

Lower Nutrient Intakes and Periodontitis: Findings from the Korea National Health and Nutrition Examination Surveys

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1. Introduction

Periodontitis is the most common chronic inflammatory disease caused by an interaction of a specific bacterial flora with the host¹. After stimulation by bacterial antigens, various reactive oxygen species (ROS) were produced by host's defense mechanism. These ROS originate from molecular oxygen and predominantly produce cellular damage (proteins, lipids, and DNA) if not neutralized by anti-oxidant substances. Their production is an essential component of the host response to a variety of insults, including bacteria² and traumas/burns³. It has recently been discovered that there is a strong relationship among increased ROS, periodontitis pathogenesis, and antioxidant support⁴. In this respect, nutrition may be of great importance since it has been implicated in a number of inflammatory diseases and conditions, including periodontitis⁵. Several reports demonstrated that a poor

diet is associated with a higher risk of periodontitis⁶. Thereby, the role of some micronutrients in the etiology and therapy of periodontitis were discussed⁵. Micronutrients are dietary compounds (vitamins and minerals) that are required in small quantities and essential for metabolism, immune response, tissue regeneration, and growth. In particular, oral soft tissue with high physiologic turnover rates are dependent on sufficient food intake. However, most studies dealt with the association between single nutrient and periodontitis⁷. Only one study examined the relationship between various nutrients and periodontitis⁸.

In this study, the Food Frequency Questionnaire (FFQ) was used to evaluate the intake of vitamins and other nutrients. Using data from a national sample of Korean adults, the aim of this study was therefore to assess the association between various nutrient intake and periodontitis in a large representative sample of the Korean population: the 2007~2010 and 2012 Korea National Health and Nutritional Examination Survey (KNHANES).

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2. Materials and Methods

2.1. Study population

Data were from the 2007~2010 and 2012 KNHANES, cross-sectional and nationwide surveys performed by the Korea Disease Control and Prevention Agency(KDCA). The survey was composed of a health interview survey, a nutrition survey, and a health examination survey. An oral health examination was included as a part of the health examination survey since the first year of the KNHANES IV in 2007). The periodontal information of the year 2011 KNHANES was not opened due to fluctuation of periodontal data. Each participant in the survey signed an informed consent form. Overall response rates were 65.8% in 2007, 74.3% in 2008, 82.8% in 2009, 81.9% in 2010, and 80.0% in 2012. For the present analysis we combined data from separate surveys sampled from 2007 through 2010 and 2012 among adults aged 25~64 years.

The number of total participants in 2007~2010 and 2012 KNHANES was 41,887 (19,059 males and 22,828 females). Among those, periodontal health status was measured for 31,908 participants (13,994 males and 17,914 females). There were 22,321 (9,766 males and 12,555 females) adults aged 25~64 years among the total participants in the 2007~2010 and 2012 KNHANES (n=41,887). The exclusion criteria were as follows: (i) those aged < 25 or > 64, (ii) those who did not participate in the periodontal examination, (iii) those who took in the nutrients more than tolerable upper intake level, and (iv) those with missing values in the health assessment or questionnaires. The final sample size was 16,555 (6,520 males and 10,035 females, 74.2% of all adults aged 25~64 years in the 2007~2010 and 2012 KNHANES) for model 2 in Table 4 and 5 and 15,847 (6,198 males and 9,649 females, 71.0% of all adults aged 25 years or older in the 2007~2010 and 2012

KNHANES) for model 3 in Table 4 and 5.

2.2. Assessment of dietary nutrients

The dietary intake for the last 12 months was assessed, and full-scale photographs were used to assist in estimating portion sizes such as tablespoons, cups, slices, units, bowls, plates and cans. Nutritional intake was analyzed using CAN-Pro 4.0 (Korean Nutrition Society, Seoul, Korea). The FFQ used by the KNHANES included 11 food groups consisting of 63 food items, and the frequency of servings was classified into nine categories: never or seldom, once a month, 2~3 times a month, one to two times a week, three to four times a week, five to six times a week, once a day, twice a day, and three or more times a day. The portion size of the food items was classified as follows: small, medium, or large¹⁰. Nutrients included in this analysis were total dietary energy, vitamin-A, -B₁, -B₂, -C, iron, phosphorus, calcium, and protein. The above mentioned nutrients were divided into three groups with estimated average requirements (EAR) and recommended intake (RI) defined by the Dietary Reference Intakes for Koreans¹¹.

