

British Journal of Medicine & Medical Research 15(10): 1-6, 2016, Article no.BJMMR.25024 ISSN: 2231-0614, NLM ID: 101570965



SCIENCEDOMAIN international www.sciencedomain.org

## Assessment of Biochemical Changes in Pregnancy Induced Hypertension (PIH) among Saudi Population at KAMC-Riyadh

Tarig Karar<sup>1</sup>, Mohamed Abdel Fattah<sup>1</sup>, Khaled Romaizan O. Alenazy<sup>1</sup>, Mohammad Jamaan Alharbi<sup>1</sup>, Fares Meshabab S. Alqahtani<sup>1</sup>, Waleed Al Tamimi<sup>2</sup>, Ikram Al Hassan<sup>2</sup>, Susan Grajo<sup>2</sup>, Ivy Alquisalas<sup>2</sup> and Shoeb Qureshi<sup>3\*</sup>

<sup>1</sup>Department of Clinical Laboratory Sciences, College of Applied Medical Sciences, King Saud Bin Abdul-Aziz University, Riyadh, Saudi Arabia. <sup>2</sup>Department of Pathology and Laboratory Medicine, National Guard Health Affairs, Riyadh, Saudi Arabia. <sup>3</sup>Research Unit, College of Applied Medical Sciences, King Saud Bin Abdul-Aziz University, Riyadh, Saudi Arabia.

## Authors' contributions

This work was carried out in collaboration between all authors. Author TK was the principal investigator of the research project, responsible for designing of the entire work. Authors TK, MAF and WAT were responsible for the entire write up. Authors KROA, MJA and FMSA were responsible for data collection. Authors IAH, SG and IA were responsible for the data management. Author SQ assisted in writing and attending to comments of the referees and rewriting some parts of the manuscript. He was responsible to correspond with the journal. All authors read and approved the final manuscript.

#### Article Information

DOI: 10.9734/BJMMR/2016/25024 <u>Editor(s)</u>: (1) Yinhua Yu, Department of Gynecology, Obstetrics and Gynecology Hospital of Fudan University, Shanghai Key Laboratory of Female Reproductive Endocrine Related Diseases, China. (2) Vijayalakshmi I. Balekundri, Sri Jayadeva Institute of Cardiovascular Sciences and Research, Bengaluru, India. <u>Reviewers:</u> (1) Ravindra Nath Sharma, St. George's University, Grenada, West Indies. (2) Anonymous, San Raffaele Scientific Institute, Milan, Italy. (3) Anonymous, University of Nairobi, Kenya. (4) Georgios Androutsopoulos, University of Patras, Rion, Greece. (5) Osredkar Josko, University Medical Centre Ljubljana, Slovenia. Complete Peer review History: <u>http://sciencedomain.org/review-history/14659</u>

Received 14<sup>th</sup> February 2016 Accepted 5<sup>th</sup> May 2016 Published 17<sup>th</sup> May 2016

Original Research Article

\*Corresponding author: E-mail: qsab2002@yahoo.co.in;

## ABSTRACT

**Background:** One of the most common complications that affect around 2-5% pregnant women is elevated blood pressure defined as gestational hypertension.

**Aims & Objective:** To compare and correlate between plasma urea, creatinine, sodium, potassium, and plasma glucose and urine protein in pre-eclamptic and normotensive groups.

**Materials and Methods:** This is quantitative retrospective chart review study. It is an hospital based case control study involving data from 199 age – matched Saudi women, discharged from Obstetrics and Gynecology Department of King Abdul Aziz medical city during the period of 2013-2014. The data were collected following standard procedures and statistical analysis was done using independent t – test and correlation.

**Results:** The levels urea, creatinine, sodium, potassium in the serum, plasma glucose and urine protein were significantly elevated in pre-eclamptic women when compared to normotensives (p= 0.000, p= 0.000, p= 0.000, p= 0.000, p= 0.000, p= 0.000, respectively). There was a significant positive correlation between urea and protein, urea and creatinine (r=0.3 P=0.002), (r=0.7 p=0.000) respectively.

**Conclusion:** The elevated values of serum creatinine, urea, urine protein, sodium, potassium and plasma glucose preclude them to be useful for consideration as consistent predictive indicator(s) for pre-eclampsia or pregnancy related hypertension.

Keywords: Preeclampsia; proteinuria; obstetrics; gynecology; normotensive.

## **1. INTRODUCTION**

One of the most common complications that affect 2-5% pregnant women is elevated blood pressure defined as gestational hypertension. Increased blood pressure usually starts after the 20th week of gestation without proteinuria [1,2]. Pregnancy-induced hypertension (PIH), which occurs in <10% of pregnancies, is a major risk factor for maternal and perinatal morbidity and mortality. PIH includes gestational hypertension as well as pre-eclampsia and eclampsia [3,4]. Measurement of blood pressure and proteinuria should be routine investigation during entire period of gestation [5].

