

A PROSPECTIVE STUDY ON PREGNANCY COMPLICATED WITH JAUNDICE AND FETOMATERNAL OUTCOME

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ABSTRACT

Objectives: The aim of the study was to determine biochemical factors and maternofetal outcome in women with jaundice during pregnancy and to study its association with jaundice in pregnancy.

Methods: It was an observational study done among 60 pregnant patients with abnormal liver functioning. Laboratory parameters were studied which mainly includes liver function tests. Demographics and the detailed clinical history were recorded. The detailed laboratory investigation was carried out to study the complications concerning biochemical parameter. SPSS (Version 22.0) was used for analysis.

Results: HELLP syndrome was the most common etiology (45%). Decreased levels of hemoglobin observed in 75%, increased total leukocyte count observed in 26%, and low platelet count observed in 34%, respectively. Among mothers, most common adverse outcome was requirement of emergency lower segment cesarean section (45%). Maternal outcomes were significantly associated with raised direct bilirubin, raised serum glutamic pyruvic transaminase, raised alkaline phosphatase, and thrombocytopenia ($p=0.03$, $p=0.01$, $p=0.02$, and $p=0.01$, respectively).

Conclusion: Hepatic dysfunction during pregnancy is associated with adverse events for both the mother and the fetus.

Keywords: Alanine transaminase, Alkaline phosphatase, Bilirubin, Liver diseases, Low birth weight, Thrombocytopenia.

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INTRODUCTION

The prevalence of liver dysfunction during pregnancies ranges between 3% and 10% [1]. Jaundice, also mentioned as hyperbilirubinemia, is presented by accumulation of bile pigments in the skin that grades in yellowing mucous membranes and the skin [2,3]. Jaundice in pregnancy may lead to adversarial maternofetal outcomes including perinatal and maternal mortality which accounts for around 60% and 14% separately [4]. However, it is caused by a number of causes, some associated and some coincidental including abnormal liver functioning distinctive to pregnancy, pre-hepatic causes, hepatic causes, and post-hepatic causes of jaundice. Abnormal liver functioning distinctive to pregnancy are HELLP syndrome, pre-eclampsia, acute fatty liver, hyperemesis gravidarum (HEG), and intrahepatic cholestasis of pregnancy. Pre-hepatic causes including hepatic pathologies (viral hepatitis), hemolytic anemia, drug-induced hepatitis, Wilson's disease and Budd-Chiari syndrome. Post-hepatic causes include CBD obstructions, pancreatitis, choledochal cyst, and gall stones. Although jaundice in pregnancy is relatively rare, it may lead to serious maternofetal complications. The rationale behind the study is to determine clinicoetiological, biochemical factors, and maternofetal outcome in women with jaundice during pregnancy.

METHODS

Post receiving Institutional Ethical Committee clearance, the observational study was conducted in department of obstetrics and gynaecology of a tertiary care hospital from July 2021 to December 2021. The sample size was calculated in SPSS using standard formula (pwr: Chi-square test [effect size=0.60, df=1, power=0.80, sig. level=0.05]) at 95% significance level, power being 80% and the minimum sample size was found to be 53. A total of 60 patients with abnormal liver functions were included in the study. All pregnant patients with abnormal liver function tests were included in this study. Patients with normal liver functioning were excluded from the study. Written informed consent was obtained from all the patients before the study. Demographic data

and a detailed clinical history were collected including age, obstetrical profile, gravida, and etiology. The detailed laboratory investigation was carried out to study the complications concerning biochemical parameter. Maternal and fetal outcomes were also recorded.

Statistical analysis

Statistical analysis was done using the software Statistical Package for the Social Sciences (SPSS) 22.0 version. Continuous variables was presented by mean±SD and categorical variables by frequency or percentages. To find the association between categorical variables, we have used Chi-square or Fisher's exact test. Independent t-test was used to find the significant difference in means of the variables. p value of <0.05 indicated a statistically significant.

RESULTS

As per Table 1, the most common age group seen the study was 26–30 years (34%) of participants, followed by 21–25 years. Although >30 years were also seen but most of females belonged to 21–30 years. Mean age of the patients was 25.2±3.75 years.

