



Rapid Review Update 1: Is there an increased risk of adverse maternal or fetal outcomes in women infected with COVID-19 during pregnancy?

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Please Note:

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Executive Summary

Background

Understanding the implications of COVID-19 infection during pregnancy may help public health practitioners to support expectant parents, and may guide obstetric care for patients with COVID-19.

This rapid review was produced to support public health decision makers' response to the coronavirus disease 2019 (COVID-19) pandemic. This review seeks to identify, appraise, and summarize emerging research evidence to support evidence-informed decision making.

This rapid review is based on the most recent research evidence available at the time of release. A previous version was completed on May 15, 2020. This updated version includes evidence available up to August 17, 2020.

In this rapid evidence review, we answer the question: Is there an increased risk of adverse maternal or fetal outcomes in women infected with COVID-19 during pregnancy?

What has changed in this version?

- Many more syntheses are emerging in the recently found evidence. Only syntheses
 published since April 2020, that included quality appraisal of included studies, were
 added to this update.
- One meta-analysis showed no difference in the rate of pre-term birth or low birthweight among pregnant women with COVID-19 infection compared to non-infected women.

Key Points

- Maternal outcomes: Overall, the available evidence shows a low risk of adverse
 maternal outcomes associated with COVID-19 infection, although most studies do not
 compare rates to those of non-infected women. The overall certainty of this evidence
 related to maternal outcomes is very low (GRADE), and findings are very likely to
 change as more evidence accumulates.
- Labour and delivery outcomes: A meta-analysis showed no difference in the rate of preterm birth among women infected with COVID-19 infection compared to non-infected women. Syntheses report rates of pre-term birth between 20-39% of cases, and a rate for cesarean deliveries among women with COVID-19 of between 48-96% (although the clinical indications for cesarean in these cases are not well described), and the limited available evidence suggests that vaginal delivery can be safe. The overall certainty of this evidence related to labour and delivery outcomes is very low (GRADE), and findings are very likely to change as more evidence accumulates.
- **Fetal and neonatal outcomes**: A meta-analysis found no difference in rates of low birthweight for infected versus non-infected pregnant women. Rates of fetal death and stillbirth are between <1-10%. In syntheses reporting on neonatal COVID-19 infection, between 0-7% of neonates were infected, although it is not known if they were infected

before or during delivery, or after delivery through exposure to infected health care workers. There is no definitive evidence of vertical transmission. The overall certainty of this evidence related to fetal and neonatal outcomes is very low (GRADE), and findings are very likely to change as more evidence accumulates.

Overview of Evidence and Knowledge Gaps

- This rapid review has focused on syntheses. Much of the available evidence comes from case reports and case series, which have an inherently high risk of bias due to the likelihood of selection bias (i.e., those who are included in the study are fundamentally different from those who were not in the study) and lack of a comparison group needed to properly identify elevated rates. Only one synthesis meta-analyzed outcomes for pregnant women with COVID-19 infection versus non-infected pregnant women, based on the results of three studies.
- Although several syntheses exist, there is a large amount of overlap of single studies
 within the reviews. For example, the same eight observed cases of maternal death were
 reported in two syntheses; the same neonatal deaths were reported in more than one
 review. Similarly, the high rate of cesarean delivery may be overestimated due to the
 same deliveries being reported in multiple reviews. Some syntheses made efforts to
 eliminate duplicated cases across studies.
- There is a large number of syntheses available at the time of this update our search located 52 completed syntheses and 41 in progress. Of the available syntheses, this update included seven, published since April 2020 and that undertook quality appraisal of included primary studies. The majority of available syntheses are of low quality, with many lacking methodological details as well as not appraising the quality of included studies. More, rigorous, primary studies are needed, and as they become available, global collaboration is needed to produce a small number of rigorous systematic reviews and meta-analyses.
- Evidence on maternal, fetal and neonate outcomes among women with COVID-19 infection is lacking, particularly for women who are infected in the first and second trimesters.
- It is not known whether COVID-19 infection resulted in clinical indication for cesarean delivery or whether clinician preference determined the method of delivery.
- It is not known whether the reported rate of cesarean delivery or pre-term delivery for women with COVID-19 infection is significantly higher than usual at these locations.

Methods

Research Question

Is there an increased risk of adverse maternal or fetal outcomes in women infected with COVID-19 during pregnancy?

