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# Vitamin D deficiency in older adults and its associated factors: a cross-sectional analysis of the Mexican Health and Aging Study

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# Abstract

**Summary**—Vitamin D deficiency was common in older adults from a country with adequate sun exposure. The variables associated with this deficiency provide insight into the next steps needed to characterize older adults with this deficiency and to treat it accordingly.

**Purpose**—The aim of this study was to describe the prevalence of and factors associated with vitamin D deficiency among Mexican older adults.

**Methods**—This was a secondary analysis of the last wave of the Mexican Health and Aging Study. Vitamin D levels along with other biomarkers were obtained from a sub-sample of Mexican adults older than 60 years. Prevalence was described by sex and age group, and a multivariate analysis was performed to test the factors associated with this condition.

**Results**—Data from 1088 adults over the age of 60 years were analyzed. The mean serum vitamin D level was  $23.1 \pm 8.1$  ng/mL and was significantly higher among men than women (25.6  $\pm$  0.6 and  $22.8 \pm 0.5$  ng/mL, respectively; p < 0.001). In total, 37.3% (n = 406) presented with vitamin D deficiency, 65% of whom were women. Low 25-(OH)-vitamin D levels were associated with female sex (OR 1.74, 95% CI 1.59–2.42), current smoking (OR 2.21, 95% CI 1.47–3.39), education (OR 1.1, 95% CI 1.06–1.13), physical activity (OR 1.74, 95% CI 1.31–2.23), and high levels of glycated hemoglobin (OR 1.16, 95% CI 1.07–1.25).

Compliance with ethical standards Conflict of interest None

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**Conclusions**—Vitamin D deficiency was highly prevalent in Mexican older adults and was associated with a number of factors, indicating the multifactorial causality of this deficiency.

#### Keywords

Vitamin D; Geriatric nutrition; Epidemiology of aging; Micronutrients

# Introduction

Vitamin D deficiency has been a recent focus of clinicians and researchers because of its potential implications for global health in different age groups. In the field of aging research, vitamin D has been studied as a result of its potential role in adverse conditions, such as sarcopenia, osteoporosis, falls, fractures, and chronic disease [1].

The general prevalence of vitamin D deficiency varies widely, ranging from 7.5 to 77% [2]. This variability depends on factors such as sex, body composition, physical activity, and vitamin D intake. Regarding environmental determinants, the differences between countries are largely due to the exposure to ultraviolet B rays (UVB) [3], which is determined by the angle of the sun in the sky at a particular geographical location.

Between 70 and 95% of vitamin D levels depend on the cutaneous synthesis of cholecalciferol, a main metabolite of hepatic 7-dehydrocholesterol [4]. Aging can affect several aspects of this synthesis; for example, there is an age-dependent decrease in 7-hydrocholesterol, and this decrease lowers the concentration of epidermal cholecalciferol, resulting in an increased need for exposure to synthesize an amount similar to that obtained by younger adults [5]. Moreover, skin changes during aging, primarily wrinkling and thinning skin, have been implicated in an altered vitamin D synthesis [6]. Finally, as older adults decline, their level of physical activity decreases, which reduces their sun exposure considerably, further worsening the availability of the main components needed for vitamin D synthesis [7].

Based on these findings, it is plausible to believe that older adults in Mexico have an advantage over those living in countries with lower sun exposure, as Mexico is located at  $19^{\circ} 6' \text{ N}, 99^{\circ} 08' \text{ O}$  and is thus recognized as having an appropriate level of UVB exposure [2].

Despite this assumption, there is little evidence on the prevalence of vitamin D deficiency and its associated factors in older adults living in developing countries [2], Mexico in particular. As Palacios stated, most studies have included randomly selected individuals from the general population, and thus, the elderly population has been under-represented, and sample sizes have not reached sufficient statistical power. Furthermore, differences in data could arise depending on the type of study [8].

Therefore, the aim of this study was to describe the prevalence of vitamin D deficiency in Mexican older adults and to determine the factors associated with this deficiency.

# Methods

#### **Design and settings**

This was a cross-sectional analysis of the third (2012) wave of the Mexican Health and Aging Study (MHAS), which was a prospective panel study conducted in Mexico. The aim and design of the MHAS have been published elsewhere [9, 10]. In brief, the three waves of the MHAS (2001, 2003, and 2012) contained a representative sample of community-dwelling Mexican older adults (50 years old) and their respective spouses (18 years old). To determine the factors that influenced aging in this population, a set of questionnaires (e.g., sociodemographic, health-related, health service access, cognitive performance, and functional status) were administered to participants in face-to-face interviews. In addition, each wave included a sub-sample in which anthropometric measures and blood samples were collected.

