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Coronavirus disease 2019 in pregnancy was associated with maternal morbidity and preterm birth

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BACKGROUND: Despite the mainly reassuring outcomes for pregnant women with coronavirus disease 2019 reported by previous case series with small sample sizes, some recent reports of severe maternal morbidity requiring intubation and of maternal deaths show the need for additional data about the impact of coronavirus disease 2019 on pregnancy outcomes.

OBJECTIVE: This study aimed to report the maternal characteristics and clinical outcomes of pregnant women with coronavirus disease 2019. **STUDY DESIGN:** This retrospective, single-center study includes all consecutive pregnant women with confirmed (laboratory-confirmed) or suspected (according to the Chinese management guideline [version 7.0]) coronavirus disease 2019, regardless of gestational age at diagnosis, admitted to the Strasbourg University Hospital (France) from March 1, 2020, to April 3, 2020. Maternal characteristics, laboratory and imaging findings, and maternal and neonatal outcomes were extracted from medical records.

RESULTS: The study includes 54 pregnant women with confirmed (n=38) and suspected (n=16) coronavirus disease 2019. Of these, 32 had an ongoing pregnancy, 1 had a miscarriage, and 21 had live births: 12 vaginal and 9 cesarean deliveries. Among the women who gave birth, preterm deliveries were medically indicated for their coronavirus disease

support was required for 13 of 54 women (24.1%), including high-flow oxygen (n=2), noninvasive (n=1) and invasive (n=3) mechanical ventilation, and extracorporeal membrane oxygenation (n=1). Of these, 3, aged 35 years or older with positive test result for severe acute respiratory syndrome coronavirus 2 using reverse transcription polymerase chain reaction, had respiratory failure requiring indicated delivery before 29 weeks' gestation. All 3 women were overweight or obese, and 2 had an additional comorbidity. **CONCLUSION:** Coronavirus disease 2019 in pregnancy was associ-

2019—related condition for 5 of 21 women (23.8%): 3 (14.3%) before 32

weeks' gestation and 2 (9.5%) before 28 weeks' gestation. Oxygen

ated with maternal morbidity and preterm birth. Its association with other well-known risk factors for severe maternal morbidity in pregnant women with no infection, including maternal age above 35 years, overweight, and obesity, suggests further studies are required to determine whether these risk factors are also associated with poorer maternal outcome in these women.

Key words: COVID-19 pneumonia, extracorporeal membrane oxygenation, intubation, maternal and neonatal outcomes, maternal morbidity, pregnancy, preterm birth, respiratory failure

Introduction

Since the beginning of the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) or coronavirus disease 2019 (COVID-19) outbreak, it has been argued that pregnant women are at increased risk of severe infection.^{1,2} Various experts and authorities rapidly published articles of advice, guidance, and information to aid professionals in caring for pregnant women with COVID-19.^{3–6} Most of the early reports, mainly from the epicenter of the pandemic in China,^{7–16} have suggested

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Click <u>Supplemental Materials</u> under article title in Contents at that pregnant women are not more severely affected than the general population.^{1,7–21} Case reports and limited case series of pregnant women from the United States,22 Iran,2,3 and South America²⁴ have contradicted these reassuring preliminary findings, however. They have instead described women with COVID-19 as patients who require invasive mechanical ventilation including extracorporeal membrane oxygenation (ECMO) and who are at risk of maternal death. These contradictory findings and the potential variation in disease severity between countries mean that additional information is urgently needed, especially from areas besides China, to determine whether or not pregnant women with COVID-19 are likely to experience severe cases with pneumonia.

We aim to present the clinical features and course of consecutive pregnant women with COVID-19 managed at a single French referral center, located in one of the main pandemic epicenters in France.

Methods Study design and patients

This single-center, retrospective study includes all consecutive pregnant women with COVID-19, regardless of gestational age at diagnosis, admitted to Strasbourg University Hospital (Strasbourg, France), a tertiary referral hospital, from March 1, 2020, to April 3, 2020. During the study period, only symptomatic women were tested using quantitative reverse transcription polymerase chain reaction (qRT-PCR) for SARS-CoV-2. Women with an ongoing pregnancy and a time from symptom onset to April 3 shorter than 14 days^{25,26} were excluded. COVID-19 was diagnosed according to the Chinese management guidelines (version 7.0),²⁶ which distinguishes confirmed cases, defined as all cases with respiratory tract





AJOG at a Glance

Why was this study conducted?

Although the available data about maternal outcomes for pregnant women with coronavirus disease 2019 (COVID-19) are reassuring, some limited case series report adverse maternal outcomes justifying further investigation. This study was conducted to report the maternal characteristics and clinical outcomes of these women.

Key findings

Among the pregnant women with COVID-19, 13 of 54 (24.1%) received oxygen support, with 3 of 54 (5.5%) requiring invasive mechanical ventilation. Among the 21 women who gave birth, 5 of 21 (23.8%) had deliveries—3 of 21 (14.3%) very preterm—indicated for severe maternal conditions related to their COVID-19 condition.

What does this add to what is known?

In contrast to most existing data, this study shows that COVID-19 in pregnancy is associated with maternal morbidity and preterm birth.

samples that test positive for SARS-CoV-2 using qRT-PCR, from suspected cases, which include an epidemiologic history (Strasbourg is a community with numerous COVID-19 cases) and at least 2 of the following clinical manifestations: (1) fever and/or respiratory syndrome, (2) imaging features of COVID-19 pneumonia, and (3) a decreased or normal total number of leukocytes and lymphocyte counts in the early stage of the disease.²⁶ The imaging features of COVID-19 pneumonia used were typical radiographic features on chest computed tomography (CT), owing to their high diagnostic sensitivity and accuracy for this diagnosis.^{26–28}

The Ethics Committee of Strasbourg University Hospital approved this study under a waiver of informed consent (reference number identification 2020: Hospital University of Strasbourg N7794).

Data collection

Maternal characteristics, laboratory findings, CT scans, management, and maternal and fetal outcomes were obtained from patients' medical records.

All laboratory tests were performed at the Strasbourg University Hospital, including the laboratory confirmation of COVID-19 using qRT-PCR to tests for SARS-CoV-2 nucleic acid on nasopharyngeal swabs.²⁹ Primer and probe sequences target 2 regions on the RdRp gene and are specific to SARS-CoV-2. Assay sensitivity is around 10 copies per reaction. For women with multiple RT-PCR assays, the diagnosis of COVID-19 was confirmed when any 1 nucleic acid test result was positive. Typical radiographic features of COVID-19 on chest CT were consolidation, ground-glass opacity, or bilateral pulmonary infiltration.²⁷ Neonatal throat and rectal swab samples were also collected to test for SARS-CoV-2 using RT-PCR assays at birth or on day 1 of life, on day 3 of life in term newborns, and on days 7 and 14 of life in preterm neonates. Skin-to-skin contact between mothers and neonates and breastfeeding were both permitted at birth.

