

*Research Article***Association between vitamin D level and metabolic syndrome in hemodialysis patients****Ahmed S. Salama, Amal K. Helmy, Hisham M. Tawfik, Rasha M. Hasan**

Department of Internal medicine, Faculty of Medicine, Minia University, Egypt

**Abstract**

**Background:** Vitamin D deficiency is common in patients with end-stage renal disease (ESRD) on maintenance hemodialysis. Metabolic syndrome has a high prevalence in hemodialysis patients. **Objectives:** In this study, the association between serum level of 25(OH) vitamin D and the prevalence of metabolic syndrome and its components in a group of ESRD patients undergoing maintenance hemodialysis was explored. **Subjects and Methods:** This cross-sectional study was carried out in the hemodialysis unit of Minia University Hospital. 90 patients on maintenance hemodialysis were enrolled. **Results:** The prevalence of metabolic syndrome was 30.6%. Reduced HDL-c emerged as the most frequent diagnosed abnormality affecting nearly 64.8% of the enrolled sample. The prevalence of vitamin D insufficiency (16-30 ng/mL) and mild deficiency (0-10 ng/mL) in hemodialysis patients enrolled in this study was 21.1% and 48.8% respectively. Both vitamin D mild deficiency and insufficiency were significantly associated with central obesity (waist circumference  $\geq 90$  cm in both sexes or BMI  $> 30$  kg/m<sup>2</sup>). Hemodialysis patients who had vitamin D mild deficiency and insufficiency were found to have a significantly lower level of HDL-c. **Conclusion:** This study concluded that there is a lack of association between serum 25(OH) vitamin D concentrations and metabolic syndrome in hemodialysis patients. However, Both vitamin D mild deficiency and insufficiency were found to be significantly associated with central obesity (waist circumference  $\geq 90$  cm in both sexes or BMI  $> 30$  kg/m<sup>2</sup>) and hemodialysis patients who had vitamin D mild deficiency and insufficiency were found to have a significantly lower level of HDL-c.

**Key words:** Hemodialysis, Metabolic syndrome, Vitamin D deficiency, Vitamin D insufficiency

**Introduction:**

The metabolic syndrome was defined by the International Diabetes Federation (IDF) criteria. Patients were diagnosed with metabolic syndrome in the presence of central obesity (waist circumference  $\geq 90$  cm in both sexes or BMI  $> 30$  kg/m<sup>2</sup>) plus at least two of the following four criteria: (1) raised triglycerides ( $\geq 150$  mg/dL) or treatment for this lipid abnormality; (2) reduced HDL-c ( $< 40$  mg/dL in men and  $< 50$  mg/dL in women) or treatment for this lipid abnormality; (3) raised blood pressure (systolic blood pressure  $\geq 130$  mmHg or diastolic blood pressure  $\geq 80$  mmHg) or treatment of previously diagnosed hypertension; and (4) raised FPG (FPG  $\geq 100$  mg/dL) or previously diagnosed type 2 diabetes. [Alberti K. G. M. M., et al. 2006]

Vitamin D deficiency plays a key role in the pathophysiology of risk factors of metabolic syndrome which affect cardiovascular system, increase insulin resistance and stimulate rennin-angiotensin-aldosterone system that cause hypertension. [Prasad P. and Kochhar A., 2016].

Metabolic syndrome has a high prevalence in hemodialysis patients and is a risk factor for cardiovascular disease, the leading cause of mortality in chronic kidney disease. [Kubrusly M., et al., 2010]. Vitamin D deficiency and insufficiency are common in patients with end-stage renal disease (ESRD) on maintenance hemodialysis. Vitamin D deficiency is associated with increased morbidity and decreased survival in patients with ESRD. [Krassilnikova M., et al. 2014]

As per National Kidney Foundation Kidney Disease Outcomes Quality Initiative guidelines, the following categories for vitamin D status were defined: severe vitamin D deficiency, 25(OH) vitamin D < 5 ng/mL; mild vitamin D deficiency, 25(OH) vitamin D between 5 ng/mL and 10 ng/mL; vitamin D insufficiency 25(OH) vitamin D between 16 ng/mL and 30 ng/mL, and vitamin D sufficiency, 25(OH) vitamin D > 30 ng/mL. [Eknayan G., et al. 2003]

**This study aimed to** identify the relationship between serum concentrations of vitamin D and metabolic syndrome in hemodialysis patients.

