# Levels of microalbuminuria in prediction of pre-eclampsia: A hospital based study

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### Abstract

**Introduction:** Pre-eclampsia (PE) is defined as the new onset of hypertension and proteinuria during the second half of pregnancy. In the Indian scenario, it accounts for 44.44% of all cases of hypertensive disorders of pregnancy. Microalbuminuria is one the classic signs of pre-eclampsia.

The aim of the study was to determine microalbuminuria as predictor of pre-eclampsia.

**Methods:** A total of 145 pregnant women in their second trimester of pregnancy were included in the study. From all these subjects, clean catch morning urine sample (5ml) was collected in a sterile container. Urinary Creatinine and urinary microalbumin was estimated by kit based method on Vitros Fusion 5.1 FS fully automated analyzer from Ortho Clinical Diagnostic USA. The statistical significance of difference between mean values was assessed by Student's unpaired t-test.

**Result:** Pregnant women who developed pre-eclamsia were having significantly higher micro albumin level and Albumin Creatinine Ratio as compared to normotensive pregnant women.

**Conclusion:** To determine microalbuminuria as predictor of pre-eclampsia, further detailed study is required in large group of population.

Keyword: Pre-eclampsia, Microalbuminuria, Albumin Creatinine Ratio, BP, Pregnancy

### Introduction

Pre-eclampsia (PE) is a clinical syndrome defined as the new onset of hypertension and proteinuria during the second half of pregnancy. This clinical condition is diagnosed when the blood pressure at or above 140/90 mmHg, occurring on two occasions at least 6 hours apart, associated with proteinuria greater than 300mg/24 hours or greater than 1 gm/l in a random sample, after 20 weeks of gestation<sup>1</sup>. In developing nations, the incidence of the disease is reported to be 4%-18% and is the second most common obstetric cause of mortality accounting for 40,000 maternal deaths annually<sup>2,3</sup>. In the Indian scenario, the incidence of preeclampsia is 5.47% in primigravida and 2.8% in multigravida. It accounts for 44.44% of all cases of hypertensive disorders of pregnancy<sup>4</sup>. Proteinuria and hypertension are the most common manifestations of pre-eclampsia. PE is not only common and dangerous for both mother and baby, but also unpredictable in onset and progression, and is incurable constituting 12% to 18% of pregnancy related maternal deaths<sup>5</sup>.

Microalbuminuria is characterized by urinary albumin excretion above normal levels in the absence of clinically detectable nephropathy<sup>6,7,8</sup>. Microalbuminuria is noted to be present if urinary albumin is within the range of 30-300 mg/24 hrs<sup>9</sup>. Microalbuminuria is one the classic signs of preeclampsia. The presence of microalbuminuria in some otherwise symptom-free patient confirms that changes in renal function are present in whom pre-eclampsia would eventually develop. Early pregnancy levels of microalbuminuria can be used as predictors of preeclampsia with high negative predictive value<sup>10</sup>. Persistent microalbuminuria indicates a high probability of damage of the glomerular filtration capacity of the kidney and is of great diagnostic relevance in pregnancy as a possible predictor of developing PE.

Many researches have been done to find tests that would predict the risk of developing PE before the classical triad of symptoms appear<sup>11</sup>. But the predictive ability of individual test has varied widely and many tests simply detect early disease. It is clear that no test reliably predicts pre-eclampsia<sup>12</sup>. Further studies are going on to detect an effective and practical early predictor of pre-eclampsia, which will be of 'gold standard'<sup>13</sup>.

### Material and Methods

The present study was conducted by Department of Biochemistry, on pregnant females attending to antenatal clinic of Obstetrics and Gynecology department in Chatrapati Shivaji Subharti Hospital, Subharti Medical College, Meerut, UP, after obtaining ethical clearance from the institutional ethical committee. A total of 145 pregnant women in their second trimester of pregnancy were included in the study. Patients with acute anxiety and stress or those undergoing severe exercise, patients with eclampsia, uncontrolled hypertension, diabetes mellitus. gestational diabetes mellitus, endocrine disorders and patient suffering from any acute or chronic illness were excluded.

All study subjects were under regular follow up until delivery. Specific note was made for the development of pre-eclampsia/eclampsia during antenatal period and/ or at the time of delivery.

From all these subjects during their second trimester of pregnancy, clean catch morning urine

sample (5ml) was collected in a sterile container and was preserved at -20°C until biochemical analysis was performed.

