

Seasonal Variation in Vitamin D levels in Saudi Patients with Type 2 Diabetes Mellitus in Western Saudi Arabia

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Abstract

Introduction: The impact of seasonal changes over serum vitamin D has been addressed in several previous studies. We conducted a cross sectional study to investigate the status of vitamin D seasonal variation in an ambulatory adult population of Jeddah, one of the western most regions of the Saudi Arabia.

Method: A cross-sectional single centre study was conducted in 4053 patients with type 2 diabetes mellitus (T2DM). Patients with T2DM attended the Diabetes Centre at King Fahad Armed Forces Hospital, Jeddah, Saudi Arabia between January 2018 and December 2018 were recruited.

Results: A total of 4059 subjects with T2DM (27.5% females and 72.5% males) with mean age of 53.9 ± 16.5 years entered into the study. The mean serum 25(OH)D levels varied across the various months from the lowest level at 46.1 ± 26.6 nmol/l in August to the highest level at 64.9 ± 30.3 nmol/l in January. The mean serum 25(OH)D levels varied significantly across the various seasons from the lowest level at 52.0 ± 28.6 nmol/l in the summer to the highest level at 62.2 ± 30.6 nmol/l in the winter ($p < 0.0001$). Females had significantly higher serum 25-OHD concentrations than the males (59.9 ± 31.8 and 53.1 ± 27.4 years respectively, $p < 0.0001$). Serum 25(OH)D levels varied significantly higher in females across the different seasons except for fall season. The subjects aged higher than 70 years had the highest serum 25(OH)D levels whereas, the subjects aged less than 30 years had the lowest values and the differences between the age groups of 70 years and below was significant. There were statistically significant wide variations in serum 25(OH)D across the various seasons in all decades of age.

Conclusions: This study indicates that seasonal changes lead to significant serum vitamin D variations with the lowest values in the summer and in the fall and the highest values the winter and in the spring.

Keywords: Seasonal Variation in Vitamin D levels

Introduction

Vitamin D is an important factor for regulation of bone metabolism. Several studies have shown a link between vitamin D deficiency and the development of many clinical conditions such as diabetes [1,2]. The primary source of vitamin D is exposure to solar UV-B radiation [3-5]. Several factors including the season, strength of the UV rays, age, place of residence, duration of UV exposure, intake of vitamin, physical activity and the amount of pigment in the skin contribute to the production of vitamin D₃ in the skin [5-11]. These observations provide a rationale for predicting the status of serum vitamin D in high risk individual [1,2]. The impact of seasonal changes over serum vitamin D has been addressed in several previous studies [12,13]. Lower serum vitamin D in winter months compared with summer has been shown [12-17].

Current studies confirm that the prevalence of vitamin D deficiency in the general world population is actually as high as 50-80%, even occurring in countries located in geographical areas which receive sunshine year-round [18]. The Middle East and the North African region in general including Saudi Arabia have very high prevalence of vitamin D deficiency even in the normal asymptomatic population [19-21]. The prevalence of type 2 diabetes mellitus (T2DM) in Saudi Arabia is one of the highest reported in the world, reaching up to 30% in a recent study.²² It has been demonstrated that vitamin D deficiency is associated with T2DM [23-26].

Despite the potentially enormous influence of vitamin D on health, there is a paucity of population-based data on the seasonality of serum vitamin D levels in patients with T2DM [25]. However, the detailed empirics of when serum 25(OH)D levels peak and trough during the year has not heretofore been well characterized with large sample size studies in Saudi Arabia. Our goal was to establish the prevalence of vitamin D seasonal variation in an ambulatory adult population of Jeddah, one of the western most regions of the Saudi Arabia.