Definitions

Fever was defined as an axillary temperature of at least 37.8°C (100°F). The severity of COVID-19 was defined according to the Chinese management guidelines (version 7.0) as mild (mild clinical symptoms and no sign of pneumonia), moderate (patients have fever and respiratory symptoms; radiologic assessments found signs of pneumonia), severe (adults meeting any of the following criteria: shortness of breath, respiratory rate \geq 30 breaths/min, oxygen saturation \leq 93% at rest, or alveolar oxygen partial pressure (PaO₂)/fraction of inspiration oxygen (FIO₂) \leq 300 mm Hg), and critical (≥ 1 of any mechanical ventilation, shock, or organ failure that needs intensive care unit [ICU] monitoring and treatment).²⁶ Acute respiratory distress syndrome (ARDS) was diagnosed according to the Berlin definition in 3 mutually exclusive categories based on degree of hypoxemia: mild (200 mm Hg<PaO₂/FIO₂ \leq 300 mm Hg), moderate (100 mm Hg<PaO₂/ FIO₂ \leq 200 mm Hg), and severe (PaO₂/ FIO₂ \leq 100 mm Hg).³⁰

Statistical analysis

Descriptive characteristics were calculated for the variables of interest. Statistical analysis included determination of rates with their 95% confidence intervals, performed with R software (R Foundation for Statistical Computing, Vienna, Austria).

Results

The study included 54 symptomatic pregnant women with COVID-19 during the study period; 38 had positive RT-PCR results for SARS-CoV-2, whereas 16 had negative RT-PCR results for SARS-CoV-2 but met the Chinese management guideline criteria for suspected COVID-19 (version 7.0).³¹ Among these 54 women, 32 had an ongoing pregnancy more than 14 days after symptom onset, 1 had a miscarriage, and 21 had live births by vaginal or cesarean delivery (Figure).

Maternal clinical characteristics

Table 1 summarizes maternal clinical characteristics. Their ages ranged from 19 to 42 years. Ten women (18.5%) had at least 1 comorbidity before pregnancy: 4 had asthma and 1 preexisting chronic hypertension, 4 were obese, and 4 had another comorbidity. No woman had a multiple pregnancy, and only 15 (27.3%) were nulliparous. The mean gestational age at diagnosis of COVID-19 was 30.4 ± 9.0 weeks, whereas the mean time from symptom onset to hospital admission was 3.5 ± 3.2 days. Fourteen (25.9%) had fever, 36 (66.7%) cough, 22 (40.7%) shortness of breath, 9 (16.7%) digestive disorders, and 20 (37.0%) anosmia or ageusia. Chest CT was performed on 12 women (22.2%) and showed typical features of COVID-19 for 10 (83.3%).

Laboratory tests

Data from laboratory tests showed that 24 women (44.4%) had lymphopenia; C-reactive protein was elevated in 22 (40.7%) and aspartate and/or alanine aminotransferases in 11 (20.4%). None had thrombocytopenia (Table 2).

Clinical course of pregnancy

Thirteen women (24.1%) received oxygen support. Two of 13 (15.4%) required high-flow oxygen, one after delivery and the other before delivery. The latter subsequently required noninvasive mechanical ventilation and has not yet given birth. In addition, 3 of the 13 women (23.1%) who received oxygen subsequently had invasive mechanical ventilation after delivery. One woman (1.9%) later required ECMO.

Three women (5.6%) received antiviral therapy (lopinavir or ritonavir 400 mg/100 mg tablet twice a day for 5 days) (Table 3).

Among the 21 women who gave birth by April 17, 2020, 9 (42.9%) had cesarean deliveries, and 12 (57.1%) had vaginal deliveries. The mean gestational age at delivery was 37.4 ± 4.7 weeks, but 5 women (23.8%) gave birth <37 weeks' gestation, 3 (14.3%) <32 weeks' gestation, and 2 (9.5%) <28 weeks' gestation. These 5 premature births were medically indicated for severe maternal conditions related to COVID-19, and 2 women were transferred to the ICU after delivery. Five women (9.3%) had postpartum hemorrhages, 4 (7.4%) required a blood transfusion, and 1 (1.9%) had arterial embolization. Five women were (9.3%) admitted to the ICU, all owing to COVID-19 respiratory symptoms. No thromboembolic event nor any other significant maternal morbidity occurred (Table 4).

Neither neonatal acidosis nor fetal or neonatal death was observed. Three neonates, 2 born at 27 weeks' gestation and 1 at 28 weeks' gestation, underwent endotracheal intubation in the labor ward and were admitted to the neonatal ICU for problems related to their extremely or very preterm birth. All 21 neonates were tested for SARS-CoV-2 using RT-PCR of throat and rectal swab

FIGURE Flowchart of the study population



* According to the Chinese management guideline for COVID-19 (version 7.0)³²

The severity of COVID-19 was defined according to the Chinese management guideline for COVID-19 (version 7.0). Adapted from China National Health Commision.²⁶

COVID-19, coronavirus disease 2019.

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specimens, and SARS-CoV-2 was never detected (Table 5).

By the end of the follow-up (April 17, 2020), 1 woman (woman 2; Supplemental Table) remained in the ICU, 27 days after her transfer there. She had a secondary infection by Klebsiella pneumoniae and has required ECMO since April 9, 2020. Another woman (woman 13; Supplemental Table) remained hospitalized with an ongoing pregnancy but was transferred back to conventional care after 6 days in the ICU. She required high-flow oxygen therapy and transient noninvasive mechanical ventilation. All the other women have been discharged from the hospital. Eighteen neonates were also discharged, healthy and with no respiratory symptoms or fever. The remaining 3 newborns were still hospitalized for care related to their prematurity.

Clinical outcomes of the 13 women who required oxygen support

The characteristics and clinical course of the 13 pregnant women (24.1%) with COVID-19 who required oxygen support are described in the Supplemental Table. All but 3 had lymphopenia ($<1.0 \times 10^9$ cells per L) or elevated concentrations of C-reactive protein (>10 mg/L). Four of them were still pregnant, whereas 9 had cesarean deliveries. Among the latter, 3 women aged \geq 35 years with positive RT-PCR results for SARS-CoV-2 had medically indicated deliveries before 29 weeks' gestation owing to severe maternal respiratory symptoms. Before delivery, all 3 women (women 1, 2, and 9) had respiratory failure even though they were receiving 5 liters of oxygen per minute. Preterm delivery was decided because physicians thought that maternal respiratory status would improve and oxygenation would be facilitated after birth.