The last wave, in 2012, assessed 18,465 participants, including 12,569 follow-up participants from 2001 on and 5896 new participants (including spouses of the selected subject, regardless of age). In this wave, anthropometrics, physical performance tests, and blood samples were additionally obtained in a sub-sample of 2089 older adults. After excluding those <60 years (n = 961), the present study evaluated 1128 participants older than 60 years old.

#### Measurements

Vitamin D—Between October and November 2012, biomarkers including serum 25-(OH)D levels were obtained from blood samples from a peripheral venipuncture obtained by trained personnel according to standardized protocols. Within 30 min of the venipuncture, the samples collected were centrifuged at 2500 rpm for 15 to 20 min to separate the serum, which was stored in two 2-mL tubes. Serum 25-(OH)D levels were measured with a chemiluminescent microparticle immunoassay (CMIA-Architect Abbott Laboratories. Abbott Park, IL, USA). The measurement interval of this CMIA ranged from 8 to 160 ng/mL, and the intra- and inter-assay coefficients of variation were <10%. The cutoff value for defining vitamin D deficiency was <20 ng/mL (<50 nmol/L) for serum 25-(OH)D levels, as suggested for sub-population studies by the American Task Force for vitamin D deficiency screening [11].

#### Independent variables

To further describe vitamin D deficiency and its related factors, variables from different domains were included. The sociodemographic characteristics included age, sex, marital status (married or not), and years of education. The health-related variables included smoking status (currently smoking), low physical activity (negative response to the following question "In the last two years, have you regularly performed exercise or other vigorous physical activity?"), self-report of six common chronic diseases (i.e., diabetes mellitus, hypertension, cardiac disease, lung disease, cancer, and stroke), geriatric conditions (i.e., falls, depression, cognitive decline, and functional dependency), anthropometry (body mass index [BMI] in kg/m<sup>2</sup>, systolic and diastolic pressure in mmHg, and waist-to-hip index), and serum biomarkers (c-reactive protein [CRP] in mg/dL, total cholesterol in

mg/dL, high-density lipoprotein cholesterol [HDL-c] in mg/dL, thyroid stimulating hormone [TSH] in mIU/mL, hemoglobin in g/dL, and glycated hemoglobin percentage).

Regarding geriatric variables, falls were considered present if the older adult answered "yes" to the question "Have you fallen in the last 12 months?". Depressive symptoms were considered present if the older adult scored 5 points on a previously validated five-item questionnaire for depression (a set of seven dichotomous items generated by specific questions related to mood) [12]. Subjects were considered to have cognitive decline if they failed two or more sub-tests of the brief version of the Cross-Cultural Cognitive Examination [13]. If the older adult required help for one of the eight activities of daily living listed, he or she was defined as having functional dependency. Trained study personnel performed standard anthropometric measurements, and the average of two measurements was used for this analysis. Regarding biomarkers, the same procedure for serum 25-(OH)D levels was conducted as previously described, and technical details for each of the biomarkers are available upon request or on the MHAS webpage http:// www.mhasweb.org/DocumentationQuestionnaire.aspx.

#### Statistical analysis

Descriptive statistics were calculated to estimate the prevalence or the relative frequency of the population that had serum 25-(OH) D levels <20 ng/mL. In addition, this prevalence was described by sex and age group (60–65, 66–75, and >75 years old). A plot of the vitamin D deficiency prevalence and age stratified by sex was generated to visualize this condition across age ranges. To assess associations, a bivariate analysis including all of the previously described variables was performed; *p* values were obtained using *t* tests or chi-square tests based on the nature of the variable (continuous or categorical, respectively). Finally, a multiple logistic regression model was fit using vitamin D deficiency as the dependent variable, and all of the independent variables were simultaneously included. All analyses were performed with the statistical package software STATA 14 @ (StataCorp, 4905 Lakeway Drive, College Station, TX 77845, USA).

#### **Ethical issues**

The MHAS was approved by the Institutional Review Boards and Ethics Committees of the University of Texas Medical Branch in the USA, the Instituto Nacional de Estadistica y Geografia (INEGI), and the Instituto Nacional de Salud Publica (INSP) in Mexico. In addition, the current analysis was registered at the Instituto Nacional de Geriatría (DI-PI-003/2016).

