Serum uric acid level in normal pregnant and preeclamptic ladies: a comparative study

Pramanik T,1 Khatiwada B², Pradhan P³

¹Department of Physiology, Nepal Medical College, Jorpati, Katmandu, Nepal, ²Deulekh, PHC, Bajhang, Nepal, ³ Department of O&G, Nepal Medical College and Teaching Hospital, Jorpati, Katmandu, Nepal

Corresponding author: Dr. Tapas Pramanik, Associate Prof., Dept. of Physiology, Nepal Medical College, Kathmandu, Nepal. Email: drpramanik@hotmail.com

ABSTRACT:

Preeclampsia is a serious pregnancy complication characterized by hypertension, proteinuria with or without pathological edema. According to some studies, serum uric acid lacks sensitivity and specificity as a diagnostic tool whereas another group of the researchers indicated uricemia as a predictor of preeclampsia in pregnant ladies. The present study was designed to assess whether serum uric acid can be used as a biochemical indicator or not in preeclamptic patients. Pre-eclamptic patients admitted in Nepal Medical College Teaching Hospital from June 2012 to June 2013 were included in this study. Age matched normal healthy pregnant ladies served as control. The record of their blood pressure and serum uric acid level was evaluated. Results showed significantly high blood pressure [SBP 149.42+12.35 vs 109.00±7.93 mm Hg; DBP 96.85±8.32 vs 72.5±7.10 mm Hg], and serum uric acid level [6.27±1.37 vs 4.27±0.61 mg/d1] in pre-eclamptic patients compared to their healthy counterparts. Uric acid is a terminal metabolite of the degradation of nucleotides, which increases their blood levels in patients with preeclampsia likely results from reduced uric acid clevel form filminished glomenular filtration, increased tubular readsorption and decreased secretion. Results of the present study indicated association of elevated serum uric acid level with preeclampsia which could be used as a biochemical indicator of preeclampsia in pregnant women.

Key words: preeclampsia, serum uric acid

INTRODUCTION:

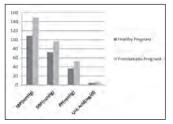
Preeclampsia is a serious pregnancy complication. It is a multi-system disorder characterized by hypertension (blood pressure ≥140/90 mm Hg), proteinuria (24-hr urinary protein ≥ 0.3 g) with or without pathological edema, beyond 20th week of gestation in previously normotensive and nonproteinuric woman. This pregnancy specific syndrome can affect virtually every organ system.1 Gestational hypertension is a common first clinical presentation of preeclampsia. Preeclampsia is associated with uricemia.2 Nevertheless; some studies reported that uricemia is not a consistent predictive factor of preeclampsia.3 Another study showed no significant difference in serum uric acid level between normal and mild preeclamptic women.4 Furthermore, Lim et al reported serum uric acid has been found to lack sensitivity and specificity as a diagnostic tool,5 whereas most of the researchers indicated uricemia as a predictor of preeclampsia.6 The present study was designed to assess/evaluate whether serum uric acid could be used as a biochemical indicator in preeclamptic patients. For this purpose, the serum uric acid level was estimated in pregnant women with and without preeclampsia admitted in Nepal Medical College Teaching Hospital.

MATERIALS AND METHODS

The present study is a case control one. The data of Preeclamptic patients admitted in Nepal Medical College Teaching Hospital from June 2012 to June 2013 was taken for this study. Pregnant ladies with blood pressure ≥140/90 mm Hg [taken on at least two occasions 6 hours apart7] and proteinuria [24-hr urinary protein ≥ 0.3g or dipstick +1 or more7] beyond 20th week of gestation were considered as pre-eclamptic clinically. Within the aforesaid period 35 cases (n=35) of pre-clapsia were admitted. Age matched normal/healthy pregnant ladies served as control (n=20). Patients with the history of urinary tract infection, renal disease, diabetes mellitus and dehydrated patients were excluded in this study. Verbal consents of the patients were taken before the collection of blood by venipuncture for diagnostic purpose. The records of their blood pressure and serum uric acid level were evaluated. Results were compared and analyzed statistically by using student's t test.8

RESULTS

The number of pre-eclamptic patients was 35 with age ranging from 19-38 years (mean 28.11 years). The range of blood pressure recorded in them was 140/90 mm Hg to 190/120 mm Hg. The number of normal pregnant ladies was 20 with age ranging from 19-30 years (mean



22.65 years). The range of blood pressure recorded in them was 90/58 mm Hg to 122/88 mm Hg.