One of these women (woman 2) needed invasive mechanical ventilation and then ECMO after delivery. All had at least 1 comorbidity: class III obesity (woman 1), overweight and gestational hypertension (woman 2), and overweight and gestational diabetes mellitus (woman 9).

Comment

Principal findings of the study

This is a descriptive study of the clinical characteristics and outcomes of 54

Women's characteristics at admission

Characteristics	All women (N=54)	Pregnant women with confirmed COVID-19 (n=38)	Pregnant women with suspected COVID-19 (n=16
Age, y	30.6±6.2	31.1±6.4	29.6±5.9
Age >35 y, n (%)	15 (27.8)	12 (31.6)	3 (18.8)
Geographic origin, n (%)			
Europe	34 (63.0)	22 (57.9)	12 (75.0)
Sub-Saharan Africa	10 (18.5)	10 (26.3)	0
North Africa	10 (18.5)	6 (15.8)	4 (25.0)
Asia	0	0	0
Non-French nationality, n (%)	21 (38.9)	18 (47.4)	3 (18.8)
BMI before pregnancy	25.3±4.7	25.6±5.1	24.2±3.3
BMI >25 kg/m², n (%)	20 (37.0)	16 (42.1)	4 (25)
BMI >30 kg/m ² , n (%)	4 (7.4)	4 (10.5)	0
Preexisting chronic hypertension, n (%)	1 (1.9)	0	1 (6.25)
Asthma, n (%)	5 (9.3)	1 (2.6)	4 (25.0)
Preexisting diabetes mellitus, n (%)	0	0	0
Other preexisting chronic disease, n (%)	4 (7.4)	3 (7.9)	1 (6.25)
Nulliparous, n (%)	15 (27.3)	9 (23.7)	6 (37.5)
Singleton pregnancy, n (%)	54 (100)	38 (100)	16 (100)
Previous cesarean delivery, n (%)	9 (16.7)	6 (15.8)	3 (18.8)
History of postpartum hemorrhage, n (%)	2 (3.7)	2 (5.3)	0
History of preterm delivery	0	0	0
Tobacco use during pregnancy, n (%)	1 (1.9)	0	1 (6.3)
Gestational diabetes, n (%)	4 (7.4)	4 (10.5)	0
Gestational hypertensive disorders, n (%)	2 (3.7)	1 (2.6)	1 (6.3)
Gestational age at the diagnosis of COVID-19, wk	30.4±9.0	29.3±8.5	31.8±6.3
Contact history in epidemic area, n (%)	54 (100)	38 (100)	16 (100)
Other family members affected, n (%)	20 (37.0)	16 (42.1)	4 (25.0)
Clinical manifestations			
Fever, n (%)	14 (25.9)	10 (26.3)	4 (25.0)
Cough, n (%)	36 (66.7)	25 (65.8)	11 (68.8)
Shortness of breath, n (%)	22 (40.7)	13 (34.2)	9 (56.3)
Diarrhea, n (%)	9 (16.7)	7 (18.4)	2 (12.5)
Fatigue, n (%)	54 (100)	38 (100)	16 (100)
Sore throat, n (%)	23 (42.6)	16 (42.1)	7 (43.8)
Anosmia or ageusia, n (%)	20 (37.0)	18 (47.4)	2 (12.5)
Acute respiratory distress syndrome, n (%)	1 (1.9)	1 (2.6)	0
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pregnant women with either laboratory-confirmed (n=38) or suspected (n=16) COVID-19; during our study, 32 pregnancies are ongoing, and

22 were completed. Oxygen support was required for 13 of 54 pregnant women (24.1%) with COVID-19 including 1 of 54 (1.9%) who required ECMO, and 3 of 21 births (14.3%) were very preterm deliveries indicated for severe COVID-19—related maternal conditions. The latter 3 women were

Women's characteristics at admission (continued)

All women	Pregnant women with	Pregnant women with
(N=54)	confirmed COVID-19 (n=38)	suspected COVID-19 (n=16)
37 (68.5)	27 (71.1)	10 (62.5)
11 (20.4)	8 (21.1)	3 (18.8)
6 (11.1)	3 (7.9)	3 (18.8)
3.5±3.2	3.6±2.8	4.0±4.2
12 (22.2)	7 (18.4)	5 (21.3)
10/12 (83.3)	7 (18.4)	3 (18.8)
7/12 (58.3)	5 (13.2)	2 (12.5)
5/12 (41.6)	4 (10.5)	1 (6.3)
1/12 (8.3)	1 (2.6)	0
• •	37 (68.5) 11 (20.4) 6 (11.1) 3.5±3.2 12 (22.2) 10/12 (83.3) 7/12 (58.3) 5/12 (41.6)	(N=54) confirmed COVID-19 (n=38) 37 (68.5) 27 (71.1) 11 (20.4) 8 (21.1) 6 (11.1) 3 (7.9) 3.5 ± 3.2 3.6 ± 2.8 12 (22.2) 7 (18.4) 10/12 (83.3) 7 (18.4) 7/12 (58.3) 5 (13.2) 5/12 (41.6) 4 (10.5)

Data are presented as mean \pm SD, unless otherwise indicated. Body mass index is weight in kilograms divided by the square of the height in meters. Other preexisting chronic disease means disease other than body mass index >25 kg/m² or preexisting chronic hypertension, diabetes mellitus, and asthma. The data include 1 woman with sickle cell anemia, 1 with chronic alcohol abuse, 1 with chronic hepatitis B, and 1 with nephropathy. Contact history of epidemic area means exposure to the relevant environment (Alsace region) or contact with a person with infection.

BMI, body mass index; COVID-19, coronavirus disease 2019; SD, standard deviation.

^a The severity of COVID-19 was defined according to the Chinese management guideline for COVID-19 (version 7.0).²⁶

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aged 35 years or older, and all had at least 1 comorbidity.

Results of the study in the context of what is known

Most of the data about the maternal, fetal, and neonatal outcomes of pregnant women with COVID-19 have been globally reassuring.^{7–21} Despite the unavailability of numerous details about maternal characteristics, including body mass index (BMI) and preexisting disease, or about the third stage of labor,^{7,11,14} the authors considered that the women's outcomes were "good"¹³ or "satisfactory"¹¹ or that there is "no evidence to suggest that COVID-19 pneumonia causes severe maternal and neonatal complications."¹⁴

Zaigham and Andersson²⁰ summarized these reports in a systematic review up to April 1, 2020, which included 108 pregnant women with COVID-19 from 18 case reports or series, mostly from China. They underlined that 3 of 108 women (3%) were admitted to ICUs and noted that severe maternal morbidity could therefore not be ruled out in COVID-19 during pregnancy,²⁰ whereas Della Gatta et al²¹ pointed out in a systematic review of 6 Chinese studies totaling 51 pregnant women^{7,9,10} that preterm—mainly indicated—deliveries (31%) were frequent in women with COVID-19.