#### Results

The study included a total of 1128 adults older than 60 years. The mean age was 69.6 years ( $\pm$ SD 7.67 years), and the mean years of education was 4.5 years ( $\pm$ SD 4.3 years). Of the total sample, 51.2% were women (n = 578), and 58% (n = 654) were married, while more than half had low levels of physical activity (59.3%). The most frequently reported chronic disease was hypertension (49%), and 42.7% (n = 482) of the older adults reported recent

falls (Table 1). The mean serum vitamin D level was 23.1 ng/mL ( $\pm$ SD 8.1 ng/mL), and 36.9% (n = 416) of the subjects had serum levels <20 ng/mL.

As shown in Table 1, vitamin D-deficient subjects were older (70.67 vs. 69.1 years, p < 0.001), were more often current smokers (14.6 vs. 9.5%, p < 0.05), had a higher mean BMI (29.1 vs. 28.1 kg/m<sup>2</sup>, p < 0.001), had a higher prevalence of diabetes (29.5 vs. 19.8%, p < 0.001), more frequently reported low activity levels (70.4 vs. 52.8%, p < 0.001), and were more dependent (23.3 vs. 14.8%, p < 0.001). In addition, some biomarkers also significantly differed between older adults with vitamin D deficiency and those without vitamin D deficiency, including systolic pressure (145.5 vs. 141.5 mmHg, p = 0.002) and CRP level (4.8 vs. 3.86 mg/dL, p = 0.008) (Table 1).

Table 2 shows the results of the stratified analysis by sex and age group. Vitamin D deficiency for the total sample significantly increased with age; however, after stratification by sex, this relationship was only present among women. In addition, vitamin D deficiency was consistently higher among women than men (Fig. 1).

In the multivariate analysis (Table 3), the multiple logistic regression model ( $R^2 = 0.115$ , p < 0.001) showed that vitamin D deficiency was independently associated with age (odds ratio [OR] 1.02, 95% confidence interval [CI] 1.01–1.05; p < 0.001), female sex (OR 1.97, 95% CI 1.59–2.42; p < 0.001), years of education (OR 1.1, 95% CI 1.06–1.13; p < 0.001), current smoking (OR 2.21, 95% CI 1.47–3.39; p < 0.001), low physical activity (OR 1.74, 95% CI 1.31–2.23; p < 0.001), BMI (OR 1.03, 95% CI 1.01–1.06; p < 0.004), systolic pressure (OR 1.02, 95% CI 1.01–1.03; p = 0.045), CRP level (OR 1.02, 95% CI 1.01–1.05; p = 0.04), TSH level (OR 1.03, 95% CI 1.01–1.07; p = 0.013), and glycated hemoglobin level (OR 1.16, 95% CI 1.07–1.25; p < 0.001).

# Discussion

Our main findings show that vitamin D deficiency (250HD < 20 ng/mL) is common in Mexican older adults, with a prevalence of 37%. Additionally, the other factors associated with this deficiency—both unique to our population and previously described in other older adult groups—included sex (female), current smoking, educational level, physical activity, and high levels of glycated hemoglobin.

Flores et al. found a lower prevalence of vitamin D deficiency after analyzing the results of the National Health and Nutrition Survey 2006 (ENSANUT 2006) to estimate vitamin D concentrations in all age groups [14]. According to these authors, 13% of Mexican older adults (>60 years) presented vitamin D deficiency; however, the sample size was small (n = 158) and not representative of older adults.

Our findings are similar to those reported for Australia, a country that is considered to have sufficient sun exposure during the year in almost all of its territories due to its geographical location  $(27^{\circ} \text{ OO' S}, 133^{\circ} \text{ OO' E})$ . According to this previous report, the prevalence of vitamin D deficiency in Australia was 31%; the lowest levels of 25-(OH)D were observed among participants living in the south (>35^{\circ} S) as a result of the lower sun exposure [15].

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Other reports from countries with different localizations show a wide range in the prevalence of vitamin D deficiency in older adults. A recent study by Klenk et al. that evaluated a group of community-dwelling individuals living in Germany reported a higher prevalence of vitamin D deficiency (79%) than that observed in our study [3]. In the USA, another country with insufficient sun exposure for at least 1 month a year as well as with a high percentage of African-Americans, who are reported to have one of the highest rates of vitamin D deficiency worldwide, the prevalence of vitamin D deficiency was 42% [16].

The heterogeneity in prevalence could occur as the result of differences in the cutoff values used to define vitamin D deficiency, with studies that use higher cutoff values tending to report higher prevalence. In our study, the cutoff value proposed by the U.S. Preventive Services Task Force (<20 ng/mL of serum 25-(OH) D) [11, 17] was used, and as comparisons were made with previous studies that used the same value, the differences in the reported prevalence can be attributed to variables other than the cutoff points.

