Fetomaternal Outcome of Pregnancy with COVID-19: An Observational Study in Tertiary Care Hospitals of Pakistan

NABILA KHAN¹, ZAHRA WASIM², AESHA SADAF RIZWAN³, AFSHAN AHMAD⁴, MUHAMMAD TAHIR⁵, IRUM BATOOL HASHMI⁶

¹Assistant Professor Gynae & Obs, BKMC/MMC Mardan.

Corresponding author: Dr. Zahra Wasim, Email: zahrawasim1973@gmail.com, Contact: +923202410300

ABSTRACT

Background: The new corona virus first appeared in Wuhan, China in December 2019 and has since spread around the world to other countries. The World Health Organization believes that this new CoV-19 epidemic is a public health emergency of international concern (PHEIC) on January 30, 2020Worldwide. The mortality rate of this viral infection ranges from 2% in Pakistan to 14.4% in Italy. Lympopenia, elevated transminase, proteinuria, increased LDH, and C-reactive protein levels are all common laboratory findings in the early stages of the disease. Covid patients have experienced a variety of complications, including extreme pneumonia, ARDS, heart defects, sepsis and septic shock, and respiratory tract super infection.

Methodology: This retrospective observational research study was carried out at the Gynecology Unit of MardanMedical Complex, Mardan and Combined Military Hospital, Risalpur for 06 months duration from April 2020 to September 2020. In a pre-constructed data collection form, biochemical and radiological parameters of medical history, test results, symptoms, pregnancy, and neonatal outcomes were noted. Patients treated in an outpatient setting were not included in the study.

Results: There were 121 patients in total, with mean age of 27 having standard deviation ± 5, having range 19-40 years. 48.3% pregnant women reported their first pregnancy(primigravida). 51.3% of SARS-Cov-2 were in their 3rd trimester while 34.7% were in their 35-40 weeks of gestational age. Common complications are gestational hypertension (PIH) (16 cases), hypothyroidism (14 cases) and gestational diabetes (GDM) 9 cases. More than half (53%) of patients are asymptomatic. Common symptoms are cough (22%) and fever (11%). The incidence of multiple organ failure was 2% as shown in table 01. Lymphopenia was common (84%). A CT scan of 24 patients showed bilateral invasion.

Conclusion: COVID-19 has a negative impact on the foetus, according to our results. Although pregnant women do not seem to be more vulnerable to COVID-19 complications than non-pregnant adults, previous research has suggested that pregnant women could be at higher risk for negative pregnancy outcomes such as preterm birth, foetal pain and respiration, symptoms, and LBW in a newborn baby.

Keywords: Corona virus, COVID-19, Pregnancy, Outcome.

INTRODUCTION

SARS CoV2 is a single-stranded RNA virus, belonging to a corona virus group, known to cause severe acute respiratory illness in humans. High transfer rates and low pathogenicity are two traits that differentiate them from other family members (SARS-CoV and MERS-CoV) (1). The new corona virus first appeared in Wuhan, China in December 2019 and has since spread around the world to other countries (2). The World Health Organization believes that this new CoV-19 epidemic is a public health emergency of international concern (PHEIC) on January 30, 2020 (3). Worldwide, the mortality rate of this viral infection ranges from 2% in Pakistan to 14.4% in Italy (4).

COVID-19 signs are mild in the majority of SARS-CoV-2 infections. Men may be at greater risk for disease than women, and the elderly and patients with comorbidities such as cardiovascular disease, respiratory disease, and diabetes are at higher risk for severity and death. (5,6). Lympopenia, elevated transminase, presence of excess protein in urine(proteinuria), increased level of Lactate Dehydrogenase (LDH), and level of C-reactive protein are all common laboratory findings in the early

stages of the disease. Covid patients have experienced a variety of complications, including extreme pneumonia, ARDS, heart defects, sepsis and septic shock, and respiratory tract super infection (7).

The prevalence of placentaprevia is reported to be 3-5 times per 1,000 people and it is statistically increasing worldwide due to risk factors for inflammation. (8) In Pakistan, the prevalence rate was 3.5%, while the proportion of women born by caesarean section rose to 65%. (9) In the past, sensitivity to the urea ratio has been observed more frequently, mainly due to the high rate of cesarean section and the unconventional late age of the pregnant mother (10). The reason for placental control is unclear, but some threats may indicate its prevalence, such as cesarean section, high mothers, increasing age and previous miscarriages. (11)

Because the immune system is less aggressive during pregnancy, the mother's body does not genetically attack other embryos. As a result, pregnant women are susceptible to various infections, including viral diseases. In addition, several physiological changes during pregnancy, such as airway swelling, increased respiratory secretions