Finally, on April 17, 2020, Chen et al¹⁵ published a case series of 118 pregnant women with COVID-19 from 50 hospitals in Wuhan, China. It reported 3 spontaneous abortions, 2 ectopic pregnancies, 4 induced abortions, 41 ongoing pregnancies, and 68 deliveries; 63 of them cesarean deliveries (93%) because of the concern about the effect of COVID-19 on the pregnancy. No neonatal asphyxia or death occurred. Among the 14 preterm deliveries (21%), half (10.5%) were related to COVID-19. Overall, 109 women (92%) had mild disease, and 9 (8%) had severe disease (hypoxemia); only 1 (0.8%) received noninvasive mechanical ventilation. The only maternal characteristics described were age, parity, and singleton or multiple pregnancy, with details about comorbidities, in no particular BMI, preexisting conditions, or pregnancy-related disease. The authors concluded that their results "do not suggest an increased risk of severe disease among pregnant women."15

Nevertheless, other authors have previously reported severe maternal complications related to COVID-19 in pregnant women, consistent with our findings.^{10,18,19} Liu et al¹⁰ retrospectively reported the clinical outcome of 13 pregnant women with COVID-19, including 3 ongoing pregnancies. Details were scarce, but the authors stated that 12 pregnant women were discharged without obvious complications. The remaining woman was admitted to the ICU with multiple organ dysfunction syndrome including ARDS, necessitating an emergency cesarean delivery. The neonate died. The woman required intubation, mechanical ventilation, and finally ECMO, but her final outcome was not reported.¹⁰

In a retrospective case series of 43 pregnant women with PCR-confirmed COVID-19 from 2 American centers, most of the women were obese, and 18 had an additional comorbidity.¹⁹ Among these 43 women, 18 gave birth, 10 by uncomplicated vaginal deliveries and 8 by cesarean deliveries for obstetrical reasons unrelated to COVID-19. Nonetheless, 2 women (4.7%) were admitted to the ICU.^{18,19} The first was a 38-yearold woman with a BMI of 38 kg/m², diabetes mellitus, and intrahepatic cholestasis of pregnancy. Severe

TABLE 2 Clinical laboratory results

Laboratory characteristics	Reference range	All women (N=54)	Pregnant women with confirmed COVID-19 (n=38) ^a	Pregnant women with suspected COVID-19 (n=16) ^a
White blood cell count ($\times 10^9$ cells per L)	4.02-11.42	8.9±3.7	8.4±3.7	10.2±3.7
Lymphocyte count ($\times 10^{9}$ cells per L)	1.24-3.56	1.6±0.7	1.5±0.7	1.7±0.7
Lymphopenia ($<$ 1.5 \times 10° cells per L), n (%)	N/A	24 (44.4)	17 (44.7)	7 (43.8)
C-reactive protein concentration (mg/L)	<5	25.7±36.9	26.4±37.6	7.5±36.9
Elevated C-reactive protein (>10 mg/L), n (%)	N/A	22 (40.7)	15 (41.7)	6 (37.5)
Elevated ALT (>45 U/L) or AST (>35 U/L), n (%)	N/A	11 (20.4)	10 (27.0)	1 (6.3)
ALT (U/L)	6—55	26.6±17.1	29.1±19.0	19.9±8.4
AST (U/L)	5—34	25.7±20.4	27.4±19.7	21.5±22.3
Hemoglobin (g/dL)	11.5—14.9	11.2±1.3	11.1±1.3	11.2±1.4
Hematocrit (%)	34.4-43.9	36.4±10.8	34.3±4.0	41.2±18.2
Thrombocytopenia <100,000/dL, n (%)	N/A	0	0	0
Blood urea nitrogen (mmol/L)	2.5-6.7	2.7±1.0	2.5±1.0	3.0±1.1
Creatinine (µmol/L)	49—90	48.4±10.2	47.3±8.3	50.9±13.7
Prothrombin ratio <80%, n (%)	N/A	2 (3.7)	1 (2.7)	1 (6.3)
Prolonged aPTT, n (%)	>1.19	6 (11.1)	5 (13.2)	1 (6.3)

Data are presented as mean \pm SD, unless otherwise indicated.

AST, aspartate aminotransferase; ALT, alanine aminotransferase; aPTT, activated partial thromboplastin time; COVID-19, coronavirus disease 2019; N/A, not applicable; SARS-CoV-2, severe acute respiratory syndrome coronavirus 2; SD, standard deviation.

^a COVID-19 was diagnosed according to the Chinese management guidelines (version 7.0),²⁶ which distinguishes confirmed cases, defined as all cases with respiratory tract samples that tested positive for SARS-CoV-2 using quantitative reverse transcription polymerase chain reaction (qRT-PCR), from suspected cases, which include an epidemiologic history (Strasbourg is a community with numerous COVID-19 cases) and at least 2 of the following clinical manifestations: (1) fever and/or respiratory syndrome, (2) imaging features of COVID-19 pneumonia, and (3) a decreased or normal total number of leukocytes and lymphocyte counts in the early stage of the disease.²⁶

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postpartum hemorrhage related to uterine atony occurred during cesarean delivery that followed failed induction of labor. Because she had fever and respiratory symptoms during delivery, a chest CT was performed and revealed groundglass opacities in the lungs. RT-PCR was subsequently positive for SARS-CoV-2. She spent only 8 hours in the ICU and was discharged 4 days later.¹⁸ The second woman admitted to the ICU was 33 years old, with a BMI of 47 kg/m², chronic hypertension, type 2 diabetes mellitus, and mild intermittent asthma. Twentyfive hours after cesarean delivery for failed induction of labor, she had coughing, respiratory distress, high fever, and reduced oxygen saturation. Although her respiratory status improved, she was admitted to the ICU after developing severe hypertension (blood pressure reaching 200/90 mm

Hg). RT-PCR results returned positive for SARS-CoV-2. After 5 days, when the report concluded, she remained hospitalized, had acute kidney injury, and still needed oxygen supplementation.¹⁸ In both cases, COVID-19 seems to have been the—or at least the only—reason for ICU admission, as the first case involved severe postpartum hemorrhage and the second case involved severe hypertension.