**Subjects and Methods**

**Patients:** This cross-sectional study was conducted from July 2016 to July 2017 in the hemodialysis unit of Minia University Hospital. 90 patients on hemodialysis were included in this study. Prior to enrollment, informed consent was obtained from all enrolled patients.

**Assessment:** All the study subjects were submitted to:

- Thorough clinical history.
- Measurement of weight, height and waist circumference. Body mass index (BMI) was

calculated by the Quetelet formula as weight in kilograms divided by height in meter squared.

- Using an appropriately sized adult cuff attached to a standard sphygmomanometer, three blood pressure readings were obtained before, during and after dialysis. The readings were then averaged.

**Laboratory evaluation:** Routine chemistry tests (lipid profile [total cholesterol, HDL-c, LDL-c, and serum triglycerides], FPG, albumin, calcium and phosphorus) were assayed using fully automated clinical chemistry auto-analyzer system Konelab 2011 (Thermo electron incorporation, Finland). Serum concentrations of parathyroid hormone (PTH) and 25(OH)-vitamin D were determined by the enzyme immunosorbent assay (EIA) method using commercial kits.

**Results**

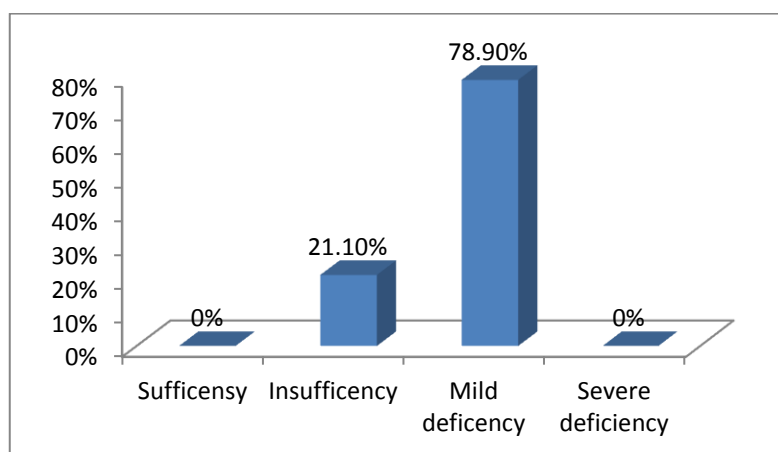
The prevalence of metabolic syndrome and its components among the hemodialysis patients enrolled in this study are presented in **table 1**. The overall prevalence of metabolic syndrome was 30.6%. The most common diagnosed abnormality was reduced HDL-c (77.8%).

**Table (1) Prevalence of metabolic syndrome and its components:**

Metabolic syndrome and its components	N (%)
Central obesity	54 (60%)
Raised triglycerides	30 (33.3%)
Reduced high-density lipoprotein cholesterol	71 (77.8%)
Raised blood pressure	47 (52.2%)
Raised fasting plasma glucose	7 (7.8%)
Metabolic syndrome	32 (35.6%)

As illustrated in **Figure 1**, the prevalence rates for vitamin D insufficiency and mild deficiency were 21.1% and 44.4% respectively.

**Figure (1): The prevalence of vitamin D sufficiency, insufficiency and deficiency:**



A comparison between demographic, clinical, and laboratory variables across the categories of vitamin D status among the hemodialysis patients enrolled in this study is shown in **table 2**. When moving from the category of vitamin D mild deficiency to vitamin insufficiency, HDL-c linearly increased (P value for trend <0.05 by t-test). A linear decreasing trend in total cholesterol was identifiable, but not reaching statistical

significance (P value = 0.07). Additionally age, triglycerides, LDL-c linearly decreased but not reaching statistical significance (P value = 0.1 in all tests). For other variables examined; namely waist circumference, BMI, FPG, systolic, and diastolic blood pressure, no linear increasing or decreasing patterns across categories of vitamin D were identified.

