prior mild cerebrovascular
85	accident while on combined oral contraceptives. Her pre-gravid body mass index (BMI)
86	was 24.7, and she denied any tobacco or illicit substance abuse. Serial surveillance with
87	obstetric ultrasound imaging and maternal echocardiography suggested that her
88	pregnancy had thus far been without complication, and she had been compliant with
89	prescribed daily prophylactic low-molecular weight heparin.
90	Upon re-presentation to the emergency department (3/24/2020, COVID disease day
91	#5), her symptoms had progressed to include fever to 101°F, worsening shortness of
92	breath prohibiting the ability to complete full sentences, and persistent non-productive
93	cough (Figure 1). Initial vital sign assessment identified significant tachypnea

(respiratory rate of 32 breaths per minute), mild tachycardia (heart rate in the low 100's
beats per minute), low normal blood pressure (mean arterial pressure in the low 70's
mmHg), and mild hypoxia (SpO <sub>2</sub> 93%) despite 4 liters of oxygen via nasal cannula.
Physical examination was notable for rhonchi and egophony throughout all lung fields. A
chest X-ray, CT pulmonary angiogram, and lung ultrasound assessment were
performed with the findings demonstrated in Figure 2. The attending emergency
department provider ordered the CT angiogram of the chest to investigate the potential
for pulmonary embolism, and the lung ultrasound imaging was performed bedside by
the attending MFM physician to further investigate the potential for COVID-19
pneumonia. Specifically, her chest X-ray identified bilateral diffuse pulmonary infiltrates,
and her chest CT scan identified bilateral airspace disease characterized by ground-
glass appearance with peripheral consolidations compatible with viral pneumonia. Her
lung ultrasound demonstrated bilateral pleural thickening and nodularity of the visceral
pleura (Figure 2). Horizontal A-lines representing normal aerated lung were absent and
replaced by multiple B-lines, pleural nodularity and thickening, and an overall "white
lung" appearance with focal areas of consolidation.
Laboratory analysis identified a normal PaO <sub>2</sub> :FiO <sub>2</sub> ratio > 300 (suggesting no evidence
of ARDS) but definite leukopenia (7,700/mcL), lymphopenia (800/mcL),
thrombocytopenia (114,000/mcL), elevated transaminases (AST 65 IU/L), and a mildly
elevated procalcitonin (0.33 ng/ml). Although the results from her prior COVID-19 RT-
PCR were pending, her clinical presentation supported the diagnosis of COVID-19
infection with pneumonia and the potential for development of Acute Respiratory

Distress Syndrome (ARDS). The emergency department physician felt that her  $SpO_2$  of 93% on 4 liters/minute (L/min) nasal cannula oxygen and  $PaO_2$ :FiO<sub>2</sub> ratio > 300 was appropriate for admission to the hospital's lower-acuity COVID floor, and her disposition was planned to that unit. However, the staff Maternal-Fetal Medicine physician recognized her  $SpO_2$  as abnormal for pregnancy and re-directed her care to the intensive care unit (ICU). The degree of her probable COVID-19 infection at this point was severe [8].

Upon arrival to a negative pressure room in the ICU, she was placed in strict isolation with airborne precautions. Reassessment of her SpO<sub>2</sub> revealed worsening hypoxia (78%) despite 4 L/min oxygen via nasal cannula. A non-re-breather mask was applied, and the inhaled oxygen rate was increased to 15 L/min. Her SpO<sub>2</sub> increased to 82%, but she complained of feeling exhausted with inspiration and the decision was made to proceed with rapid sequence intubation with planned mechanical ventilation for respiratory failure and critical illness severity (approximately 9.5 hours after presentation to the emergency department). Continuous electronic fetal monitoring was initiated and demonstrated a 3-minute prolonged fetal heart rate deceleration to the 80 beat-perminute range that spontaneously resolved following intubation. During this brief period, the consideration for emergent delivery was entertained. Maximal ventilatory assistance was applied, however, and both the maternal and fetal status improved - prompting initiation of both antenatal corticosteroid administration and magnesium sulfate for the dual benefit of fetal neuroprotection and control of maternal bronchospasm (6-gram bolus over 20 minutes intravenously followed by 2 grams per hour IV). She required

large dosages of intravenous benzodiazepines and narcotics for sedation, and her mean arterial blood pressures (MAPs) dropped below 65 mmHg requiring continuous infusion of norepinephrine.