Moreover, Yan et al¹⁶ reported a retrospective series expanded from 4 previous small case series,^{7,9,10} totaling 116 pregnant women with COVID-19 pneumonia from 25 hospitals in China. Preterm delivery before 34 and 37 weeks' gestation occurred in 2% and 21.2% of cases: 8 of 116 women (6.9%) were admitted to the ICU, 6 of 116 (5.2%) required noninvasive and 2 of 116 (1.7%) invasive ventilation, and 1 of 116

(0.9%) required ECMO. Although these results were consistent with ours, the authors surprisingly concluded that "there is no evidence that pregnant women with COVID-19 are more prone to develop severe pneumonia, in comparison to nonpregnant patients."¹⁶

Interestingly, since April 22, 2020, 3 different teams from distinct parts of the world have also alerted healthcare professionals about the possible detrimental effect of COVID-19 among pregnant women. Hirshberg et al²² described the clinical course of 5 American women, all requiring mechanical ventilation, including ECMO for one. All had at least 1 of the following comorbidities: overweight or obesity, chronic kidney disease, hypertension, insulin-dependent diabetes, obstructive sleep apnea, and mild asthma.²² Finally, maternal deaths have

Management of COVID-19 pneumonia

	All women (N=54)	Pregnant women with confirmed COVID-19 (n=38)	Pregnant women with suspected COVID-19 (n=16)
Management of COVID-19 pneumonia (before and aft	er delivery)		
Oxygen support (nasal cannula), n (%)	13 (24.1)	10 (26.3)	3 (18.8)
High-flow oxygen, n (%)	2 (3.7)	1 (2.6)	1 (6.3)
Noninvasive mechanical ventilation, n (%)	1 (1.9)	1 (2.6)	0
Invasive mechanical ventilation, n (%)	3 (5.6)	1 (2.6)	2 (12.5)
Extracorporeal membrane oxygenation, n (%)	1 (2.6)	1 (2.6)	0
Antiviral therapy, n (%)	3 (5.6)	3 (7.9)	0
Antibiotic therapy, n (%)	4 (7.4)	4 (10.5)	0
Corticosteroid, n (%)	0	0	0
Hydroxychloroquine or chloroquine, n (%)	0	0	0
Management of COVID-19 pneumonia before delivery			
Oxygen support (nasal cannula), n (%)	13 (24.1)	10 (13.2)	3 (18.8)
High-flow oxygen, n (%)	1 (1.9)	0	1 (6.3)
Noninvasive mechanical ventilation, n (%)	1 (1.9)	0	1 (6.3)
Invasive mechanical ventilation, n (%)	0	0	0
Extracorporeal membrane oxygenation, n (%)	0	0	0
Antiviral therapy, n (%)	0	0	0
Antibiotic therapy, n (%)	2 (3.7)	2 (5.3)	0
Corticosteroid, n (%)	0	0	0
Hydroxychloroquine or chloroquine, n (%)	0	0	0
Management of COVID-19 pneumonia after delivery			
Oxygen support (nasal cannula), n (%)	0	0	0
High-flow oxygen, n (%)	1 (1.9)	1 (6.3)	0
Noninvasive mechanical ventilation, n (%)	0	0	0
Invasive mechanical ventilation, n (%)	3 (5.6)	1 (2.6)	2 (12.5)
Extracorporeal membrane oxygenation, n (%)	1 (2.6)	1 (2.6)	0
Antiviral therapy, n (%)	3 (5.6)	3 (7.9)	0
Antibiotic therapy, n (%)	2 (3.7)	2 (5.3)	0
Corticosteroid, n (%)	0	0	0
Hydroxychloroquine or chloroquine, n (%)	0	0	0

Five women had more than 1 oxygen support procedure: 1 (woman 2; Supplemental Table) had invasive mechanical ventilation and then extracorporeal membrane oxygenation; 2 (women 3 and 4; Supplemental Table) had invasive mechanical ventilation; 1 (woman 7; Supplemental Table) received high-flow oxygen. The last one (woman 13; Supplemental Table) received high-flow oxygen and then required noninvasive mechanical ventilation.

COVID-19, coronavirus disease 2019.

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been reported in pregnant women with COVID-19.^{23,24} Amorim et al²⁴ reported the cases of 9 women; the details were unfortunately limited, as the sources were mainly local media or the Brazilian or Mexican ministries of

health. An Iranian case series reported the clinical outcomes of 9 pregnant women with severe COVID-19. Seven of them died, 3 aged 35 years or older.²³ Nevertheless, no details were provided about any comorbidities related to these maternal deaths, including BMI.²³ Thus, because of the considerable methodological limitations of the latter 2 reports, their principal information is that maternal deaths do occur in pregnant women with COVID-19.^{23,24}

Maternal and neonatal outcomes of the pregnant women with confirmed or suspected COVID-19

Characteristics	All women (N=54)	Pregnant women with confirmed COVID-19 (n=38)	Pregnant women with suspected COVID-19 (n=16)
Miscarriage, n (%)	1 (1.9)	1 (2.6)	0
Abortion, n (%)	0	0	0
Termination of pregnancy, n (%)	0	0	0
Ongoing pregnancy, n (%)	32 (59.3)	20 (52.6)	12 (75.0)
Preeclampsia, n (%)	3 (5.6)	2 (5.3)	1 (6.3)
Fetal growth restriction, n (%)	1 (1.9)	1 (2.6)	0
Preterm premature rupture of membranes, n (%)	0	0	0
Stillbirth, n (%)	0	0	0
Gestational age at delivery (n=21)	37.4±4.7	36.9±5.2	39.0±0.8
Delivery before 37 weeks' gestation, n/N (%)	5/21 (23.8)	5/17 (29.4)	0
Delivery before 32 weeks' gestation, n/N (%)	3/21 (14.3)	3/17 (17.6)	0
Delivery before 28 weeks' gestation, n/N (%)	2/21 (9.5)	2/17 (11.8)	0
Cesarean delivery, n/N (%)	9/21 (42.9)	7/17 (41.1)	2/4 (50)
Cesarean delivery related to COVID-19, n/N (%)	7/21 (33.3)	6/17 (35.3)	1/4 (25)
Prelabor cesarean delivery, n/N (%)	8/21 (38.1)	6/17 (35.3)	2/4 (50)
Cesarean delivery during labor, n/N (%)	1/21 (4.8)	1/17 (5.9)	0
Vaginal delivery, n/N (%)	12/21 (57.1)	10/17 (58.8)	2/4 (50)
Operative vaginal delivery, n/N (%)	2/21 (9.5)	1/17 (5.9)	1/4 (25)
Spontaneous vaginal delivery, n/N (%)	10/21 (47.6)	9/17 (52.9)	1/4 (25)
Third- or fourth-degree perineal lacerations, n/N (%)	1/21 (4.8)	0	1/4 (25)
PPH, defined by blood loss ${\geq}500$ mL, measured with a graduated collector bag, n/N (%)	5/21 (23.8)	4/17 (23.5)	1/4 (25)
Estimated total blood loss (n=21)	1060±182	1100±183	900
Additional uterotonics for excessive bleeding, n/N (%)	4/21 (19.0)	4/17 (23.5)	0
Blood transfusion, n/N (%)	4/21 (19.0)	4/17 (23.5)	0
Arterial embolization or surgery for PPH, n/N (%)	1/21 (4.8)	1/17 (5.9)	0
Acute respiratory distress syndrome, n/N (%) ^a	1/21 (4.8)	0/17 (0)	0
Thromboembolic events, n (%)	0	0	0
Kidney failure, n (%)	0	0	0
Admission to ICU, n (%)	5 (9.3)	3 (7.9)	2 (12.5)
ICU length of stay (d), median (IQR) (n=5)	9.2 (1.0-6.5)	13.0 (1.5—13.0)	4.0 (1.0-4.5)
Total hospital length of stay (d), median (IQR)	4.0 (3-5.5) (n=30)	3.0 (3.0–6.0) (n=19)	4.0 (4.0-4.8) (n=11)