²Classified Gynaecologist, CMH Risalpur

³Assistant Professor Gynaecology and Obstetrics Shahida Islam Medical and Dental College/ Shahida Islam teaching Hospital Lodhran

⁴Gynaecologist DHQ hospital Batkhela Malakand

⁵Assistant Professor of Oncology Allied Hospital, Faisalabad/Faisalabad Medical University Faisalabad

⁶Assistant Professor Obs/Gynae, Gomal Medical College Dera Ismail Khan.

and oxygen demand, increased diaphragmatic and decreased functional function, are factors that have so far worsened the obstetric prognosis in pregnant women suffering from viral pneumonia. Literature report. Limited data published in China on COVID-19 showed the worst birth outcomes, including preterm birth and mortality at birth during multiple pregnancies (12).

There is anxiety among pregnant women, and our limited ability to answer questions affects women's health in some way. Due to the lack of guidelines regarding the effects of SARS-CoV-2 on pregnancy, clinical medical records of patients with pregnancy SARS-CoV-2 are collected and reported to the third HMC treatment center for further evaluation and management. This research looked at the outcomes of foetal mothers in COVID-19-positive pregnant women.

METHODOLOGY

This retrospective observational research study was carried out at the Gynecology unit of tMardanMedical Complex, Mardan and Combined Military Hospital, Risalpurfor 06 months duration from April 2020 to September 2020after receiving ethical approval from the Research and Ethics Committee of the hospitals. A total of 121 pregnant women and their infants were treated with SARS CoV-2 RNA identified by RT PCR. Patients treated in an outpatient setting were not included in the study. In a pre-constructed data collection form, biochemical and radiological parameters of medical history, test results, symptoms, pregnancy, and neonatal outcomes were registered. Analyze your data using SPSS version 20.

RESULTS

There were 121 patients in total, with mean age of 28 having standard deviation \pm 7, having range 17-42 years. 48.3 percent pregnant women reported their first pregnancy(primigravida). 51.3% of SARS-Cov-2 were in their 3rd trimester while 34.7% were in their 35-40 weeks of gestational age. Common complications are gestational hypertension (PIH) (16 cases), hypothyroidism (14 cases) and gestational diabetes (GDM) 9 cases.

Table I. Demographic and clinical characteristics of pregnant women with COVID-19 (N=81)

MONION WILL COVID TO (N=01)	
General characteristics	Percentage
Mean age 27 years (range 19-40)	=
Nulliparous	48.3
Multiparous	51.7
Sign and symptoms	
Asymptomatic	53
Cough	22
Fever	11
Shortness of breath	4
Anosmia	3
Multi-organ failure	2
Fever, headache, severe pneumonia	3
Hemoptysis	2

More than half (53%) of patients are asymptomatic. Common symptoms are cough (22%) and fever (11%). The incidence of multiple organ failure was 2% as shown in

table 01. Lymphopenia was common (84%). A CT scan of 24 patients showed bilateral invasion.

Most common was mild to moderate disease in about 92.6% of patients while in about 7.4% patients there were only severe disease. In about 5 patients due to critical disease required mechanical ventilation. Patients suffered from a serious illness before giving birth and received mechanical ventilation after giving birth. Patients with mild to moderate form of diseases (92.6%) received treatment in corona ward while severe patients(7.4%) patients received treatment in Intensive care unit (ICU).

All of the patients (100%) were given zinc, vitamin C, and vitamin D supplements, as well as Sodium Ceftriaxone metronidazole covers, Cefixime, and Metronidazole is given to all caesarean section patients. Azithromycin, moxifloxacin, and colistin are some of the antibiotics used in severe diseases. Methylprednisolone, Inj. remdisivir and Inj. tocilyzumab. 36 (29.6%) patients with low molecular weight heparin to prevent thrombosis and Tab. Rebroxaben was used in 6 (7.4%) severe patients after delivery. In 30 patients, ivermectin was used as a tablet (24.7 percent). There was no difference in results between those who received ivermectin care and those who did not. In the report, 106 out of 121 women (87.65%) were discharged, with seven of them suffering from moderate or severe disease. There was only one death: that of the mother. In the sample population, there were three spontaneous abortions. In the study, 78 (64.2%) of 121 patients gave birth. A Caesarean section was performed on 63 (80.8 percent), a hysterectomy was performed on 3 (2.4 percent), and a regular vaginal delivery was performed on 12 (15.3) percent).. A total of 12 (15.3%) patients had an abortion. A hysterectomy was performed for chronic hypertension with combination HELLP syndrome and pre-eclampsia (PE). Indications for cesarean section are anterior cesarean section (31), fetal depression (9), preterm labor (4), eclampsia (5), severe PE (3), HELLP syndrome (3), cervix No good condition of the cervix (3), caesarean section (3), preterm birth with breech birth (2). Current pregnancy 39 (32.09%).