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Over the next 8 hours, the ventilator settings were increased to 100% FiO<sub>2</sub> and PEEP (positive end-expiratory pressure) of 10 cm H<sub>2</sub>0, without complete resolution of her hypoxia (SpO<sub>2</sub> ~ 90-93%). A volume control ventilation modality with automatic pressure augmentation (VC+) was employed to minimize volu- and baro-trauma with the following settings: respiratory rate of 20 breaths per minute, tidal volume of 6 ml/kg, FiO<sub>2</sub> of 100%, inspiratory-to-expiratory time of 1:1.3, and PEEP of 10 cm H<sub>2</sub>0. The PaO<sub>2</sub>:FiO<sub>2</sub> ratio remained below 150 signifying severe ARDS per American-European Consensus Conference criteria [9]. Surprisingly, the fetal heart rate remained reassuring and signified a certain degree of maternal stability. Per recommendations by the infectious disease specialist, the following medications were initiated: ceftriaxone, azithromycin, oseltamivir, and hydroxychloroquine. Marked improvement in her oxygenation was not witnessed until the team manually placed her in a prone position per the PROSEVA study protocol [9]. The initial plan included utilization of a mechanical rotating bed designed for prone ventilation, but manual prone positioning was preferred due to the ability to more quickly return her to supine positioning for performance of CPR or emergent delivery. Manual pronation required a collaborative effort involving the intensive care and obstetric team members to establish invasive hemodynamic monitoring (central venous access and arterial line access), secure her airway, cushion and support her gravid abdomen, and maintain continuous tocodynamometry and

electronic fetal heart rate monitoring (Figure 3). This allowed for gradual diminishment of the FiO<sub>2</sub> and PEEP requirements.

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Her care throughout the following week involved twice daily multi-disciplinary "huddles" to elicit input from the Infectious Disease specialists, Pulmonology and Critical care teams, Maternal-Fetal Medicine teams, Anesthesia Critical Care specialists, Cardiology, CardioThoracic surgeons, Neonatology, and Obstetric nursing teams. Plans were established regarding staff exposure mitigation, emergency preparedness, delivery timing, neonatal resuscitation, nutritional support (oral gavage feedings), venous thromboembolism prophylaxis (subcutaneous heparin twice daily), and adjunctive measures including the potential for inhaled pulmonary vasodilators (epoprostenol) and extracorporeal membrane oxygenation (ECMO). Her COVID-19 RT-PCR result returned positive on hospital day 4 (COVID disease day #8), and strict isolation with airborne droplet precautions was maintained. Consent and regulatory permission was obtained to allow for initiation of a 10 day course of remdesivir – a promising antiviral agent targeting a wide array of RNA viruses including SARS/MERS-CoV [7]. Daily lung ultrasound assessments revealed a lack of visual improvement to her sonographic findings. She was rotated between the prone and left lateral decubitus positions each day with a gradual tolerance toward longer durations out of the prone position on hospital days 6 and 7. This tolerance combined with a gradual reduction in FiO<sub>2</sub> and PEEP requirements negated the need to move forward with delivery or to institute inhaled pulmonary vasodilators or ECMO.