Fig. 1 Blood pressure and uric acid level in preeclamptic patients and healthy pregnant ladies.

Results presented in table-1 and fig-1. indicated significantly high blood pressure in pre-eclamptic patients compared to healthy pregnant ladies [SBP 149,42±12.35 vs 109.00±7.93 mm Hg; DBP 96.85±8.32 vs 72.5±7.10 mm Hg] Serum uric acid level was also significantly high in pre-eclamptic patients compared to their healthy counterparts [6.27±1.37 vs 4.27±0.61 mg/dl] (Table-1)

DISCUSSION:

Although the number of pre-celamptic patients in the chosen study period was 35, the number of normal pregnant ladies that served as control was limited to 20, as the uric acid level in most of the normal pregnant ladies was found to be consistently within normal lower range (about 4 mg/dl in our study). Punthumapol and Kittichotpanich(2008) also compared uric acid levels of 68 pre-celamptic patients with 36 normal pregnant women.4 Present study recorded high level of serum uric acid level in precelamptic patients (6.27 mg/dl vs 4.27 mg/ dl) which corroborated the findings of previous study.⁹

High blood pressure (\geq 140/90 mm Hg), proteinuria (24-hr urinary protein \geq 0.3g) and/or pathological edema, beyond 20th week of gestation in previously normotensive and non-proteinuric woman are the characteristic features of preeclampsia. Present study recorded elevated blood pressure (149.42)96.85 mmHg) in precelamptic patients, whereas blood pressures in normal pregnant ladies were within normal range (109.00/72.50 mm Hg).

Constant high blood pressure increases the level of vasoconstrictors like thromboxane A_2 , angiotensin II, endothelium I and decreases the level of vasodilators like prostaglandin I₂, prostaglandin E2, NO etc.¹⁰ As a result, there was increase in peripheral resistance and further increase of blood pressure.

Endothelial dysfunction in precelampsia increases capillary permeability leading to edema. Proteinuria due to leakage of protein from glomerular capillaries causes loss of protein resulting in decreased plasma colloidal osmotic pressure. Reduced plasma colloidal osmotic pressure causes edema in the victims of precelampsia.

As mentioned earlier, present study recorded high level of serum uric acid level in preeclamptic patients (6.27 mg/ dl) vs 4.27 mg/dl). Uric acid is formed by the breakdown of purines and by direct synthesis from 5-phosphoribosyl phosphate (5-PRP) and glutamine. The normal blood uric acid in humans is approximately 4 mg/dl. In the kidney, uric acid is filtered, reabsorbed and secreted. Normally, 98% of the filtered uric acid is reabsorbed and the remaining 2% makes up approximately 20% of the amount excreted. Remaining 80% comes from the tubular secretom^{11,12}

Uric acid is a terminal metabolite of the degradation of nucleotides, which increases their blood levels in patients with precelampsia-eclampsia, increasing its synthesis by damage and death of trophoblastic cells and proliferation.¹³

In pre-ecalamptic patients significant increase of pulse pressure was noticed (52.57 mm Hg vs 36.50mm Hg). Elevation of pulse pressure has shown to induce endothelial dysfunction in small vessels and is a possible antecedent of atherosclerosis¹⁴. It indicated reduction in arterial compliance.¹⁵ Elevated pulse pressure indicates hyperdymanic circulation that exerts more shearing force of blood on endothelium causing loss of more endothelium. Metabolism of this nucleoprotein of shredded endothelium may produce more uric acid.