Search

On May 7, 2020, and again on August 17, 2020, the following databases were searched:

- Pubmed's curated COVID-19 literature hub: <u>LitCovid</u>
- Trip Medical Database
- World Health Organization's Global literature on coronavirus disease
- Joanna Briggs Institute COVID-19 Special Collection
- COVID-19 Evidence Alerts from McMaster PLUS™
- Public Health +
- COVID-19 Living Overview of the Evidence (L·OVE)
- McMaster +
- Cochrane COVID-19 Special Collections
- Cochrane Rapid Reviews <u>Question Bank</u>
- Oxford COVID-19 Evidence Service
- Oxford COVID-19 Evidence Service: Current Questions Under Review
- Guidelines International Network (GIN)
- CovidReview
- Prospero Registry of Systematic Reviews
- NCCMT COVID-19 Rapid Evidence Reviews
- MedRxiv preprint server
- PubMed database

A copy of the search strategy is available on request.

What has changed in the methods for this version?

Only syntheses published since April 2020, that included quality appraisal of included studies, were added to this update. Single studies previously included were removed in this update, given the large number of syntheses now available.

Study Selection Criteria

The search results were first screened for recent guidelines and syntheses. English- and French-language, peer-reviewed sources and sources published ahead-of-print before peer review were included. In the update, only recent syntheses that included quality appraisal of included studies were added.

	Inclusion Criteria	Exclusion Criteria
Population	Pregnant women	
Intervention	COVID-19 infection	
Comparisons		
Outcomes	Maternal outcomes including: Mortality; Clinical features and symptoms Fetal/Neonatal outcomes including: Miscarriage/stillbirth/mortality; Birthweight; COVID-19 infection; Vertical transmission Labour and delivery outcomes including: Preterm; Cesarean.	

Data Extraction and Synthesis

Data relevant to the research question, such as study design, setting, location, population characteristics, interventions or exposure and outcomes were extracted when reported. We synthesized the results narratively due to the variation in methodology and outcomes for the included studies.

Appraisal of Evidence Quality

We evaluated the quality of included evidence using critical appraisal tools as indicated by the study design below. Quality assessment was completed by one reviewer and verified by a second reviewer. Conflicts were resolved through discussion.

Study Design	Critical Appraisal Tool
Synthesis	Assessing the Methodological Quality of Systematic Reviews (AMSTAR)
	AMSTAR 1 Tool

Completed quality assessments for each included study are available on request.

The Grading of Recommendations, Assessment, Development and Evaluations (<u>GRADE</u>) approach was used to assess the certainty in the findings based on eight key domains.

In the GRADE approach to quality of evidence, **observational studies**, as included in this review, provide **low quality** evidence, and this assessment can be further reduced based on other domains:

- High risk of bias
- Inconsistency in effects
- Indirectness of interventions/outcomes
- Imprecision in effect estimate
- Publication bias

and can be upgraded based on:

- Large effect
- Dose-response relationship
- Accounting for confounding.

The overall certainty in the evidence for each outcome was determined taking the characteristics of the available evidence into account (observational studies, some not peer-reviewed, unaccounted-for potential confounding factors, imprecise estimates, lack of valid comparison groups for most studies). A judgement of 'overall certainty is low (or very low)' means that the findings are likely or very likely to change as more evidence accumulates.

Findings

Summary of Evidence Quality

This version adds seven new completed syntheses and removes three singles studies, for a total of 18 publications included in this review. Case reports and case series were excluded from this update, due to the risk of bias inherent in these study designs and the availability of more rigorous evidence.

Outcome	Evidence found		Overall certainty in evidence
Maternal mortality	Completed syntheses	3	Very low
Maternal clinical characteristics	Completed syntheses	7	Very low
Fetal/neonatal mortality	Completed syntheses	10	Very low
Birthweight	Completed syntheses	2	Low
Newborn COVID-19 infection	Completed syntheses	6	Low
Vertical transmission	Completed syntheses	5	Very low
Pre-term birth	Completed syntheses	6	Low
Cesarean	Completed syntheses	11	Very low

Warning

Given the need to make emerging COVID-19 evidence quickly available, many emerging studies have not been peer reviewed. As such, we advise caution when using and interpreting the evidence included in this rapid review. We have provided an assessment of the overall certainty of the evidence and a summary of the quality of the evidence as low, moderate, or high to support the process of decision making. Where possible, make decisions using the highest quality evidence available.