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On hospital day 8 / COVID disease day #13 (32.0 weeks' gestation), the continuous tocodynamometry and fetal heart rate tracing began to demonstrate regular uterine contractions with persistent late decelerations. This prompted a "huddle" and mobilization of all teams to prepare for urgent but non-emergent delivery via repeat cesarean section. Her ICU ventilator circuit was maintained and transported with her to the operating suite to minimize exposure during transport. Bedside echocardiography was utilized to monitor her intravascular volume status to help guide fluid resuscitation. Repeat laboratory assessment revealed a mild coagulopathy (elevated INR 1.7, PT 20.2 sec, aPTT 35.1 sec), and preparations were made for prevention of massive hemorrhage including procurement of blood products and uterotonics (oxytocin and misoprostol) in the operating room. A vertical midline skin incision was made to optimize exposure and minimize vascular injury in case hemorrhage was encountered. Delayed cord clamping was intentionally not employed, and the neonatology team utilized an adjacent operating room for neonatal resuscitation to minimize staff and newborn exposure to the mother. The patient tolerated the repeat low-transverse cesarean delivery quite well without postpartum hemorrhage or respiratory compromise. She returned to the ICU where her status continued to improve over the next several days. A "de-brief" was held among all team members to review opportunities for improvement. Umbilical cord blood gas analysis revealed a normal pH (7.2), PCO2 (63 mmHg), PaO2 (21 mmHg), base deficit of 3, and the male newborn transitioned to extra-uterine life without complication and was extubated on day of life (DOL) 3. His amniotic fluid and nasopharyngeal swabs were sent for COVID-19 RT-PCR analysis on DOL 1 and 2 (24 hours apart), and both results returned negative. The need to maintain strict isolation

and airborne droplet precautions prohibited the patient's husband from visiting either his wife or his newborn son for a total of 14 days. Currently (hospital day 17 / COVID disease day #22), she is improving but continues on synchronized intermittent mandatory ventilation (SIMV) with an FiO2 of 35%, no PEEP requirement, and daily attempts of spontaneous breathing trials.

**Discussion:** 

This case highlights the rapidity of COVID-19 infection in pregnancy with development of severe COVID-19 ARDS within 10 hours of admission, and the importance of considering physiologic maternal adaptations in delineating an algorithmic approach. The maternal physiologic adaptations to pregnancy not only leave the woman more vulnerable to cell-mediated viral infections such as COVID-19 but also more susceptible to rapid cardio-pulmonary decompensation due to the reduced cardiac and pulmonary reserves. Such considerations may not be forefront in the minds of the intensive care team members, and these physiologic alterations must be emphasized by the obstetric providers as they assist in serving in a "quarter-back" role leading the implementation of the algorithmic approach. Such approach must entail input from multiple disciplines and establish a framework for optimal team dynamics utilizing daily "huddles" or other open means of direct communication. This planning should occur prior to any patient's arrival where the myriad of team members establish a consensus regarding the optimal imaging investigations, laboratory studies, COVID testing, fetal assessment, and

admission locations for these women. The team's safety must also remain a priority ensuring appropriate personal protective equipment, staffing (nurse-to-patient ratio), facilities equipped to minimize exposure, and mobile or hand-held equipment with easy cleaning / disinfecting. An example of our management algorithm is included for reference but should be individualized to one's own institution (Figure 4).

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One of the most difficult yet crucial aspects of the management approach is the determination of delivery timing. The physiologic adaptations to labor, delivery, and the immediate postpartum period include maximization of the maternal cardiac output, autotransfusion of up to 500 mL of blood volume back into the intravascular compartment, a catecholaminergic surge, release of inflammatory mediators within the endothelium, and considerable fluid shifts between the interstitial, intracellular, and intravascular compartments. In the setting of severe systemic infection, these physiologic changes can serve to exacerbate the dysregulated inflammatory cascade leading to a higher potential for endothelial dysfunction, pulmonary edema, myocardial edema and cardiac dysfunction [11]. Thus, the decision to proceed toward delivery should be deferred in the setting of severe and critical maternal COVID-19 infection until maternal cardiopulmonary stability can be achieved unless the pregnancy has reached full term, fetal status is non-reassuring, or the maternal status is so dire that evacuation of the uterus is likely to facilitate improvement in cardio-pulmonary function [4]. Consideration for administration of antenatal corticosteroids prior to anticipated preterm birth is controversial in the setting of severe maternal COVID-19 infection. Evidence from treatment studies for SARS suggested that high dosages of corticosteroids posed a risk

for severe side effects that drastically affected prognosis, but shorter courses of low-to-moderate dosages may be considered in the care for the critically ill COVID-19 patient [12]. The decision regarding administration of magnesium sulfate for fetal neuroprotection prior to 32 weeks' gestation should proceed per standard indications in that this agent may provide an additional benefit of bronchodilation in the setting of bronchospasm following intubation. Caution is advised to minimize fluid overload with the administration of magnesium sulfate due to the potential for development of additional pulmonary edema, and we recommend restricting the total volume of infused intravenous fluids to 125 ml/hour or less. Delayed cord clamping and immediate skin-to-skin maternal contact should be avoided [4]. Table 1 represents our approach to delivery considerations including timing, location, and medications.

