Association between Vitamin D Level in Blood and Periodontitis in Korean Elderly

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This study identified an effective control method for periodontitis by investigating the association between blood levels of vitamin D and periodontitis in Korean elderly based on raw data from the fifth Korea National Health & Nutrition Examination Survey of 2010 (KNHANES). In this study, 1,021 adults over 65 years of age were evaluated based on data from the KNHANES. Periodontal disease was assessed using community periodontal index (CPI), with CPI codes ≥ 3 defined as periodontitis. Blood levels of vitamin D were measured from blood samples and divided into four groups (first quartile: ≤ 13.23 ng/ml, second quartile: $13.24 \sim 16.95$ ng/ml, third quartile: $16.96 \sim 21.58$ ng/ml), and fourth quartile > 21.59 ng/ml). Using multiple logistic regression analyses, the variables were adjusted for general characteristics, oral health-related characteristics, health-related characteristics, and bone mineral density. The statistical analysis was performed using the SAS (ver. 9.2). The results of this study are as follows: the prevalence of periodontitis was 42.6% in Korean elderly. After adjusting for general, oral health-related, and health-related, the risk of periodontitis in the first quartile group was 1.74 times (95% confidence interval [CI], $1.02 \sim 2.98$) higher than that of the fourth quartile group ($p=0.041$). After adjusting for general, oral health-related, and health-related characteristics as well as bone mineral density, the risk of periodontitis in the first quartile group was 1.73 times (95% CI, $1.02 \sim 2.96$) higher than that of the four quartile group ($p=0.042$). There was a significant relationship between blood vitamin D level and periodontitis in Korean elderly. For the prevention of periodontitis, factors related to vitamin D should be considered along with other risk factors.

Key Words: Bone density, Elderly, Korean, Periodontitis, Vitamin D

Introduction

Periodontal disease is a chronic inflammatory condition caused by various factors, including hereditary, local or systematic factors, in addition to pathogenic bacterial factors. In the earliest stage, the inflammation forms only on the gingiva (or gums); however, as it progresses, the depth of the gum pocket increases and the tooth loosens due to periodontal tissue loss and alveolar bone resorption. It is a common oral disease that leads to tooth loss in adults. According to the Korea National Health & Nutrition Examination Survey (KNHANES), the prevalence of periodontitis among adults over 19 years of age was 22.7%, 27.7%, and 29.2% in 2012, 2013, and 2014, res-

pectively whereas the prevalence of periodontitis among elderly people over 65 years of age was 42.4%, 48.5%, and 44.9%, respectively¹⁾.

Vitamin D is a crucial substance that aids in bone growth and retention and maintains the homeostasis of minerals²⁾. Vitamin D deficiency leads to decreased calcium absorption in the small intestine and kidneys by 10~15%, subsequently lowering blood calcium levels. To compensate for this deficiency, the secretion of parathyroid hormone increases. Chronic vitamin D deficiency causes decreased blood calcium levels and increased levels of parathyroid hormone, which activates 1,25(OH)D and osteoclasts. To compensate for deficient blood calcium levels, calcium is released from the calcified bone matrix,

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which prevents bone calcification, decreases bone mineral density, and increases the risk of osteoporotic bone fracture³⁾. Periodontal disease is mainly characterized by the loss of connective tissue and alveolar bone; under the premise that decreased bone mineral density accompanies the loss of alveolar bone, studies have reported the correlations between osteoporosis and alveolar bone resorption⁴⁾, between overall bone mineral density and alveolar bone height^{5,6)}, and between bone mineral density and tooth loss⁷⁻¹¹⁾. A side from regulating calcium homeostasis, vitamin D is also involved in immunity by binding cellular vitamin D receptors, converting to 1,25(OH)₂D₃ as it moves into immunocytes such as macrophages and dendritic cells¹²⁾. Vitamin D also exerts positive effects on periodontal health: higher blood levels of vitamin D reduce gingival bleeding¹³⁾ and increase the antibacterial activity of oral gingival epithelial cells¹⁴⁾.

Bone mass is established through the process of the formation, resorption, and reformation of bone tissues; in old age, bone resorption occurs more frequently than bone formation, leading to reduced bone mass¹⁵⁾. Particularly after menopause, women experience a rapid bone loss due to the effect of hormonal changes¹⁶⁾. In regard to bone health, vitamin D plays a significant role in affecting bone growth or retention and bone mineral density.