Unfittingly implanted placenta is suggested as the main reason of gestational hypertension in pregnant women. Consequently correctly embedded placenta facilitates best transferring of oxygen from mother to fetus; otherwise failure in proper growth of placenta in uterus leads to development of pre-eclampsia and eclampsia [6,7]. Moreover a pregnant woman may have intercurrent diseases associated with gestational hypertension, that may become worse and be a potential risk to the pregnancy.

PIH causes important changes of biochemical parameters such as increased levels of blood alucose. urea. creatinine. uric acid. dehydrogenase transaminases, lactate and increased of with level proteinuria hypoalbuminemia which are used as indicator of disease severity. Biochemical changes also include alteration of lipid level, because hypertension is associated with peripheral vascular diseases, so gestational hypertension appears to be crucial justification in changes of lipid [8].

There is a paucity of literature on PIH from Riyadh, Saudi Arabia, hence the present study was undertaken to compare and correlate the biochemical changes that arise from PIH and early detection of these changes offer chance to avoid complications.

#### 2. MATERIALS AND METHODS

#### 2.1Study Design

This is quantitative retrospective chart review, an hospital based case controlstudy.

#### 2.2 Study Area and Duration

This study was conducted in King Abdul-Aziz Medical City (KAMC) complex in Riyadh, Saudi Arabia, during the period from September 2015 to November 2015.

#### 2.3 Study Subjects

One hundred and ninety nine pregnant women were selected from all women who attended obstetrics department from 2013-2014; they were categorized into two groups according to the following criteria:

#### 2.3.1 Inclusion criteria

The first group included pregnant women of all ages with a diagnosis of pre-eclampsia, while the control group included pregnant women without gestational hypertension.

#### 2.3.2 Exclusion criteria

Women diagnosed with hypertension or diabetes mellitus before pregnancy were excluded. Women diagnosed with gestational diabetes were also excluded from second group.

#### 2.3.3 Sample size

The sample size is 199 subjects based on previous auditing institutional report and on the calculated prevalence of gestational hypertension in Riyadh region.

The sample size and confidence level computed were 200 and 95% respectively.

### 2.4 Sampling Technique

The sample included pregnant women of different age groups discharged from KAMC-CR with diagnosis of gestational hypertension and other groups of pregnant women without gestational hypertension, at any point in time during 2013 and 2014. Blood and urine samples from both the study groups, pre-eclamptic and normotensive women were collected after 20<sup>th</sup> week of pregnancy within second trimester.

## 2.5 Ethical Consideration

Patient's confidentiality was strictly observed throughout the study by using Medical Record Number. Data access was restricted to the investigators. The study was approved by Institutional Review Board of King Abdullah International Medical Research Center, National Guards, Riyadh.

#### 2.6 Data Management and Analysis Plan

All information and data collected from the patients was entered into Excel file. A backup soft copy as well as a hard copy was dated, saved and secured after each data entry update. A designated study binder and a dedicated USB flash memory were kept with the principal investigator.

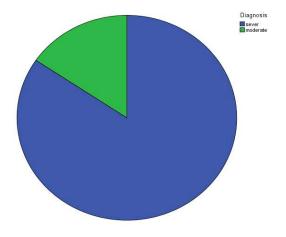
#### 2.7 Statistical Analysis

Statistical analysis of study variables was carried out using SPSS software. Descriptive and

demographic data was expressed as mean  $\pm$  standard deviation and percentage. Independent t and correlation tests were used, wherever applicable.

#### 3. RESULTS

This study was conducted on 149 pregnant women diagnosed with pre-eclampsia as test group and 50 women with normal pregnancy as control group. Age of the test group was matched with the control group. In the present study 14.8% (n=22) of test group was diagnosed with severe pre-eclampsia and 85.2% (n=127) of test group was diagnosed with moderate preeclampsia as illustrated in Fig. 1.



# Fig. 1. Frequency of severe and moderate pre-eclampsia among all patients

#### 3.1 Independent t-test

Table 1 Comparison of means of serum levels of urea, Creatinine, Sodium, Potassium, Plasma Glucose and urine protein was done between test and control groups. Table 2 Comparison of means of serum levels of urea, Creatinine, Plasma Glucose and urine protein between types of pre-eclampsia against test.

## 3.2 According to Correlation Analysis

Fig. 2 shows moderate positive correlation between levels of urea in mmol/l and levels of protein in mg/dl (r=0.3P=0.002).

