

Maternal and perinatal outcome in HELLP syndrome at VIMS, Ballari

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ABSTRACT

Objective: The aim of this study was to ascertain the presentation, diagnosis, severity and complications of HELLP syndrome and evaluation of the maternal and fetal outcome. **Methods:** Pregnancy induced hypertension (PIH) between 1st February 2016 and 31st January 2017 was included in the study. Diagnosed cases of HELLP syndrome were classified according to Tennesse criteria after assessing inclusion and exclusion criteria. The analysis of the data was done on all the patients diagnosed with HELLP syndrome. **Results:** The incidence of HELLP syndrome during the period was 1.14% of total deliveries and 3.82% of pregnancy induced hypertension (PIH). Majority (50%) patients belong to age group of 21 to 25 years. Mean age was 22.5 years. Most of them were primigravida (55%). Majority (52.5%) was in 36-40 weeks gestation and mean gestational age was 33.6 weeks. Head ache (56.25%) was the most common imminent symptom. Most of symptoms were nonspecific like malaise (50%), edema (45%), vomiting (20%) and epigastric pain (7.5%). Out of 80 patients of HELLP syndrome 19 delivered by LSCS and 61 delivered vaginally. Ascites (26.25%), PPH (25%) and placental abruption (22.5%) were the most common maternal complications in HELLP syndrome followed by acute renal failure (18.75%), pulmonary edema (12.5%), DIC (6.25%) and cerebrovascular accidents (6.25%). Maternal mortality in our study was 11.25%. Perinatal mortality was 41.25%. **Conclusion:** The reason for higher morbidity in our study is delay in identifying the problem by referring doctors. Earlier diagnosis and intervention improves maternal and perinatal outcome.

Keywords: HELLP, maternal mortality, perinatal mortality.

The HELLP syndrome is a serious complication in pregnancy characterized by hemolysis, elevated liver enzymes and low platelet count occurring in 0.5 to 0.9% of all pregnancies and in 10–20% of cases with severe preeclampsia. The HELLP syndrome is a serious condition associated with substantial risk for the mother and her fetus. A wide range of complications may arise

and the condition represents diagnostic and therapeutic problems; timing and method of delivery are important¹.

The term HELLP syndrome was coined by Weinstein in 1982 and defined by the presence of all of the three following criteria: hemolysis (characteristic peripheral blood smear, serum lactate dehydrogenase ≥ 600 IU/L, total serum bilirubin ≥ 1.2 mg/ml), elevated liver enzymes

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(serum aspartate aminotransferase ≥ 70 IU/l), and low platelet count ($<100,000/l$). Partial HELLP syndrome (PHS) was defined by the presence of one or two features of HELLP, whereas in complete HELLP all three features are present².

The diagnosis of HELLP syndrome may become very challenging because the patients may present with vague symptoms like nausea, vomiting, headache, malaise or flu-like symptoms. This leads to misdiagnosis of HELLP syndrome with various mild conditions like viral hepatitis to serious life-threatening conditions like acute fatty liver of pregnancy³. This study was undertaken with the objective to ascertain the presentation, diagnosis, severity, and complications of HELLP syndrome and evaluation the maternal and fetal outcome.

Materials and methods

After the approval of Institutional Ethical Committee and proper counseling and consent of the patients, this prospective observational study was conducted at Department of OBG, VIMS, Bellary in diagnosed cases of HELLP syndrome from 1st February 2016 to 31st January 2017. Diagnosed cases of HELLP syndrome were classified according to Tennessees criteria after assessing inclusion and exclusion criteria. Mainly women with eclampsia and severe pre-eclampsia were investigated. The analysis of the data was done on all the patients diagnosed with HELLP syndrome. Associated symptoms in the mother noted. Maternal morbidity in terms of pulmonary edema, ascites, acute renal failure, DIC, placental abruption, vision loss and maternal mortality noted. The perinatal outcome in terms of the IUFD, stillbirths, NICU admissions and early neonatal death were also noted.

Inclusion criteria

1. Women with HELLP syndrome using the Tennessees criteria

Criteria -

Complete HELLP Syndrome

- Platelet count $\leq 1,00,000/cu.mm$
- Serum LDH ≥ 600 IU/L
- Elevated liver enzyme – AST ≥ 70 IU/L

Partial HELLP Syndrome - If one or two of the above criteria present.

2. Any gestational age
3. Puerperal cases were also included

Exclusion Criteria

- 1) Known case of hepatic disease, hemolytic anaemia,
- 2) platelet disorders.
- 2) Chronic hypertension in pregnancy.
- 3) Chronic renal diseases.