Estimation of serum/urinary Creatinine and urinary microalbumin was done using kit method by Vitros Fusion 5.1 FS, fully automated analyzer from Ortho Clinical Diagnostic USA. The cut-off value of Albumin Creatinine Ratio (ACR) from Microalnbuminuria was taken as  $30-300 \mu g/mg$  of Creatinine.

Data were expressed as mean±SD. The statistical significance of difference between mean values assessed by Student's unpaired t-test. The "p" value of <0.05 was considered as significant.

### Results

In this clinical study all 145 pregnant women in 12-36 weeks of gestation age were followed up till delivery. Out of this total number, 107 pregnant women remained normotensive till their delivery. Their systolic BP, diastolic BP, Microalbumin and ACR were in the normal reference range (Table 1). Remaining 39 pregnant women developed pre-eclampsia in their third trimester of pregnancy. Their mean systolic BP was diastolic BP 151.15±10.61, was 99.76±4.92, Microalbumin level was 66.59±48.76 and ACR was 81.14±54.29 respectively. Pregnant women who developed pre-eclamsia were having significantly higher micro albumin level and ACR as compared to normotensive pregnant women (p value <0.05). It was a further notice that the women who developed preeclampsia were mostly in the age group of 25 to 36 years.

# Table 1: Values of different parameters innormotensive and pre-eclamptic pregnant women

|                            | Normotensive | Pre-Eclampsia | p<br>value |
|----------------------------|--------------|---------------|------------|
| Number (%)                 | 107(73.79)   | 39(26.89)     | < 0.0      |
| Mean gestational age       | 19.5±2.84    | 28.75±3.6     | 5          |
| Mean Systolic<br>BP(mmHg)  | 113.34±12.48 | 151.15±10.61  |            |
| Mean diastolic<br>BP(mmHg) | 78.5±4.68    | 99.76±4.92    |            |
| Microalbuminuri<br>a       | 19.4±5.71    | 66.59±48.76   |            |
| Mean ACR                   | 24.46±11.85  | 81.14±54.29   |            |

\*p value<0.05 considered as significant

| Table 2: | Age wise | e distribution ( | of parameters |
|----------|----------|------------------|---------------|
|----------|----------|------------------|---------------|

| Age   | Mean      | Mean      | Microalbu  | Mean level  |
|-------|-----------|-----------|------------|-------------|
| grou  | Systolic  | diastolic | -minuria   | of ACR      |
| р     | BP        | BP        |            |             |
|       | (mmHg)    | (mmHg)    |            |             |
| 18-26 | 102±11.24 | 84±6.1    | 22.34±9.6  | 23.46±15.85 |
| 27-36 | 136±17.35 | 95±12.8   | 57.59±48.7 | 71.14±44.29 |
|       |           |           | 6          |             |

Table 3: Occurrence of pre-eclampsia in primi andmultigravida

| Pre-eclampsia | Primigravida | Multigravida |
|---------------|--------------|--------------|
| 39(26.89)     | 16(41.02)    | 23(58.97)    |

### Discussion

Various studies have been conducted to see the association of micro albumin with outcome of pregnancy. In our study, we have taken 145 pregnant females in age group of 19-36 years and found that levels of microalbumin and ACR were significantly higher in women who developed pre-eclampsia in her later weeks of gestation. It was further noticed that the occurrence of preeclampsia was higher in multigravida as compared to primigravida and blood pressure was also higher in multigravida.

It has been established in recent years that preeclampsia is associated with widespread vascular dysfunction both in placenta and mother<sup>14</sup>. Various biochemical markers found to be raised in case of micro-vascular damage and among these microalbumin is commonly used because of its cost effectiveness and ease of estimation. Salako et al. found that microalbuminuria might be a good predictor of this condition with a high sensitivity but a low positive predictive value<sup>15</sup>. Bar et al. have reported in their study that microalbuminuria early in the third trimester of pregnancy is a good predictor of hypertensive complications in pregnancy<sup>16</sup>. Poonet et al. proposed that the appearance of clinical protienuria in preeclampsia (determined during 11-13 weeks of pregnancy) in 55% of normal pregnancies and in only pregnancies 75% of that complicated by preeclampsia<sup>17</sup>. Gangaram et. al reported in their study about the comparison of diagnostic value of the micro ACR with that of 24-h urine protein test when screening for proteinuria among pregnant women with hypertension. They proposed that the value of tow tests in the context was same and concluded that measurement of the micro ACR may be a good substitute for a random urine protein test<sup>18</sup>. Chhabra et al found that values of micro albumin around 18 weeks of gestation seemed useful, especially in primigravida<sup>19</sup>. K.K. Fatema et al reported in their study that patients having microalbuminuria will develop preeclampsia eventually<sup>20</sup>. It is clear that more clinical studies are needed to establish the usefulness of micro-albumin in prediction of pre-eclampsia.