Vitamin D deficiency in the elderly increases the risk of various diseases, including osteoporosis, osteopenia, and cardiovascular diseases¹⁷⁾. Due to indoor life styles, loss of appetite, and chronic diseases, it is easy to become deficient in vitamin D¹⁸⁾. Clinical diseases that have been associated with vitamin D deficiency were identified mainly through epidemiological studies, but most did not adjust for disruption variables such as diet, everyday vitamin D levels, body mass index, age, or physical activity^{19,20)}. In addition, many research subjects in previous studies were post-menopausal women^{5,7,9)}. While studies have assessed the correlation between bone mineral density and periodontal diseases and the correlation between vitamin D and bone tissue, few have focused on the association with periodontal diseases by considering vitamin D and bone mineral density.

In this context, the present study was conducted to provide baseline data for the prevention and treatment of

periodontitis by investigating the correlation between blood levels of vitamin D level with bone mineral density and periodontitis among elderly people over 65 years of age and by identifying the risk factors for periodontitis.

Materials and Methods

1. Subjects

The present study used raw data from the first-year survey of the fifth Korea National Health and Nutritional Examination Survey (KNHANES). The first survey was conducted from January to December 2010. A total of 8,058 people participated in more than one form of survey among the health, health examination, and nutrition surveys. To investigate the association between periodontal diseases and bone mineral density, this study selected 1,021 elderly people over 65 years of age whose periodontal tissue, bone mineral density, and blood vitamin D level were measured in response to dental exam and health survey questions. Prior to this study, the researcher obtained approval from the Institutional Review Boards at Kosin University Gospel Hospital (KUGH-2015-10-005-002).

2. Methods

The independent variables of this study included socio-demographic characteristics (five questions), oral health behavior characteristics (four questions), general health behavior characteristics (five questions), blood vitamin D level, and bone mineral density. The dependent variable was the periodontitis status as determined during the dental exam.

1) Blood vitamin D levels

In addition to the health examination questions, blood levels of vitamin D level were measured in regular blood tests and divided into four quartile groups as follows: first quartile (Q1), vitamin D level ≤ 13.23 ng/ml; second quartile (Q2), $13.24 \sim 16.95$ ng/ml; third quartile (Q3), $16.96 \sim 21.58$ ng/ml; and fourth quartile (Q4), > 21.59 ng/ml (Q4). Because there is no consensus optimal blood vitamin D level, this study divided the vitamin D levels into quartile groups in order to examine these levels in

elderly people in the KNHANES, to identify problems with vitamin D levels, and to determine the optimal level for the prevention of periodontitis based on previous study findings.

2) Bone mineral density

The subjects were divided based on overall femoral T-scores from bone mineral density tests collected as part of the health exam items. The t-score was the number derived from dividing the difference between participant bone mineral density score and the highest bone mineral density score of a young adult in the same sex group by standard deviation. $T \geq -1.0$ was categorized as normal range; $-1.0 > T > -2.5$ and $T \leq -2.5$ were considered osteopenia and osteoporosis, respectively²¹⁾.

3) Periodontitis

The condition of the periodontal tissue was assessed using the community periodontal index (CPI). After dividing the oral cavity into sextants (1st and 2nd maxillary molars on the left, maxillary central incisor, 1st and 2nd maxillary molars on the right, 1st and 2nd mandibular molars on the left, mandibular central incisor, and 1st and 2nd mandibular molars on the right), the researcher measured around the 10 teeth for standard testing in terms of the bleeding status, plaque attachment, and depth of the periodontal pocket using the periodontal probe designed by the World Health Organization. The highest score of the sextants was recorded. Normal periodontal tissue was healthy tissue (CPI 0); bleeding gum on probing was considered bleeding periodontal tissue (CPI 1); periodontal tissue with plaque attachment was considered periodontal tissue with plaque retention (CPI 2); periodontal tissue with a 4~5 mm gum pocket depth was considered shallow pocket periodontal tissue (CPI 3); and periodontal tissue with over 6 mm periodontal pocket depth was considered deep pocket periodontal tissue (CPI 4). Based on the clinical definition of periodontist for CPI values above 3²²⁾, this study also classified subjects with a CPI value 3 or above in one of the sextants as having periodontitis.