Fig. 3 shows strong positive correlation between levels of urea in mmol/land levels of creatinine in  $\mu$ mol/l(r=0.7 P=0.000).

Fig. 4 shows moderate positive correlation between levels of urea in mmol/l and levels of potassium in mmol/l(r=0.3 P=0.001).

#### Table 1. Comparison of means of serum levels urea, creatinine, sodium, potassium, plasma glucose and urine protein between test and control group

Variable	Test	Control	р.
	(n=149)	(n=50)	value
Urea	3.7±1.6	2.2±0.6	0.000
Creatinine	57.72±18.69	46.68±5.426	0.000
Sodium	135.54±1.761	136.34±1.636	0.005
Potassium	4.221±.3989	3.928±.3064	0.000
Glucose	5.419±1.7405	4.590±0.7413	0.000
Protein	348±50	19.10±9.9	0.000

The table shows the mean ± standard deviation and probability (P). P value ≤ 0.05 is considered significant

#### Table 2. Comparison of means of serum levels of urea, creatinine, plasma glucose and urine proteins between types of preeclampsia against test

Variable	Sever preeclampsia (n=127)	Moderate preeclampsia (n=22)	<i>p</i> . Value
Glucose	5.6±1.8	4.6±1.1	0.003
Protein	385±53	136±31	0.000
Urea	3.8±1.7	3.5±.74	0.223
Creatinine	57.7±20.	57.7±6.4	0.990

The table shows the mean ± standard deviation and probability (P). P value ≤ 0.05 is considered significant

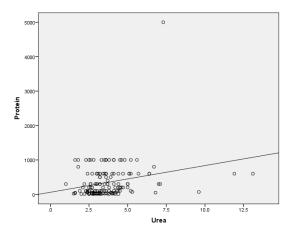


Fig. 2. A scatter plot shows the correlation between serum levels of urea in mmol/land urine levels of protein in mg/dl (r=0.3 p=0.002)

## 4. DISCUSSION

Being given to understand that proteinuria is an important sign of pre-eclampsia and the minimum criteria for the diagnosis of preeclampsia are hypertension plus proteinuria [9] it is recommended that urinary dipstick testing (visual or automated) may be used for screening proteinuria the of when suspicion of preeclampsia is low [10]. Accordingly results of present study revealed significant increase in proteinuria in preeclamptic women compared with normotensive women. Furthermore our results observed significant increase in proteinuria among pregnant women diagnosed with severe pre-eclampsia compared with pregnant women diagnosed with moderate preeclampsia. This observation is supported by the known criteria for diagnosis of severe preeclampsia [11] where the increase of BP (< 140 systolic and < 90 diastolic) and proteinuria (0.3 g/day) after 24 week of gestation.

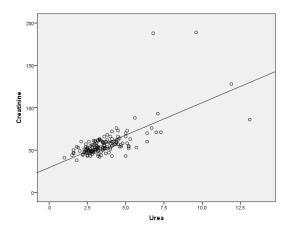
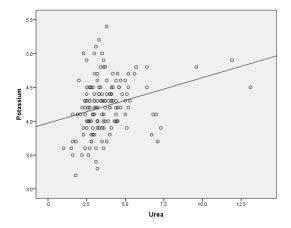
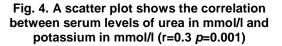


Fig. 3. A scatter plot shows the correlation between serum levels of urea in mmol/l and creatinine in µmol/l (r=0.7 *p*=0.000)





Creatinine and urea are non-protein nitrogenous metabolites that are removed from the body by the kidney through glomerular filtration and also used as indicators of kidney function and other conditions. Although measurement of urea and creatinine may have little value in the prediction of preeclampsia, estimation of these parameters in serum during pregnancy helps to diagnose kidney function especially at women with preeclampsia signs [12]. Previous study by Egwuatu et al. [13] showed increased creatinine level in preeclamptic women. Furthermore Magna and Sitikantha [8] also observed significant elevation in serum creatinine and small insignificant increase in serum urea level in pre-eclamptic when compared to normotensive women. These studies support our observation on the significant increase in serum urea and creatinine in pregnant women with pre-eclampsia compared to normotensive pregnant women, besides lack of correlation between urea and creatinine level and severity of eclampsia for both.

It is a well-known fact that electrolytes play an important role in the pathogenesis of hypertension. Dietary sodium restriction is one of the prime treatments of high blood pressure [14]. Therefore estimation of serum electrolytes in PIH provides a very useful index for the study of physiological and pathological changes during pregnancy. In the present study, a significant decrease in plasma sodium levels was seen in the preeclamptic patient compared to normotensive. Our findings are in accordance with those reported by others [14,15].