Methods

A cross-sectional single centre study was conducted in 4059 patients with T2DM attending the Diabetes Centre at King Fahad Armed Forces Hospital, Jeddah, Saudi Arabia between January 2018 and December 2018 were recruited. Eligible patients were 20 years or older. Exclusion criteria were known hepatic or renal disease, metabolic bone disease, malabsorption, hypercortisolism,

malignancy, immobility for more than one-week, pregnancy, lactation, and medications influencing bone metabolism. The serum concentration of 25(OH)D was measured by competitive protein binding assay using kits (Immunodiagnostic, Bensheim, Germany). Glycosylated hemoglobin (HbA1c) was measured by the high performance liquid chromatography method (Bio-Rad Laboratories, Waters, MA, USA). The total number of cohort were separated on basis of age values into six groups: 20-29 years, 30-40 years, 40-49 years, 50-59 years, 60-70 years and ≥ 70 years. The study was approved by the ethical committee board of King Fahad Armed Forces Hospital.

Statistical Analysis

Data are presented as means \pm standard deviation (SD) or numbers (%). Quantitative variables were compared between two groups by using the Student's test. Differences in categorical variables were analyzed using the chi-square test. The data were analyzed by one-way analysis of variance (ANOVA), to estimate the significance of different between groups. The relationship between continuous variables was assessed using coefficients of correlation. P value < 0.05 indicates significance. The statistical analysis was conducted with SPSS version 23.0 for Windows.

Results

A total of 4059 subjects with T2DM (2944 (27.5%) females and 1115 (72.5%) males) with mean age of 53.9 \pm 16.5 years entered into the study (Table 1). The mean age of females and males were non-statically significant different (54.1 \pm 16.5 and 53.8 \pm 16.5 years respectively, $p=0.6$). The mean and median 25-OHD concentrations were 58.0 \pm 30.8 and 52.0 respectively.

Variable	Values	
Total	4059	
Age (years)	53.9 \pm 16.5	
Gender	Male	1115 (27.5)
	Female	2944 (72.5)
HbA1c (%)	7.7 \pm 1.9	
25-hydroxyvitamin D (nmol/L)	58.0 \pm 30.8	

Table 1: Patient characteristics [mean \pm standard deviation or number (%)].

The mean serum 25(OH)D levels varied across the various months from the lowest level at 46.1 \pm 26.6 nmol/l in August to the highest level at 64.9 \pm 30.3 nmol/l in January (Figure 1). The mean serum 25(OH)D levels

varied significantly across the various seasons from the lowest level at 52.0 ± 28.6 nmol/l in the summer to the highest level at 62.2 ± 30.6 nmol/l in the winter ($p < 0.0001$). Serum 25(OH)D levels in each season varied across sexes. Females had significantly higher serum 25-

OHD concentrations than the males (59.9 ± 31.8 and 53.1 ± 27.4 years respectively, $p < 0.0001$). Serum 25(OH)D levels varied significantly higher in females across the different seasons except for fall season (Figures 2 A and 2B).

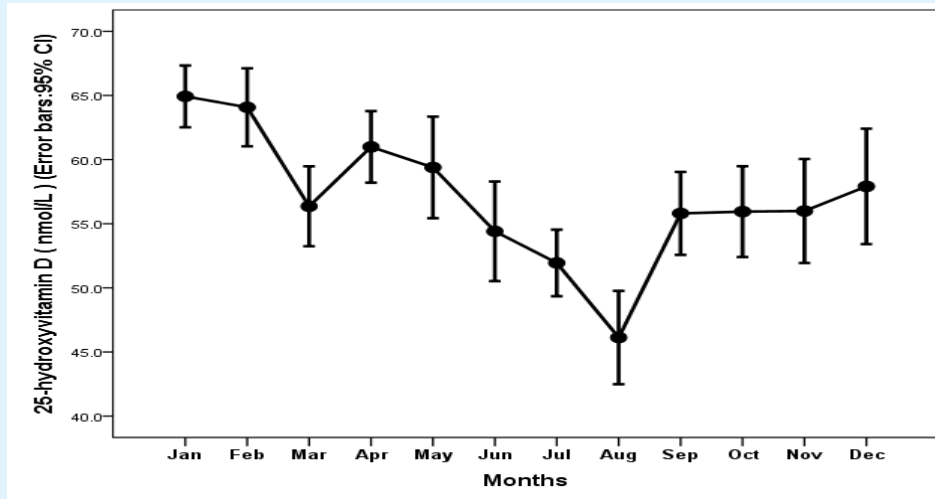


Figure 1: Monthly mean levels of 25-hydroxyvitamin D (CI, Confidence interval).