When attempting to defer delivery and achieve resolution of the acute maternal illness with supportive care, several adjunctive therapies should be considered. Emerging evidence suggests that antiviral agents including hydroxychloroquine and remdesivir may demonstrate efficacy in treating the SARS-CoV-2 virus, but neuraminidase inhibitors such as oseltamivir have no proven benefit [10,13]. Although the safety of these agents in pregnancy has not been definitely determined and their efficacy remains controversial, the pharmacokinetic properties and mechanisms of action may support their judicious use while we await further clinical trials. Non-invasive modes of ventilation such as "CPAP" or "BiPAP" are not recommended for managing acute hypoxemic respiratory failure due to their increased likelihood of failure with need for more urgent transition to invasive ventilation [14]. Rapid sequence endotracheal

intubation should be performed per routine but with consideration for a slightly smaller
endotracheal tube size due to the potentially edematous and narrowed airway calibers
in pregnancy. Oxygenation and ventilatory goals include consideration for the
physiologic mild respiratory alkalosis of pregnancy, the diminished functional residual
volume, a higher PEEP requirement, and potential for less lung compliance with higher
innate plateau pressures due to diaphragmatic compression by the gravid uterus and
chest wall compression by enlarged breast tissue. Physiologic tidal volumes in
pregnancy are greater than the target value of 6ml/kg ideal body weight utilized in the
ARDS Network study [15]. This coupled with the decreased chest wall / diaphragmatic
compliance present a challenge to the "lung protective" strategy for mechanical
ventilation in pregnant patients. Our clinical observation suggests a 5 cm $\rm H_20$ difference
in plateau pressures prior to and immediately following evacuation of the gravid uterus.
Therefore, it seems reasonable to increase tidal volume and/or PEEP to meet goal
PaCO <sub>2</sub> and oxygenation targets remaining mindful not to allow alveolar plateau
pressures to exceed 35 cm H <sub>2</sub> 0.
The prone position can help overcome some of these issues. Prone ventilation has
been found to significantly improve oxygenation in the setting of ARDS, and its
feasibility and safety in pregnancy have been documented [16,17]. Lastly, veno-venous
ECMO is a proven life-saving salvage therapy for severe, reversible respiratory failure,
and its benefit among critically ill pregnant women has been reported [18].

Consideration for ECMO cannulation should be entertained among a multi-disciplinary team of experienced providers in situations where the patient's oxygenation is so

301	severely compromised as to require maximal ventilatory support early in the disease
302	process (less than 7 days of mechanical ventilatory support). Often therapeutic
303	anticoagulation is required, and the postpartum period appears to be a potentially
304	tenuous timepoint for initiation of ECMO with 100% maternal mortality in a recent case
305	series [18].
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307	In summary, this case of rapid clinical decompensation and development of severe
308	(PaO2:FiO2 < 150) COVID-19 related ARDS in a woman at 31 weeks' gestation
309	highlights many of the physiologic and management considerations for the care of
310	critically ill pregnant women with COVID-19. Few contemporary resources exist to guide
311	the multi-disciplinary team through decisions regarding optimal maternal-fetal
312	surveillance, intensive care procedures, and delivery timing. This detailed case reviews
313	the thought process, team-based strategy, and algorithmic approach to this emerging
314	disease's diagnosis and management.
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### Figure Legend

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- 3 Figure 1. Clinical course, major symptoms, and outcomes from illness onset in this
- 4 patient with COVID-19 related critical illness and severe ARDS (acute respiratory
- 5 distress syndrome.

