Prevalence and Symptomatic Correlation of Vitamin D Deficiency in Individuals Visiting for an Annual Health Check-up – A Cross-sectional Study

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Abstract

Vitamin D plays a crucial role in electrolytic reabsorption and calcium homeostasis, thereby maintaining skeletal and extra-skeletal integrity. In addition, the literature establishes its potency to prevent insulin-dependent diabetes, cardiovascular diseases, malignancies, and various autoimmune disorders. Therefore, determining vitamin D deficient states and its correlation with systemic factors plays a notable role in clinical medicine. The present cross-sectional study was conducted to determine the Prevalence and symptomatic correlation of vitamin D deficiency in individuals visiting an annual health check-up at a quaternary care hospital, Lucknow, Uttar Pradesh, India. The study stipulated the prevalence rate of serum vitamin D deficiency levels was 91 % among healthy individuals. The overall mean value of serum vitamin D was 17.54 ng/mL. With 91.2 % symptomatic individuals, the mean age of patients was 48.08 ± 10.33 years, with the majority of males (61.53 %). The mild, moderate, and severe grades of vitamin D deficiency were seen in 32.53 %, 53.01 %, and 14.45 % of study participants, respectively. On applying Pearson correlation, there is a strong negative correlation between the total symptom score (12.98) and serum vitamin D levels with an Rvalue of -0.788. The study reveals the higher prevalence rate of vitamin D deficiency among apparently healthy participants. The associated symptoms among moderate and severe vitamin D deficient study participants indicate the alarming stage in the country. The study concluded the need for comprehensive public health programs concerning the advantages of sun exposure and sufficient intake of fortified resources in the 21st-century lifestyle.

Keywords: Bone pain, Cholecalciferol, Depression, Muscle pain, Prevalence, Symptoms, Vitamin D deficiency.

Introduction

Vitamin D, as a micronutrient, plays a variety of major and minor roles in maintaining the overall systemic health of individuals, including electrolytic reabsorption, muscle contraction, calcium, and phosphorous homeostasis, and even affects the immune system, especially T - cell-mediated immunity. Out of primary variants, vitamin D2 and D3, the active form cholecaliferol (1, 25 – dihydroxy vitamin D3), plays a noteworthy function in scientific medicine, thereby maintaining skeletal integrity. Due to its physiological functions, vitamin D is potent in preventing inflammatory bowel diseases, insulin-dependent diabetes, cardiovascular diseases, malignancies, and various autoimmune disorders and infectious diseases [1 - 4].

The ultraviolet rays from the sunlight are the primary source that endogenously synthesizes cholecalciferol on the skin surface. Animal sources, including dairy products or fatty fish and plant sources such as mushrooms, even contribute to vitamin D sources to a lower extent.

The reduced sunlight exposure attributed to melanin pigmentation, types of clothes worn by the individuals, and more importantly, solar zenith angle, latitude, seasonal variations, atmospheric pollution, insufficient intake of sources rich in vitamin D are the major etiologic factors associated with vitamin D deficiency states. From the clinically apparent manifestations ranging from rickets in children to periosteal reactions in adults; to the subclinical manifestations, vitamin D deficiency may lead to death in severely deficient individuals [5, 6].

The studies in literature reported 30 - 50 % vitamin D deficient states in all age groups of countries like Europe, the United States, Canada, and New Zealand. In the past two decades. even in intense sunlight-prone countries like India, Iran, China, Saudi Arabia, etc., the studies have reported an estimated prevalence of vitamin D deficiency between 30 93% of the population. Health care professionals play a major role in the evaluation of vitamin D deficient states in the different age groups of individuals and its systemic health implications [5, 7]. There are many studies on the Prevalence of vitamin D deficiency; very few published however, on its symptomatic correlation in the literature. Therefore, the present study was attributed to determine the Prevalence and symptomatic correlation of vitamin D deficiency in individuals visiting for an annual health checkup at a quaternary care hospital. Lucknow, Uttar Pradesh, India.