Data are presented as mean±SD, unless otherwise indicated. Denominator is the number of pregnancies ended beyond the first trimester (n=21). In all preterm births, medically indicated premature delivery was decided because of severe maternal respiratory symptoms related to COVID-19.

COVID-19, coronavirus disease 2019; ICU, intensive care unit; IQR, interquartile range; PPH, postpartum hemorrhage; SD, standard deviation.

^a Acute respiratory distress syndrome was diagnosed according to the Berlin definition (Acute Respiratory Distress Syndrome: The Berlin Definition).³⁰

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NII women (N=54)	5	Pregnant women with
		suspected COVID-19 (n=16)
800±899	2810±982	2913±364
2/21 (9.5)	1/17 (5.9)	1/4
0	0	0
3/21 (14.3)	3/17 (17.6)	0
9/21 (14.3)	3/17 (17.6)	0
3/21 (14.3)	3/17 (17.6)	0
/21 (4.8)	1/17 (5.9)	0
/21 (4.8)	1/17 (5.9)	0
0	0	0
0	0	0
	/21 (9.5) 0 /21 (14.3) /21 (14.3) /21 (14.3) /21 (4.8) /21 (4.8) 0	/21 (9.5) 1/17 (5.9) 0 0 /21 (14.3) 3/17 (17.6) /21 (14.3) 3/17 (17.6) /21 (14.3) 3/17 (17.6) /21 (14.3) 3/17 (17.6) /21 (14.3) 1/17 (5.9) /21 (4.8) 1/17 (5.9) 0 0

Data are presented as mean±SD, unless otherwise indicated. Denominator was the number of pregnancies ended beyond the first trimester (n=21). Small for gestational age was defined as birthweight less than the 10th percentile for gestational age and sex based on the French intrauterine growth curve.

COVID-19, coronavirus disease 2019; CPAP, continuous positive airway pressure; ICU, intensive care unit; SARS-CoV-2, severe acute respiratory syndrome coronavirus 2; SD, standard deviation. Sentilhes et al. Coronavirus disease 2019 in pregnancy can cause severe maternal morbidity. Am J Obstet Gynecol 2020.

Clinical and research implications

Our case series confirms that COVID-19 during pregnancy may be responsible for severe maternal morbidity that may require very or extremely preterm elective delivery. In our series, 3 of 54 women (5.5%) had such medically indicated premature births; 13 of 54 (24.1%) required oxygen support, including invasive mechanical ventilation or ECMO for 3 of 54 women (5.5%); and 5 of 54 (9.2%) were admitted to the ICU. Interestingly, except for 1 of 2 women who required ECMO and for whom very few details were available,¹¹ the most severe cases reported came from countries outside China, that is, from the United States,¹⁸ France (our series), South America,²⁴ and Iran,²³ where maternal characteristics and the prevalence of preexisting conditions may differ from those in China. Most of the French and American women had >1 maternal characteristics that are known to be associated with severe maternal morbidity in pregnant women, including an age of \geq 35 years, overweight or obesity, gestational hypertension or diabetes, and preexisting asthma.³² These new findings should alert physicians and help them to manage and follow up pregnant women with COVID-19.

Another interesting finding of our study is the high rate of postpartum hemorrhage (9.3%) and blood transfusions (7.4%) observed in pregnant women with COVID-19, much higher than that reported in the general population of pregnant women.³¹ The literature thus far unfortunately fails to detail these women's third stage of labor.⁷⁻²¹ Despite the overall low numbers and the single limited experience they come from, our findings warrant further investigation to examine whether COVID-19 might increase the risk of blood loss after delivery. Hemostasis changes may occur in these women, especially as clinically significant coagulopathy with antiphospholipid antibodies has been reported in nonpregnant patients with COVID-19 pneumonia.33

Finally, all the neonates we tested were negative for SARS-CoV-2, consistent with previous reports⁷⁻²¹; however, no conclusion can be drawn from our results about the possibility of intrauterine vertical transmission.

Strengths and limitations of the study

Our retrospective study is one of the largest series of pregnant women with confirmed and suspected cases of COVID-19, and it provides the most detailed look at maternal characteristics and maternal outcomes. It is one of the first to suggest that COVID-19 may be responsible for severe maternal morbidity that can require medically indicated extremely preterm delivery. Nevertheless, several limitations of our study must be underlined. The first is its retrospective design, common to all the studies that have thus far assessed maternal outcome in pregnant women with COVID-19. All the flaws of retrospective analyses apply. Moreover, the reported rates should be interpreted cautiously; an overestimation of severe cases is possible as Strasbourg University Hospital is the referral center for the management of pregnant women with COVID-19 in the Alsace region. Moreover, generalizability is limited to regions with COVID-19 prevalence similar to that in the region where our hospital is located, which is one of the principal pandemic hot spots in France (along with the Paris metropolitan region). Moreover, similar to authors,^{14,15} previous we have included suspected cases with typical chest CT findings but with RT-PCR testing negative for SARS-CoV-2, in the Chinese accordance with

management guideline (version 7.0).²⁶ Given the frequency of false negatives for COVID-19 cases, owing to low virus titers, inappropriate swabbing sites, and sampling at late disease stages,³⁴ as well as the limited testing capacity in France during our study period (which also explains why only symptomatic women were tested during the study period), limiting inclusion to laboratory-confirmed cases would likely result in missing some COVID-19 cases.¹⁴ Nonetheless, all 5 premature women with births, including the 3 with the most severe illness in our study, those who required indicated extremely or very preterm cesarean delivery, had positive RT-PCR findings for SARS-CoV-2. Finally, even though our series is one of the largest so far reported, the small sample size should be taken into account when interpreting the results.