TableII. Maternal outcome of pregnant ladies with COVID-19 (N=121)

Maternal outcome	Frequency	Percentage
Mild to moderate disease	112	92.6
Severe disease	9	7.4
Delivered	78	64.2
Discharged	106	87.7
Ongoing pregnancy	39	32.1
Maternal death	2	1.2

The SARS-CoV-2 test was performed with a neonatal pharyngeal swab, and 6 were positive. There were 70 births, 4 endometrial deaths and 3 neonatal deaths. Neonatal death was caused in more premature infants as shown in Table 3. The mean length of continuity was 13.3 (standard deviation \pm 6.5) (range 6–31) days. The average number of days it took to start breastfeeding was 13.5 (SD \pm 6.44) (range 5–25).

Table III. Pregnancy an	d neonatal outcome of	patients with	COVID-19 (N=121)
-------------------------	-----------------------	---------------	------------	--------

Pregnancy and neonatal outcome	Frequency	Percentage
Delivery	78	64.2
Normal vaginal delivery (NVD)	12	15.1
Caesarean section	63	80.8
Hysterectomy	3	2.4
Abortion	4	3.7
Live birth	70	90.4
Preterm birth	12	15.4
Neonatal death	3	3.8
Neonatal COVID infection	6	7.6

DISCUSSION

Physiologic changes occur in pregnant women's different body systems like immune, cardiopulmonary and coagulation thus made them more prone tom infection. Some research studies reported no exacerbations of Covid pneumonia or any kind of CT scan features in Covid infected pregnant women. (13). Results of one of previous study on pregnant Covid infected woman patient symptoms and laboratory findings concluded fever (68%) and cough(34%) as common symptoms while increase level of C-reactive protein(70%) and lymphopenia(59%) were common laboratory findings. (14). In comparison to above studies results, our study patients reported cough(22%) and fever(11%) only while most of the patients had lymphopenia(84%) while only 7.6% patients were done with CT scan.

Maternal and fetaloutcomes: Many researches in china was done on pregnant women having Covid infection. One of the case series study done on three pregnant women having covid positive PCR test reports gave birth to one preterm low birthweight(LBW) baby while two normal babies (15). Another study of same perspective reported low birthweight baby(LBW) healthy baby in Covid positive 30 weeks pregnant women (16). In comparison of our study results with previous studies, 15.8% of pregnant Covid positive patients reported preterm deliveries.

Study of Liu et. Al. done on 13 covid positive pregnant reported positive PCR test results in two women before 28 weeks while rest of eleven patients got infection after 28 weeks. Among these thirteen Covid positive Covid pregnant women ten patients had caesarean sections, five of which were emergency caesarean sections, three patients had Foetal depression while only one had premature fluid rupture and one have still to born (17). In our study, 49.1 percent were infected in the third trimester, 52 patients were born in the study period, six were seriously ill and received intensive treatment, 3 were artificially ventilated, and one was died during study period.

Chen H et al. identified two cases of foetal distress, four cases of preterm birth (1 case of maternal preeclampsia, one case of PROM, and two cases with a history of caesarean section or stillbirth dead) and two cases of LBW in their study (with maternal preeclampsia) There was one case of PROM among nine pregnant women who had a positive Covid test (18). Another previous study done on 17 pregnant Covid positive women reported premature birth of 3 babies (19). Results of

previous study reported death of neonate in Covid positive severely ill pregnant woman and the reason of death identified was uterine asphyxia(20).In comparison of our study results with above mentioned studies, six cases of foetal distress, three cases of IUD were identified while in eight cases there were termination of pregnancy earlier.

There were multiple negative pregnancy outcome in study carried out by Zhu H. et al on nine pregnant women with Covid positive clinical signs and symptoms in these nine women started in different time frames like in four pregnant women these symptoms started nbefore delivery, while on the day of delivery two of women got infected and three women got infection after delivery. Among these nine pregnant covid infected women gave birth to six premature babies and two babies were of SGA (small gestational age) (21).

N. Li and colleagues compared case-control study of 16 pregnant women with confirmed COVID-19 pneumonia and 18 pregnant women with suspected COVID-19 pneumonia who give birth in the third trimester in two cohort studies in 2019 and 2020(22).