As per Table 2, high number of patients had low level of Hb (67%), total leukocyte count between 7000 and 11000 cells/ μ L (60%), platelets >150000 cells/ μ L (34%), total bilirubin <2 mg/dL (54%), direct bilirubin >0.2 mg/dL (59%), serum glutamic oxaloacetic transaminase >70 U/Lit (74%), serum glutamic pyruvic transaminase (SGPT) >70 U/Lit (74%), alkaline phosphatase >180 IU/Lit (65%), lactate dehydrogenase >600 U/Lit (74%), albumin >2.5 gm/dL (72%), total protein >8.5 gm/dL (54%), and urea >6.8 mg/dL (60%), while creatinine (mg/dl) (54%).

As per Table 3, among 60 cases showed, adverse maternal and fetal outcomes were observed in 27 and 29 cases, respectively. Low birth weight was observed as most common adverse fetal outcomes in 35% of cases. Among mothers, most common adverse outcome was requirement of emergency cesarean section lower segment cesarean section (45%).

Table 1: Age-wise distribution of the study participants (n=60)

| Age groups | Frequency (%) |
|-------------|---------------|
| <20 years | 8 (17) |
| 21–25 years | 20 (33) |
| 26–30 years | 22 (34) |
| >30 years | 10 (16) |

Table 2: Details of laboratory parameters in the study participants (n=60)

| Parameters | Category | Frequency (%) |
|--|----------------|---------------|
| Hb (g%) | <7 | 40 (67) |
| | >7 | 20 (33) |
| Total leukocyte count (cells/ μ L) | <7000 | 10 (13) |
| | 7000–11,000 | 34 (57) |
| | >11,000 | 16 (30) |
| | >150,000 | 20 (34) |
| Platelets (cells/ μ L) | <50,000 | 08 (13) |
| | 51,000–150,000 | 32 (53) |
| | >150,000 | 20 (34) |
| Total bilirubin (mg/dL) | <2 | 32 (54) |
| | >2 | 28 (46) |
| Direct bilirubin (mg/dL) | <0.2 | 25 (41) |
| | >0.2 | 35 (59) |
| SGOT (U/Lit) | <70 | 16 (26) |
| | >70 | 44 (74) |
| SGPT (U/Lit) | <70 | 16 (26) |
| | >70 | 44 (74) |
| Alkaline phosphatase (IU/Lit) | <180 | 21 (35) |
| | >180 | 39 (65) |
| Lactase dehydrogenase (U/Lit) | <600 | 16 (26) |
| | >600 | 44 (74) |
| Albumin (gm/dL) | <2.5 | 17 (28) |
| | >2.5 | 43 (72) |
| Total protein (gm/dL) | <8.5 | 28 (46) |
| | >8.5 | 32 (54) |
| Urea (mg/dL) | <6.8 | 24 (40) |
| | >6.8 | 36 (60) |
| Creatinine (mg/dL) | <1.4 | 28 (46) |
| | >1.4 | 32 (54) |

Table 3: Maternal and fetal outcomes

| Outcome | Frequency (%) |
|--------------------------|---------------|
| Maternal outcomes | |
| LSCS | 27 (45) |
| ICU admission | 17 (29) |
| Blood transfusion | 13 (27) |
| Maternal mortality | 01 ((2) |
| Fetal Outcomes | |
| LBW | 21 (35) |
| IUGR | 19 (32) |
| Preterm | 07 (13) |
| Neonatal death | 01 (2) |

As per Table 4, raised serum total bilirubin level, direct bilirubin level, and thrombocytopenia were significantly associated with adverse fetal outcomes and low hemoglobin. Maternal outcomes were significantly associated with raised direct bilirubin, raised SGPT, raised alkaline phosphatase and thrombocytopenia, and low hemoglobin ($p < 0.05$).

