ISSN 2063-5346



SERUM LEVEL OF VITAMIN D AND HYPERTENSIVEDISORDER IN PREGNANCY

Dr. BALSAM NASER ABDULHUSSEN¹, DR. ZAINAB TAREQ ALYASSEN², DR. KHOLOD ALSALAMY³

Article History, Accepted, 01.02.2025 Accepted, 10.04.2025	Article History: Received: 01.02.2023	Revised: 07.03.2023	Accepted: 10.04.2023
--	---------------------------------------	---------------------	----------------------

Abstract

Introduction: Hypertensive disorders of pregnancy, including gestational hypertension, preeclampsia (PE), and eclampsia, are among the major complications that account for approximately 14% of maternal mortality. The aim of this study is to determine the association between serum level of vitamin D and hypertensive disorder in pregnancy that include. Patients and Methods: This is a prospective case control study carried out in AL Basrah maternity &child hospital in the period from January 2021 till the 31 of January 2022. The study included 110 pregnant women who were divided in two groups; First group; (Cases) included women diagnosed with hypertensive disorders of pregnancy. Second group; (controls) normotensive pregnant women. Result: All 110 pregnant women enrolled in the study had either deficient (49 (98.0%) for cases and 58 (96.7%) for controls) or insufficient level of vitamin D (1 (2.0%) for cases and 2 (3.3) for controls). in our study we found there's NO association between serum level of Vitamin D and hypertensive disorder in pregnancy. Conclusion: in our study we found there's NO association between serum level of Vitamin D and hypertensive disorder in pregnancy.

Keywords: Serum, vitamin D, hypertensive, disorder, pregnancy

^{1,2,3}College of Medicine, University of Basra

DOI:10.31838/ecb/2023.12.s1-B.379

Introduction:

Hypertensive disorders of pregnancy, including gestational hypertension, preeclampsia (PE), and eclampsia, account for approximately 14% of maternal mortality and complicate about 10% of pregnancies worldwide. PE is a major cause of maternal and perinatal morbidity and mortality, affecting 2% to 8% of pregnancies. Complications of PE include eclampsia, disseminated intravascular coagulation, HELLP syndrome. intrauterine growth restriction (IUGR), and fetal death ¹⁻⁵. PE has been hypothesized to be a two-stage disorder. In the first stage, reduced placental perfusion leads to defective placental implantation. In the second stage, reduced vascularization at the placental site activates a maternal inflammatory response, leading to generalized endothelial dysfunction and hypertension. A modified version this hypothesis proposes of that maternal factors in combination with inflammatory changes normal in pregnancy can lead to endothelial dysfunction with or without reduced perfusion 6-10 placental Hypovitaminosis D has been associated with PE. Vitamin D's roles include pro-inflammatory modulating responses, decreasing oxidative stress in PE, promoting angiogenesis through VEGF gene modulation. and decreasing blood pressure through the renin-angiotensin system (RAS). Vitamin D status is determined by measuring circulating 25hydroxyvitamin D [25(OH)D] levels ¹¹⁻ ¹⁶. Vitamin D is thought to play a significant role in PE as an immune modulator, helping to mount an appropriate maternal immune response to the placenta and preventing the release of anti-angiogenic factors. 1,25(OH)2D, the active form of vitamin D, has been shown to suppress T cell proliferation and down-regulate proinflammatory cytokines. Vitamin D also promotes angiogenesis in endothelial progenitor cells, possibly increasing VEGF expression and pro-MMP-2 activity ¹⁷⁻²¹. The aim of this study is to determine the association between serum level of vitamin D and hypertensive disorder in pregnancy that include preeclampsia and gestational hypertension.

Method:

This is a prospective case control study carried out in AL Basrah maternity &child hospital in the period from January 2021 till the 31 of January 2022. The study included 110 pregnant women who were divided in two groups; First group; (Cases) included women diagnosed with hypertensive disorders of pregnancy. Second group; (controls) normotensive pregnant women. Both cases and controls were selected randomly from patients admitted to the labour ward and from obstetric outpatient in ALBasrah maternity and child hospital. All pregnant women in this study (cases and control) serum level of Vitamin D were measured at admission. Special printed questionnaire formula was used for each pregnant women including age, parity, gestational age, mode of delivery, neonatal outcome, admission to NICU, educational state, if she takes supplement of vitamin D during ANC or not. Vitamin D is considered adequate when 25(OH)D levels are above 50 nmol/L, as defined by the Institute of Medicine. A level between (30-50)nmol/L is considered insufficient, and less than 30 nmol/L, deficient (22).