An estimated 70% of cases in both groups had other maternal complications (primary GDM, preeclampsia, hypothyroidism), a much higher proportion than in the control group (33%). In three confirmed cases there were ceased periods among which two were because of PROM and one was because of bleeding from placentaand three suspected cases (16.9 percent, one because of preeclampsia / preeclampsia and one because of placenta previa). This represents 5.8% of the control community. PIH, hypothyroidism and GDM were the most common comorbidities with 13.58%, 11.11% and 7.40%% respectively in our research.

Delivery: The majority of caesareans were administered 24/28 (85.7%), 19/24 (79%) emergency caesareans, and 4/24 limited caesareans in a prospective follow-up report (16.6 percent) (23). Pathological CTG, birth failure, and PROM are all reasons for a caesarean section (birth failure, maternal needs and severe sepsis). two case-control studies by Zhang L. et al. performed during pandemic in China, comparing the pregnancy results of sixteen covid positive patients among which one was severe and 45 uninfected women, all of whom were delivered by caesarean section (24). The amount of blood lost during the procedure did not vary significantly, according to the researchers. In our study there were more caesarean

section as compared to normal vaginal delivery with 64.2 % C-section and 15.14% normal vaginal delivery (NVD).

Neonatal outcome: Shortness of breath in six patients, fever in two patients and thrombocytopenia with impaired liver function in two patients were among the clinical symptoms identified by Zhu H. et al. in his study which he carried out on six newborns with a paediatric critical illness score (PCIS) of less than 90. There was one case of illness (n = 1) and one case of vomiting (n = 1). Four newborns had gastrointestinal signs such as bloating, milk rejection, food aversion, and gastric bleeding, while seven had imaging anomalies (infection, n = 4, neonatal respiratory distress syndrome, n = 2, and pneumothorax, n = 1). 8 days after birth, the baby experienced failure of many body systems and organs and diffuse clotting in between vessels, and died 9 days later after receiving a blood transfusion (21). In our study, only 6 cases were found to be positive for RT-PCR. A very premature birth resulted in 3 newborns. The rest is fine.

CONCLUSION

COVID-19 has a negative impact on the fetus, according to our results. Although pregnant women do not seem to be more vulnerable to COVID-19 complications than nonpregnant adults, previous research has suggested that pregnant women could be at higher risk for negative pregnancy outcomes such as preterm birth, foetal pain and respiration, symptoms, and LBW in a newborn baby. More clinical research studies arethe need of time to determine the true impact of Covid pandemic on Fetomaternal outcomes, as well as to direct the most effective gynecologist's recommendations. Only a multi-faceted evaluation of current virus features, epidemiology, disease immunopathology, future preventive and treatment methods, and awareness of current clinical findings may help determine the exact impact of Covid infection before, during as well as after pregnancy.

REFERENCES

- Zhao S, Lin Q, Ran J, Musa SS, Yang G, Wang W, Lou Y, et al. Preliminary estimation of the basic reproduction number of novel coronavirus (2019-nCoV) in China, from 2019 to 2020: A datadriven analysis in the early phase of the outbreak. International journal of infectious diseases. 2020;92:214-7.
- R Gajbhiye, D Modi and S. Mahale, "Pregnancy outcomes Newborn complications and Maternal-Fetal Transmission of SARS-CoV-2 in women with COVID-19: A systematic review", medRxiv, Jan 2020.
- A Schwartz and AL Graham, "Potential maternal and infant outcomes from (Wuhan) coronavirus 2019-nCoV infecting pregnant women: lessons from SARS MERS and other human coronavirus infections", Viruses, vol. 12, no. 2, pp. 194, Feb 2020
- R Mushtaq, K Parveen, A Siraj, M Jannat and H Ali, "OUTCOME OF TWENTY PREGNANT WOMEN WITH COVID-19 INFECTION-A CASE SERIES FROM PAKISTAN", Pakistan Armed Forces Medical Journal, vol. 70, no. 2, pp. 572-77, Sep 2020.
- Wu Z, McGoogan JM. Characteristics of and important lessons from the coronavirus disease 2019 (COVID-19) outbreak in China: summary of a report of 72 314 cases from the Chinese Center for Disease Control and Prevention. Jama. 2020 Apr 7;323(13):1239-42
- IstitutoSuperiore di Sanità. Report about the Characteristics of Death Patients Positive to COVID-19 in Italy (Based on Data Updated on 17th March 2020).