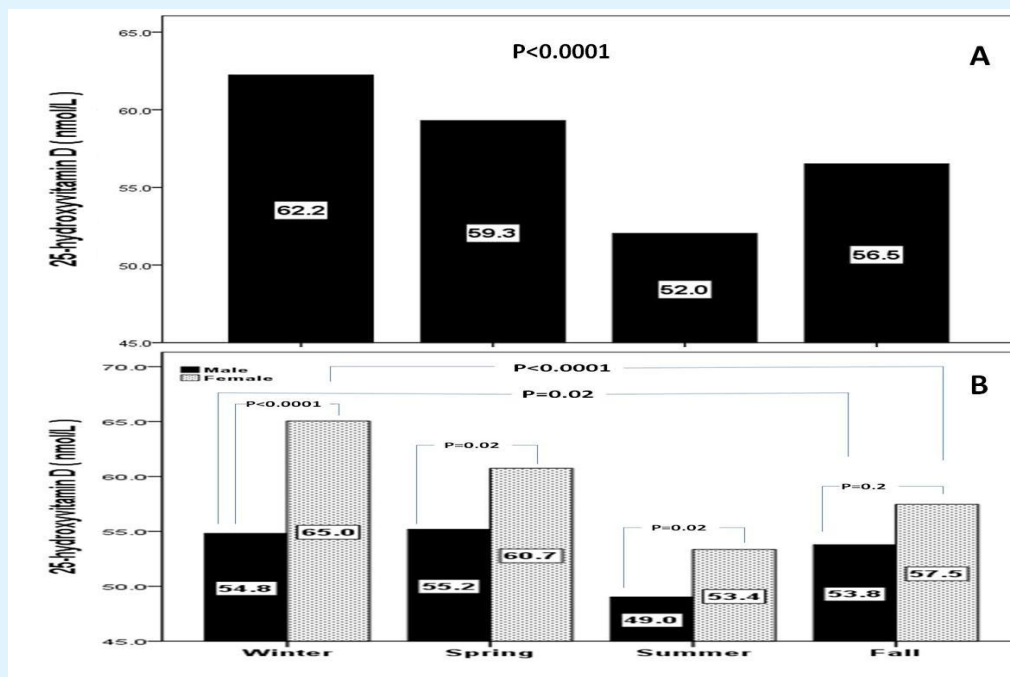


Figure 2: Serum 25-hydroxyvitamin D values in the study population across various seasons (A) and according sex (B).

Serum 25(OH)D levels were positively increased with age (Figure 3 and 4). The subjects aged higher than 70 years had the highest serum 25(OH)D levels whereas, the subjects aged less than 30 years had the lowest values

and the differences between the age groups of 70 years and below was significant (Figure 5). There were statistically significant wide variations in serum 25(OH)D across the various seasons in all decades of age (Table 2).

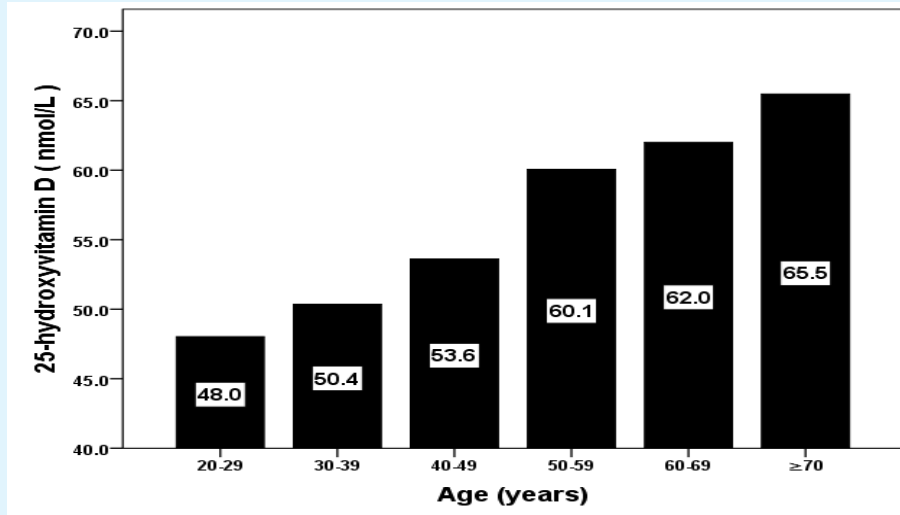


Figure 3: Serum 25-hydroxyvitamin D values in the study population across various seasons according age.