**Table (2) A comparison between demographic, clinical, and laboratory variables across the categories of vitamin D status:**

Variables	Vitamin D status		P value
	Insufficiency (n= 19)	Mild deficiency (n= 71)	
	Mean ± SD	Mean ± SD	
Age	42.0±10.3	48.0±13.6	0.1
Waist circumference (cm)	93.1±11.9	93.6±10.0	0.8
BMI (kg/m <sup>2</sup> )	20.9±4.9	27.2±6	0.8
Cholesterol	178±34	187±42.6	0.07
Triglycerides	124.3±41.3	147.4±62.0	0.1
HDL	42.3±7.1	38.9±7.4	0.04*
LDL	101±34.4	118.1±42.3	0.1
FBS	80.7±27.3	79.2±21.8	0.2
SBP	134.7±20.4	128.7±22.1	0.3
DBP	83.1±14.1	80.2±11	0.3

The association of vitamin D status with components of the metabolic syndrome among the hemodialysis patients enrolled in this study is explored using logistic regression as shown in **table 3**. Both vitamin D deficiency and insufficiency were significantly associated with central obesity.

**Table (3): The association of vitamin D status with components of the metabolic syndrome:**

	Vitamin D status			
	Insufficiency		Deficiency	
	OR (95% CI)	P value	OR (95% CI)	P value
	<b>Adjusted for age and sex</b>			
<b>Central obesity</b>	0.1 (0.01-0.8)	0.02*	8.1 (1.2-54)	0.02*
<b>Raised TG</b>	0.0 (0.1-2.7)	0.4	1.8 (0.3-9.2)	0.4
<b>Reduced HDL-c</b>	0.1 (0.3-68.3)	0.2	0.1 (0.01-2.0)	0.2
<b>Raised BP</b>	0.2 (0.05-1.2)	0.08	4.0 (0.8-19.0)	0.08
<b>Raised FPG</b>	0.1 (0.01-1.04)	0.05	8.9 (0.9-83.9)	0.05

**Discussion**

Our study showed that prevalence of metabolic syndrome was 30.6%. Similar results were reported by Alswat K. A., et al., 2017. In his study, the prevalence of metabolic syndrome in hemodialysis patients was 38.2 %.

In our study, the prevalence of vitamin D insufficiency and mild deficiency in hemodialysis patients was 21.1% and 28.8% respectively. These results were close to that reported by Marinelli A., et al., 2013 who reported that prevalence of vitamin D insufficiency and deficiency was 23% and 33% of the patients on chronic hemodialysis respectively.

In our study, it was found that hemodialysis patients who had vitamin D mild deficiency and insufficiency were found to have a significantly lower level of HDL-c. Kim S., et al. 2012 found that Serum 25(OH) vitamin D concentration was inversely associated with the risk of having reduced HDL-C among Korean population. Yasein N., et al., 2010 reported that vitamin D deficiency and insufficiency were significantly associated with higher level of serum triglycerides and serum LDL among osteoporotic postmenopausal Jordanian female patients.

In our study, both vitamin D mild deficiency and insufficiency in hemodialysis patients were significantly associated with central obesity (waist circumference  $\geq 90$  cm in both sexes and BMI > 30 kg/m<sup>2</sup>). Shafinaz I. S., and Moy F. M., 2016 found a that higher Body Mass Index and larger waist circumference were significantly associated

with lower serum 25(OH) vitamin D level among multi-ethnic adults in Kuala Lumpur, Malaysia.

In our study, we found a lack of association between serum 25(OH) vitamin D concentrations and metabolic syndrome in hemodialysis patients. Bonakdaran S., et al., 2016 found no significant difference in serum 25(OH) vitamin D concentrations between individuals with or without metabolic syndrome in the Iranian population. In contrast to our results, Ahmadi, F., et al., 2016 reported that both vitamin D insufficiency and deficiency were significantly associated with metabolic syndrome in Iranian patients on maintenance hemodialysis.