Sodium transport is altered across the cell membrane and this leads to the accumulation of sodium in the extravascular spaces and a decrease in the plasma sodium levels. The serum sodium levels tend to decline in cases of preeclampsia as the disorder increases in severity [15]. These studies corroborates our observation on the significant increase in the serum levels potassium in PIH cases as compared to that in the normal normotensive pregnant. However; these results are not in accordance with the observation of Indumati et al. [14]. Hence, this increase might be due to renal impairment or using of corticosteroid therapy during pregnancy.

Results of present study demonstrated significant increase in blood glucose in preeclamptic women compared to normotensive. These results confirm a previous study which observed that a minor degree of glucose intolerance is associated with a higher occurrence of preeclampsia in non-diabetic pregnant women [16].

#### 5. CONCLUSION

The elevated values of urea, creatinine, sodium, potassium in the serum,plasma glucose and urine protein precludes them to be useful for consideration as consistent predictive indicator(s) for preeclampsia or pregnancy related hypertension. Furthermore the elevated values for plasma glucose and urine protein could be used to differentiate between moderate and sever preeclampsia. Therefore it was recommended that all pregnant women should be screened for preeclampsia at the first prenatal visit and periodically throughout the remainder of the pregnancy.

#### CONSENT

It is not applicable.

## **COMPETING INTERESTS**

Authors have declared that no competing interests exist.

#### REFERENCES

- Abrams Elizabeth, Julienne Rutherford. Framing postpartum hemorrhage as a consequence of human placental biology: An evolutionary and comparative perspective. Am Anthropol. 2011;113(3): 417-30.
- 2. Guancario P. Journal of clinical Endocrinology and Metabolism. 2006; 91(4):1223-1238.
- 3. Why mothers die 2000-2002. The sixth report of the Confidential Enquiries into Maternal Deaths in the United Kingdom. London: RCOG Press; 2004.
- 4. Roberts JM, Pearson G, Cutler J, Lindheimer M. Summary of the NHLBI Working Group on Research on Hypertension during Pregnancy. Hypertension. 2003;41:437-45.
- 5. Cross JC. The genetics of pre-eclampsia: A feto-placental or maternal problem? Clinical Genetics. 2003;64(2):96-103.
- Kocijancic DM, Plesincac S, Plecas D, Aksam S, Kocijancic A. Correlation of biochemical parameters and neonatal outcome in patients with gestational hypertension. Clinical and Experimental Hypertension. 2013;35:6-10.

Karar et al.; BJMMR, 15(10): 1-6, 2016; Article no.BJMMR.25024

- 7. Dutta DC. Hypertensive disorders in pregnancy. In: Textbook of Obstetrics, Ed. Konar HL, 5th edition, New Central Book Agency, Kolkata. 2001;234-55.
- Magna Manjareeka, Sitikantha Nanda. Elevated levels of serum uric acid, serum creatinine or urea in pre-eclamptic women. International Journal of Medical Science and Public Health. 2013;2(1):43-47.
- 9. Amita Sharma, Pandey Kiran, Bhagoliwal Ajai. Spot urine protein/creatinine ratio, A quick and accurate method for diagnosis of pre-eclampsia. Journal of Obstetrics and Gynecology. 2013;3:609-612.
- Sogc. Clinical Practice Guideline. Diagnosis, evaluation, and management of the hypertensive disorders of pregnancy: JOGC MAI 2014 No. 307. 2014;416-438.
- 11. Lana K. Wagner diagnosis and management of preeclampsia. American Family Physician. 2004;70(12):2317–24.

- 12. Lalitha Devi Seerla, Syed Abdul Jaweed, Jyothinath Kothapalli. Is non protein nitrogenous compound have role in preeclampsia. National Journal of Laboratory Medicine. 2014;3:23-26.
- 13. Egwuatu VE. Plasma concentration of urate, urea and creatinine in Nigerian primigravidae with pre-eclampsia. Trop Geogr Med. 1986;38:11-15.
- 14. Indumati V, Kodliwadmath MV, Sheela MK. The role of serum electrolytes in pregnancy induced hypertension. Journal of Clinical and Diagnostic Research. 2011;5(1):66-69.
- 15. Searcy RL. Diagnostic Biochemistry, New York: McGraw-Hill Book Company. 1969;469:476.
- Khan KS, Dava S. Plasma glucose and pre-eclampsia. International Journal of Gynecology and Obstetrics. 1996;53(2): 111–116.

© 2016 Karar et al.; This is an Open Access article distributed under the terms of the Creative Commons Attribution License (http://creativecommons.org/licenses/by/4.0), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Peer-review history: The peer review history for this paper can be accessed here: http://sciencedomain.org/review-history/14659