The clinical data included were value of pulse, blood pressure, respiratory rate and urine output. While the laboratory evaluations recorded were serial measurement of complete blood cell count, coagulation profile (PT, APTT), liver function tests (Serum bilirubin, ALT, AST, and LDH), and renal function tests (Blood Urea, Serum Creatinine, and Serum Uric Acid). Disseminated intravascular coagulation (DIC) was diagnosed by presence of low platelets, prolonged PT and APTT, INR and D-dimer.

Abdominal ultrasonography was done when a subcapsular liver hematoma was suspected. Ophthalmoscopic examination was done in cases of severe pre-eclampsia and eclampsia. Magnesium Sulphate was used for seizure prevention and control. The diastolic BP was maintained between 90 and 100 mm Hg using Labetalol (oral or parental) or Nifedepine (oral). Blood and blood products (FFP & platelets) were used to correct coagulation abnormalities or anemia as needed. Antenatal steroid was given for fetal lung maturity if gestational age is less than 32 weeks. Fetal surveillance was done by modified Bio Physical Profile and Doppler. Decision to prolongation of pregnancy or termination was judged by duty doctor and Unit chiefs depending on maternal and fetal condition. Patients were managed in High Dependency Unit. Patients were shifted to ICU if mechanical ventilation was required or was hemodynamically unstable despite the treatment.

Statistical analysis data obtained was collected and analyzed statistically by proportions and tests of significance. Statistical significance was set at $p < 0.05$.

Results

During the study period, there were a total of 6983 deliveries including 2094 women with PIH. A total of 80 cases of HELLP syndrome were diagnosed and managed. Incidence of HELLP syndrome was 1.14 % of the total deliveries and 3.82 % of the PIH patients. Severe Pre-eclampsia were 65(81.25%), eclampsia 10(12.5%) and normotensive were 5 (6.25%) cases. In PIH patients

Table1: Shows various maternal complaints in HELLP syndrome

Complaints	Complete HELLP	Partial HELLP	No	Percentage
Head ache	25	20	45	56.25 %
Malaise	24	16	40	50%
Edema	20	16	36	45%
Vomiting	10	6	16	20%
Haematuria	12	3	15	18.75%
Convulsions	6	4	10	12.5%
Epigastric pain	5	1	6	7.5%
Visual Symptoms	2	0	2	2.5%

45(2.14%) were complete and 35(1.67%) partial HELLP syndrome. Majority (91%) was of lower socio-economic

Table 2: Admission parameters in our patients were divided according to Mississippi classification

Admission Parameters	Class 1	Class 2	Class 3
Platelet count (/μL) (Mean ± SD)	0.39 ±0.07	0.88 ± 0.21	1.02 ± 0.16
LDH (IU/L) (Mean ±SD)	1665 ± 655	1011.17 ± 422.4	900.59 ± 347.8
AST (IU/L) (Mean ±SD)	468.4 ± 68.3	178.6 ± 41.03	169.24±69.3

status and was emergency admission (80%). Nearly 2/3 belonged to rural places. Fifty percent belong to age group of 21 to 25 years and twenty six percent to 18 to 20

Table 3: Maternal complications in HELLP syndrome

Maternal Complications	Complete HELLP	Partial HELLP	Total
Ascites	15	6	21(26.25%)
PPH	14	6	20 (25%)
Placental abruption	12	6	18(22.5%)
Acute Renal Failure	12	3	15(18.75%)
Jaundice	8	6	14(17.5%)
Pulmonary Edema	6	4	10 (12.5%)
DIC	4	1	5 (6.25%)
Intra Cranial Bleeding	4	1	5 (6.25%)
Sepsis	3	1	4(5%)
Visual Disturbance	2	0	2 (2.5%)
Post Partum Collapse	2	0	2 (2.5%)

years. The mean age was 22.5 years. Primigravida were 44% and Gravida 2 were 25%. Post partum cases were 3 (3.75%). But, there is no statistical significance of maternal age and gravidity between complete and partial HELLP Syndrome (P = 0.33).

Majority (52.5%) of patients were between 36-40 weeks of gestation. Mean gestational age in this study is 33.6 weeks. Again, there is no statistical significance of gestational age among the different classes of HELLP syndrome (P=0.2333).

Out of total 80 patients of HELLP syndrome, 9 had spontaneous vaginal deliveries and 7 patients were taken for caesarian straight away. Remaining 64 patients were induced, out of these 52 delivered vaginally and 12 patients underwent emergency LSCS for various obstetric indications. Atonicity was the cause of post partum haemorrhage (PPH) and majority were treated by uterotonics and two cases of PPH following LSCS were controlled by Bakri balloon (Table 3).