3. Data analysis

With regard to the data from the KNHANES, a complex sampling analysis method that utilizes cluster sampling variables and estimated variance with individual weight was employed to maintain the rolling survey sampling method and perform a more accurate data analysis.

Frequency and cross over analyses were performed on variables such as sociodemographic characteristics, health behavior, oral health behavior, periodontitis status, blood vitamin D level, and bone mineral density. To examine the risk of periodontitis, multiple logistic regression analysis was performed on independent variables by adjusting for the sociodemographic characteristics, health behavior characteristics, oral health behavior characteristics, and bone mineral density. The data were analyzed using SAS (ver. 9.2; SAS Institute, Cary, NC, USA) and tested for significance at $p < 0.05$.

Results

1. The prevalence of periodontitis based on research subjects sociodemographic characteristics

Examination of the prevalence of periodontitis based on the research subjects' general characteristics showed the following: by sex, periodontitis was more prevalent among men (46.5%) than women (41.5%); by education level, periodontitis was most prevalent among subjects with a middle school education (51.5%); by income level, periodontitis was most prevalent among the low middle class (44.4%); by area of residence, periodontitis was more prevalent in rural areas (50.0%) than in urban areas (38.7%), which showed a statistically significant difference ($p=0.024$; Table 1).

2. The prevalence of periodontitis based on oral health-related characteristics

Examination of the prevalence of periodontitis based on the research subjects' oral health-related characteristics showed the following: by average number of daily brushings, periodontitis was most prevalent among subjects who brushed less than once per day (49.5% twice 40.5%, three times or more 37.9%; $p=0.040$) by oral hygiene product use, periodontitis was highly prevalent

Table 1. Comparison of Periodontitis Prevalence by General Characteristics (n=1,021)

Variable	Normal	Periodontitis	p-value
Gender			0.490
Male	251 (53.5)	218 (46.5)	
Female	324 (58.5)	228 (41.5)	
Education			0.064
Elementary school	368 (55.8)	292 (44.2)	
Middle school	74 (48.5)	62 (51.5)	
High school	76 (61.3)	64 (38.7)	
University or over	51 (72.7)	23 (27.3)	
Income			0.965
Low	136 (57.2)	115 (42.8)	
Middle-low	123 (55.6)	102 (44.4)	
Middle-high	150 (57.9)	111 (42.1)	
High	157 (58.3)	110 (41.7)	
Living area			0.024
Urban	427 (61.3)	298 (38.7)	
Rural	148 (50.0)	148 (50.0)	

Values are presented as n (%).

Table 2. Comparison of Periodontitis Prevalence by Oral Health Related Characteristics (n=1,021)

Variable	Normal	Periodontitis	p-value
The frequency of tooth-brushing			0.040
Less or once	147 (50.5)	144 (49.5)	
Twice	263 (59.5)	198 (40.5)	
Three time or more	165 (62.1)	104 (37.9)	
Use of oral hygiene devices			0.042
Yes	89 (67.1)	49 (32.9)	
No	485 (56.1)	397 (43.9)	
Oral health check-up			0.788
Yes	91 (58.7)	64 (41.3)	
No	484 (57.2)	382 (42.8)	
Missing tooth			0.149
Yes	303 (54.9)	264 (45.1)	
No	272 (60.3)	182 (39.7)	

Values are presented as n (%).

among those who did not use oral hygiene products (43.9%, $p=0.042$). By dental checkup, the prevalence of periodontitis was high among those who had not had any dental checkups within the past year (42.8%); by tooth loss, the prevalence of periodontitis was high among subjects who reported to have lost teeth (45.1%); however, the difference was not statistically significant (Table 2).

Table 3. Comparison of Periodontitis Prevalence by Health Related Characteristics (n=1,021)

Variable	Normal	Periodontitis	p-value
Drinking			0.383
Yes	396 (55.2)	321 (44.8)	
No	172 (60.0)	120 (40.0)	
Smoking experience			0.841
Current smoking	71 (55.8)	64 (44.2)	
Past smoking	161 (59.1)	137 (40.9)	
No	336 (57.0)	240 (43.0)	
BMI			0.484
Underweight	17 (64.4)	9 (35.6)	
Normal	227 (57.6)	167 (42.4)	
Overweight	147 (60.6)	109 (39.4)	
Obesity	184 (53.6)	159 (46.4)	
Regular physical activity			0.684
Yes	108 (55.1)	96 (44.9)	
No	400 (56.8)	312 (43.2)	
Walking practice			0.489
Yes	386 (57.3)	303 (42.7)	
No	121 (60.6)	90 (39.4)	

Values are presented as n (%).