2.3. Periodontal examination

Oral health examination was conducted by dentists in the survey. The World Health Organization (WHO) community periodontal index (CPI) was used to assess periodontitis¹². Periodontitis was defined as a CPI greater than or equal to "code 3", which indicates that more than one site had a 3.5 mm pocket or larger in the index teeth. The index teeth numbers were 11, 16, 17, 26, 27, 31, 36, 37, 46, and 47, respectively.

2.4. Assessment of covariates

Confounders in this study were total energy intake and major socio-demographic factors and included

survey year, age, income, education, occupation, marital status, and residence. Mediators considered in the relationship between nutrients and periodontitis were oral health behaviors (frequency of daily toothbrushing and regular dental check-up), general health behaviors (smoking), somatic health problems (diabetes), and a measure for inflammation (White Blood Cell : WBC count). We assumed these mediators as intermediate variables in the causal pathway between nutrients and periodontitis. Except for diabetes and WBC count, information on covariates was collected during the health interview. Monthly household income was adjusted for the number of household member and categorized into four groups: < 25 %, 25~49 %, 50~75 %, and > 75 % of total equivalised income in the survey. Education level was categorized into four groups: below primary school, middle school, high school, and college or higher. Occupation was divided into white collar, blue collar, and others. Marital status was categorized as married and single. Participants' residence was presented as urban and rural. Information on the frequency of daily toothbrushing was collected based on the following question, "How often did you brush your teeth yesterday?" (times/day). Participants were asked whether they had visited dental clinic for regular check-up during a year prior to the interview (yes or no). Smoking status was divided into two categories: current smokers and no smoker. Fasting plasma glucose was measured and grouped as normal (fasting plasma glucose level <100 mg/dl), prediabetes (fasting plasma glucose level <126 mg/dl), and diabetes (fasting plasma glucose level ≥126 mg/dl or diabetic medication). WBC count was measured with blood specimens.

2.5. Statistical analysis

Individual sample weights and the complex sample

design including stratification and primary sampling units were considered in the analysis. Due to the significant interactions between age and nutrients intake and between gender and nutrients, separated analyses according to age group (25~44 years and 45~64 years) and gender were performed.

As seen in Fig. 1, the role of confounders can be determined by examining the association of confounders with periodontitis (A in Fig. 1). The basic association between nutrients and periodontitis is evaluated (B in Fig. 1). Adjusted odds ratio of periodontitis adjusting for sociodemographic variables represents the unconfounded total impact of shift work on periodontitis in this study. The relationship of mediators with nutrients (C in Fig. 1) and periodontitis (D in Fig. 1) should also be examined to determine the role of mediators. This role of mediators was evaluated with the percentage (%) excess odd explained, which can be calculated as $[(OR_{(\text{adjusted for sociodemographic factors})} - OR_{(\text{adjusted for sociodemographic factors} + \text{mediators})}) / (OR_{(\text{adjusted for sociodemographic factors})} - 1)]$ in this study¹³⁾.

This % excess odd explained represents the degree to which a mediator explains the relationship between shift work and periodontitis.

In this study, the characteristics of study subjects by periodontitis status were presented with frequency distributions for the categorical variables and mean (and standard deviation) for continuous variable (Table 1). Chi-square tests for categorical variables and t tests for continuous variables were used to assess the associations of periodontitis with confounders and mediators (Table 1). The association between nutrients intake and periodontitis was presented with frequency distribution and survey year and age-adjusted ORs of periodontitis according to age groups (Table 2 for male and Table 3 for female). Series of multiple logistic regression analysis were used to estimate the adjusted

odds ratio (AOR) of periodontitis after adjusting for mediator variables by gender (Table 4 for 25~44 years and Table 5 for 45~64 years). The model adjusting for confounders were base model (Model 1 in the Table 4 and 5) in this analysis. IBM SPSS Statistical Software (version 22.0, IBM Corp, Armonk, NY, 2013) was used for all analysis. All reported P values are 2-tailed. P values $\leq .05$ were considered as statistically significant.

