The Study of Serum Uric Acid as a Biochemical Indicator for Maternal and Fetal Outcome in Gestational Hypertension

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Abstract
Introduction: Hypertensive pregnancy disorders are among the most common medical disorders during pregnancy and are considered a significant cause of maternal and fetal morbidity and mortality.

Aim: The aim of the study was to study of serum uric acid as a biochemical indicator for maternal and fetal outcome in patients with gestational hypertension (GHT).

Materials and Methods: This prospective study was conducted in the Department of Obstetrics and Gynaecology, Government Theni Medical College and Hospital, Tamil Nadu, from June 2018 to June 2019 in 200 pregnant women with GHT. Informed consent was obtained from the patients. Results were analyzed statistically and discussed below.

Results: Out of 200 pregnant females, the mean age was 24.2 ± 4.5 years. Ninety-four patients were prim gravida, 67 patients were gravida 2, and 30 patients were gravida 3. Ninety-six had GHT with associated abnormalities, 90 were diagnosed with preeclampsia, and 47 had non-severe preeclampsia. In contrast, 43 had severe pre-eclampsia, eight patients had developed eclampsia, while the remaining six subjects were diagnosed with HELLP syndrome. About 66% had serum uric acid level ≤6 mg/dl and 34% had serum uric acid level ≥6 mg/dl. In subjects with normal uric acid level, 31.6% and 11.3% of subjects had non-severe preeclampsia and severe preeclampsia. In a subject with increased uric acid level, 46 patients had severe proteinuria. Preterm delivery was more commonly seen in women with serum uric acid higher than 6 mg/dl.

Conclusion: Our study concludes that the measurement of serum uric acid levels after 20 weeks of gestation is an excellent diagnostic and prognostic tool to assess fetal outcomes. This study shows that the estimation of serum uric acid levels in pregnancies complicated by hypertension and preeclampsia helps assess the severity of the disease.

Key words: Gestational hypertension, Maternal, Fetus, Uric acid

INTRODUCTION
Hypertensive pregnancy disorders (HDP) are among the most common medical disorders during pregnancy and are considered a significant cause of maternal and fetal morbidity and mortality. In developing countries, HDP ranks second only to anemia with approximately 7–10% of all pregnancies complicated by some form of hypertensive disorder and lead to various maternal and fetal complications.[1] In India, the incidence of preeclampsia, as recorded from hospital statistics, varies widely from 5% to 15%, while eclampsia is about 1.5%.[2] Strangely, the exact etiopathogenesis for HDP, including preeclampsia and eclampsia, remains obscure and presents an exciting mystery in obstetric practice.

The diagnosis of preeclampsia is based on the presence of the following clinical features: Blood pressure (BP) 140/90 mmHg after 20 weeks of gestation AND proteinuria 2300 mg/24-h or +1 with the dipstick.[3] There may be other associated abnormalities reported which increase the likelihood of preeclampsia occurrence such as elevated serum creatinine (new-onset), platelet count <100,000/pL, hepatic enzyme abnormalities, persistent headache with or without cerebral or visual disturbances, and epigastric pain. Preeclampsia with the presence of seizures is known as eclampsia.[4]

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The elevated uric acid level in maternal blood, presumably due to decreased renal excretion, is frequently found in women with preeclampsia. Of the several hypothesized factors for elevated uric acid in patients with preeclampsia, the following appear to be most intriguing: Abnormal renal function, increased tissue breakdown, acidosis, and increased activity of the enzyme xanthine oxidase/dehydrogenase.\[5\]

An association of uric acid elevation in clinically evident pre-eclampsia has been known since 1917.\[6\] Several studies have correlated the rise in uric acid with the severity of the preeclampsia.\[7\] Although hyperuricemia does correlate with maternal morbidity, there is an even stronger association of increased uric acid with risk for small or low birth weight infants and with overall fetal mortality.\[8-10\]

**Aim**

The aim of the study was to study of serum uric acid as a biochemical indicator for maternal and fetal outcome in patients with gestational hypertension (GHT).

**MATERIALS AND METHODS**

This prospective study was conducted in the Department of Obstetrics and Gynaecology, Government Theni Medical College and Hospital, Tamil Nadu, from June 2018 to June 2019 in 200 pregnant women with GHT.

**Inclusion Criteria**

All pregnant women suffering from GHT, preeclampsia, and eclampsia >20 weeks of gestation were included in the study.

**Exclusion Criteria**

Normotensive pregnant females, patients with hypertension at 20 weeks of gestation (chronic hypertension), patients with chronic renal disease, patients suffering from diabetes mellitus, diagnosed patients of hepatic dysfunction, patients suffering from gout, patients with epilepsy, patients with thyroid dysfunction, patients with cardiac disease, patients with leukemia’s, patients with pancreatitis, and patients with hemolysis were excluded from the study.

After obtaining informed written consent from all the study subjects or their attendants (in case of unconscious patients, or patients unable to provide consent due to their condition), relevant data were documented in a pre-defined datasheet. Maintaining all aseptic precautions, blood samples were collected from all subjects to estimate serum uric acid concentration and other parameters.

The data were analyzed using the Statistical Package for the Social Sciences (IBM SPSS Inc.). Categorical variables were presented as percentages, and continuous variables were presented as mean/median. The association between categorical variables was tested using the Chi-square test. A P-value (two-tailed) of <0.05 was considered statistically significant.