3. The prevalence of periodontitis based on health-related characteristics

Examination of the prevalence of periodontitis based on the research subjects' health-related characteristics showed the following: by alcohol consumption status, the prevalence of periodontitis was high among subjects who drank alcohol (44.8%); by smoking status, the prevalence was high in the order of current smokers (44.2%), non-smokers (43.0%), and previous smokers (40.9%); by body mass index, the prevalence was high in the order of obesity (46.4%), normal weight (42.4%), overweight (39.4%), and underweight (35.6%). By level of activity, the prevalence was high among subjects who reported engaging in a moderate level of exercise (44.9%), followed by those who walked (42.7%) (Table 3).

4. The prevalence of periodontitis based on blood levels of vitamin D and bone mineral density

Examination of the prevalence of periodontitis based on the research subjects' blood vitamin D level showed the following: the prevalence of periodontitis was highest among subjects with vitamin D levels below 13.23 ng/ml (49.4%), followed by levels between 16.96~21.58 ng/ml

(44.9%), above 21.59 ng/ml (39.2%), and between 13.24 ~ 16.95 ng/ml (38.8%). There was no statistically significant difference. The prevalence of periodontitis based on bone mineral density was highest among subjects with a normal range (44.5%), followed by those with osteoporosis (41.7%) and osteopenia (40.1%). There was no statistically significant difference (Table 4).

5. Blood levels of vitamin D level and their correlation to periodontitis

To examine the correlation between blood levels of vitamin D and periodontitis, logistic regression analysis was performed by adjusting for disruption variables separately. Model 1 was adjusted for general characteristics; Model 2 for oral health-related characteristics; Model 3 for health-related characteristics; Model 4 for general characteristics and health-related characteristics; Model 5 for health-related and oral health-related characteristics; Model 6 for general, health-related, and oral health-related characteristics; and Model 7 for general, health-related, and oral health-related characteristics, as well as bone mineral density. When the disruption variables were not adjusted, the risk for periodontitis decreased with reduced blood vitamin D levels; however, the difference was not statistically significant. In Model 3, the risk for periodontitis in the Q1 group increased 1.68 times (1.00 ~ 2.83) compared to that in the Q4 group. The difference was statistically significant ($p=0.047$). In Model

Table 4. Comparison of Periodontitis Prevalence by Vitamin D Level and Bone Mineral Density

Variable	Normal	Periodontitis	p-value
Vitamin D level ^a			0.208
Q1	113 (50.6)	109 (49.4)	
Q2	124 (61.2)	96 (38.8)	
Q3	132 (55.1)	102 (44.9)	
Q4	206 (60.8)	139 (39.2)	
Bone mineral density ^b			0.552
Normal	318 (55.5)	258 (44.5)	
Osteopenia	226 (59.9)	165 (40.1)	
Osteoporosis	31 (58.3)	23 (41.7)	

Values are presented as n (%).

^aQ4: > 21.59 ng/ml, Q3: 16.96 ~ 21.58 ng/ml, Q2: 13.24 ~ 16.95 ng/ml, Q1: ≤ 13.23 ng/ml. ^bNormal: T-score ≥ -1.0, osteopenia: -2.5 < T-score < -1.0, osteoporosis: T-score ≤ -2.5.

