An evaluation of association of vitamin D insufficiency with gestational hypertension in pregnant women

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ABSTRACT

Background: Preeclampsia (PE) is a disease specific to pregnancy affecting many bodily systems. This is characterized by high blood pressure and proteinuria after the 20th week of pregnancy. The objective of this study was to evaluate an association of vitamin D insufficiency with gestational hypertension in pregnant women.

Methods: This was a cross-sectional study. A total of 104 pregnant women were included in the study. The diagnosis of gestational hypertension was confirmed using the “Report of the American College of Obstetricians and Gynaecologists’ Task Force on Hypertension in Pregnancy” criteria. Based on these criteria, patients with systolic blood pressure ≥140 mmHg and/or diastolic blood pressure ≥90 mmHg (measured after a period of rest of four hours, twice daily) and proteinuria (≥300 mg protein/24 h) were diagnosed as gestational hypertension.

Results: Vitamin D deficiency was found among 78.9% (90/114) women. There was no significant (p >0.05) difference in age of women between vitamin D deficient and sufficient. Vitamin D level was significantly (p = 0.0001) lower among the women between vitamin D deficient (15.93±4.66) and sufficient (35.70±3.25). There was a significant (p >0.05) association of Vitamin D level with parity, family history of hypertension, gestational age and fasting blood glucose. There was significant (p = 0.00011) difference in BMI between Vitamin D deficient (15.93±4.66) and sufficient (35.70±3.25). Tobacco consumption was also associated with the level of Vitamin D. There was significant negative correlation between BMI and Vitamin D level (r = -0.56, p = 0.00011). There was no significant (p >0.05) association of Vitamin D level with parity, family history of hypertension, gestational age and fasting blood glucose. There was significant (p = 0.00011) difference in BMI between Vitamin D deficient (15.93±4.66) and sufficient (35.70±3.25). Tobacco consumption was also associated with the level of Vitamin D. There was significant negative correlation between BMI and Vitamin D level (r = -0.56, p = 0.00011). There was no significant (p >0.05) association of Vitamin D level with parity, family history of hypertension, gestational age and fasting blood glucose. There was significant (p = 0.00011) difference in BMI between Vitamin D deficient (15.93±4.66) and sufficient (35.70±3.25). Tobacco consumption was also associated with the level of Vitamin D. There was significant negative correlation between BMI and Vitamin D level (r = -0.56, p = 0.00011). There was no significant (p >0.05) association of Vitamin D level with parity, family history of hypertension, gestational age and fasting blood glucose. There was significant (p = 0.00011) difference in BMI between Vitamin D deficient (15.93±4.66) and sufficient (35.70±3.25). Tobacco consumption was also associated with the level of Vitamin D. There was significant negative correlation between BMI and Vitamin D level (r = -0.56, p = 0.00011). There was no significant (p >0.05) association of Vitamin D level with parity, family history of hypertension, gestational age and fasting blood glucose. There was significant (p = 0.00011) difference in BMI between Vitamin D deficient (15.93±4.66) and sufficient (35.70±3.25). Tobacco consumption was also associated with the level of Vitamin D. There was significant negative correlation between BMI and Vitamin D level (r = -0.56, p = 0.00011). There was no significant (p >0.05) association of Vitamin D level with parity, family history of hypertension, gestational age and fasting blood glucose. There was significant (p = 0.00011) difference in BMI between Vitamin D deficient (15.93±4.66) and sufficient (35.70±3.25). Tobacco consumption was also associated with the level of Vitamin D. There was significant negative correlation between BMI and Vitamin D level (r = -0.56, p = 0.00011). There was no significant (p >0.05) association of Vitamin D level with parity, family history of hypertension, gestational age and fasting blood glucose. There was significant (p = 0.00011) difference in BMI between Vitamin D deficient (15.93±4.66) and sufficient (35.70±3.25). Tobacco consumption was also associated with the level of Vitamin D. There was significant negative correlation between BMI and Vitamin D level (r = -0.56, p = 0.00011). There was no significant (p >0.05) association of Vitamin D level with parity, family history of hypertension, gestational age and fasting blood glucose. There was significant (p = 0.00011) difference in BMI between Vitamin D deficient (15.93±4.66) and sufficient (35.70±3.25). Tobacco consumption was also associated with the level of Vitamin D. There was significant negative correlation between BMI and Vitamin D level (r = -0.56, p = 0.00011).