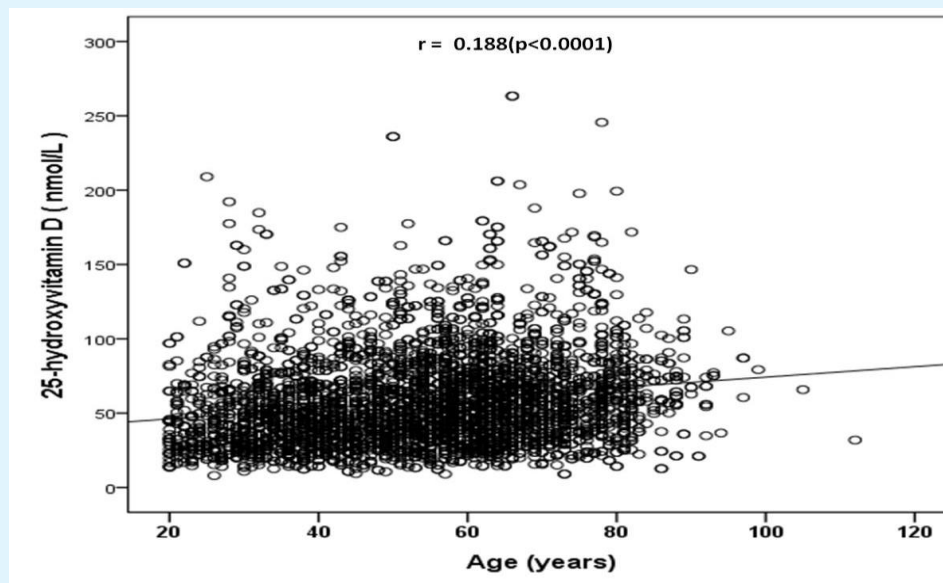


Figure 4: Correlation of 25-hydroxyvitamin D concentration and age.

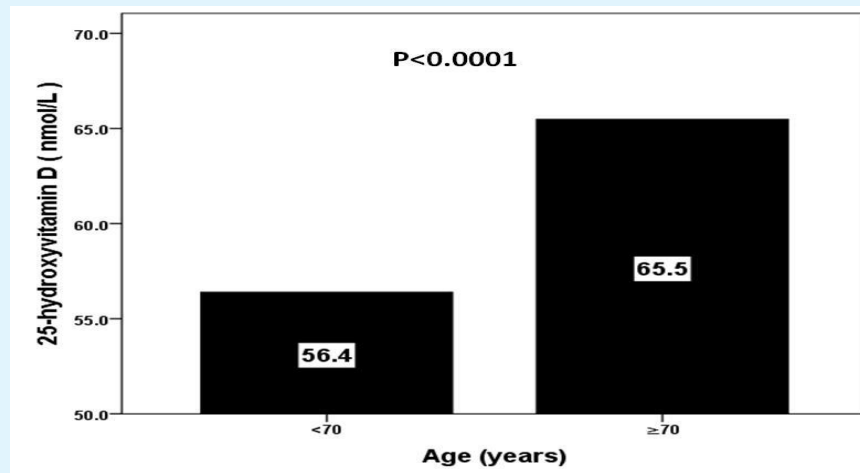


Figure 5: Serum 25-hydroxyvitamin D values in the study population according age above and below 70 years.