Table 1: Syntheses

Reference	Date Released	Review Question [Population (P), Exposure (E), Comparison (C), Outcomes (O)]	Description of Included Studies	Summary of Findings	Quality Rating: Synthesis	Quality Rating: Included Studies
New evidence repor	ted September	3, 2020				
Akhtar, H., Patel, C., Abuelgasim, E., & Harky, A. (2020). COVID-19 (SARS-CoV-2) infection in pregnancy: A systematic review. Gynecologic and Obstetric Investigation. Epub ahead of print.	Jul 30, 2020 (Search completed May 22, 2020)	P: pregnant women and their fetuses/neonates E: COVID-19 infection C: no comparator O: maternal COVID-19 symptoms, pregnancy complications, neonatal COVID- 19 symptoms, neonatal health status, delivery timing and type, vertical transmission	This review includes 22 case reports and case series, reporting on 156 pregnant women aged 22 to 42 with COVID-19 infections.	Outcomes reported were: 108 births (includes 4 sets of twins), 13 women still pregnant at time of reporting, 41 missing outcome data 10 fetal deaths 27 pre-term births Maternal outcomes reported: 19 vaginal and 66 cesarean deliveries; others unknown Most received nasal oxygen therapy 11 required mechanical ventilation Most had low lymphocytes Most had unilateral or bilateral pneumonia on CT scan Symptoms were most commonly fever (53%), cough (32%), malaise (13%) and myalgia (11%) 8 maternal deaths Fetal/neonatal outcomes included: Intrauterine distress (14%) and premature rupture of membranes (8%) Symptoms were most commonly shortness of breath (6%), gastrointestinal symptoms (4%) and fever (3%) The above outcomes were not compared to pregnant women without COVID-19 infection.	Moderate	High
				Outcomes related to vertical transmission were not reported.		

de Melo, G.C. & de Araujo, K.C.G.M. (2020). COVID-19 infection in pregnant women, preterm delivery, birth weight, and vertical transmission: a systematic review and meta-analysis. Cadernos de Saúde Pública, 36(7), e00087320.	Jul 17, 2020 (Search completed May 4, 2020)	P: pregnant women and their fetuses/neonates E: COVID-19 infection C: no COVID-19 infection O: pre-term delivery, neonatal health status, birth weight, delivery timing and type, vertical transmission	This review includes 38 studies reporting on a total of 520 pregnant women in their third trimesters with COVID-19 infections. Most studies were from China, with other studies from Italy, Peru, India, USA, Portugal, South Korea. • 2 cohort studies • 11 crosssectional studies • 4 case control studies • 11 case reports • 10 case series reports	 Birth outcomes were: 433 newborns (366 cesarean deliveries) Delivery outcomes for remaining pregnancies not reported In a meta-analysis of 3 studies, 60 pregnant women with COVID-19 infection, as compared to 219 without COVID-19 infection, found: Odds of pre-term delivery not statistically significant (OR = 2.25, 95% CI 0.96, 5.31) Association of COVID-19 infection and birthweight not statistically significant (MD = -124.16g, 95% CI -260.54, 12.22) 1-2% of neonates tested positive for COVID-19 infection shortly after birth 	Moderate	Very low
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Khalil, A., Kalafat, E., Benlioglu, C., O'Brian P. Morris Completed	P: pregnant women and their	This review includes 86	Maternal socio-demographic and medical risk factors:	Moderate	High
O'Brien, P., Morris, E., Draycott, T., Magee, L. A. (2020). SARS-CoV-2 infection in pregnancy: A systematic review and meta-analysis of clinical features and pregnancy outcomes. EClinicalMedicine. Epub ahead of print.	fetuses/neonates E: COVID-19 infection C: no comparator O: maternal COVID-19 symptoms, maternal mortality, neonatal COVID- 19 infection, neonatal health status, delivery timing and type, vertical transmission	studies with 17 studies included in the quantitative synthesis (2567 pregnancies, with 73.9% in the last trimester of pregnancy). There were five reports from national registries (UK, Netherlands, China, France and Brazil), while the rest were regional reports from other countries (Italy, USA, Spain).	 50.8% were minorities (Black, Asian and Other) 38.2% were obese 32.5% had co-morbid conditions Meta-analysis of pooled outcomes showed: 48.3% of deliveries were by cesarean section 21.8% of births were pre-term (before 37 weeks) and medically required in 18.4% of cases 19.0% of deliveries were for COVID-19 related reasons Maternal ICU admission occurred in 7.0% of cases 3.4% required intensive interventions in ICU maternal death occurred in 0.9% of cases ICU admissions were higher in women with co-morbid health conditions (beta = 0.007, p=0.05) and in those with maternal age > 35 years (beta=0.007, p<0.01) 0.9% of births were still births Neonatal death occurred in 0.6% of cases 1.4% of neonates were SARS-CoV-2 positive after delivery. The above outcomes were not compared to pregnant women without COVID-19 infection. 		