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- 7 Figure 2. Patient's chest imaging including chest CT (A) with ground glass opacities and
- 8 peripheral consolidation bilaterally (arrows), chest X-ray (B) demonstrating bilateral
- 9 diffuse pulmonary infiltrates (arrow), normal (different healthy patient) lung ultrasound
- 10 (C) with characteristic normal A-lines (arrow), and the patient's abnormal lung
- ultrasound images (D, E, F) demonstrating abnormal B-lines (arrow, D), pleural
- thickening and nodularity (arrow, E), and focal sub-pleural consolidation (arrow, F) with
- underlying "white lung" (arrow, F). G represents a schematic demonstrating the 12
- anatomic locations to thoroughly evaluate the lung with sonography.

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- 16 Figure 3. A depiction of pad placement and body positioning to achieve manual prone
- ventilation in a pregnant woman with severe COVID-19 related ARDS. When the patient
- is supine, six to eight standard hospital bed pillows are placed across the patient's face,
- 19 upper chest and arms, lateral abdomen on each side, pelvis, and upper legs. A
- 20 bedsheet is draped over these pillows and then rolled together with the bedsheet
- beneath the patient's back on each side to create a "sandwich". The rolled sheets on
- 22 each side are grasped by the team members and used to roll the patient onto her side
- and then prone such that the pillows remain in position as shown.

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25	Figures 4A, 4B, and 4C. Example management algorithm for the pregnant patient with
26	COVID-19.
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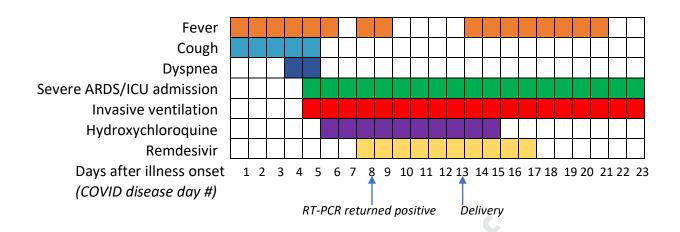
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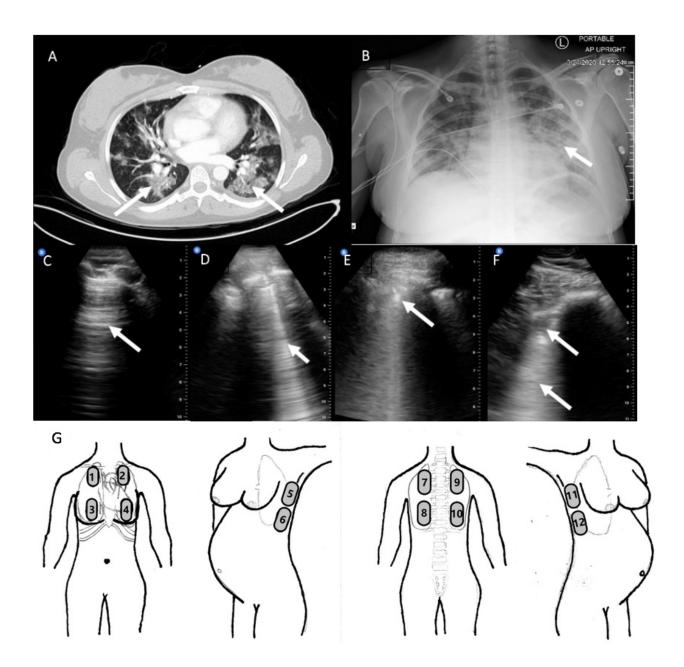
		Delivery Considerations
GA < 24 weeks	Non-Critically ill	If previable PTL – can deliver in COVID unit or LDR
GA < 24 weeks	Critically ill	<ul> <li>Avoid delivery in an UNSTABLE mother</li> <li>If previable PTL – deliver in ICU, main OR if D&amp;C required</li> </ul>
GA 24-34 weeks	Severe but Non-Critically ill	<ul> <li>Attempt to delay delivery and stabilize/ treat mother</li> <li>Betamethasone if imminent delivery within a week</li> <li>MgSO4 for fetal neuroprotection if GA &lt; 32 weeks (if benefits outweigh risk of pulmonary edema)</li> <li>Consider delivery for NRFHTs (category 3 or persistent category 2 fetal tracing) if stable mother</li> <li>Imminent need for SVD – move to LDR</li> <li>Imminent need for C/section – move to L&amp;D OR</li> </ul>
GA 24-34 weeks	Critically ill	<ul> <li>Avoid delivery in UNSTABLE mother</li> <li>Attempt to delay delivery &amp; stabilize / treat mother</li> <li>Case by case determination of delivery for maternal or fetal benefit if stable mother</li> <li>Betamethasone ONLY if HIGH risk for imminent delivery within a week</li> <li>MgSO4 for fetal neuroprotection if GA &lt; 32 weeks (if benefits outweigh risk of pulmonary edema)</li> <li>Imminent need for SVD – deliver in ICU</li> <li>Imminent need for C/section – move to Main OR</li> <li>Perimortem c/section – proceed in ICU</li> </ul>
GA ≥ 34 weeks	Severe but Non-Critically ill	<ul> <li>Attempt to delay delivery and stabilize / treat mother</li> <li>Case by case determination of delivery for maternal or fetal benefit if stable mother</li> <li>Consider delivery for NRFHTs if stable mother</li> <li>Avoid late preterm betamethasone</li> <li>Imminent need for SVD – move to LDR</li> <li>Imminent need for C/section – move to L&amp;D OR</li> </ul>
GA ≥ 34 weeks	Critically ill	<ul> <li>Avoid delivery in UNSTABLE mother</li> <li>Case by case determination of delivery for maternal or fetal benefit if stable mother</li> <li>Avoid late preterm betamethasone</li> <li>Imminent need for SVD – deliver in ICU</li> <li>Imminent need for C/section – move to Main OR</li> </ul>