Table-1: Blood pressure and uric acid level in preeclamptic patients and healthy pregnant ladies

Volunteers	SBP (mmHg)	DBP (mmHg)	PP (mmHg)	Uric Acid (mg/dl)
	Mean ± SD	Mean ± SD	Mean ± SD	Mean ± SD
Healthy pregnant (n=20)	109.00±7.93	72.50±7.10	36.5 <u>+</u> 7.07	4.27 <u>+</u> 0.61
Preeclamptic pregnant (n=35)	149.42 <u>+</u> 12.35*	96.85 <u>+</u> 8.32*	52.57 <u>+</u> 7.41*	6.27 <u>+</u> 1.37*

* = p ≤ 0.05; SBP = Systolic Blood Pressure; DBP = Diastolic Blood Pressure; PP = Pulse Pressure

Uricemia in preeclampsia likely results from reduced uric acid clearance from diminished glomerular filtration, increased tubular reabsorption and decreased secretion.1 Hyperuricemia in preeclampsia was once thought to result solely from reduced renal clearance, but levels of uric acid are now also thought to increase through increased uric acid production caused by trophoblast breakdown, cytokine release and ischemia. Uric acid can promote endothelial dysfunction, damage and inflammation, which leads to oxidation. 3

Results of the present study indicated that preeclampsia is associated with elevated serum uric acid level which could be used as a biochemical indicator of preeclampsia in pregnant women.

ACKNOWLEDGEMENTS

Authors are grateful to Prof. P Roychowdhury, Head, Dept of Physiology, and Dr. SB Rizyal, Principal, Nepal Medical College, for their help and support.

REFERENCES

- Lindheimer MD, Conrad K, Karumanchi SA: Renal physiology and disease in pregnancy. In Alpern RJ, Hebert SC, editors. Seldin and Giebisch's the kidney: physiology and pathophysiology (4th ed.). New York: Elsevier 2008a: 2339.
- Wu Y, Xiong X, Fraser WD, Luo ZC. Association of uric acid with progression to precelampsia and development of adverse conditions in gestational hypertensive pregnancies. *Amer J Hypertens* 2012; 25:711-17.
- Martin AC, Brown MA. Could uric acid have a pathogenic role in pre-eclampsia? Nat Rev Nephrol 2010; 6: 744-8.
- 4. Punthumapol C, Kittichotpanich B. Serum calcium,

magnesium and uric acid in preecclpsia and normal pregnancy. J Med Assoc Thai 2008; 91: 968-73.

- Lim KH, Friedman SA, Ecker JL, Kao L, Kilpatrick SJ. The clinical utility of serum uric acid measurements in hypertensive diseases of pregnancy. *Am J Obstet Gynecol* 1998; 178: 1067-71.
- Kang DH, Finch J, Nakagawa T, Karumanchi SA, Kanellis J, Granger J, Johnson RJ. Uric acid, endothelial dysfunction and pre-eclampsia: searching for a pathogenetic link. *J Hypertens* 2004; 22: 229-35.
- James DK, Steer PJ, Weiner CP, Gonik B, editors. High risk pregnancy management options (3rded.). New Delhi: Elsevier 2006; 414.
- Maharjan BK. Methods in biostatistics for medical students and research workers. 5th ed. New Delhi: Jaypee brothers medical publishers 1989; 141-46.
- Taefi A, Jamal AS, Delavari H. The role of serum uric acid in preeclapsia. J Family Reprod Health 2008; 2:159-62.
- Walsh SW. Preeclampsia: an imbalance in placental prostacyclin and thromboxane production. Amer J Obstetr Gynecol 1985; 152: 335-40.
- Barret KE, Berman SM, Boitano S, Brooks HL, editors. Ganong's Review of Medical Physiology (24th ed.). Mc Graw Hill Lange 2012. pp 13.
- Hames BD, Hooper NM. Instant notes on biochemistry. 2nd ed New Delhi : Viva Books Pvt Ltd 2003; 381.
- Vazquez-Rodriguez JG, Rico-Trejo El. Role of uric acid in preeclampsia-eclampsia. *Ginecol Obstet Mex* 2011; 79: 292-7.
- Dart AM, Kingwell BA. Pulse pressure A review of mechanisms and clinical relevance. J Amer Coll Cardiol 2001; 37: 975-84.
- Ryan SM, Waack BJ, Weno BL, Heistad DD. Increases in pulse pressure impairs acetylcholine induced vascular relaxation. *Amer J Physiol* 1995; 268: H 359- 63.