## Conclusion

Our study was conducted in a small group of pregnant females, so to establish the association between microalbuminuria and occurrence of preeclampsia a further detailed study required in a large group of population along with measurement of other established marker.

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### References

- 1. Arias F. Practical Guide to high risk pregnancy and delivery. 2nd edition. India: Mosby;1992.
- Cunningham FG, Leveno KJ, Bloom SL, Hauth JC, Gilstrap LC, Wenstrom KD. Williams Obstetrics, 22nd edn. Bengaluru, Karnataka. McGraw Hill, 2005.
- Barker P, Kingdom J. Pre-eclampsia: Current Perspective and Management, 1st edn. Canada: RCOG Press, 2004. pp. 25–35.
- Prakash J, Pandey LK, Singh AK, Kar B. Hypertension in pregnancy: hospital based study. J Assoc Physicians India. 2006;54:273–8.
- Dr Liaquat Ali, Coordinator, Biomedical Research Group, Bangladesh Institute Of Research And Rehabilitation In Diabetes, Endocrine And Metabolic Disorders(BIRDEM), Dhaka.
- Mogensen CE, et. al. "Early Change in Kidney Function, Blood Pressure and Stages in Diabetic Nephropathy," in Prevention and Treatment of Diabetic Nephropathy, H. Keen and M. Legrain, ed. Boston, MTP Press, 1983;57-83,99-105.
- Mogensen CE, Christensen CK, Vittinghus E. The stage in diabetic renal disease. With emphasis on the stage of incipient diabetic nephropathy. Diabetes. 1983;32(Suppl 2):64-78.
- Viberti GC, Mackintosh D, Bilous RW, Pickup JC, Keen H. Proteinuria in diabetes mellitus: Role of spontaneous and experimental variation of glycemia. Kidney Int. 1982;21(5):714-20.
- Rodriguez MH, Masaki DI, Mestman J, Kumar D, Rude R. Calcium/creatinine ratio and microalbuminuria in the prediction of pre-eclampsia. Am J Obstet Gynecol. 1988;159(6):1452-5.
- Beaulieu MD. Prevention of preeclampsia. In: Canadian Task Force on the Periodic Health Examination. Canadian Guide to Clinical Preventive Health Care. Ottawa: Health Canada, 1994, pp136-43.
- 11. Robson SC, Edmond DK. Hypertensive and renal disease in pregnancy. Dewhurst's Textbook of Obstetrics and Gynaecology for Postgraduates 6th ed. United Kingdom: Blackwell Science Ltd; 1999. p.166-18.
- Lijima T, Sujuki S, Sekijuka K, Hishika T, Yagme M, Jinde K et al. Follow-up study on urinary type IV collagen in patients with early stage diabetic nephropathy. J Clin Lab Anal. 1998; 12(6): 378-382.
- Granger JP, Alexander BT, Llinas MT, Bennett WA, Khalil RA. Pathophysiology of hypertension during preeclampsia linking placental ischemia with endothelial dysfunction. Hypertension. 2001;38(3 Pt 2): 718-22.
- Salako BL, Olayemi O, Odukogbe AT, Adedapo KS, Aimakhu CO, Alu FE, et al. Microalbuminuria in pregnancy as a predictor of preeclampsia and eclampsia. West Afr J Med. 2003;22(4):295-300.
- Bar J, Hod M, Erman A, Friedman S, Gelerenter I, Kaplan B, et al. Microalbuminuria as an early predictor of hypertensive complications in pregnant women at high risk. Am J Kidney Dis. 1996;28(2):220-5.
- Poon LC, Kametas N, Bonino S, Vercellotti E, Nicolaides KH. Urine albumin concentration and albumin-tocreatinine ratio at 11(+0) to 13(+6) weeks in the prediction of pre-eclampsia. BJOG. 2008;115(7):866-73.

- 17. Gangaram R, Naicker M, Moodley J. Comparison of pregnancy outcomes in women with hypertensive disorders of pregnancy using 24-hour urinary protein and urinary microalbumin to creatinine ratio. Int J Gynaecol Obstet. 2009;107(1):19-22.
- Chhabra S, Gandhi D. Prediction of pregnancy induced hypertension/preeclamsia by detecting microalbuminuria. J Obstet Gynaecol India. 2002;52:56-60.
- Fatema K, Khatun M, Akter S, Ali L. Role of urinary albumin in the prediction of preeclampsia. Faridpur Med. Coll. J. 2011;6(1):14-18.