- Chowdhury L, Jahan I, Sharmin MI, Ferdaushi J, Tasnim T, Nahar S. Fetomaternal Outcome of Pregnancy with COVID-19: An Observational Study in A Tertiary Care Hospital of Bangladesh. Journal of Bangladesh College of Physicians and Surgeons. 2021 Mar 9;39(2):100-5.
- 8. Khursheed F, Shaikh F, Das CM, Shaikh RB. Placenta previa: an analysis of risk factors. Medical Channel 2010; 16(3):417-9.
- Tabassum R, Raheel R, Bhutto A, Riaz H, Hanif F. The risk factor associated with placenta previa in patients presented to civil hospital Karachi- a case report control study. Medical Channel 2010; 16(2):276-9.
- Shoukat A, Zafar F, Asghar S, Nighat, Ayoub A, Ambreen N, et al. Frequency of placenta previa with previous Csection:[internet].2011.Available from: http://pjmhsonline. com/frequency_of_placenta_previa_wit.htm.
- Hung TH, Hsieh CC, Hsu JJ, Chiu TH, Lo LM, Hsieh TT. Risk factors for placenta previa in an Asian population. International Journal of Gynecology and Obstetrics 2007; 97,26–30
- Zaigham M, Andersson O. Maternal and perinatal outcomes with COVID-19: A systematic review of 108 pregnancies. ActaObstetGynecolScand 2020; 99(1): 823-29.
- Wong SF, Chow KM, Leung TN, Ng WF, Ng TK, Shek CC, Ng PC, Lam PW, Ho LC, To WW, Lai ST. Pregnancy and perinatal outcomes of women with severe acute respiratory syndrome. American journal of obstetrics and gynecology. 2004 Jul 1;191(1):292-7.
- Zaigham M, Andersson O. Maternal and Perinatal Outcomes with COVID-19:a systematic review of 108 pregnancies. ActaObstetGynecol Scand. 2020.
- Chen S, Huang B, Luo DJ, Li X, Yang F, Zhao Y, Nie X, Huang BX. Pregnancy with new coronavirus infection: clinical characteristics and placental pathological analysis of three cases. Zhonghuabing li xuezazhi= Chinese journal of pathology. 2020 Mar 1;49(5):418-23.
- Wang X, Zhou Z, Zhang J, Zhu F, Tang Y, Shen X. A case of 2019 Novel Coronavirus in a pregnant woman with preterm delivery. Clinical infectious diseases. 2020 Feb 28.
- Liu Y, Chen H, Tang K, Guo Y. Clinical manifestations and outcome of SARS-CoV-2 infection during pregnancy. The Journal of infection. 2020 Mar 4.
- Chen H, Guo J, Wang C, Luo F, Yu X, Zhang W, Li J, Zhao D, Xu D, Gong Q, Liao J. Clinical characteristics and intrauterine vertical transmission potential of COVID-19 infection in nine pregnant women: a retrospective review of medical records. The lancet. 2020 Mar 7;395(10226):809-15.
- Chen R, Zhang Y, Huang L, Cheng BH, Xia ZY, Meng QT. Safety and efficacy of different anesthetic regimens for parturients with COVID-19 undergoing Cesarean delivery: a case series of 17 patients. Canadian Journal of Anesthesia/Journal canadiend'anesthésie. 2020 Jun;67(6):655-63.
- Zhang B, Liu S, Tan T, Huang W, Dong Y, Chen L, Chen Q, Zhang L, Zhong Q, Zhang X, Zou Y. Treatment with convalescent plasma for critically ill patients with severe acute respiratory syndrome coronavirus 2 infection. Chest. 2020 Jul 1;158(1):e9-13.
- Zhu H, Wang L, Fang C, Peng S, Zhang L, Chang G, Xia S, Zhou W. Clinical analysis of 10 neonates born to mothers with 2019-nCoV pneumonia. Translational pediatrics. 2020 Feb;9(1):51.
- Duran P, Berman S, Niermeyer S, Jaenisch T, Forster T, Gomez Ponce de Leon R, De Mucio B, Serruya S. COVID-19 and newborn health: systematic review. RevistaPanamericana de SaludPública. 2020 May 29;44:e54.
- Maternal COVID-19 infection, clinical characteristics, pregnancy, and neonatal outcome: A prospective cohort study.doi: 10.1016/j.ejogrb.2020.07.008
- Zhang L, Jiang Y, Wei M, Cheng BH, Zhou XC, Li J, Tian JH, Dong L, Hu RH. Analysis of the pregnancy outcomes in pregnant women with COVID-19 in Hubei Province. Zhonghuafuchankezazhi. 2020 Mar 7;55(3):166-71.