Materials and Methods

The present cross-sectional study was conducted in the department of Preventive health, at 330 bedded quaternary care Apollo Medics hospital in Lucknow, Uttar Pradesh. The approval from Medical Ethics Committee was taken before commencing the study. The written consent was obtained from all the participants. The study was conducted between February 2019 and April 2020 for the duration of 14 months.

Study Population

A total of 100 study participants reported for the annual health check-up to the department with prescribed serum vitamin D levels tests, and 91 met the inclusion criteria. Out of which, 56 were males, and 35 were females.

Inclusion Criteria

- 1. Patients between the age of 18 80 years.
- 2. Both symptomatic and asymptomatic patients, prescribed with serum vitamin D level test.
- 3. Patients agreed to participate and sign for written consent.

Exclusion Criteria

- 1. Patients with a history of vitamin D supplements.
- 2. Patients with diagnosed hepatic or renal disorders.
- 3. Patients with diagnosed metabolic bone disorders.
- 4. Pregnant and lactating females.
- 5. Patients with diagnosed other systemic disorders like diabetes, hypercortisolism, etc.

Data Collection and Questionnaire

The patients visiting the department of Preventive health were enrolled after meeting the inclusion criteria. All the study participants were explained about the methodology, and consent were obtained. All questions were asked in the local language per the understanding of the study participants.

A structured questionnaire was used to record the demographic details and associated symptoms of the study participants. The demographic details include name, age, gender, weight, height, address, educational qualifications, socioeconomic status, etc. In the present study, symptoms were recorded as follows:

- 1. Group 1 Bony Symptoms: Joint pain, back pain associated with the spine, etc.
- 2. Group 2 Muscular Symptoms: Myalgia, muscle weakness, muscle cramps, etc.
- 3. Group 3 Generalized Symptoms: Weakness, lethargy, morning stiffness, etc.
- 4. Group 4 Psychogenic symptoms: Depression, schizophrenia, mood swings, etc.

To maintain the uniformity, the symptoms were further graded according to severity (1 to 9), where the score 1-3 was categorized as mild, 4-6 as moderate, and 7-9 as severely symptomatic.

Sample Collection and Processing

The blood samples were collected from the study participants between 8:00 to 10:00 am for testing in NABH certified pathology laboratory at Apollo Medics hospital, Lucknow, Uttar Pradesh. The tests were performed using fully automatic machinery under the guidance of a senior pathologist. The deficient vitamin D levels were classified as –

- 1. Mild = 20 29 ng/mL.
- 2. Moderate = 10 19 ng/mL.
- 3. Severe = < 10 ng/mL.

After obtaining demographic data, questionnaire records, and blood samples, the data was taken into a required format for further analysis. The descriptive data was mentioned in mean \pm standard deviation and percentages. To determine the significance of obtained data, the Chi-square test was used with a p-value less than 0.05. The data recorded were tabulated and analyzed using software version 21 (SPSS Inc., Chicago, IL, USA).

Results

The Prevalence of study participants with vitamin D deficiency was 91 % in the present study population. Out of 91 study participants enrolled in the study, 56 (61.53 %) were males, and 35 (38.46 %) were females. The majority of males and females had moderate levels of deficiency, followed by mild levels. The present study reveals no significant difference in both genders in terms of vitamin D deficient severity with a p-value of 0.707. In the present study, the majority of study participants belong to the 5th decade of life, followed by the 4th decade. The mean age of patients was 48.08 ± 10.33 years in the present group of study participants (Table 1, Table 2, Figure 1).