DISCUSSION

HELLP syndrome is defined as increased blood pressure with proteinuria or end-organ dysfunction in absence of proteinuria seen post 20 weeks of gestation [5]. Although, in the present study, HELLP syndrome was the most common syndrome seen (44%) tracked by acute fatty liver of pregnancy (AFLP) (32%), preeclampsia (28%), cholestasis of pregnancy (28%), and HEG (12%). Likewise, in study by Reddy *et al.*, HELLP syndrome was most common and was perceived in 33.3% patients, followed by

Table 4: Association of laboratory parameters with adverse maternal and fetal outcomes

| Parameters | Category | Adverse fetal outcome | Adverse Maternal Outcome |
|--|----------|-----------------------|--------------------------|
| Hb (g%) | <7 | 0.02* | 0.01* |
| Total leukocyte Count (cells/ μ L) | >11000 | 0.11 | 0.21 |
| Platelets (cells/ μ L) | <50000 | 0.001* | 0.001* |
| Total bilirubin (mg/dL) | >2 | 0.01* | 0.43 |
| Direct bilirubin (mg/dL) | >0.2 | 0.03* | 0.01* |
| SGOT (U/Lit) | >70 | 0.32 | 0.01* |
| SGPT (U/Lit) | >70 | 0.31 | 0.11 |
| Alkaline phosphatase (IU/Lit) | >180 | 0.45 | 0.001* |
| Lactase dehydrogenase (U/Lit) | >600 | 0.31 | 0.22 |
| Albumin (gm/dL) | <2.5 | 0.11 | 0.32 |
| Total protein (gm/dL) | <8.5 | 0.17 | 0.76 |
| Urea (mg/dL) | >6.8 | 0.34 | 0.23 |
| Creatinine (mg/dL) | >1.4 | 0.12 | 0.41 |

AFLP in 22.2% and intrahepatic cholestasis of pregnancy in 11.1% patients [6]. Inconsistently, in a study conducted by Suresha *et al.* and Allen *et al.*, HELLP syndrome was the second mutual etiology after eclampsia and preeclampsia, correspondingly [1,7]. Satia *et al.*, reported viral hepatitis (62%) as the most common etiology monitored by cholestasis of pregnancy (24%) [8]. The pathogenesis of HELLP is still not strong but is supposed to involve abnormalities in placental vasculature and faults in maternal vascular endothelial cells, which consequences in poor perfusion [5]. In developing countries, anemia is a public health problem particularly during pregnancy [9]. Change in blood count is a common spectacle in pregnancy [10–12]. Liver dysfunction was quite obvious in high number of patients due to increased level of bilirubin. SGOT, SGPT, and level of bilirubin were high in 70%, 70%, and 44% of cases, respectively. However, in aalike study, Ronceglia *et al.* reported judiciously increased bilirubin level of 1–10 mg% and hypoglycemia [13]. Shinde *et al.* also reported elevated bilirubin level in the pregnant patients with jaundice compared to non-pregnant patients [14]. In their study, serum bilirubin levels between 11 and 15 mg/dl and between 16 and 25 mg/dl were recorded in 38.4% and 19.2% of pregnant patients, respectively, but among non-pregnant patients, 36.5% had serum bilirubin between 6 and 10 mg/dl [14]. Most common adversarial neonatal outcome was low birthweight (35%) followed by intrauterine growth retardation (32%). About 40% of the neonates were preterm and rate of neonatal death was originate to be 12%. In the present study, percentage of fetal deaths due to jaundice among total perinatal deaths was 12%. Parveen *et al.* also reported low birth weight as the most common adverse fetal outcomes [15]. This can be attributed to the low levels of hemoglobin among mothers, as it limits the oxygen supply to the fetus which results in restriction of intrauterine growth and low birth weight [16]. Bora *et al.* reported the significant association between fetal birth weight and anemia (mild and severe) [17].

CONCLUSION

Overall, hepatic abnormalities during pregnancies consequently have the adverse effect on both maternal and neonatal outcomes. Maternal anemia, thrombocytopenia, coagulopathy, and hyperbilirubinemia are also evident. Early diagnosis of these patients is crucial and may reduce the adverse outcomes of both mother and the new-born.

CONFLICT OF INTEREST

None declared.

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None.

REFERENCES

- Suresh I, Vijaykumar TR, Nandeesh HP. Predictors of fetal and maternal outcome in the crucible of hepatic dysfunction during pregnancy.