Results:

A total of 110 women were involved in this study. The first group the cases were 50 pregnant women diagnosed with hypertension in pregnancy including gestational hypertension, and preeclampsia (PE), Pregnancy induced hypertension, defined as blood pressure greater than 140/90 mmHg on two consecutive occasions >6 h apart occurring after 20 weeks of pregnancy, complicates approximately 10% of all pregnancies worldwide. Pre-eclampsia (PE) is hypertension and proteinuria (protein in urine ≥ 0.3 g/24 h (1+ dipstick) on two occasions ≥ 6 h apart) or edema, the second group were controls included 60 normotensive pregnant women. Their age ranged from 15-42 years old, and their mean age was 27.1±6.31 years. The majority of them were those between 21-40 years old (86.0% and 86.7% among

case and control respectively). Regarding the parity, they range from 0-9, median equal to 2 children. Fiftyeight percent of mothers with hypertension have 1-4 children while mothers in the control group most of having their first children them There is no significant (46.7%). difference between both groups since the P- value = 0.148. Most of the women in the case group and control group had school education (62.0% and 66.7% respectively). There is no difference in educational status (Pvalue=0.062) as showed in (Table 1).

Table 1: The sociodemographic characteristics among both studied group

	Case		P-value		
Varibles		Control			
Age					
≤20	6 (12.0%)	6 (10.0%)			
21-40	43 (86.0%)	52 (86.7%)	0.890*		
≥41	1 (2.0%)	2 (3.3%)			
Parity					
Nuliparas	15 (30.0%)	28 (46.7%)			
Para 1-4	29 (58.0%)	24 (40.0%)	0.148**		
>4	6 (12.0%)	8 (13.3%)			
Educational status					
Illiterate	8 (16.0%)	2 (3.3%)			
School education	31 (62.0%)	40 (66.7%)	0.062 *		
Higher education	11 (22.0%)	18 (30.0%)			
Total	50 (100.0%)	60 (100.0%)			
	*fisher-exact	test			
	**chi-secure	test			

The mode of delivery, most of the women in both groups had a normal vaginal delivery (P-value = 0.265). Regarding the perinatal outcome, 24.0% of children born to women with hypertension need NCU admissions in comparison to 6.7% in normal mothers. There is a significant difference between groups (P-value=0.048). The women in both groups showed no significant difference regarding the gestational age at delivery (P-value= 0.563). Regarding the association between hypertension in pregnancy and vitamin D3 levels, the majority of women in both groups had vitamin levels less than 30 nmol/l (98.0% and 96.7% among case and control respectively). P- value = 0.669 so there is no significant association between hypertension in pregnancy and vitamin D3 levels.

Vit D3 level	Case	Control	P-value
Defecient < 30 nmol/l	49(98.0%)	58 (96.7%)	
Insuffecient (30-50) nmol/l	1 (2.0%)	2 (3.3%)	0.716*
Normal > 50 nmol/l	0 (0.0%)	0(0.0%)	-
Total	50 (100.0%)	60 (100.0%)	
*fisher-exact test			

The case group involved women with gestational hypertension and preeclampsia, Table 4 showed the vitamin D3 level among women with gestational hypertension and those with preeclampsia. The serum vitamin D3 level looked to be lower among women with preeclampsia than its level in women with Gestational

	Case		P-value	
Varibles		Control		
Mode of delivery				
NVD	39 (78.0%)	40 (66.7%)	0.265**	
S/C	11 (22.0%)	20 (33.3%)		
Admission to NCU				
Yes	12 (24.0%)	4 (6.7%)	0.048*	
No	38 (76.0%)	56 (93.3%)		
Gestational age				
$\geq 37 wk$	25 (50.0%)	26 (43.3%)		
38-40 wk	25 (50.0%)	34 (56.7%)	0.563*	
>40 wk	0 (0.0%)	0 (0.0%)		
Total	50 (100.0%)	60 (100.0%)		
*fisher-exact test				
**chi-sequre test				

hypertension, still there is no significant difference (P-value = 0.073).