Berbari, E., Fares, J., Murad, M.H. (2020). Clinical presentation and outcomes of pregnant women with coronavirus disease 2019: A systematic review and meta-analysis. Epub ahead of print. E: COVID-19 infection C: no comparator O: maternal COVID-19 treatment, maternal age ranging from 30 to 40 weeks. Outcomes, delivery timing and type, neonatal outcomes Most were from China, with one each from USA, Korea, Iran, neonatal outcomes Women) with maternal age ranging from 25 to 34 years and the gestational age ranging from 30 to 40 weeks. Most were from China, with one each from USA, Korea, Iran, Honduras. **Nonatal outcomes** **37.7% received antibiotic therapy **67.5% received antibiotic therapy	(2020). Clinical presentation and outcomes of pregnant women with coronavirus disease 2019: A systematic review and meta-analysis.	zo20 (Search completed Apr 30, 2020) nd nen tus A riew ysis.	C: no comparator O: maternal COVID-19 treatment, maternal outcomes, delivery timing and type, neonatal	ranging from 25 to 34 years and the gestational age ranging from 30 to 40 weeks. Most were from China, with one each from USA, Korea, Iran,	 73.3% received oxygen therapy 1 maternal death occurred 76.3% delivered by cesarean section Neonatal outcomes 37.7% were preterm Neonatal intensive care admission occurred in 63.7% of cases 3 fetal deaths occurred 2 newborns tested positive for COVID-19 The above outcomes were not compared to 	Moderate	Good
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Trippella, G.,	Jun 18,	P: pregnant	This review	Outcomes reported:	Moderate	High
Ciarcià, M., Ferrari,	2020	women and their	includes 37 case	• 248 newborns (60 vaginal and 179		
M., Buzzatti, C.,	(Search	fetuses/neonates	reports and case	cesarean deliveries)		
Maccora, I., Azzari,	completed	E: COVID-19	series, reporting	• 48 (23%) pre-term births of 208 reporting		
C., Chiappini, E.	Apr 18,	infection	on a total of 275	gestational age		
(2020). <u>COVID-19 in</u>	2020)	C: no comparator	pregnant women	3 first-trimester abortions		
pregnant women		O: maternal	with COVID-19	2 stillbirths; 1 fetal death		
and neonates: A		COVID-19	infections. 181 of	33 women still pregnant at time of		
systematic review		symptoms, birth	the cases were	reporting or missing outcome data		
of the literature with		weight, Apgar	from China,			
quality assessment		scores, neonatal	with 94 from other	Maternal outcomes included		
of the studies.		health status,	countries.	36 received nasal oxygen therapy		
Pathogens, 9(6),		COVID-19 test		5 required mechanical ventilation		
485.		results, delivery		• 29% had low lymphocytes		
		timing and type,		• 95% had unilateral or bilateral pneumonia		
		vertical		on CT scan		
		transmission		Symptoms were most commonly fever		
				(58%), cough (36%), malaise (14%) and		
				shortness of breath (10%)		
				Fetal/neonatal outcomes included		
				• Intrauterine distress (13%) and premature		
				rupture of membranes (8%)		
				The above outcomes were not compared to		
				pregnant women without COVID-19		
				infection.		
				Of 191 neonates tested for the virus that		
				causes COVID-19, 14 (7%) tested positive.		