3. Results

Analysis results showed that those who had low intake of nutrients had greater odds of having periodontitis than those who consumed recommended intake of nutrients. Table 1 shows that significant differences in the distribution of periodontitis were found with regards to all variables except for regular oral check-up among males and except for current smoking among females (all P values < 0.001).

Table 2 shows the relationship of periodontitis with nutrients among males. Those who took in phosphorus under EAR level had a significant survey year-, age-, and total energy intake-adjusted OR (AOR=1.86). Those who ingested calcium and protein under EAR level showed a significant great OR (AOR=1.25 for calcium and AOR=1.52 for protein) of periodontitis. In 25~44 years old group, these significant odds ratios were marked with vitamin B2 (AOR=1.27), iron (AOR=1.45), phosphorus (AOR=2.68), and protein (AOR=1.97). The associations between nutrients and periodontitis had a pronounced tendency in 25~44 years old group.

Table 3 shows the relationship of periodontitis with nutrients among females. Those who took in nutrients under EAR level had a significant adjusted OR (AOR=1.32 for vitamin A, 1.21 for vitamin B1, 1.24 for vitamin B2, 1.21 for vitamin C, 1.25 for iron, 1.33 for

phosphorus, and 1.27 for protein) except for calcium. In 25~44 years old group, these significant odds ratios were marked with vitamin A (AOR=1.26) and vitamin C (AOR=1.32). In 45~64 years old group, vitamin A (AOR=1.39), vitamin B1 (AOR=1.22), vitamin B2 (AOR=1.30), vitamin C (AOR=1.19), iron (AOR=1.27), and phosphorus (AOR=1.32).

Table 4 shows series of AOR of periodontitis by nutrients after adjusting for mediators among 25~44 years old group. The base model (Model 1) was the model adjusted for confounders (survey year, age, education, occupation, income, marital status, residence, and total energy intake). Analysis results revealed that those who ingested phosphorus and protein under EAR level had significantly higher AOR than those who ingested more than RI level (AOR=2.19 for phosphorus and 1.67 for protein) among males. The magnitude of AOR of periodontitis by nutrients attenuated with adjustment of a marker of inflammation (WBC count) except for phosphorus. The % excess odd explained for WBC count were 8.3% for vitamin A, 23.5% for vitamin B₁, 0.0% for vitamin B₂, 5.0% for vitamin C, 18.5% for iron, -10.1% for phosphorus, 21.7% for calcium, and 6.0% for protein among males and 21.4% for vitamin A, 5.6% for vitamin B₁, 18.2% for vitamin B₂, 9.1% for vitamin C, 0.0% for iron, 36.7% for phosphorus, -25.0% for calcium, and 6.7% for protein among females. The magnitude of AOR of periodontitis by nutrients had a higher tendency with adjustment of somatic problem and health behaviors than a marker of inflammation among both gender.

Table 5 shows series of AOR of periodontitis by nutrients after adjusting for mediators among 45~64 years old group. Analysis results revealed that those who ingested vitamin A under EAR level had significantly higher AOR than those who ingested more than RI level (AOR=1.34) among females. The

% excess odd explained for WBC count were 18.2% for vitamin A, 40.0% for vitamin B₁, 28.6% for vitamin B₂, 22.2% for vitamin C, 80.0% for iron, 20.0% for phosphorus, 0.0% for calcium, and 50.0% for protein among males and -11.8% for vitamin A, 7.7% for vitamin B₁, -11.1% for vitamin B₂, -15.4%

for vitamin C, 0.0% for iron, 4.2% for phosphorus, 0.0% for calcium, and 33.3% for protein among females. The magnitude of AOR of periodontitis by nutrients had a higher tendency with adjustment of somatic problem and health behaviors than a marker of inflammation among both gender.