Table 4: Serum vitamin D3 among cases

PIH	S. vit D	<i>P-value</i>	
Preeclampsia	11.39 ± 3.11	0.073*	
Gestational hypertention	13.56 ± 4.82		
*two sample independent t-test			

The women in the study were asked about vitamin D3 supplementation during the current pregnancy, and most of them mention no history o vitamin D3 supplement intake (84.0% and 83.3% among case and control respectively). There is no difference between the case and control (P-value =0.925).

Table 5: The distribution of studied group in relation to the vitamin D3 supplementation: -

			P-value
Supplemen T	Cases	Control	
Yes	8 (16.0%)	10 (16.7%)	
No	42 (84.0%)	50 (83.3%)	0.925*
Total	50 (100.0%)	60 (100.0%)	
*fisher-exact test			

Discussion:

Vitamin D deficiency has emerged as a public health problem worldwide due to its important role in health and disease. Evidence from clinical and epidemiological studies support а possible relationship between low vitamin D level and hypertension, and there are some biological mechanisms as well. However, epidemiological studies are always vulnerable to multiple confounding factors that cannot be always controlled. Several studies have reported high a associations between Vitamin D and hypertensive disorder in pregnant women (Rose Mary J. Vatakencherry L Saraswathy Karen and

M.OCallaghan and Mairead kiely ^{(23,} ²⁴⁾. This is in contrast to our study. Our results revealed there's no difference between normotensive pregnant women and pregnant women with hypertensive disorders regarding the level of vitamin D. The explanation for this finding is based on the fact that all the women in this study wear hijab and all their time spend indoors with less sun exposure because of elevated temperature in summer, and covering their bodies completely for cultural reasons that makes sun exposure difficult. Although in this study no between association vitamin D deficiency and hypertensive disorder among pregnant women was found,

over 90% of pregnant women in this study were vitamin D deficient in both cases and control. Vitamin D deficiency or insufficiency is not expected to be a cause of hypertensive disorder, but still vitamin D status becomes crucial for maternal health. fetal skeletal growth, and optimal maternal and fetal outcomes. Pregnant have poor vitamin women D supplementation practices and usually consume vitamins that are necessary for maternal and neonate health such as folic acid and iron ^[25, 26] so more efforts must be done by the health personals to encourage more vitamin D supply to all pregnant women

Conclusion:

There is no association between serum level of VIT D and hypertensive disorder in pregnancy found in this study.

References:

- World Health Organization . Fact Sheet – Maternal Mortality

 http://www.who.int / mediacentre / factsheets / fs348/en/. Accessed 23 July 2016.
- Peters RM , Flack JM . Hypertensive disorders of pregnancy . Jobstet Gynecol Neonatal Nurs . 2004 : 33 (2) : 209 – 20 .
- 3. Robert JM ,Balk JL, Bodnar LM, Belizan JM,BergelE, MartinezA. Nutrient involvement in preeclampsia.J Nutr.2003,133(5SuppL): 1684S_92S.
- 4. Zhang J,ZeislerJ ,Hatch MC,Berkowitaz G,Epidemeology of pregnancy _induced hypertenstion .Epidemiol Rev.1997,19(2):218-32.

- 5. Duley L . the global impact of pre eclampsia and eclampsia and eclampsia . 2009; 33 (3): 130 7.
- 6. Srinivas SK , Edlow AG , Neff PM , Sammel MD , Andrela CM , Elovitz MA.Rethinking IUGR in preeclampsia : dependent or independent of maternal Purswani et al . BMC P BMC Pregnancy and Childbirth (2017) 17:231
- Redman CW. Immunological aspects of pre-eclampsia. Baillieres Clin Obstet Gynaecol. 1992;6(3):601–15.
- 8. Wei SQ, Audibert F, Luo ZC, Nuvt AM, Masse B, Julien P, Fraser WD, Group MS. Maternal plasma 25hydroxyvitamin D levels. angiogenic and factors. preeclampsia. Obstet Am J Gynecol. 2013;208(5):390 e391-6.
- 9. Woodham PC, Brittain JE. Baker AM, Long DL, Haeri S, Camargo CA Jr, Boggess KA, Stuebe AM. Midgestation 25maternal serum hydroxyvitamin D level and soluble fms-like tyrosine kinase 1/placental growth factor ratio as predictors of severe preeclampsia. Hypertension. 2011;58(6):1120-5.
- 10. Roberts JM, Hubel CA. The two stage model of preeclampsia: variations on the theme. Placenta. 2009;30(Suppl A):S32–7.
- Skull SA, Ngeow JY, Biggs BA, Street A, Ebeling PR. Vitamin D deficiency is common and unrecognized among recently arrived adult immigrants from the horn of Africa. Intern Med J. 2003;33(1-2):47-51
- 12. Nesby-O'Dell S, Scanlon KS, Cogswell ME, Gillespie C,