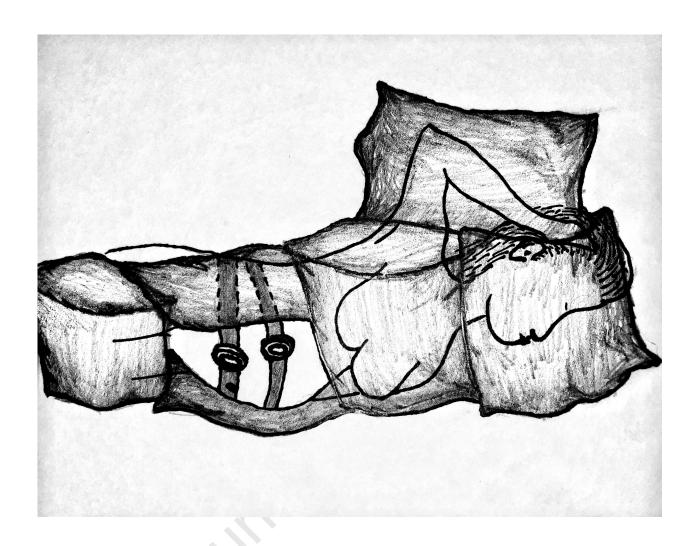
Perimortem c/section – proceed in ICU



Critical Care Goals					
MAP	> 65 mmHg	<ul> <li>First assess if fluid responsive with passive leg raise or bolus LR 500 mL to see if MAP raises &gt; 65 mmHg</li> <li>Start norepinephrine @ 5 mcg/min (up-titrate to 10 mcg/min) for MAP &lt; 65 mmHg</li> <li>Ensure CEFM if GA &gt; 24 weeks</li> </ul>			
SpO <sub>2</sub>	> 94%	<ul> <li>Increase PEEP to 10-24 cm H<sub>2</sub>0</li> <li>Consider VC+ modality</li> <li>Consider prone positioning</li> <li>Ensure finger is warm or place monitor on forehead</li> </ul>			
PaO <sub>2</sub>	> 80 mmHg	<ul> <li>Increase PEEP to 10-24 cm H<sub>2</sub>O</li> <li>Increase I:E ratio</li> <li>Consider prone positioning</li> </ul>			
PaCo <sub>2</sub>	< 40 mmHg	<ul> <li>Increase ventilatory / respiratory rate to 20-25 bpm</li> <li>Consider higher tidal volume than 6 ml/kg ideal body weight</li> <li>Ensure no "auto-PEEP" – keep plateau pressure &lt; 35 cmH<sub>2</sub>0</li> </ul>			
рН	7.3-7.5	<ul> <li>First assess if acidemic or alkalemic</li> <li>Then assess which is more out-of-range (PO<sub>2</sub> or PCO<sub>2</sub>)</li> <li>If metabolic acidosis, assess anion gap &amp; ensure appropriate ventilatory compensation (Bicarb x 1.5) + 8 = PCO<sub>2</sub></li> </ul>			
Bicarb	16-22 mmHg	Consider addition of IV bicarb if low AND pH is < 7.1			
Anion Gap	6-15	Correct for hypoalbuminemia (add 2.5 to gap for every 1 g/dl albumin below level of 2.5 g/dl)			
PiP	< 35 mmHg	<ul> <li>Check peak inspiratory pressure on vent &amp; ensure &lt; 40 cm H<sub>2</sub>0</li> <li>Consider VC+ modality</li> </ul>			
UOP	> 20 ml/kg/hr	Place foley and ensure strict Is/Os + daily weights			
Skin	No break-down	Evaluate skin front & back daily (esp under fetal monitors)			
VTE	prophylaxis	Consider institution of Heparin 7,500 U BID in 2 <sup>nd</sup> trimester & 10,000 U BID in 3 <sup>rd</sup> trimester if delivery is not imminent			
Peptic Ulcer	prophylaxis	Consider H2 blockade			
CEFM	Category 1-2	<ul> <li>Delivery for category 3 if GA &gt; 28 weeks</li> <li>Worsening category 2 may signal worsening maternal status</li> </ul>			