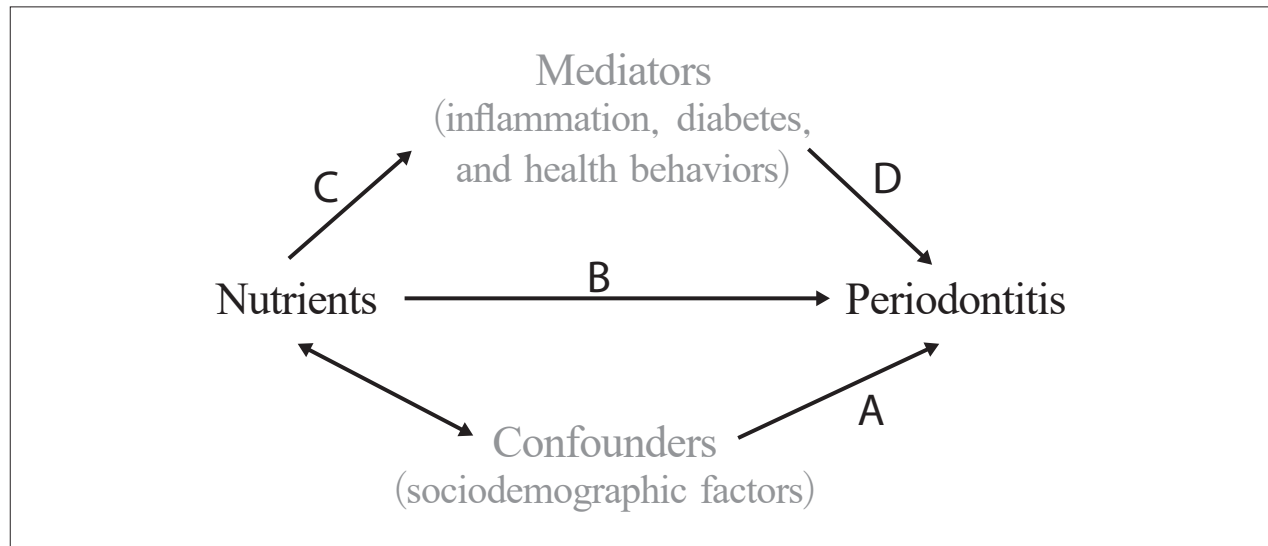


Figure 1. A model for the relationship between nutrients and periodontitis and the roles of confounders and mediators in the relationship.