de Sousa, Á.F.L., de	Jun 11,	P: pregnant	This review	Maternal outcomes:	High	Very low
Carvalho, H.E.F., de	2020	women and their	includes 49	84% of women were in the third		,
Oliveira, L.B.,	(Search	fetuses/neonates	studies reporting	trimester of pregnancy. Of those that		
Schneider, G.,	completed	E: COVID-19	on a total of 755	delivered at the time of the study		
Camargo, E.L.S.,	May 26,	infection	pregnant women	148 (65%) delivered by cesarean		
Watanabe, E.,	2020)	C: no comparator	and 598 infants	section		
Fronteira, I. (2020).	•	O: maternal	(born at time of	 In 103 of these, infection was 		
Effects of COVID-19		COVID-19	study) and	listed as the cause for		
Infection during		treatment,	consisted of:	performing the C- section		
Pregnancy and		maternal	• 21 case	but no further details were		
Neonatal Prognosis:		outcomes,	reports	provided		
What Is the		delivery timing	• 19 cross-	8 maternal deaths were reported		
Evidence?		and type,	sectional			
International		neonatal	descriptive	Neonatal outcomes:		
Journal of		outcomes	studies	 598 babies were born 		
Environmental			• 7 cross-	 493 (82%) of infants were tested for 		
Research and Public			sectional	SARS-CoV-2, of whom 9 (2%) tested		
<i>Health, 17</i> (11), 4176.			analytical	positive*		
			studies	• 101 (20%) were premature		
			1 case-control	• 28 (6%) were underweight		
			study	10 neonatal deaths and one		
			 1 cohort study 	spontaneous abortion were reported		
				 2 had high rates of IgG and IgM 		
			Women were	antibodies specific to the virus but		
			from China (635),	were asymptomatic.		
			USA (60), Italy	, ,		
			(42) Iran (10) and	*the authors report various scenarios in		
			one from each of	which infants tested positive which makes it		
			Asia, Honduras,	difficult to establish route of transmission		
			Australia, Turkey,	(vertical transmission, vs contact route of		
			Spain, Peru,	transmission due to exposure to mother		
			Switzerland and	and or healthcare workers providing care		
			Canada.	[directly and/or indirectly]).		
				The outcomes were not compared to		
				pregnant women without COVID-19		
				infection.		

Juan, J., Gil, M. M., Rong, Z., Zhang, Y., Yang, H., & Poon, L. C. (2020). Effect of coronavirus disease 2019 (COVID-19) on maternal, perinatal and neonatal outcome: systematic review. Ultrasound in Obstetrics & Gynecology, 56(1), 15–27.	May 19, 2020 (Search completed Apr 20, 2020)	P: pregnant women and their fetuses/neonates E: COVID-19 infection C: no comparator O: maternal COVID-19 symptoms, pregnancy complications, neonatal COVID- 19 symptoms, neonatal health status, delivery timing and type, vertical transmission	This review includes 24 case reports and case series, reporting on a total of 324 pregnant women with COVID-19 infections. The majority of the cases originated from China, with other cases from Australia (1) Canada/France (1), Korea (1) Iran (2 case reports and 1 case series), Italy (1 case series and 1 case report), Peru (1), Spain (2), Sweden (1),	Delivery outcomes: 219 births (most were cesarean deliveries; gestational age ranged from 28 to 41 weeks) 4 neonatal deaths 4 miscarriages or abortions 72 women still pregnant at time of reporting Maternal outcomes included Most required ICU admission Few required mechanical ventilation 43.1% had low lymphocytes Most had unilateral or bilateral pneumonia on CT scan The above outcomes were not compared to pregnant women without COVID-19 infection. One case suggested vertical transmission	Moderate	High
		transmission	1 case report), Peru (1), Spain (2), Sweden (1), Turkey (1)	infection.		
			and USA (1 case series and 4 case reports).			