Table 5. The Association between Vitamin D Level and Periodontitis

Classification	Unadjusted	Model 1	Model 2	Model 3	Model 4	Model 5	Model 6	Model 7
Vitamin D level								
Q4	1	1	1	1	1	1	1	1
Q3	1.26 (0.83 ~ 1.92)	1.24 (0.81 ~ 1.89)	1.22 (0.79 ~ 1.87)	1.19 (0.76 ~ 1.89)	1.15 (0.73 ~ 1.83)	1.17 (0.74 ~ 1.86)	1.11 (0.70 ~ 1.77)	1.11 (0.70 ~ 1.75)
p-value	0.272	0.324	0.364	0.442	0.538	0.506	0.644	0.656
Q2	0.98 (0.60 ~ 1.61)	0.97 (0.59 ~ 1.60)	1.00 (0.61 ~ 1.64)	1.07 (0.65 ~ 1.77)	1.05 (0.64 ~ 1.75)	1.09 (0.66 ~ 1.80)	1.06 (0.64 ~ 1.75)	1.05 (0.64 ~ 1.73)
p-value	0.948	0.909	0.993	0.796	0.836	0.731	0.811	0.848
Q1	1.51 (0.91 ~ 2.52)	1.56 (0.93 ~ 2.61)	1.53 (0.91 ~ 2.57)	1.68 (1.00 ~ 2.83)	1.74 (1.03 ~ 2.95)	1.70 (1.01 ~ 2.89)	1.74 (1.02 ~ 2.98)	1.73 (1.02 ~ 2.96)
p-value	0.107	0.088	0.107	0.047	0.037	0.046	0.041	0.042

Values are presented as odds ratio (95% confidence interval). Model 1: adjusted by general characteristics, Model 2: adjusted by oral health behaviors, Model 3: adjusted by health behaviors, Model 4: adjusted by general characteristics, health behaviors, oral health behaviors, Model 5: adjusted by general characteristics, health behaviors, oral health behaviors, bone mineral density.

4, the risk for periodontitis in the Q1 group increased 1.74 times (1.03~2.95) compared to that in the Q4 group, a statistically significant difference ($p=0.037$). In Model 5, the risk for periodontitis in the Q1 group increased 1.70 times (1.01~2.89) compared to that in the Q4 group, a statistically significant difference ($p=0.046$). In Model 6, the risk for periodontitis in the Q1 group increased 1.74 times (1.02~2.98) compared to that in the Q4 group, a statistically significant difference ($p=0.041$). Finally, in Model 7, the risk for periodontitis in the Q1 group increased 1.73 times (1.02~2.96) compared to that in the Q4 group, a statistically significant difference ($p=0.042$, Table 5).

Discussion

Periodontal disease, which is caused by bacterial infection and lowered immunity of the host, progresses to alveolar bone resorption and periodontal tissue loss²³. The prevalence of periodontal disease increases with age.

According to Kim and Lee²⁴, 57.9% of the elderly population perceives their dental health to be poor. According to Jang and Nam²⁵, the number of sextants with periodontal pockets increases with age.

Nutritional elements are related not only to oral health such as dental caries or periodontal diseases but also to systematic diseases such as diabetes, cardiovascular disease, and osteoporosis²⁶. In particular, vitamin D helps with bone growth and retention, maintains the homeostasis of calcium in the bone, and controls immune system with a natural inoculation effect⁹. The present study presumed that a low level of vitamin D in the blood increases the prevalence of periodontitis and investigated the correlation of periodontitis to blood levels of vitamin D level in elderly people over 65 years of age using data on bone mineral density, blood vitamin D level, and periodontitis from the 2010 KNHANES. In this study, the vitamin D levels were divided into quartiles: Q1 had vitamin D levels below 13.23 ng/ml; Q2 had levels between 13.24~16.95 ng/ml; Q3 had levels between 16.96~21.58 ng/ml; and Q4 had levels above 21.59 ng/ml. Logistic regression analysis was performed by adjusting for disruption variables in order to examine the blood vitamin D level in

elderly people assessed by the KNHANES and to examine the size and significance of correlation of periodontitis with blood levels of vitamin D based on the vitamin D level problem and previous study findings.

Examination of blood levels of vitamin D revealed the following: 21.1% of subjects had vitamin D levels below 13.23 ng/ml, 22.1% had levels between 13.24~16.95 ng/ml, 23.2% had levels between 16.96~21.58 ng/ml, and 33.6% had levels above 21.59 ng/ml. According to the data from the United States (US) National Health and Nutrition Examination Survey (NHANES), 46% of American adults had vitamin D deficiency, with levels below 20 ng/ml²⁷. As for South Korea, the 2010 KNHANES reported that 65.9% of adults over 20 years were vitamin D deficient. In addition, 7.4% of subjects had vitamin D levels below 10 ng/ml, which is a seriously deficient state. Compared to the US, the vitamin D levels among the Korean people are a markedly low. As for the elderly, 44.9% of men and 76.6% of women had Vitamin D levels below 20 ng/ml. The reason for such a high percentage in elderly women is that the rate of cholesterol transformation to estrogen decreases after menopause, which leads to increased cholesterol levels in the blood²⁸. In return, increased fat tissues lower levels of vitamin D in the blood²⁹; therefore, it is speculated that women have lower blood vitamin D levels than men.