Parameters	20-29	30-39	40-49	50-59	60-69	70≤
Numbers	323 (8.0)	622 (15.3)	618 (15.2)	874 (21.5)	884 (21.8)	738 (18.2)
Winter	40.0 ±28.5	53.5 ±29.3	56.6 ±26.7	61.6 ±30.1	70.1 ±27.7	74.1 ±33.4
Spring	48.5 ±31.9	52.4 ±29.1	55.0 ±25.3	65.2 ±31.8	59.1 ±33.7	66.6 ±35.6
Summer	48.9 ±34.0	45.7 ±20.5	52.1 ±33.3	50.5 ±26.0	57.2 ±30.6	54.7 ±27.3
Fall	41.5 ±28.2	47.2 ±26.9	48.0 ±25.9	62.3 ±29.1	58.7 ±33.9	64.6 ±30.5
Total	48.0 ±30.7	50.4 ±27.1	53.6 ±28.3	60.1 ±29.9	62.0 ±31.7	65.5 ±32.7
P value	0.6	0.02	0.06	<0.0001	<0.0001	<0.0001

Table 2: Variations in serum 25-hydroxyvitamin D by mean±SD in different decades of age across various seasons [mean±standard deviation or number (%)].

Discussion

The findings of this study indicated variations in mean serum 25(OH)D levels across the various seasons and in relation to age and sex. The serum 25(OH)D level in the summer months particularly in young was at the lowest level versus other seasons. The results of this study are in disagreement with several published studies that addressed the relation between the seasonal changes and the status of vitamin D where the lowest 25(OH)D levels were in winter [12,13,17,27]. These observations indicate that seasonal changes should not consider the exclusive cause of vitamin D variations but many other factors are also contributed to the changes in serum vitamin D over the different seasons [28-30]. Variation between countries regarding dietary vitamin D intake, food fortification regimen and supplement use may also contribute to the differences. Also, differences in laboratory methods for measuring 25(OH)D can contribute to variations between the studies. In addition,

seasonal changes can substantially influence the cutaneous production of 25(OH)D. Moreover, the causes of low 25(OH)D levels in summer could be due to changing life style with people adopting a more sedentary life, little exposure to sunlight because of the hot weather in summer, reduced outdoor activity, changes in dietary habits. It is possible that a patient with sufficient 25(OH)D serum levels at the end of winter may be deficient at the end of summer. In our study, there was a decrease in 25(OH)D serum levels of almost 22% from March to August (56.4±31.3 and 46.1±26.6 nmol/l respectively). These monthly patterns are in contrast to the results of other studies [31-33].

In this study, lower age was related to low serum vitamin D in winter months and so should be considered as a risk factor of vitamin D deficiency in the winter. Despite the low ability of skin in synthesis of vitamin D in elderly population, serum 24-OHD in older group of this study was close to normal indicating low synthesis of 1, 24 dihydroxycholecalciferol by ageing and resultant

higher 25-OHD levels [6,34,35]. We found that females had significantly higher serum 25-OHD concentrations than the males. In Canada, gender differences were significant, with deficiency prevalence being 30% for males and 24% for females [36]. In this regard, a large meta-analysis across the world also showed that 25(OH)D levels were higher in females than in males [37]. On the contrary, in Iranian subjects, females had slightly lower levels of 25(OH)D than males (20.6 versus 23.2 ng/mL) [38]. Another explanation of the discrepancy could be a greater awareness of osteoporosis and the need for calcium and vitamin D intake among postmenopausal women.

We had several limitations. study was done at only one centre and was done at one point of time. The study sample confined to patients with T2DM but without comparable groups. A larger sample size would have improved precision in some subgroups and a less homogenous population regarding skin type and life-style habits with a broader range of exposure might have provided different estimates. In addition, we have lack of information regarding the dietary intake of vitamin D of the participants. Interpreting and taking action on serum 25(OH)D values analyzed during different seasons of the year may pose a problem to the clinician. In conclusion, this study indicates that seasonal changes lead to significant serum vitamin D variations with the lowest values in the summer and in the fall and the highest values the winter and in the spring. Although, seasonal change in itself cause significant reduction of serum vitamin D in geographic region of this study, and may lead to serum vitamin D reduction in subjects who are at risk of vitamin D deficiency.

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