Conclusions

COVID-19 may be associated with severe maternal outcomes with hypoxemic respiratory failure despite oxygen support and can require an indicated-and sometimes very or extremely pretermdelivery. Healthcare providers should be aware that many of the standard risk factors associated with severe maternal morbidity without COVID-19, such as maternal age above 35 years, overweight or obesity, and preexisting and/or gestational hypertension or diabetes, may also increase the risk of severe maternal morbidity of pregnant women with COVID-19.

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SUPPLEMENTAL TABLE Clinical characteristics and outcomes of the 13 women with confirmed or suspected COVID-19 who required oxygen support

	Woman 1	Woman 2	Woman 3	Woman 4	Woman 5	Woman 6	Woman 7	Woman 8	Woman 9	Woman 10	Woman 11	Woman 12	Woman 13	
Characteristics	Women w	ho had vag	inal or cesa	arean delive	ery					Ongoing pre	gnancies			n/N (%)
Baseline characteristics										_				
Date of admission	March 16, 2020	March 16, 2020	March 20, 2020	March 20, 2020	March 20, 2020	March 25, 2020	March 26, 2020	March 27, 2020	March 28, 2020	March 19, 2020	March 26, 2020	March 29, 2020	April 1, 2020	
Age, y	36	42	32	33	41	24	42	36	36	34	36	32	21	34.2±6.3
Gestational age at admission, wk+d	27+2	26+5	37+2	38+1	40+3	39+2	35+0	33+5	28+2	34+0	34+5	36+0	25+0	32.5±5.1
Non-French nationality	No	Yes	No	No	Yes	No	No	No	Yes	Yes	No	No	Yes	5/13 (38.5)
Body mass index before pregnancy	42	25.7	26.2	24.2	29.4	19.0	25.0	31.1	26.2	24.2	27.5	24.2	27.0	27.1±5.3
Preexisting chronic basic disease	No	No	No	Asthma	No	No	No	No	No	Chronic hepatitis B	No	No	No	2/13 (15.4)
Primiparous	No	Yes	No	No	Yes	No	No	No	No	No	Yes	No	No	3/13 (23.1)
Gestational complication before admission	No	PIH	No	No	No	Preeclampsia	No	No	GDM	FGR	No	No	No	4/13 (30.8)
Other family members affected	Yes	No	No	No	No	No	No	No	No	Yes	No	No	No	2/10 (20)
Time from symptom onset to hospital admission, d	5	0	8	0	0	0	3	6	5	1	7	5	5	3.5±2.9
Clinical manifestations														
Fever	Yes	Yes	Yes	No	Yes	Yes	No	No	Yes	Yes	No	Yes	Yes	9/13 (69)
Cough	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	12/13 (92)
Shortness of breath	Yes	Yes	Yes	Yes	Yes	Yes	No	No	Yes	Yes	Yes	Yes	Yes	11/13 (85)
Diarrhea	No	No	No	No	No	No	Yes	No	No	No	No	Yes	Yes	3/13 (23)
Fatigue	Yes	No	Yes	No	No	No	Yes	No	Yes	No	No	Yes	Yes	6/13 (46)
Sore throat	No	Yes	No	No	No	No	No	No	No	No	No	Yes	Yes	3/13 (23)
Anosmia or ageusia	No	No	No	No	No	No	No	Yes	No	Yes	Yes	Yes	No	4/13 (31)
Acute respiratory distress syndrome	No	Yes	No	No	No	No	No	No	No	No	No	No	No	1/13 (7.7)
distress syndrome Sentilhes et al. Coronavirus disease	e 2019 in pregn	ancy can cause	e severe materi	ıal morbidity.	Am J Obstet O	Gynecol 2020.								(contin

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OBSTETRICS **Original Research**

SUPPLEMENTAL TABLE

Clinical characteristics and outcomes of the 13 women with confirmed or suspected COVID-19 who required oxygen support (continued)

	Woman 1	Woman 2	Woman 3	Woman 4	Woman 5	Woman 6	Woman 7	Woman 8	Woman 9	Woman 10	Woman 11	Woman 12	Woman 13	
Characteristics	Women w	/ho had vag	inal or ces	arean delive	ery		_			Ongoing pro	egnancies		-	n/N (%)
Chest CT	Yes	Yes	Yes	No	Yes	Yes	Yes	Yes	Yes	No	No	Yes	Yes	10/13 (77)
Typical signs of viral infection on CT^d	Yes	Yes	Yes	N/A	No	No	Yes	Yes	Yes	N/A	N/A	Yes	Yes	8/10 (80)
Laboratory characteristics														
RT-PCR positive for SARS-CoV-2	Yes	Yes	Yes	No	No	No	Yes	Yes	Yes	Yes	Yes	Yes	No	9/13 (69)
White blood cell count $(\times 10^9 \text{ cells per L})$	5.68	7.33	3.32	9.6	19.1	10.3	10.9	5.1	5.3	7.8	11.4	9.1	4.37	8.4±4.1
Lymphocyte count (×10 ⁹ cells per L)	0.8	0.9	0.8	0.9	0.95	0.6	0.70	1.3	0.8	NP	1.8	1.7	0.8	1.0±0.4
C-reactive protein concentration (mg/L)	48.5	13.5	72	13.2	27.9	61.8	7.4	4.3	93.6	7.3	24	17.7	65.7	35.1±29.6
ALT (U/L)	55	48	22	14	13	15	36	76	26	12	75	12	12	32.0±24.0
AST (U/L)	65	49	45	23	29	30	76	71	36	15	64	18	13	42.1±22.2
Hemoglobin (g/dL)	12.4	10.1	11.6	8.8	13.10	10.7	9.4	11.7	11.6	10.9	12.9	10.4	9.6	11.0±1.3
Creatinine (µmol/L)	38.2	48.7	49.8	39.6	47.4	84.8	44.6	46.3	39.2	39.8	NP	59.9	35.2	47.8±13.4
Treatment before delivery														
Oxygen support (nasal cannula)	Yes	Yes	Yes	Yes	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes	12/13 (92)
Oxygen (L/min)	5	5	2	4	15 ^b	N/A	4	4	5	1	1	1	4	_
High-flow oxygen therapy	No	No	No	No	No	No	No	No	No	No	No	No	Yes	1 (8)
Noninvasive mechanical ventilation	No	No	No	No	No	No	No	No	No	No	No	No	Yes	0
Invasive mechanical ventilation	No	No	No	No	No	No	No	No	No	No	No	No	No	0
Admission to the ICU	No	No	No	No	No	No	No	No	No	No	No	No	Yes	1 (8)
Antenatal steroids for fetal lung maturity	Yes	Yes	No	No	No	No	No	No	Yes	No	No	No	Yes	4 (31)
Delivery	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No	No	No	No	9 (69)
Gestational age at delivery, wk+d	27+5 ^c	27+2 ^c	37+2	38+1	40+3	39+2	35+0 ^c	34+0 ^c	28+4 ^c	N/A	N/A	N/A	N/A	35±4.8
Sentilhes et al. Coronavirus diseas	e 2019 in pregr	iancy can caus	e severe mater	nal morbidity.	Am J Obstet (Gynecol 2020.								(continued)

