



## ASSOCIATION OF VITAMIN D DEFICIENCY WITH HYPOTHYROIDISM AMONG POPULATION ATTENDING NORTH BENGAL MEDICAL COLLEGE AND HOSPITAL

### Physiology

<b>Anup Kumar Debsinha</b>	Senior Resident, Department of Physiology, North Bengal Medical College and Hospital, Darjeeling
<b>Krishna Chandra Sarkar*</b>	Associate Professor, Department of Physiology, North Bengal Medical College and Hospital, Darjeeling *Corresponding author
<b>Shiuli Roy (Adak)</b>	Associate Professor Department of Biochemistry, North Bengal Medical College and Hospital, Darjeeling
<b>Arup Kumar Banerjee</b>	Assistant Professor, Department of Biochemistry, North Bengal Medical College and Hospital, Darjeeling
<b>Anil Kumar</b>	Professor, Department of Physiology, Faculty of Medicine and Health Sciences, SGT University, Budhera, Gurugram.

### ABSTRACT

**Background:** Vitamin D deficiency is health problem which is prevalent globally. People require sufficient levels of Vitamin D in their blood for regulation of calcium metabolism in body for normal physiological actions. Very few governments and national or international organization have given much importance on the vitamin D deficiency or declared it a public health problem.

**Objectives:** There is a high prevalence of hypothyroidism and calcium deficiency in Indian population. Vitamin D deficiency is one of the major causes of calcium deficiency in our body. With this fact we found an opportunity to study the association between hypothyroidism and Vitamin D deficiency if any, among population of sub-Himalayan region of West Bengal attending North Bengal Medical College and Hospital.

**Material and Methods:** 120 young and adult individuals between the age group 15 to 45 years comprising 60 number as control (n=60; 26 male and 34 female) and 60 as case (n=60; 24 male and 36 female) were taken in the study. Serum fT<sub>3</sub>, fT<sub>4</sub>, TSH and 25(OH) vitamin D levels were estimated using ELISA method. Patients having serum TSH > 10 mIU/ml or serum TSH levels between 6 to 10 mIU/ml along with decreased serum fT<sub>3</sub> &/or fT<sub>4</sub> levels were diagnosed as hypothyroid cases. Statistical analysis was done by SPSS.

**Results:** Serum 25(OH) Vitamin D was significantly lower in hypothyroid patients (13.30 ± 3.84 ng/ml) compared to controls (32.36 ± 5.88 ng/ml), p < 0.001. There was insignificantly decreased levels of Serum 25(OH) Vitamin D in females than male patients (t = 1.38, p > 0.05).

**Conclusion:** The study indicates that patients with hypothyroidism are frequently associated with vitamin D deficiency which suggests that they need routine screening and vitamin D supplementation along with thyroid hormone replacement therapy.

### KEYWORDS

Vitamin D Deficiency, TSH, Hypothyroidism.

#### Introduction:

Deficiency of Vitamin D is recognized throughout the world as one of the major problem of public health<sup>[1]</sup>. Almost one billion persons around the world have deficiency or insufficiency of vitamin D<sup>[1]</sup>. Over a period of last 20 years many articles published; indicate that low levels of vitamin D exist among patients of hypothyroidism. Prevalence of vitamin D deficiency is widespread in India which is around 50-90%<sup>[3]</sup>. Vitamin D deficiency is still widely prevalent among the elderly according to recent report and it also exists as an unrecognized serious problem in many populations globally<sup>[1]</sup>. It is an established fact that vitamin D maintains normal blood levels of calcium and phosphate required for bone and teeth formation and plays important role in the different physiological functions of the body.<sup>[13]</sup>, as well as different immunoregulatory and anti-inflammatory properties<sup>[19,20,21]</sup>. Deficiency of vitamin D is associated with the skeletal disorders, cardiovascular diseases<sup>[4,5]</sup>, diabetes mellitus<sup>[6,7]</sup>, multiple sclerosis<sup>[8,9]</sup>, hypertension<sup>[10]</sup>, atherosclerosis<sup>[11]</sup>, infectious diseases<sup>[12]</sup>. Vitamin D maintains normal blood levels of calcium and phosphate which in turn plays an important role in the different physiological functions of the body.<sup>[13]</sup>.

Vitamin D binds to its receptors (VDR) and mediates its effect through activation of VDR responsive genes<sup>[14]</sup>, which is also a common site action of thyroid hormone and thus affects the action of one another<sup>[3]</sup>. As a result Vitamin D deficiency is likely to increase similar problems which are associated with hypothyroidism<sup>[15,16]</sup>.

According to our knowledge, no recent studies have evaluated vitamin D levels among healthy and hypothyroid young or middle-aged population in the Sub Himalayan region of West Bengal. We took the opportunity to assess the association between vitamin D deficiency

and hypothyroidism among population who attended North Bengal Medical College and Hospital for one year period.