Conclusions: The present study demonstrates that vitamin D plays a role in the etiology and pathophysiology of gestational hypertension. Among the population having the risk of vitamin D deficiency, the risk of pregnancy induced hypertension may be decreased through vitamin D supplementation.

Keywords: Gestational hypertension, Insufficiency, Pregnant women

INTRODUCTION

Preeclampsia (PE) is a disease specific to pregnancy affecting many bodily systems. This is characterized by high blood pressure and proteinuria after the 20th week of pregnancy. This complicates 2-8% of pregnancies. This increases maternal and fetal mortality and morbidity. Different factors such as angiogenic, endothelial dysfunction, syncytiotrophoblastic microparticles and inflammatory activation play an important role in the progression of preeclampsia. The maternal diets are associated with the etiology of preeclampsia.

Vitamin D has received great attention as a possible etiological factor in pre-eclampsia. Vitamin D deficiency can affect the health of both mother and fetus by increasing the production of inflammatory cytokines and stimulating the activity of T-regulatory cells. It results in...
poor bone mineralization in infants, low birth weight and other adverse pregnancy outcomes.³

Epidemiological studies have shown the importance of vitamin D deficiency in the development of preeclampsia.⁴ The importance of vitamin D deficiency in immunomodulation and placental development has been highlighted in various studies. Thus, they put the emphasis on vitamin D deficiency regarding its probable role in the physiology of preeclampsia.⁵ ⁶

The objective of the present study was to evaluate of association of vitamin D insufficiency with gestational hypertension in pregnant women.

METHODS

This was a cross-sectional study conducted in the Department of Obstetrics and Gynecology, Integral Institute of Medical Sciences, Lucknow. The study was approved by the Ethical Committee of the Institute. The consent was taken from each participant before including in the study.

A total of 104 pregnant women were included in the study. The diagnosis of gestational hypertension was confirmed using the Report of the American College of Obstetricians and Gynecologists criteria. Based on these criteria. Patients with systolic blood pressure ≥140 mmHg and/or diastolic blood pressure ≥90 mmHg and proteinuria ≥300 mg protein/24 h) were diagnosed as gestational hypertension.⁷

The acceptance criteria for gestational hypertensive women for the study was if they had normal blood pressure during the first 20 weeks of gestation, no previous history of metabolic disorders, twin pregnancy, recurrent miscarriage, fetal growth retardation, placenta abruption, thrombophilia, renal disease, chronic hypertension or diabetes mellitus as well as no history of antioxidant intake and no medication for hypertension.

The interval between the first day of the mother’s last menstrual period and expected date of delivery was defined as gestational age.

None of the patients were on any medication at the time of blood sample collection. Demographic and clinical data were collected during routine obstetric visits. Fasting venous blood specimen was taken from the antecubital vein and collected in vacutainers with no additives. The serum separator tube specimens were allowed to clot. Then centrifuged for 10 minutes at 3,000 g to separate the serum and stored at -70°C until analyzed.

Total vitamin D assays were analyzed with the fully automatic Advia Centaur XP immunoassay systems. Results were automatically calculated to the standard deviation.

The SPSS 16.0 version (Chicago, Inc., USA) was used to analyze the data. Normal distribution was tested using the Shapiro-Wilk test and variability coefficients. The data with normal distribution was analyzed using parametric methods. Two independent groups were compared using the Student unpaired t-test. Categorical variables were compared using the Pearson Chi-Square test. Quantitative data was expressed as a mean±SD (standard deviation). Categorical values were expressed as numbers (n) and percentages (%). A p-value <0.05 was considered significant.

RESULTS

Vitamin D deficiency was found among 78.9% (90/114) women (Figure 1).

![Figure 1: Distribution of Vitamin D deficiency.](image)

There was no significant (p >0.05) difference in age of women between vitamin D deficient and sufficient. Vitamin D level was significantly (p = 0.0001) lower among the women between Vitamin D deficient (15.93±4.66) and sufficient (35.70±3.25). There was no significant (p>0.05) association of Vitamin D level with parity, family history of hypertension, blood pressure,
gestational age and fasting blood glucose. There was significant ($p = 0.0001$) difference in BMI between Vitamin D deficient (30.10±4.95) and sufficient (24.04±2.75). Tobacco consumption was also associated with the level of Vitamin D (Table 1).