Despite having sufficient sun exposure, the prevalence of vitamin D deficiency in Mexico was high and could be explained by other associated factors. In fact, Portela et al. [18–20] reported that the prevalence of vitamin D deficiency is linked with factors such as sex, age, genetics, diet, adiposity, and comorbidities.

Our results, as well as those from Oliveri et al. [21], Portela [18], and Linnebur [22], demonstrate a strong correlation between female sex and vitamin D deficiency. These results could be explained by the variations in hormone secretion between the sexes, an association that merits exploration in future research. With respect to age, as shown by Choi et al. [23] and Daly et al. [15], lower levels of vitamin D were observed in older people in our sample, as expected [24]. Regarding education, low levels of education were surprisingly related to better vitamin D status; this finding is consistent with Forrest et al. Our hypothesis concerning this finding is that less educated people may have more sun exposure as a result of performing activities related to agriculture and livestock; however, these findings should be further studied in future research.

In our study, vitamin D deficiency occurred among older people with low physical activity, functional dependency, and comorbidities such as diabetes. These associations have been previously reported [25–27] and can be explained by the "reverse causality" phenomenon described by Thacher [1]. This phenomenon suggests that ill people are less likely to participate in more outdoor activities and consequently have less sun exposure than those in better health conditions. Additionally, as vitamin D3 (cholecalciferol) is stored in adipose tissue and a vitamin D reserve is generated during summer, older adults who are not exposed to sunlight during this season are at a higher risk of having vitamin D deficiency.

In this sense, it is conceivable that a higher BMI would be correlated with a better vitamin D status; nonetheless, our results as well as those by Linnebur [22] have demonstrated that serum 25(OH)D levels are lower among elderly individuals with a higher BMI than among those with a normal BMI (19.0 vs. 26.2 ng/mL). This relationship could result from the sedentary lifestyle that characterizes individuals with a higher BMI, as they may practice less outdoor activities and thus have reduced sun exposure. As Bell et al. reported, in people

with obesity, vitamin D is sequestered in fat tissue, reducing its release into the circulation and therefore its bioavailability [28].

Although causal mechanisms were not studied in this work, it is reasonable to suggest that vitamin D deficiency is part of a complex homeostatic imbalance that is characterized by the presence of multimorbidity and other physical impairments and an increased risk of frailty, disability, and dependence; in turn, these factors promote a deficient nutritional status, which, in addition to a low sun exposure, contributes to permanent vitamin deficiency.

This study on the prevalence of vitamin D deficiency and its associated factors provides an overview of the health conditions of older adults in Mexico. The findings thus support the design of strategies and interventions based on a population's specific characteristics and needs to ensure that better results in the clinical setting and in public health can be achieved. Based on our results, interventions that focus on modifiable determinants of vitamin D deficiency, such as those promoting adequate sun exposure as well as a healthy diet and physical activity, need to be addressed to reduce the prevalence of vitamin D deficiency in this population.

Some limitations should be considered to interpret our results in context. This study used a cross-sectional design, which limited the possibility of interpreting the causality of the associations. In addition, some variables that were not available in our dataset could have been associated with vitamin D deficiency (such as place of residence, nutritional factors, and other hormones). In particular, vitamin D levels show well-described seasonal variations, and data on these changes were not available in our study, indicating the need for further studies on this topic [29].

Another possible limitation of our study is the use of self-report questionnaires, which may have introduced recall bias. However, different epidemiological studies that have aimed to characterize large populations have used self-report, as it is a methodology that has demonstrated effectiveness in achieving results and reducing time and costs [30].

An important strength of our study is that it is the first population-based study to report on the prevalence of vitamin D deficiency and its determinants in older Mexican adults and is one of the few studies in Latin America that has focused on this population group [2, 31]; furthermore, the quality of the method used to determine the levels of vitamin D as well as the sample size diminished the variability in our results.

These findings present opportunities for new research in the field, as there is little evidence on the determinants and mechanisms of vitamin D deficiency in the older Mexican population. In this respect, we consider it important for future research, especially studies aiming to explain causality, to measure the percentage of fat mass rather than BMI, parathyroid hormone levels, environmental and lifestyle factors that account for more substantial variations in serum 250HD levels [32], characteristics of solar exposure, levels of pollution, types of clothes, and race and ethnicity to better determine the role of these components in vitamin D reserve.