#### Subjects and Methods:

A total number of 120 subjects with age group between 15-45 years were included in our cross-sectional study. A control group (will be called Group I) with 60 healthy euthyroid subjects including 34 women and 26 men was formed. Similarly a case group (will be called Group II) included 60 subjects including 36 women and 24 men of recently diagnosed hypothyroidism having serum TSH level higher than 5.0 mIU/ml with normal or lower serum levels of fT<sub>3</sub> and/or fT<sub>4</sub> was taken. There was no history of treatment for hypothyroidism in the past.

#### Statistical Analysis:

Statistical analysis was carried out using Statistical package for social sciences (SPSS) software.

- The mean, Standard Deviation (SD), differences between mean values tested by using independent simple student's t-test.
- The association between study variables was assessed using Pearson's correlation analysis presented by correlation coefficient "r".
- Results considered significant or non-significant when P < 0.05 or > 0.05, respectively.

**Results:** Age and sex distribution in both the studied groups and the mean values ± S.D of all studied parameters are shown in Table (1).

Regarding age and sex, there was no statistical difference (P > 0.05) between both the groups.

**Table 1: Demographical and clinical characteristics of the participants used in the study.**

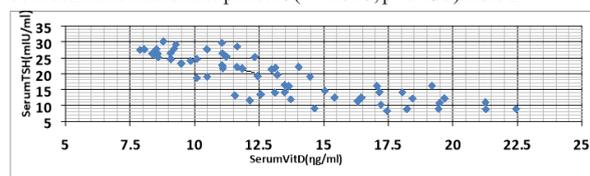
Parameters : Mean ± SD	Group I (n=60)	Group II (n=60)	t - test/ p - value
Sex	26 M (43%), 34 F (57%)	24 M (40%),36 F(60%)	p>0.05
Age (years)	34.43 ± 9.214	36.27 ± 8.634	t= 1.125, p= 0.263
BMI (kg/m <sup>2</sup> )	19.33 ± 1.99	19.59 ± 1.70	t=0.757, p= 0.45
Serum 25(OH)Vit D (ng/ml)	32.357 ± 5.884	13.301 ± 3.846	t= 20.997, p= 0.000
Serum TSH mIU/ml	3.383 ± 1.228	18.948 ± 6.776	t= 17.505, p = 0.000
Serum T3 ng/ml	1.585 ± .283	1.332 ± .238	t= 5.281, p = 0.000
Serum T4 ng/dl	1.126 ± .399	1.874 ± 1.264	t= 4.368, p = 0.000

By using t-test it was revealed that serum 25(OH) D level was significantly lower in hypothyroid subjects than in controls (t=20.997, p=0.000) as illustrated in table 1. According to the sex distribution, serum 25(OH) vitamin D levels were insignificantly decreased in females than those of male in both control and hypothyroid patients (t=.138, p>0.05) respectively, table 2.

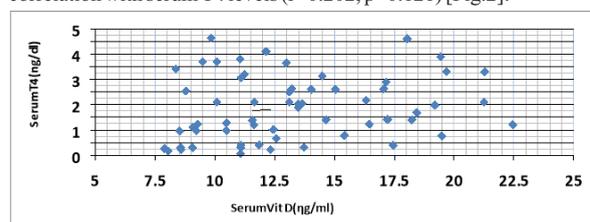
**Table 2: Mean ±SD of serum T3, T4, 25(OH) vitamin D, and TSH levels in hypothyroid patients according to sex.**

Parameters(Mean ±SD)	Male = 24	Female = 36	t-test/p- value
T <sub>3</sub> (ng/ml)	1.314 ± .251	1.345 ± .232	t= .488, p= 0.628
T <sub>4</sub> (ng/dl)	2.135 ± 1.328	1.699 ± 1.207	t= -1.317, p= .193
25(OH)vit D (ng/ml)	13.358 ± 4.069	13.217 ± 3.567	t= .138, p= 0.891
TSH (mIU/ml)	18.659 ± 6.701	19.140 ± 6.914	t= .267, p= 0.790

TSH level was significantly higher in hypothyroid subjects than that of controls (t=17.505, p=0.000) table 1. There was a non-significant difference of serum TSH between male and female patients (t=0.267, p=0.790) table 2. When serum T4 levels in hypothyroid patients were compared according to sex, we noticed a non-significant difference between male and female patients (t=-1.317, p=0.193) table 2.

**Figure 1: Correlation between serum TSH and 25(OH) vitamin D levels in hypothyroid patients.**

Regarding control group, serum 25(OH) vitamin D levels were non-significantly correlated to serum TSH levels (r= -0.142, p=0.281) and serum T4 levels (r=0.178, p=0.173). In hypothyroid patients, serum 25(OH) vitamin D levels had a significant negative correlation with serum TSH levels (r=-0.835, p=0.000) [Fig.1] but a non-significant correlation with serum T4 levels (r=0.202, p=0.121) [Fig.2].