**Table 1: Comparison of Vitamin D status with basic characteristics of women.**

| Basic characteristics | Vitamin D status (ng/dl) | p-value |
|-----------------------|--------------------------|---------|
|                       | $\leq 30$ | $> 30$ |
| Age in years, mean±SD | (n=90) | (n=24) |
| Vitamin D level       | 26.83±6.49 | 25.79±6.50 | 0.48 |
| Parity, no. (%)       |         |         |       |
| Nullipara             | 51 (56.7) | 15 (62.5) | 0.60 |
| Multipara             | 39 (43.3) | 9 (37.5)  |       |
| Family history of hypertension, no. (%) | |         |       |
| Present               | 19 (21.1) | 2 (8.3)   | 0.12 |
|Absent                | 71 (78.9) | 22 (91.7) |       |
| Tobacco habit         |         |         |       |
| Present               | 71 (78.9) | 9 (37.5)  | 0.0001* |
| Absent                | 19 (21.1) | 15 (62.5) |       |
| BMI, mean±SD          |         |         |       |
| Systolic blood pressure, mean±SD | 30.10±4.95 | 24.04±2.75 | 0.0001* |
| Diastolic blood pressure, mean±SD | 121.50±11.04 | 117.17±9.56 | 0.06 |
| Gestational age in weeks, mean±SD | 72.51±11.55 | 68.25±9.55 | 0.10 |
| Fasting blood glucose mean±SD | 26.43±1.48 | 25.71±1.48 | 0.06 |

*Significant

There was significant negative correlation between BMI and Vitamin D level ($r = -0.56, p = 0.00011$) (Figure 2).

**DISCUSSION**

Pathogenesis of gestational hypertension is complex, and a vitamin D deficiency is one of the factors in the etiology of gestational hypertension. Vitamin D is being considered a major role in the synthesis and regulation of genes. They are effective in the development of placenta.\(^8\) Deep trophoblast invasion and remodeling of the vascular structures in the placental bed play a role in the development of gestational hypertension.\(^8\) Vitamin D deficiency has been advocated to be a predisposing factor for the peripheral vascular phase modulation. This results in inadequate placental development and the development of gestational hypertension.\(^10\)

It has been reported that the production of vitamin D in the decidua placenta and the maternal kidneys increases during pregnancy.\(^11\) Because, vitamin D level is being dependent on the sunlight exposure and dietary factors, there may be little or no change in the level of vitamin D during pregnancy.\(^12\)

In the present study, Vitamin D deficiency was found among 78.9% women. The studies conducted in USA, Australia, the Middle East and South Asia, vitamin D deficiency was found to be in 26-98% of pregnant women. Vitamin D deficiency has been reported to be 66-100% among the women with a dark skin color.\(^13,14\)

In this study, there was significant ($p=0.0001$) difference in BMI between Vitamin D deficient (30.10±4.95) and sufficient (24.04±2.75). Similar finding was reported in the study by Bakacak et al.\(^2\)

Based on the literature, establishing the relationship between gestational hypertension and vitamin D is complicated. There are various studies on the role of vitamin D on the development and pathophysiology of gestational hypertension. A low vitamin D level in the second trimester has been emphasized in previous studies to be an indicator of gestational hypertension.\(^15,16\)

In the present study, there was significant ($p = 0.0001$) difference in BMI between Vitamin D deficient (30.10±4.95) and sufficient (24.04±2.75). Tobacco consumption was also associated with the level of Vitamin D. Bakacak et al reported similar findings.\(^2\)

The studies on the role of vitamin D in the development of gestational hypertension and vitamin D have been emphasized to play an immunomodulatory and anti-inflammatory role in many systems.\(^17\) Vitamin D has been suggested to play an important role in signal and gene regulations in the development of placental trophoblasts in the placental growth.\(^18,8\)

**CONCLUSION**

The present study demonstrates that vitamin D plays a role in the etiology and pathophysiology of gestational hypertension. Among the population having the risk of vitamin D deficiency, the risk of pregnancy induced hypertension may be decreased through vitamin D supplementation.

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