- Gastroenterol Res 2017;10:21-7.
2. International Classification of Diseases. 10th Revision, Clinical Modification (ICD-10-CM). Available from: <https://www.cdc.gov/nchs/icd/icd10cm.htm> [Last assessed on 2021 Jan 28].
 3. Cunningham G, Leveno KJ, Bloom SL, Hauth JC, Rouse DW, Spong CY. Hepatic, Gallbladder, and Pancreatic Disorders. In: Williams Obstetrics. 3rd ed. New York: McGraw-Hill; 2010.
 4. Choudhary N, Sen S, Varalakshmi K. A prospective study on pregnancy complicated with jaundice with special emphasis on fetomaternal outcome. *Int J Reprod Contracept Obstet Gynecol* 2017;6:5081-88. doi: 10.18203/2320-1770.ijrcog20175029.
 5. Sharma AV, John S. Liver Disease in Pregnancy. In: Stat Pearls Florida: Stat Pearls; 2019. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK482201> [Last accessed on 2019 Jul 30].
 6. Reddy MG, Prabhakar GC, Sree V. Maternal and fetal outcome in jaundice complicating pregnancy. *J NTR Univ Health Sci* 2014;3:231-33. doi: 10.4103/2277-8632.146598.
 7. Allen AM, Kim WR, Larson JJ, Rosedahl JK, Yawn BP, McKeon K, *et al*. The epidemiology of liver diseases unique to pregnancy in a US community: A population-based study. *Clin Gastroenterol Hepatol* 2016;14:287-94.e1. doi: 10.1016/j.cgh.2015.08.022, PMID 26305066.
 8. Satia MN, Jandhyala MA. Study of fetomaternal outcomes in cases of jaundice at a tertiary care center. *Int J Reprod Contracept Obstet Gynecol* 2016;5:2352-57.
 9. Black RE, Victora CG, Walker SP, Bhutta ZA, Christian P, de Onis M, *et al*. Maternal and child undernutrition and overweight in low-income and middle-income countries. *Lancet* 2013;382:427-51.
 10. Stephen G, Mgongo M, Hussein Hashim T, Katanga J, Stray-Pedersen B, Msuya SE. Anaemia in pregnancy: Prevalence, risk factors, and adverse perinatal outcomes in Northern Tanzania. *Anemia* 2018;2018:1846280. Article ID 1846280. doi: 10.1155/2018/1846280, PMID 29854446.
 11. Bakrim S, Motiaa Y, Ouarour A, Masrar A. Hematological parameters of the blood count in a healthy population of pregnant women in the Northwest of Morocco (Tetouan-M'diq-Fnideq provinces). *Pan Afr Med J* 2018;29:205. doi: 10.11604/pamj.2018.29.205.13043, PMID 30100959.
 12. Lata I. Hepatobiliary diseases during pregnancy and their management: An Update. *Int J Crit Illn Inj Sci* 2013;3:175-82. doi: 10.4103/2229-5151.119196, PMID 24404454.
 13. Roncaglia N, Trio D, Roffi L, Ciarla I, Tampieri A, Scian A, *et al*. Intrahepatic cholestasis in pregnancy: Incidence, clinical course, complications. *Ann Ostet Ginecol Med Perinat* 1991;112:146-51. PMID 1812797.
 14. Shinde NR, Patil TB, Deshpande AS, Gulhane RV, Patil MB, Bansod YV. Clinical profile, maternal and fetal outcomes of acute hepatitis E in pregnancy. *Ann Med Health Sci Res* 2014;4(Suppl 2):S133-9. doi: 10.4103/2141-9248.138033, PMID 25184080.
 15. Parveen T, Begum F, Akhter N. Feto-maternal outcome of jaundice in pregnancy in a tertiary care hospital. *Mymensingh Med J* 2015;24:528-36. PMID 26329951.
 16. Stangret A, Wnuk A, Szewczyk G, Pyzlak M, Szukiewicz D. Maternal hemoglobin concentration and hematocrit values may affect fetus development by influencing placental angiogenesis. *J Matern Fetal Neonatal Med* 2017;30:199-204. doi: 10.3109/14767058.2016.1168395, PMID 27076257.
 17. Bora R, Sable C, Wolfson J, Boro K, Rao R. Prevalence of anemia in pregnant women and its effect on neonatal outcomes in Northeast India. *J Matern Fetal Neonatal Med* 2014;27:887-91. doi: 10.3109/14767058.2013.845161, PMID 24041147.