Previously reported e	vidence					
Elshafeey, F., Magdi, R., Hindi, N., Elshebiny, M., Farrag, N., Mahdy, S., Nabhan, A. (2020). A systematic scoping review of COVID-19 during pregnancy and childbirth. International Journal of Gynaecology and Obstetrics 150(1), 47–52.	Apr 24, 2020 (Search completed Apr 19, 2020)	P: pregnant women and their fetuses/neonates E: COVID-19 infection C: no comparator O: maternal COVID-19 symptoms, pregnancy complications, neonatal COVID- 19 symptoms, neonatal health status, delivery type, vertical transmission	Scoping review of 33 studies from around the world, published between 8 December 2019 and 19 April 2020, reporting on clinical characteristics, maternal and perinatal outcomes of 385 pregnant women with COVID-19. Study designs included case control, case reports, case series.	Included cases were predominantly mild: 368 (95.6%) mild; 14 (3.6%) severe; and 3 (0.8%) critical. 17 women were admitted to intensive care, including 6 who were mechanically ventilated and 1 maternal mortality. 252 women gave birth: 69.4% cesarean and 30.6% vaginal births. Outcomes for 256 newborns included 4 RT-PCR positive neonates, 2 stillbirths, and 1 neonatal death. No evidence of vertical transmission. Limited data suggest that pregnant women have a clinical presentation and severity similar to non-pregnant adults, and adverse maternal and perinatal outcomes were rare.	Moderate	Not reported
Yang, Z., Wang, M., Zhu, Z., & Liu, Y. (2020). Coronavirus disease 2019 (COVID-19) and pregnancy: a systematic review. The Journal of Maternal-Fetal & Neonatal Medicine. Epub ahead of print.	Apr 20, 2020 (Search completed Mar 26, 2020)	P: pregnant women and their fetuses/neonates E: COVID-19 infection C: no comparator O: maternal COVID-19 symptoms, pregnancy complications, neonatal COVID- 19 symptoms, neonatal health status, delivery timing and type, vertical transmission	Systematic review of 18 studies from around the world, published between 1 January 2020 and 26 March 2020, reporting on maternal, fetal, and neonatal outcomes of 114 pregnant women infected with (COVID-19). Study designs included case control, case reports, case series.	91% had cesarean delivery. In terms of fetal and neonatal outcomes, stillbirth (1.2%), neonatal death (1.2%), preterm birth (21.3%), low birth weight (<2500 g, 5.3%), fetal distress (10.7%), and neonatal asphyxia (1.2%) were reported. No direct evidence of intrauterine vertical transmission. The clinical characteristics of pregnant women with COVID-19 infection are similar to those of non-pregnant adults. Fetal and neonatal outcomes appear good in most cases. Limitation: Available data only include pregnant women infected in their third trimesters.	Moderate	Low

Della Gatta, A.N., Rizzo, R., Pilu, G., & Simonazzi, G. (2020). Coronavirus disease 2019 during pregnancy: A systematic review of reported cases. American Journal of Obstetrics and Gynecology, 223(1), 36–41.	Apr 17, 2020 (Search completed Mar 16, 2020)	P: pregnant women and their fetuses/neonates E: COVID-19 infection C: no comparator O: maternal COVID-19 symptoms, pregnancy complications, neonatal COVID- 19 symptoms, neonatal health status, delivery timing and type, vertical transmission	Systematic review of 6 studies from around the world, published prior to 16 March 2020, reporting on clinical outcomes for 51 pregnant women infected with COVID-19. Study designs included case series, case reports, and retrospective studies.	39% had preterm birth; 96% had cesarean delivery with unclear indications. No evidence of vertical transmission. 1 fetal death occurred in a critically ill patient. Clinical outcome has been generally favorable for both mothers and neonates.	Moderate	Low to moderate
Parazzini, F., Bortolus, R., Mauri, P.A., Favilli, A., Gerli, S. and Ferrazzi, E. (2020). Delivery in pregnant women infected with SARS-CoV-2: A fast review. International Journal of Gynecology & Obstetrics, 150(1), 41–46.	Apr 10, 2020 (Search completed Mar 31, 2020)	P: pregnant women and their fetuses/neonates E: COVID-19 infection C: no comparator O: maternal COVID-19 symptoms, pregnancy complications, neonatal COVID- 19 symptoms, neonatal health status, delivery timing and type, vertical transmission	Rapid review of 13 studies from around the world, published between 1 January to 31 March 2020, of clinical maternal characteristics, mode of delivery and neonatal outcomes for 64 pregnant women infected with COVID-19. Study designs included case reports and retrospective clinical series.	Pneumonia was present in 80.3% women, oxygen support was needed by 82.9%, and 6.5% were admitted to a critical care unit (among limited cases for which the information was available). 39.6% had preterm birth; 48.4% had caesarian delivery due to worsening of maternal conditions. In all reported cases the 5-minute Apgar score was greater than 7 and generally 9 or 10. Zero or low rate of vertical or peripartum transmission through cesarean delivery; no data available for vaginal delivery. Risk of transmission during breastfeeding is unknown. Overall, risk of adverse maternal and fetal outcomes is low.	Moderate	Not reported