**Figure 2: Correlation between serum T4 and 25(OH) vitamin D levels in hypothyroid patients.**

## Discussion:

### Summary of findings:

In the present study of one year duration; serum vitamin D level was estimated among 120 subjects in the age group of 35.35 ± 8.93 years of North Bengal Medical College and Hospital consisting of two groups;

one Euthyroid and one hypothyroid group. Vitamin D deficiency was considered when serum 25(OH) Vitamin D levels were below 20ng/ml, insufficiency when between 20-30ng/ml and sufficient when levels were above 30ng/ml<sup>[17]</sup>.

If we consider the prevalence of vitamin D deficiency in hypothyroid group, vitamin D levels in male were below 20ng/ml in 23(95.8%) cases. The mean vitamin D level in male among hypothyroid subjects was 13.21 ± 3.57ng/ml and below 20ng/ml in 34(94.4%) female. No male or female in hypothyroid group were found with sufficient vitamin D level (> 30 ng/ml). In control group, we found a non-significant correlations between serum 25(OH) vitamin D and serum T<sub>3</sub> levels.

### Comparative other similar studies:

Prevalence of Vitamin D deficiency was noted in 92% autoimmune hypothyroidism cases was the observation of few other workers, in a study which is relevant to our finding also<sup>[29]</sup>.

In hashimoto's thyroiditis, a serum 25(OH) vitamin D was significantly lower than the controls and with the duration of Hashimoto's thyroiditis; severity of vitamin D deficiency increases.<sup>[30]</sup>

Some researchers also observed no deficiency of vitamin D in early stages of autoimmune thyroid diseases<sup>[31]</sup>.

Serum levels that vitamin D are significantly more decreased in females than males was the observation of some workers<sup>[28]</sup> but in our study this was insignificantly decreased.

Some experimental study reported that the possibility of low thyroid hormones may be contributed by deficiency of Vitamin D in female patients with graves' disease of new on set<sup>[33]</sup>

Vitamin D deficiency in 40% of women and around 20% of men (p < 0.005) was found in a study in Japan including 200 euthyroid patients with Graves' disease<sup>[32]</sup>. The discrepancies between these studies can be explained by differences in the selection of patients, dietary vitamin D intake, and exposure to sunlight, and seasonal variations.

We observed in this study that there was a significant negative correlation between serum TSH levels and serum 25(OH) Vitamin D levels on Pearson's correlation analysis (r= - 0.835, p<0.005). In this study, the hypothyroid patients had significantly lower level of serum 25(OH) vitamin D as compared to controls (p<0.005) and the results are in close agreement with that of study conducted by Amal Mohammed Husein Mackaway et al.; suggesting that the deficiency of serum vitamin D levels was significantly associated with degree and severity of hypothyroidism<sup>[23]</sup>.

We observed that 95% hypothyroid patients had serum 25(OH) Vitamin D levels below 20ng/ml(vitamin D deficiency) and 5% had serum 25(OH) Vitamin D levels between 20 to 30ng/ml having Vitamin D insufficiency . But no subject in the hypothyroid group had sufficient levels (>30ng/ml) of Vitamin D. On the other hand, healthy controls had higher levels of vitamin D.

Our results revealed decreased serum 25 (OH) vitamin D levels in females than those of males in both the controls and hypothyroid patients. Similar type of observations were found in other studies where serum vitamin D levels were significantly more decreased in females than male controls and patients<sup>[22,24,25,26,27]</sup>. One of the probable cause may be; males are engaged in more outdoor activities where as females mostly do indoor activities as well as difference in dietary intake.

### Limitation of the study

In our study, the sample size was too small. The study was limited in its ability to conclude that vitamin D status is directly related to the pathogenesis of hypothyroidism. We could not measure the serum PTH levels and clarify its effect on vitamin D and hypothyroidism.

Further studies with a larger number of subjects are needed to determine whether hypothyroidism is caused by vitamin D deficiency or vice versa. Many study including ours also pointing to the involvement of vitamin D in the pathogenesis of hypothyroidism. So, screening vitamin D levels in patients with thyroid diseases encourage the advisability of vitamin D supplementation.

**Conclusion:**

There is increased incidence and prevalence of vitamin D deficiency in India and other countries due to rapid economic development, dietary changes, life style modification, like inadequate sunlight exposure<sup>[18]</sup>. However, data from epidemiologic studies in India are still limited. If we consider the high prevalence of vitamin D deficiency in India, there is a need for further research to identify the major factors responsible for vitamin D deficiency, despite of abundance of sunlight in our country. Vitamin D deficiency affects all age groups. Strategies like mass awareness programme about adequate exposure to sunlight, dietary fortification with vitamin D should be undertaken irrespective of the socio-economic status. This strategy may be adopted and implemented for prevention and control of vitamin D deficiency as well as hypothyroidism as a part of National Disease Prevention and Control programme. It will not be unjustified if vitamin D is combined with thyroid hormones which are given as medication to hypothyroid patients.

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