Age Distribution	Mild	Moderate	Severe	Total	Percentage (%)
(years)	(20 - 29 ng/ml)	(10 -19 ng/ml)	(1 -9 ng/ml)	(n=91)	8 ()
21 - 30	2	2	1	5	5.49
31 - 40	3	7	5	15	16.48
41 - 50	13	14	4	31	34.06
51 - 60	14	17	1	32	35.16
61 – 70	3	4	0	7	7.69
71 - 80	0	0	1	1	1.09
Total	35	44	12	91	100

Table 1. Age-wise Distribution of Study Participants according to the Severity of Vitamin D Deficient States

Table 2. Gender Wise Distribution of Study Participants

Gender Distribution	Mild	Moderate	Severe	Total	Percentage (%)	
	(20 - 29 ng/ml)	(10 -19 ng/ml)	(1 -9 ng/ml)	(n=91)		
Female	15	15	5	35	38.46	
Male	20	29	7	56	61.53	

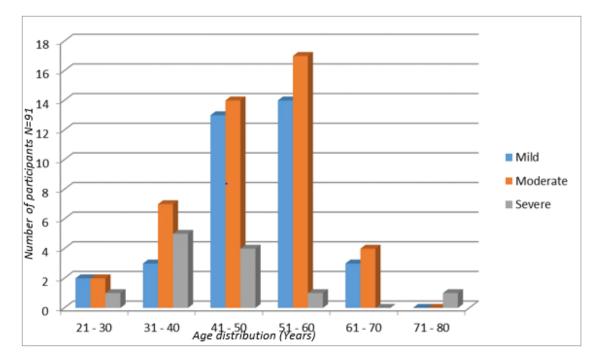


Figure 1. Age-wise Distribution of Symptomatic Study Participants

Out of 91 vitamin D deficient study participants, 83 (91.2 %) were symptomatic. Among 8 (8.79 %), asymptomatic patients only mild vitamin D deficiency was seen. On applying chi-square test, the maximum numbers of individuals were in the moderate vitamin D deficiency category with p-value of 0.00 (Table 3).

Symptoms	Mild	Moderate	Severe	Total	Percentage (%)
	(20 - 29 ng/ml)	(10 -19 ng/ml)	(1 -9 ng/ml)	(n=91)	
Present	27	44	12	83	91.2
Absent	8	0	0	8	8.79

Table 3. Vitamin D Deficiency - Severity Wise Distribution of Individuals

Bony symptoms such as joint pain or back pain were seen in the majority of study participants (75), followed by muscular (72) and generalized symptoms (71), while psychological symptoms were seen in 17 study participants only. On applying ANOVA test, there is a statistically significant difference between the occurrence of different symptoms while the psychological symptoms were found less compared to other symptoms (P-Value of < .00001) (Table 4).

Table 4. Symptoms Wise Distribution of Study Participants based on Severity

	Study Participants (n)				
Symptoms	Mild Symptoms	Moderate Symptoms	Severe Symptoms (Score 7-9)		
	(Score 1-3)	(Score 4-6)			
Bony Symptoms	23	33	19		
Muscular Symptoms	20	30	22		
General Symptoms	15	29	27		
Psychological Symptoms	8	6	3		

The overall mean value of serum vitamin D is 17.54 ng/mL, and the mean value of the total symptom score is 12.98. The mild, moderate, and severe levels of serum vitamin D levels were seen in 32.53 %, 53.01 %, and 14.45 % of study participants, respectively. On applying

the Pearson correlation coefficient formula, there is a strong negative correlation between the total symptom score and serum vitamin D levels with an R-value of -0.788 (p-value = 1.84) (Figure 2).

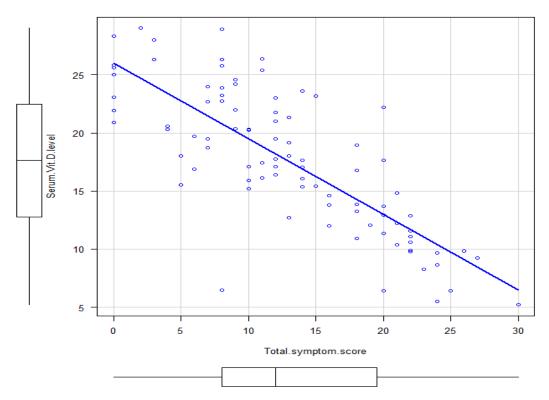


Figure 2. Correlation between Serum Vitamin D Levels and Total Symptom Score

All the symptoms have a moderate negative correlation with serum vitamin D levels except for psychological symptoms where there is a weak negative correlation. The mean serum values of vitamin D based on symptoms are listed in Table 5.