Andersson, O. (2020). Maternal	Apr 6, 2020 (Search completed Apr 4, 2020)	P: pregnant women and their fetuses/neonates E: COVID-19 infection C: no comparator O: maternal COVID-19 symptoms, pregnancy complications, neonatal COVID- 19 symptoms, neonatal health status, delivery timing and type, vertical transmission	Systematic review of 18 studies from around the world, published between 8 December 2019 and 4 April 2020, of clinical manifestations and maternal and perinatal outcomes for 108 pregnant women with labconfirmed COVID-19 infection. Study designs included case reports and case	68% presented with a fever at admission. 91% had cesarean delivery. The majority of mothers were discharged without any major complications. However, severe maternal morbidity as a result of COVID-19 and perinatal deaths were reported, with 3% of women requiring admission to maternal ICU, and 1 case of perinatal death. No clear evidence for vertical transmission.	Moderate	Low
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Table 2: In-progress Syntheses

Title	Anticipated	Description of Document			
	Release Date				
Previously reported evidence					
Bahri, N., Dashti, S., Najafi, T.F., &	Mar 1, 2021	This review will assess the possibility of vertical transmission of COVID.			
Tohidinik, H.R. (2020). Assessment of the					
possibility of vertical transmission of					
COVID-19: a systematic review and meta- analysis protocol. <i>PROSPERO</i>					
CRD42020173886.					
Fen, L.Y., Lei, G.X., Han, N.C., Mei, Y.N.P.,	Dec 31, 2020	This review will address four questions: 1) What are the maternal and neonatal			
& Amin, Z. (2020). A systematic review of	Dec 31, 2020	serological results (i.e. IgM and IgG) in reported COVID-19 pregnancies; 2) What are the			
neonates and COVID-19. PROSPERO		immunological profiles of mothers and neonates in reported COVID-19 pregnancies; 3)			
CRD42020183500.		What is the effect of gestational age at maternal COVID-19 infection on neonatal			
		outcomes and 4) What are the neonatal outcomes of perinatally acquired infection			
		compared to postnatally acquired infection?			
Foratori-Junior, G.A., Mosquim, V., & de	Sep 24, 2020	This review will look to synthesize evidence on the symptoms of COVID in pregnant			
Carvalho Sales-Peres, S.H. (2020). COVID-		women and in their newborn children as well as the impacts on delivery and the			
19 and its relation with pregnancy and		possibility of vertical transmission.			
neonates: a systematic review.					
PROSPERO CRD42020177354.					
Ramos, M. (2020). COVID-19 in pregnant	May 31, 2020	The research question for this review is, "What evidence is available about COVID-19 in			
women: a systematic review. PROSPERO		pregnant women?" The primary outcomes are hospitalization, mortality, medical			
CRD42020179843.		complications and readmission and secondary outcomes include clinical management and delivery arrangements.			
Poon, L., Yang, H., del Mar Gil, M., &	May 25, 2020	The research question looks to answer, "What is the effect of SARS-CoV-2 infection			
Juan, J. (2020). Maternal, fetal and	Widy 23, 2020	(COVID-19 disease) in pregnancy, mother, fetus and newborn".			
neonatal characteristics and clinical		(COVID To alcoado) in programoy, mound, rotae and nowborn.			
outcomes in cases of COVID-19 infection					
during pregnancy: a systematic review.					
PROSPERO CRD42020181557.					
Novoa, R., Muñoz, W.Q., Melendez,	Apr 30, 2020	This review will look to answer two questions: 1) What are the clinical characteristics of			
P.A.L., & Laveriano, W.V. (2020). Maternal		COVD-19 infection in pregnant women; and 2) What are the perinatal outcomes of			
clinical characteristics and perinatal		COVID-19 infection in pregnant women?			
outcomes of pregnant women infected by					
coronavirus (COVID-19). A systematic					
review. PROSPERO CRD42020176534.					

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