Acute renal failure (ARF) patients had gradual rise of renal parameters (blood urea and serum creatinine) for 7-14 days with oliguria followed by improvement in renal parameters in the 3rd week. Initially these patients were managed using colloid plasma substitute with intravenous (IV) furosemide. They were monitored with serial renal function test (RFT) and serum electrolytes. If values persistently

increasing, such patients were subjected to dialysis. In our study four number of ARF patients underwent dialysis and recovered (Table 3). Ten patients had pulmonary oedema, managed with oxygen, IV furosemide and ionotropes (dopamine or noradrenaline). All these patient required ventilator support, seven gradually recovered and three patient died. On ophthalmoscopic examination one case had retinal detachment and regained vision 24 hours after delivery. In other case, fundoscopy showed papilloedema, and gained vision over a period of one week. All Maternal complications were more in complete HELLP syndrome. But there was no statistical significance.

There were a total of 9(11.25%) maternal deaths. Most of them were complete HELLP syndrome. Eclampsia accounted for 7 (77.77%) and severe pre-eclampsia for 2 (22.22%) of maternal death. Pulmonary edema, DIC and cerebrovascular accident each accounted for 33.33% causes of maternal death. All these cases were

Table 4: Perinatal outcome in HELLP syndrome

Perinatal outcome	Complete HELLP	Partial HELLP	Total No (%)	P Value
IUD	15	6	21(26.25%)	0.421
Still Birth	4	1	5 (6.25%)	0.352
Live Baby	22	32	54 (67.5%)	0.921
Preterm	16	20	36 (45%)	0.843
LBW				
[Mean in Kgs]	2.27	2.49	2.38	0.167
NICU admission	22	8	30 (37.5%)	0.095
Early neonatal death	5	2	7 (8.75%)	0.561
Take home baby	27	20	47 (58.75%)	0.565

emergency admissions referred from periphery in a moribund status.

Perinatal mortality was 33(41.25%). Majority of live baby and take home baby were in partial HELLP

Table 5: Perinatal Complications among live born

Complications	Complete HELLP	Partial HELLP	Total[n=54] [Live born]
Low APGAR score at 5 min	6	2	8 (14.8%)
IVH	3	2	5 (9.25%)
Hypoxic encephalopathy	3	2	5 (9.25%)
Icterus	7	5	12 (22.22%)

syndrome. But none of the perinatal outcome is statistically significant (Table 4). There was no correlation between perinatal complications and HELLP Syndrome severity (Table 5).

Discussion

Abildgaard U et al¹ reported the HELLP prevalence to range from 10-20%. Abbade et al² reported incidence of 12.9% . Rekha Gurumurthy et al⁴ found incidence of HELLP to be as high as 22.22%. In our study 1.14 % of the total delivered patients developed HELLP syndrome, among them 0.64 % had complete and 0.5% had partial HELLP. In PIH patients incidence was 3.82% among them 2.14% had complete and 1.67% had partial HELLP.

In Jayashree Mulik et al⁵ most of patients in HELLP syndrome were in 21-25 year age group; mean age was 24.5 yrs while in Aswini Mallappasera et al³ mean

maternal age was 21.4 yrs. Our study is in full agreement with Jayashree Mulik et al⁵.

Jayashree Mulik et al⁵ series most of the patients with HELLP were primigravida (85.2%); which is a little higher than those reported by Ashwini Mallappasera et al³(57.89%) and Campos A et al¹⁰ (64.3%). In our study also most of patients with HELLP were primigravida (44%).

Gasem et al⁷ and Gupta et al⁸ reported mean gestational age of 32.4 and 32.2 weeks. Jayashree Mulik et al⁵ reported maximum patients of gestation age between 33-36 wks (mean 34.6 weeks). This early presentation results in higher chances of prematurity. In our study majority were between 36-40 weeks of gestation. Mean gestational age was 31.5 weeks.

Headache, edema feet, puffiness of face, blurring of vision and abnormal weight gain were the major presenting symptoms in both HELLP and partial HELLP respectively. This finding is almost similar to all other studies^{3,4,5,6}. Almost all the deranged lab test results reported by peers in other studies were corroborated with similar findings in our study^{3,4,5,6}.

The nonspecific presentation of HELLP syndrome overlaps with a number of medical disorders. Therefore, the differential diagnosis of HELLP syndrome should include any of the conditions such as upper respiratory infection, hepatitis, cholecystitis, pancreatitis, acute fatty liver of pregnancy (AFLP), or immune thrombocytopenic purpura (ITP). In the absence of hypertension or proteinuria, these cases are misdiagnosed as having other conditions. Conversely, women with other conditions such as TTP, HUS, systemic lupus erythematosus, sepsis, or catastrophic antiphospholipid antibody syndrome may be erroneously diagnosed as having HELLP syndrome. Hence the most experienced clinician can face a difficult diagnostic challenge. Therefore, sincere efforts should be made to attempt to accurate diagnosis, so that management may be different among these conditions. It is important to emphasize that these patients have a variety of atypical signs and symptoms, none of which are diagnostic of severe preeclampsia. Hence, pregnant women with probable preeclampsia presenting with atypical symptoms should have a complete blood count, a platelet count, and liver enzyme determinations

irrespective of maternal BP findings³.