Mean Serum Vitamin D Levels (ng/ml)					
Symptoms	No Symptoms (Score 0)	Mild Symptoms (Score 1-3)	Moderate Symptoms (Score 4-6)	Severe Symptoms (Score 7-9)	Correlation of Serum Vit D with Symptom Score (r value)
Bony Symptoms	20.72	22.52	15.94	11.61	- 0.6478
Muscular Symptoms	20.15	15.95	16.42	18.28	- 0.5669
General Symptoms	22.39	19.55	19.43	10.8	- 0.6933
Psychological					
Symptoms	17.96	19	15.58	7.17	- 0.2775

Table 5. The Mean Serum Vitamin D Levels based on Symptom Score

Discussion

In the current scenario, Vitamin D deficiency is considered one of the major medical worldwide. conditions The physiological variants of vitamin D2 and D3 play a significant role in clinical medicine ranging from maintaining skeletal and extra-skeletal integrity to its screenplay in supporting T cellmediated immunity. The noteworthy task of vitamin D in preventing various clinical and subclinical infections and disorders such as cardiovascular diseases, inflammatory bowel malignancies, diabetes, metabolic disease, disorders, etc., thereby proves its importance as a micronutrient. The major risk factors associated with vitamin D deficiency include inadequate sun exposure depending on geographical location, seasonal variations, atmospheric pollution, skin colour, types of clothing worn by population, and insufficient intake of vitamin D in terms of animal (fatty fish, fortified eggs) and plant (mushrooms) sources [2, 3, 8, 9].

With today's healthcare system and changing lifestyle in the 21st century, the deficient states of vitamin D are seen in approximately 1 billion populations, according to global estimation. The Prevalence of vitamin D deficiency in the present study population is 91 %. A study has conducted by [10], in her review published in 2004, stated that the community and hospital-based studies in India have a prevalence rate of 50 % to 94% and 37 % to 99 %, respectively, among healthy individuals. The authors further concluded that modernization, cultural practices, pollution, unplanned pregnancies, and increased use of sunscreen lotions are the major causes leading to vitamin D deficiency among all age groups. The above review is in accordance with the present study [10].

The reported Prevalence of vitamin D deficiency in healthy Indians is as high as 70 % to 100% from all parts of the country, which is again in accordance with the present study. A

similar study on vitamin D deficiency performed in the same city in 2004 on healthy individuals stated 78.3 % deficient states. Another hospital-based cross-sectional study by [12] concluded a higher vitamin D deficiency prevalence rate among patients visiting the medical outpatient department in Mahad. The similarity between both the studies can be due to environmental factors, changing indoor lifestyles, insufficient nutritional intake, and lack of awareness among the population regarding the sources enriched in vitamin D [11, 12].

In the present study, the mean age of study participants is 48.08 ± 10.33 years, with the majority of individuals belonging to the 4th and 5th decade of life. The study published by [13], stated that the potential risk of vitamin D deficiency is in direct association with increasing age. The reduced potential of vitamin D synthesis by the human body with age and lack of residential care leading to the insufficient synthesis of vitamin D in the body can be the associated factors suggesting a high prevalence rate of vitamin D deficiency in elderly individuals [13, 14]. The studies contradicting the present study was published by [15], stating the higher Prevalence of vitamin D deficiency in the adult population. The authors further supported the study by concluding that the increase in awareness regarding vitamin D supplements among the younger generation and changes in lifestyle such as clothing habits, reduced outdoor activities, etc [15]. Most of the studies published in the literature stated no significant variation in vitamin D deficiency among different age groups. This can be attributed to the similar age study participants included in the study thereby, further suggesting the need for comprehensive research on vitamin D deficiency among all age groups, ranging from children to the geriatric population [7].