Sedation	Lowest achievable	<ul> <li>Goal is to achieve RASS of 0 (alert &amp; calm) while on mechanical ventilation</li> <li>May need to increase sedation with propofol, fentanyl, &amp; midazolam</li> <li>May need paralytic (cisatracurium) esp when proning</li> </ul>			







READINESS	RECOGNITION	RESPONSE	REPORTING		
<ul> <li>Pre-Hospital</li> <li>Awareness</li> <li>Testing</li> <li>Transport</li> <li>Therapies</li> </ul>	<ul> <li>Presentation</li> <li>Signs Symptoms</li> <li>Physiologic considerations</li> </ul>	<ul> <li>Bed Placement</li> <li>Nurse: patient</li> <li>Capabilities</li> <li>Isolation</li> </ul>	<ul> <li>Internal</li> <li>Debrief</li> <li>Iris reporting</li> <li>QA</li> </ul>		
Hospital     Staffing     Bed space     Equipment     PPE     Preparedness / simulation	<ul> <li>Work-up</li> <li>Labs</li> <li>Imaging</li> <li>Ancillary teams</li> <li>Point people / champions</li> </ul>	<ul> <li>Multi-disciplinary</li> <li>Communication</li> <li>Huddles</li> <li>Assign "Captain"</li> <li>Delivery preparedness &amp; decision tree</li> </ul>	External     Regional HD     State ODH     National - CDC,     SMFM registry		
	<ul> <li>Logistics</li> <li>Timely triage</li> <li>Timely dispo</li> <li>Communication</li> <li>Minimizing exposure</li> </ul>	<ul> <li>Treatment</li> <li>Medications</li> <li>Ventilation/Oxy</li> <li>Positioning</li> <li>Surveillance</li> <li>Family / support</li> <li>Care for self</li> </ul>			

# Possible COVID-19

Fever, nonproductive cough, SOB, tachypnea, HA, diarrhea

Evaluation in Emergency Dept (airborne/droplet precautions)

#### Work-up:

- CXray / CT chest
- Lung US
- CBC w/diff
- CMP
- Lactate
- CRP
- Procalcitonin
- D-dimer
- Rapid Influenza
- Respiratory viral panel
- COVID-19 PCR
- ABG
- Blood cultures
- Sputum cultures
- FHT (CEFM if > 24 weeks)

#### Suspected COVID-19

- CXray / CT chest: bilateral groundglass opacities, peripherally located consolidation
- Lung-US: multiple B-lines, nodular & thickened appearance of pleura (esp visceral), white lung, sub-pleural consolidations
- Leukopenia (<9k), lymphopenia, thrombocytopenia
- Mild transaminitis
- Elevated D-dimer > 1.0
- Low or normal procalcitonin
- Rapid influenza and resp path panel negative
- Elevated CRP
- Elevated lactate > 2