Hollis BW, Looker AC, Allen C, Doughertly C, Gunter EW, Bowman BA. Hypovitaminosis D prevalence and determinants among African American and white women of reproductive age: third National Health and nutrition examination survey, 1988-1994. Am J Clin Nutr. 2002;76(1):187–92.

- Finch PJ, Ang L, Colston KW, Nisbet J, Maxwell JD. Blunted seasonal variation in serum 25hydroxy vitamin D and increased risk of osteomalacia in vegetarian London Asians. Eur J Clin Nutr. 1992;46(7):509–15.
- Bodnar LM, Catov JM, Simhan HN, Holick MF, Powers RW, Roberts JM. Maternal vitamin D deficiency increases the risk of preeclampsia. J Clin Endocrinol Metab. 2007;92(9):3517–22.
- 15. Gray TK, Lester GE, Lorenc RS. Evidence for extra-renal 1 alphahydroxylation of 25hydroxyvitamin D3 in pregnancy. Science. 1979;204(4399):1311–3.
- 16. Thorne-Lyman A, Fawzi WW. Vitamin D during pregnancy and maternal, neonatal and infant health outcomes: a systematic review and metaanalysis. Paediatr Perinat Epidemiol. 2012;26(Suppl 1):75–90.
- Noyola-Martinez N, Diaz L, Avila E, Halhali A, Larrea F, Barrera D. Calcitriol downregulates TNF-alpha and IL-6 expression in cultured placental cells from preeclamptic women. Cytokine. 2013;61(1):245–50.
- 18. Grundmann M, Haidar M, Placzko S, Niendorf R. Darashchonak N, Hubel CA, von Versen-Hoynck F. Vitamin improves the angiogenic D properties of endothelial

progenitor cells. Am J Physiol Cell Physiol. 2012;303(9):C954–62.

- 19. Merchant SJ, Narumiya H, Zhang Y, Guilbert LJ, Davidge ST. The effects of preeclampsia and oxygen environment on endothelial release of matrix metalloproteinase-2. Hypert Pregnancy. 2004;23(1):47–60.
- 20. Cardus A, Parisi E, Gallego C, Aldea M. Fernandez E. 1.25-Valdivielso JM. Dihydroxyvitamin D3 stimulates vascular smooth muscle cell proliferation through a VEGF-mediated pathway. Kidney Int. 2006;69(8):1377-84.
- 21. Irani RA, Xia Y. The functional role of the renin-angiotensin system in pregnancy and preeclampsia. Placenta. 2008;29(9):763–71.
- 22. Holick Sunlight MF. and vitamin D for bone health and prevention of autoimmune diseases. cancers. and cardiovascular disease. Am J Clin Nutr. 2004;80(6 Suppl):1678S-88S.
- 23. Jounal of Family Medicine and Primay Care : Wolters Kluwer – Medknow Publications Rose Mary J . Vatakencherry and L Saraswathy2019 – 2061 : (6) 8 -2067
- 24. Nutrients 2018 Mar 10(3) 294 Karen M.OCallaghan and Mairead Kiely.
- 25. Roberts JM, Balk JL, Bodnar LM, Belizan JM, Bergel E, Martinez A. Nutrient involvement in preeclampsia. J Nutr. 2003;133(5 Suppl 2):1684S–92S.
- 26. Halhali A, Tovar AR, Torres N, Bourges H, Garabedian M, Larrea F. Preeclampsia is associated with low circulating

levels	of	in	sulin-like	growth	
factor		Ι	and	1,25-	
dihydroxyvitamin D in maternal					
and		um	bilical	cord	

compartments. J Clin Endocrinol Metab.2000;85(5):1828–33.