The current study stated majority of males (61.53%) are in the present deficient population. [13] included only males in the

study determining the Prevalence of osteoporosis in the aged 50 years and above population. He concluded the severity of vitamin D deficiency and osteoporosis is more relevant in females as per literature, but the present study stated that the males are even equally deficient in vitamin D [13]. Another case-control study conducted on 120 males suffering from acute myocardial infarction seeking treatment in AIIMS, Delhi, concluded the deficiency of vitamin D is strongly associated with cardiovascular diseases in males [16]. The above-stated studies showed higher Prevalence in males as per the present study, while various studies in literature also showed the greater Prevalence of vitamin D deficiency among pregnant and postmenopausal women [10].

The mean serum levels of vitamin D in the present study participants are 17.54 ng/mL. In the present study, the mild (20-29 ng/mL), moderate (10-19)ng/mL) and severe (<10ng/mL) levels of serum vitamin D levels were seen in 32.53 %, 53.01 % and 14.45 % of study participants respectively. The serum levels < 20 ng/mL were seen in 67.46 % of study participants, which is in accordance with the below studies published in the literature. A review published by Aparna P involving the community and hospital-based vitamin D deficiency prevalence studies considers the serum 25 (OH) D levels less than 20 ng/mL as deficient [10]. Other hospital-based studies published by [12, 17] among the Indian population stated that similar levels of serum vitamin D levels (< 20 ng/mL) were seen in 65.4 % and 70 % of study participants, respectively. A similar study conducted by Roy A et al. among AIIMS, Delhi study participants were stating < 10 ng/mL and 10 - 30 ng/mLserum vitamin D levels among 79.2 % and 19.2 % cases, respectively, enrolled in the study [16]. Another Delhi based hospital study in consonance with the present study, published by [18] among healthy individuals (above 50 years) stated that the severe vitamin D deficient levels of < 20 ng/mL were seen among 91.2% study participants and serum levels between 20–30 ng/mL were seen in 6.8 % participants only [18].

The present study revealed that general (78.02%), bony (82.41%), and muscular (79.12%) symptoms were seen in the majority of the study participants, which is in consonance with the study published in literature [19, 20]. [19] conducted a study on 276 individuals with musculoskeletal pain and further concluded that 63.4 % of participants were diagnosed with vitamin D deficiency. [20] conducted a study on 150 participants diagnosed with musculoskeletal pain, in which 93 % of patients had vitamin D deficiency. Another published research by Stewart and Hirani concluded that depressive symptoms are strongly associated with clinical vitamin D deficiency, which is not in accordance with the present study stating that only 18.68 % of study complain participants of psychological symptoms [21]. A strong association between vitamin D deficiency and musculoskeletal symptoms was seen in present study participants, while psychological symptoms showed a weak correlation. Though the majority (91%) seems to be symptomatic, these people didn't present to the hospital for these symptoms. The present study is single centered along with the limited number of study participants enrolled. In the future, we recommend the multi-centred study in the same geological location, along with a large study population. The study also reveals the need for a validated questionnaire and clinical scoring system to evaluate the signs and symptoms associated with vitamin D deficiency.

Conclusion

The present study concluded that vitamin D deficiency is heading towards the silent epidemic. This stresses on increase in awareness regarding the advantages of routine sunlight exposure and intake of fortified rich sources. Previously, the increased burden on health care professionals was more associated with compromised individuals, but in the present 21st century, the alarming prevalence rate of 91 % among healthy individuals has raised a question on the modernization, insufficient dietary intake, and lastly, on the country's health care system. With the known consequences of its deficiency, the present condition needs due attention through comprehensive public health programs.

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Conflict of Interest

We have no conflict of interest.

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