#### Not Likely COVID-19

- Lack of CXray, CT, or US findings
- + influenza or resp viral pathogens
- Elevated procalcitonin & CRP w/ imaging findings that are focal
- No leukopenia, normal D-dimer, CRP, and lactate

# Suspected COVID-19

#### Not Critically-ill

- SpO<sub>2</sub> > 95% on 4 L NC or less
- Resp Rate < 26
- Lactate < 2

Admit - COVID Unit

Airborne/droplet

precautions

Cont pulse ox

Hydroxychloroquine

400 mg BID on day

1,200 mg BID x 4

Azithromycin 500

mg daily x 5 days

ceftriaxone 2 gm gD

Consider remdesivir Daily ABG, CBC,

Strict Is/Qs & daily

Strict PPE

FHT daily

days

Consider

CMP, lactate

- Able to complete sentences
- MAP ≥ 70 mmHg

< 24 Weeks

# ≥ 24 Weeks

- Admit COVID Unit
- precautions

- Hydroxychloroquine 1,200 mg BID x 4 days
- Azithromycin 500 mg daily x 5 days
- 2 gm daily
- Consider remdesivir
- Daily ABG, CBC, CMP, lactate
- Strict Is/Qs & daily weights
- Consider delivery as salvage for critical

#### Critically-ill

- SpO2 < 95% on 4 L NC or more
- Resp Rate ≥ 26 or pH < 7.3, PCO<sub>2</sub> > 40, PO<sub>2</sub> < 80 mmHg
- Lactate ≥ 2
- Not able to complete sentences
- MAP < 70 mmHa

< 24 Weeks

#### ≥ 24 Weeks

- Airborne/droplet
- Strict PPE
- Cont pulse ox
- Twice daily EFM w/ daily NST
- 400 mg BID on day
- Consider ceftriaxone
- Daily Lung US

#### Admit - ICU

- Airborne/droplet precautions
- Strict PPE
- Cont pulse ox & cardiac monitoring
- FHT daily
- Hydroxychloroquine 400 mg BID on day 1,200 mg BID x 4 days
- Azithromycin 500 mg daily x 5 days
- Consider ceftriaxone 2 gm daily
- Consider remdesivir
- Q 12 hr. ABG, CBC. CMP, lactate
- Strict Is/Qs & daily weights + Foley
- Intubate / mech ventilation if SpO2 < 95% on > 10 L
- Avoid midazolam in 1st trimester
- Consider prone positioning rotating q 8 hrs.
- May need PEEP > 10 cm H<sub>2</sub>0
- Consider inhaled epoprostenol if PaO<sub>2</sub> < 80 mmHg
- Norepinephrine to keep MAPs > 65
- Avoid direct supine
- Daily Lung US

- Admit ICU
- Airborne/droplet precautions + PPE
- Cont pulse ox & cardiac monitoring
- CEFM
- Hydroxychloroquine 400 mg BID on day 1,200 mg BID x 4 days
- Azithromycin 500 mg daily x 5 days
- Consider ceftriaxone 2 gm daily
- Consider remdesivir
- Q 12 hr. ABG, CBC. CMP, lactate
- Strict Is/Qs & daily weights + Foley
- Intubate / mech ventilation if SpO2 < 95% on > 10 L
- Consider prone positioning rotating q 8 hrs.
- May need PEEP > 10 cm H<sub>2</sub>0
- Consider inhaled epoprostenol if PaO<sub>2</sub> < 80 mmHg
- Norepinephrine to keep MAPs > 65
- Avoid direct supine
- If ≥ 34 wks, consider delivery
- Daily Lung US

#### **CRediT Author Statement**

William Schnettler: Conceptualization, Writing – Original Draft, Visualization, Supervision.

Yousef Al Ahwel: Writing - Review & Editing, Resources, Methodology

Anju Suhag: Conceptualization, Writing – Review & Editing, Validation, Resources,

Methodology, Supervision