Jayashree Mulik et al⁵ also reported increased rate of platelet and blood products transfusion in patient with HELLP syndrome. In our study more than 50% patients with HELLP and partial HELLP received blood transfusion. FFP was given to 55.55% patient of HELLP and 25% patients with partial HELLP. Our study is in full agreement with Gupta et al⁸. The early termination of pregnancy in HELLP syndrome was very high in several studies, including ours^{4,5,6}. In most of studies ascites, pulmonary edema, placental abruption, PPH, DIC, ARF, cerebral haemorrhage, renal complications, and retinal detachment were seen. Our study is in full agreement with the study of Rekha Gurumurthy et al⁴, Jayashree Mulik et al and Sowjanya Kumari et al⁶ in maternal complications^{4,5,6}.

Jayashree Mulik et al⁵ reported 3.8%, Sowjanya Kumari et al⁶ reported 4.5% and Isler CM et al¹⁰ reported 7.8% of maternal mortality^{5,6,10}. Cerebral haemorrhage, pulmonary edema, cardiac failure, renal failure and multi organ dysfunction were the cause of death in their series. In our study a total of 9(11.25%) maternal death occurred. In our study CVA, DIC and PPH equally contributed to maternal death. All cases were complete HELLP syndrome. All cases were referred from outside in a moribund status. High mortality in our series is because of delay in identifying the problem by referring doctors and late referral.

Sowjanya Kumari et al⁶ and Gupta T et al⁹ reported 13.53% and 16% IUD⁶ where as in Rekha Gurumurthy et al⁴ series incidence of intra uterine death (IUD) was 33.3% which was high. Abruption was the cause in 88% of these IUD. In our study 21(26.25%) were IUD and 5 were still birth. Abruption was associated with (12) 57.14% of IUD and 60% of still birth. Our study is comparable to Rekha Gurumurthy et al⁴.

Rekha Gurumurthy et al⁴ reported 16.6% early neonatal death due to sepsis, respiratory distress and birth asphyxia. Sowjanya Kumari et al⁶ reported 9% early neonatal death due to birth asphyxia and prematurity. In our study early neonatal death accounted for 8.75% similar to Sowjanya Kumari et al. Causes were birth asphyxia (28.5%), intra ventricular hemorrhage (14.2%), respiratory distress syndrome (28.5%) and meconium aspiration syndrome (28.5%).

The perinatal mortality in Sowjanya Kumari et al⁶ is 35.33% is comparable to our study i.e., 41.25%. This high mortality is associated with preterm birth and its complications and abruption of placenta. These infants have high rates of respiratory distress syndrome, bronchopulmonary dysplasia, intracerebral hemorrhage and necrotizing enterocolitis. The reason for higher morbidity in our study is delay in identifying the problem by referring doctors. Most of the cases were referred to us at a late stage, biochemical changes identified HELLP syndrome and termination of pregnancy was planned upon admission to the hospital.

Conclusion

HELLP syndrome is associated with poor outcome for the mother and fetus. Nonspecific symptoms and signs results in miss diagnosis and inappropriate management. Anticipation of the problem and early identification of HELLP syndrome will reduce the complications. Definitive treatment for HELLP syndrome is delivery of the baby. Timely diagnosis and management of HELLP syndrome by induction and delivery by vaginal route or by cesarean section is beneficial and prevents complications in mother and fetus.

Early registration and regular ANC plays a major role in early diagnosis and to reduce complications like HELLP syndrome. Early detection, prompt referral, better transport facility, appropriate and timely intervention, availability of life saving facilities like mechanical ventilators, dialysis equipment and blood products like FFP, Platelets, Packed cell transfusion at tertiary care centers will significantly reduce the maternal and fetal morbidity and mortality. HELLP syndrome should be treated at tertiary care centers. The obstetricians at any level should be attentive, alert and need to improve quality care and make efforts for early identification even at its atypical presentation and provide skilled management techniques. Doctors should promote early and regular antenatal care at community health centers and ensure availability of new born care that will provide better maternal and perinatal outcome.

The global mortality rate of HELLP syndrome is very high. That's why it's critical for expecting mothers to be aware of the condition and its symptoms so that they can receive early diagnosis and treatment. Doctors should enhance their skills in antenatal care to identify the high

risk factors at primary and tertiary care centers. This study uncovers only the tip of the iceberg as this is only a hospital based study. HELLP syndrome in non PIH cases was not evaluated.

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