Examination of periodontitis status based on blood levels of vitamin D showed that the group with high vitamin D level (Q4) had a higher prevalence of periodontitis compared to that of the group with a low vitamin D level (Q1), but the difference was not statistically significant. This finding is consistent with the report by Lee and Roh³⁰ that the relationship between blood levels of vitamin D among men in their 50s and periodontal disease was not significant. However, this result conflicts with the finding by Jimenez et al.³¹ who reported that the risk for periodontal disease decreased by 9% with a 10 nmol/L increase in blood vitamin D among men between 40 and 75 years of age. It is also contrary to the finding of Dietrich et al.¹³, who reported that the group of subjects over 50 years with higher blood vitamin D level has less attachment tissue loss. Such conflicting results may be explained by the following. Previous studies employed

different measurement criteria for blood levels of vitamin D level, and Jimenez et al.³¹⁾ determined the periodontitis status based solely on the history of being diagnosed with periodontitis. Considering that the correlation between vitamin D level and attachment tissue loss among women in the study by Dietrich et al.¹³⁾ was in significant, sex difference may explain the conflicting results.

To examine the size and significance of the correlation between periodontitis and blood level of vitamin D, logistics regression analysis was performed by differently adjusting for each disruption variable. When health-related characteristics were adjusted, the Q1 group's risk for periodontitis was 1.68 times (1.00~2.83) higher than that of the Q4 group; when general and health-related characteristics were adjusted, the Q1 group's risk increased by 1.74 times (1.03~2.95). When health-related and oral health-related characteristics were adjusted, the risk increased by 1.70 times (1.01~2.89); when general, oral health-related, and health-related characteristics were adjusted, the risk increased by 1.74 times (1.02~2.98). When all disruption variables were adjusted, the Q1 group's risk for periodontitis increased by 1.73 times (1.02~2.96), which demonstrates that a low level of vitamin D in the blood increases the risk of periodontitis. Meanwhile, there was no significant correlation between periodontitis and blood levels of vitamin D among research subjects under 64 years of age, although the details are not presented in this paper. The study findings matched those of Dietrich et al.³²⁾ that a higher level of vitamin D in the blood was related to a lower level of attachment tissue loss among men and women over 50 years of age, while there was no significant correlation between attachment tissue loss and blood vitamin D level among subjects ages between 20 and 49 years. Considering the fact that 58.5% of the research subjects were elderly women, these findings can be interpreted as the result of the skin's decreased capability for vitamin D synthesis with age as the skin becomes thinner and the number of keratinocytes or fibrocytes reduces while vitamin D levels in the blood decrease due to increased serum cholesterol levels³⁰⁾.

Dietrich et al.¹³⁾ reported that higher blood levels of vitamin D contribute to reduced gingival bleeding. According to the study, the presence of gingival bleeding

indicates a gingival infection; the consumption of optimal levels of vitamin D and calcium can prevent tooth loss among elderly people because the progression of gingivitis can lead to periodontal tissue destruction³³⁾. Findings from previous studies suggest an optimal blood vitamin D level 36~40 ng/ml for the prevention of periodontitis³⁴⁾.

The present study is limited in that other factors influencing blood levels of vitamin D such as exercise, calcium intake, hormone supplementation, and systematic disease were not considered, and that the correlation to periodontitis cannot be interpreted in terms of a causal relationship due to the cross-sectional nature of the study design. To rectify these limitations, epidemiological research needs to be conducted in a follow-up clinical study that can evaluate the factors influencing blood levels of vitamin D and the dose-response relationship to periodontitis and identify the causal relationships. Despite these limitations, the present study is significant in that it utilized highly reliable data from the KNHANES, which represents the entire population. The results of this study provide evidence for the necessity of vitamin D control for periodontitis prevention and treatment by presenting the correlation between blood levels of vitamin D and periodontitis.

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