Our study has certain limitations. Firstly, it is a cross-sectional study and direct associations between the variables of interest could not be determined. Additionally, the lack of association between vitamin D status and metabolic syndrome can be attributed to the relatively small number of patients undergoing hemodialysis who were included. Furthermore, we could not consider factors such as amounts of sunlight exposure, calcium intake, and vitamin D intake – which could have affected the serum levels of 25(OH)D – because of the limited data. Finally, measurements were performed only once for each participant, and serial measurements over a year would be required for more accurate studies.

**Conclusion**

This study concluded that there is a lack of association between serum 25(OH) vitamin

D concentrations and metabolic syndrome in hemodialysis patients. However, Both vitamin D mild deficiency and insufficiency were found to be significantly associated with central obesity (waist circumference  $\geq 90$  cm in both sexes or BMI  $> 30$  kg/m<sup>2</sup>) and hemodialysis patients who had vitamin D mild deficiency and insufficiency were found to have a significantly lower level of HDL-c.

### Recommendations

Inclusion of more hemodialysis patients is recommended for more information about the relationship between vitamin D level and metabolic syndrome in hemodialysis patients. Serial measurements of 25(OH) vitamin D over a year would be required for more accurate studies.

### References

- 1- Ahmadi, F., Damghani, S., Lessan-Pezeshki, M., Razeghi, E., Maziar, S., & Mahdavi-Mazdeh, M. (2016). Association of low vitamin D levels with metabolic syndrome in hemodialysis patients. *Hemodialysis International*, 20(2), 261-269.
- 2- Alberti, K. G. M. M., Zimmet, P., & Shaw, J. (2006). Metabolic syndrome—a new world-wide definition. A consensus statement from the international diabetes federation. *Diabetic medicine*, 23(5), 479-484.
- 3- Alswat, K. A., Althobaiti, A., Alsaadi, K., Alkhalidi, A. S., Alharthi, M. M., Abuharba, W. A., & Alzaidi, A. A. (2017). Prevalence of Metabolic Syndrome Among the End-Stage Renal Disease Patients on Hemodialysis. *Journal of clinical medicine research*, 9(8), 687.
- 4- Bonakdaran, S., Fakhraee, F., Karimian, M. S., Mirhafez, S. R., Rokni, H., Mohebati, M., Mazidi, M., Mousavid, M., A. Fernsf, G. & Ghayour-Mobarhan, M. (2016). Association between serum 25-hydroxyvitamin D concentrations and prevalence of metabolic syndrome. *Advances in medical sciences*, 11(2), 219-223.
- 5- Eknayan, G., Levin, A., & Levin, N. W. (2003). Bone metabolism and disease in chronic kidney disease. *American Journal of Kidney Diseases*, 42, 1-20.
- 6- Kim, S., Lim, J., Kye, S., & Joung, H. (2012). Association between vitamin D status and metabolic syndrome risk among Korean population: based on the Korean National Health and Nutrition Examination Survey IV-2, 2008. *Diabetes research and clinical practice* 96(2), 230-236.
- 7- Krassilnikova, M., Ostrow, K., Bader, A., Heeger, P., & Mehrotra, A. (2014). Low dietary intake of vitamin D and vitamin D deficiency in hemodialysis patients. *Journal of nephrology & therapeutics*, 4(3).
- 8- Marinelli, A., Pistolesi, V., Rossi, V., Battista, M., Buono, A., Della Grotta, F., & Di Napoli, A. (2013). Severe 25-OH vitamin D deficiency in patients on chronic hemodialysis. *Giornale italiano di nefrologia: organo ufficiale della Societa italiana di nefrologia*, 31(5).
- 9- Prasad, P., & Kochhar, A. (2016). Interplay of vitamin D and metabolic syndrome: a review. *Diabetes & Metabolic Syndrome: Clinical Research & Reviews*, 10(2), 105-112.
- 10- Shafinaz, I. S., & Moy, F. M. (2016). Vitamin D level and its association with adiposity among multi-ethnic adults in Kuala Lumpur, Malaysia: a cross sectional study. *BMC public health*, 16(1), 232.
- 11- Yasein, N., Shroukh, W., & Hijawi, R. (2015). Serum vitamin D and the metabolic syndrome among osteoporotic postmenopausal female patients of a family practice clinic in Jordan. *Advances in clinical and experimental medicine: official organ Wroclaw Medical University*, 24(2), 245-250.