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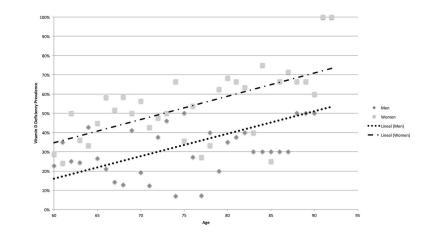
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# Table 1

# General description of the population by vitamin D deficiency status

	Total ( <i>n</i> = 1128)	Deficient, <20 ng/mL ( $n = 416$ )	Not deficient, 20 ng/mL ( $n = 712$ )	p value
Age, mean (SD)	69.6 (7.67)	70.67 (8.01)	69.1 (7.4)	0.001
Women, $n(\%)$	578 (51.2)	271 (65.1)	307 (43.1)	< 0.001
Marital status, n(%)	654 (57.9)	212 (50.9)	442 (62.1)	< 0.001
Years of education, mean (SD)	4.5 (4.26)	5.3 (4.5)	4.1 (4.05)	< 0.001
Current smoker, $n(\%)$	129 (11.4)	61 (14.6)	68 (9.5)	0.009
Low physical activity, n (%)	669 (59.3)	293 (70.4)	376 (52.8)	< 0.001
Diabetes mellitus, n (%)	264 (23.4)	123 (29.5)	141 (19.8)	< 0.001
Hypertension, n (%)	554 (49.1)	214 (51.4)	340 (47.7)	0.232
Cardiac disease, $n(\%)$	43 (3.8)	17 (4.1)	26 (3.6)	0.713
Lung disease, n(%)	68 (6.04)	33 (7.93)	35 (4.93)	0.041
Cancer, $n(\%)$	31 (2.75)	15 (3.6)	16 (2.2)	0.178
Stroke, $n(\%)$	26 (2.3)	9 (2.1)	17 (2.3)	0.809
Falls, $n(\%)$	482 (42.7)	181 (43.5)	301 (42.2)	0.686
Depression, n (%)	427 (37.9)	166 (40.2)	261 (36.7)	0.246
Cognitive decline, $n(\%)$	99 (8.78)	36 (8.6)	63 (8.8)	0.911
Functional dependency, $n(\%)$	203 (18)	97 (23.3)	106 (14.8)	< 0.001
BMI, mean (SD)	28.4 (5.3)	29.1 (5.6)	28.1 (5.1)	0.001
Systolic pressure, mean (SD)	143 (21.1)	145.5 (21.6)	141.5 (20.7)	0.002
Diastolic pressure, mean (SD)	77.8 (11.2)	78.4 (11.2)	77.5 (11.1)	0.21
Waist-to-hip index, mean (SD)	0.96 (0.07)	0.95 (0.08)	0.96 (0.07)	0.037
CRP, mean (SD)	4.2 (5.79)	4.8 (7.3)	3.86 (4.64)	0.008
Total cholesterol, mean (SD)	198 (45.3)	201.5 (48.7)	195.9 (43.1)	0.046
HDL-c, mean (SD)	41.1 (10.5)	41.8 (11.4)	40.6 (9.9)	0.066
TSH, mean (SD)	2.85 (5.2)	3.4 (7.25)	2.52 (3.59)	0.006
Hemoglobin, mean (SD)	14.8 (2.12)	14.8 (1.99)	14.8 (2.1)	0.927
Glycated hemoglobin	0.39 (0.48)	0.46 (0.49)	0.34 (0.47)	0.002

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	Total $(n = 416)$	Total $(n = 416)$ 60–65 years $(n = 146)$ 66–75 years >75 years $(n = 104)$ p value	66–75 years	>75 years ( <i>n</i> = 104)	<i>p</i> value
Men, $n$ (%)	145 (26.3)	68 (27.9)	43 (22.1)	34 (30.1)	0.255
Women, <i>n</i> (%) 271 (46.8)	271 (46.8)	78 (36.7)	123 (52.1)	70 (54.2)	0.001
Total, $n(\%)$	416 (36.9)	146 (32.1)	166 (38.5) 104 (42.9)	104 (42.9)	0.029

#### Table 3

Multiple logistic regression model. Only significant variables with vitamin D deficiency as the dependent variable are presented

	OR (95% CI)	p value
Age	1.02 (1.01–1.05)	< 0.001
Women	1.97 (1.59–2.42)	< 0.001
Years of education	1.1 (1.06–1.13)	< 0.001
Current smoker	2.21 (1.47-3.39)	< 0.001
Low physical activity	1.74 (1.31–2.23)	< 0.001
BMI	1.03 (1.01–1.06)	0.004
Systolic pressure	1.02 (1.01–1.03)	0.045
CRP	1.02 (1.01–1.05)	0.04
TSH	1.03 (1.01–1.07)	0.013
Glycated hemoglobin	1.16 (1.07–1.25)	< 0.001

 $R^2 = 0.1151, p < 0.001$ , for the model