SUPPLEMENTAL TABLE

Clinical characteristics and outcomes of the 13 women with confirmed or suspected COVID-19 who required oxygen support (continued)

	Woman 1	Woman 2	Woman 3	Woman 4	Woman 5	Woman 6	Woman 7	Woman 8	Woman 9	Woman 10	Woman 11	Woman 12	Woman 13	
Characteristics	Women w	ho had vag	inal or cesa	arean delive	ery					Ongoing pre	gnancies			n/N (%)
Spinal or epidural anesthesia	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	N/A	N/A	N/A	N/A	9/9 (100)
Indicated cesarean delivery because of respiratory symptoms related to COVID-19	Yes	Yes	Yes	Yes	No	No	Yes	Yes	Yes	N/A	N/A	N/A	N/A	7/9 (78)
Indicated cesarean delivery for other reasons	No	No	No	No	No	Yes ^a	No	No	No	N/A	N/A	N/A	N/A	1/9 (11)
Cesarean delivery during labor	No	No	No	No	No	Yes	No	No	No	N/A	N/A	N/A	N/A	1/9 (11)
Vaginal delivery	No	No	No	No	Yes	No	No	No	No	N/A	N/A	N/A	N/A	1/9 (11)
PPH, defined by blood loss $\geq\!500$ mL, measured with a graduated collector bag	No	No	No	No	No	No	Yes	No	No	N/A	N/A	N/A	N/A	1/9 (11)
Blood transfusion	No	No	No	No	No	No	Yes	No	No	N/A	N/A	N/A	N/A	1/9 (11)
Treatment after delivery														
Oxygen support (nasal cannula)	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes	N/A	N/A	N/A	N/A	8/9 (89)
Oxygen (L/min)	5	N/A	4	1	1	4	4	2	3	N/A	N/A	N/A	N/A	
High-flow oxygen therapy	No	No	No	No	No	No	Yes	No	No	N/A	N/A	N/A	N/A	1 (8)
Noninvasive mechanical ventilation	No	No	No	No	No	No	No	No	No	N/A	N/A	N/A	N/A	0
Invasive mechanical ventilation	No	Yes	No	Yes	Yes	No	No	No	No	N/A	N/A	N/A	N/A	3/9 (33)
Extracorporeal membrane oxygenation	No	Yes	No	No	No	No	No	No	No	N/A	N/A	N/A	N/A	1/9 (11)
Admission in ICU	Yes	Yes	No	Yes	Yes	No	No	No	No	N/A	N/A	N/A	N/A	4/9 (44)
Antiviral therapy	Yes	Yes	Yes	No	No	No	No	No	No	N/A	N/A	N/A	N/A	3/9 (33)
Antibiotic therapy	Yes	Yes	Yes	No	No	No	Yes	No	Yes	N/A	N/A	N/A	N/A	4/9 (44)
Discharge from hospital at the end of follow-up	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No	11/13 (85)
Sentilhes et al. Coronavirus disease	2019 in pregn	ancy can caus	e severe materi	ıal morbidity.	Am J Obstet C	Synecol 2020.								(continued)

SUPPLEMENTAL TABLE

Clinical characteristics and outcomes of the 13 women with confirmed or suspected COVID-19 who required oxygen support (continued)

	Woman 1	Woman 2	Woman 3	Woman 4	Woman 5	Woman 6	Woman 7	Woman 8	Woman 9	Woman 10	Woman 11	Woman 12	Woman 13			
Characteristics	Women w	omen who had vaginal or cesarean delivery										Ongoing pregnancies				
Neonatal outcome																
Sex	Male	Male	Female	Female	Female	Female	Male	Male	Male	N/A	N/A	N/A	N/A	N/A		
Birthweight (g)	1080	1085	2220	3370	2840	2520	2590	2320	1340	N/A	N/A	N/A	N/A	2152±811		
Small for gestational age	No	No	No	No	No	No	No	No	No	N/A	N/A	N/A	N/A	0		
5-min Apgar score <7	No	No	No	No	No	No	No	No	No	N/A	N/A	N/A	N/A	1/9 (11)		
Admission to neonatal ICU	Yes	Yes	No	Yes	No	No	No	Yes	Yes	N/A	N/A	N/A	N/A	5/9 (55)		
Attempted CPAP in the first 24 h of life	No	No	No	No	No	No	No	No	No	N/A	N/A	N/A	N/A	0		
Breastfeeding	Yes	No	No	Yes	Yes	Yes	Yes	Yes	No	N/A	N/A	N/A	N/A	6/9 (67)		
Discharge from hospital at the end of follow-up	No	No	Yes	Yes	Yes	Yes	Yes	No	No	N/A	N/A	N/A	N/A	5/9 (55)		

Data are presented as mean±SD, unless otherwise indicated. The body mass index is the weight in kilograms divided by the square of the height in meters. Reference ranges of the different laboratory results are provided in Table 2. Small for gestational age was defined as birthweight less than the 10th percentile for gestational age and sex based on French intrauterine growth curves.

ALT, alanine aminotransferase; AST, aspartate aminotransferase; COVID-19, coronavirus disease 2019; CPAP, continuous positive airway pressure; CT, computed tomography; FGR, fetal growth restriction; GDM, gestational diabetes mellitus; ICU, intensive care unit; IV/A, not applicable as the woman has not given birth; NP, not performed; PIH, pregnancy-induced hypertension; PPH, postpartum hemorrhage; SD, standard deviation.

^a Cesarean delivery for preeclampsia; ^b High concentration oxygen mask; ^c Preterm birth was decided because of severe maternal respiratory symptoms related to COVID-19; ^d Typical radiographic features of COVID-19 on chest CT ground-glass opacity, or bilateral pulmonary infiltration.²⁷

Sentilhes et al. Coronavirus disease 2019 in pregnancy can cause severe maternal morbidity. Am J Obstet Gynecol 2020.