**RESULTS**

In this study, 200 pregnant patients were included in the mean age of 24.2 ± 4.5 years. Ninety-four patients were prim gravida, 67 patients were gravida 2, and 30 patients were gravida 3 [Figure 1]. The mean BMI was 24.1 ± 3.3 kg/m².

The earliest presentation of GHT in this study was in 25th week of gestation (24 weeks 5 days) while the latest presentation was in the 40th week of gestation (39 weeks 3 days) [Figure 2].

In 200 patients, 96 had GHT with associated abnormalities. Ninety of the 200 subjects were diagnosed with preeclampsia, 47 had non-severe preeclampsia while 43 had severe preeclampsia. Eight of the 200 subjects developed eclampsia while the remaining six subjects were diagnosed with HELLP syndrome and were considered to be in imminent risk of eclampsia [Figure 3].

<table>
<thead>
<tr>
<th>Complications</th>
<th>≤6</th>
<th>≥6</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Intra-uterine death</td>
<td>0</td>
<td>10</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>Intra-uterine growth retardation</td>
<td>13</td>
<td>12</td>
<td>0.128</td>
</tr>
<tr>
<td>Intra-cranial hemorrhage</td>
<td>0</td>
<td>0</td>
<td>n/a</td>
</tr>
<tr>
<td>Pulmonary edema</td>
<td>0</td>
<td>0</td>
<td>n/a</td>
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<tr>
<td>Abruption</td>
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<td>2</td>
<td>0.046</td>
</tr>
<tr>
<td>Pre-term/low birth weight</td>
<td>66</td>
<td>36</td>
<td>0.701</td>
</tr>
<tr>
<td>HELLP syndrome</td>
<td>1</td>
<td>5</td>
<td>0.009</td>
</tr>
<tr>
<td>Spontaneous expulsion</td>
<td>0</td>
<td>1</td>
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</tr>
<tr>
<td>Maternal death</td>
<td>0</td>
<td>0</td>
<td>n/a</td>
</tr>
</tbody>
</table>

**Figure 1: Distribution of obstetrics score**
Out of 200 patients, 66% had serum uric acid level ≤6 mg/dl, and 34% had serum uric acid level ≥6 mg/dl [Figure 4].

In comparing gestational age with serum uric acid level 28 patients with gestational age <34, 14 patients between 34 and 36, 91 patients greater than 36 had serum uric acid level ≤6 mg/dl. Twenty-one patients with gestational age <34, 11 patients between 34 and 36, 35 patients greater than 36 had serum uric acid level ≥6 mg/dl [Figure 5].

Out of 200 patients, 46 patients with serum uric acid level greater than 6 mg/dl had severe proteinuria [Figure 6].

In subjects with normal uric acid level, 31.6% and 11.3% of subjects had non-severe preeclampsia and severe preeclampsia ($P < 0.0001$) [Figure 7].

In abnormal uric acid level patients, intrauterine death and HELLP syndrome are observed as statistically significant maternal and fetal complications [Table 1].

**DISCUSSION**

Uric acid is filtered by the renal glomeruli, absorbed by the first part of the proximal convoluted tubule, with a further secretion and reabsorption phase. Ten percent of the filtered urate is excreted in the urine. During pregnancy, uric acid clearance increases from 6–12 mL/min to 12–20 mL/min, with a 25% decrease in blood concentration.

Lim et al. recently documented that mean serum uric acid levels were significantly higher in women with transient hypertension, preeclampsia, and superimposed preeclampsia. In Redman’s study, the best correlation to predict perinatal mortality was the serum uric acid level at 28–32 weeks’ gestation. A serum uric acid level of ~420/1 mol/L only predicted perinatal mortality of 6–9% in their group with term pregnancies. Plouin et al. documented a poor perinatal outcome (including stillbirths and neonatal deaths) in pregnancies...
complicated by preeclampsia and predicted by serum uric acid levels.

In our study, most hypertensive mothers were in the age group of 20–35 years. Number Indian authors noticed similar age incidence.[21,22]

In his study, Robert et al.[21] showed that 6% of patients had HELLP syndrome, and 16% had eclampsia. In our study, subjects with normal uric acid level, 31.6% and 11.3% of subjects had non-severe preeclampsia and severe preeclampsia, respectively.

Magann et al.,[23] there is a positive correlation between increasing/raised SUA level and increased incidence of PNM, stillborn. According to Robert et al.,[21] the preterm risk is increased in groups of patients with hypertension proteinuria hyperuricemia. Even in the absence of proteinuria, hyperuricemia is a significant risk factor for preterm birth.

**CONCLUSION**

Our study concludes that the measurement of serum uric acid levels after 20 weeks of gestation is an excellent diagnostic and prognostic tool to assess fetal outcomes. This study shows that the estimation of serum uric acid levels in pregnancies complicated by hypertension and preeclampsia helps assess the severity of the disease and identify life-threatening maternal and fetal complications as a result of preeclampsia/eclampsia. Since serum uric acid is a very simple analytical tool that can be readily performed at any biochemical laboratory, this is a very cost-effective method to gauge GHT complications and improve maternal and perinatal outcomes.

**REFERENCES**

Uric acid is as important as proteinuria in identifying fetal risk in women with gestational hypertension. Hypertension 2005;46:1263-9.


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