Table 1. Characteristics of subjects in the 2007~2010 and 2012 KNHANES

		Male				Female			
		Total N	Healthy % (95% CI)*	Periodontitis % (95% CI)*	<i>p</i> -value [†]	Total N	Healthy % (95% CI)*	Periodontitis % (95% CI)*	<i>p</i> -value [†]
Age group	25~44 yr	4349	76,6 (74,8~78,2)	23,4 (21,8~25,2)	<0,001	5930	87,8 (86,5~88,9)	12,2 (11,1~13,5)	<0,001
	45~64 yr	4278	49,6 (47,3~51,9)	50,4 (48,1~52,7)		5682	65,8 (63,9~67,6)	34,2 (32,4~36,1)	
Education	≤ primary school	919	44,6 (40,3~49,0)	55,4 (51,0~59,7)	<0,001	2214	59,1 (56,4~61,8)	40,9 (38,2~43,6)	<0,001
	middle school	947	45,7 (42,0~49,6)	54,3 (50,4~58,0)		1391	67,4 (64,2~70,4)	32,6 (29,6~35,8)	
	high school	3153	64,4 (62,1~66,5)	35,6 (33,5~37,9)		4345	79,9 (78,2~81,5)	20,1 (18,5~21,8)	
	> college or higher	3483	73,8 (71,7~75,8)	26,2 (24,2~28,3)		3542	88,8 (87,5~90,1)	11,2 (9,9~12,5)	
Occupation	White collar	3993	70,5 (68,5~72,4)	29,5 (27,6~31,5)	<0,001	3965	82,3 (80,7~83,8)	17,7 (16,2~19,3)	<0,001
	Blue collar	3296	56,8 (54,4~59,1)	43,2 (40,9~45,6)		2101	67,4 (64,6~70,0)	32,6 (30,0~35,4)	
	Others	1135	65,3 (61,8~68,7)	34,7 (31,3~38,2)		5396	78,1 (76,6~79,5)	21,9 (20,5~23,4)	
Income	I	2090	61,3 (58,7~63,8)	38,7 (36,2~41,4)	<0,001	2819	74,7 (72,6~76,7)	25,3 (23,3~27,4)	<0,001
	II	2112	62,3 (59,6~64,8)	37,7 (35,2~40,4)		2861	75,8 (73,7~77,8)	24,2 (22,2~26,3)	
	III	2125	66,9 (64,4~69,4)	33,1 (30,6~35,6)		2881	79,6 (77,7~81,4)	20,4 (18,6~22,3)	
	IV	2152	69,5 (66,8~71,1)	30,5 (27,9~33,2)		2861	81,8 (79,7~83,8)	18,2 (16,2~20,3)	
Marital status	Married	6912	60,8 (59,0~62,6)	39,2 (37,4~41,0)	<0,001	9373	77,0 (75,7~78,3)	23,0 (21,7~24,3)	0,001
	Single	1698	77,3 (74,8~79,7)	22,7 (20,3~25,2)		2230	81,0 (78,8~83,0)	19,0 (17,0~21,2)	

		Male				Female			
		Total N	Healthy	Periodontitis	<i>p</i> -value [†]	Total N	Healthy	Periodontitis	<i>p</i> -value [†]
			% (95% CI)*	% (95% CI)*			% (95% CI)*	% (95% CI)*	
Residence	Urban	6809	66.6 (64.8~68.3)	33.4 (31.7~35.2)	<0.001	9331	79.0 (77.7~80.3)	21.0 (19.7~22.3)	<0.001
	Rural	1818	57.0 (53.4~60.6)	43.0 (39.4~46.6)		2281	72.5 (69.3~75.4)	27.5 (24.6~30.7)	
Diabetes	Normal	5321	68.1 (66.2~69.9)	31.9 (30.1~33.8)	<0.001	8493	79.8 (78.5~81.0)	20.2 (19.0~21.5)	<0.001
	Prediabetes	1906	60.1 (57.1~62.9)	39.9 (37.1~42.9)		1580	70.5 (67.5~73.3)	29.5 (26.7~32.5)	
	Diabetes	747	41.7 (37.4~46.1)	58.3 (53.9~62.6)		630	61.9 (57.3~66.2)	38.1 (33.8~42.7)	
Toothbrushing (times/day)	< 2	1275	57.4 (54.1~60.6)	42.6 (39.4~45.9)	<0.001	798	71.0 (67.0~74.8)	29.0 (25.2~33.0)	<0.001
	≥ 2	7277	66.0 (64.3~67.6)	34.0 (32.4~35.7)		10746	78.3 (77.1~79.5)	21.7 (20.5~22.9)	
Regular oral check-up	No	6025	64.6 (62.9~66.3)	35.4 (33.7~37.1)	0.614	8294	76.9 (75.5~78.3)	23.1 (21.7~24.5)	0.002
	Yes	2531	65.2 (62.7~67.7)	34.8 (32.3~37.3)		3254	80.1 (78.3~81.8)	19.9 (18.2~21.7)	
Current smoking	No	2919	73.0 (70.8~75.0)	27.0 (25.0~29.2)	<0.001	10674	77.9 (76.6~79.1)	22.1 (20.9~23.4)	0.634
	Yes	5611	60.5 (58.7~62.4)	39.5 (37.6~41.3)		862	77.1 (73.6~80.2)	22.9 (19.8~26.4)	
Total energy intake (kg)			2.42 (2.38~2.45) [‡]	2.33 (2.29~2.37) [‡]	<0.001 [§]		1.72 (1.70~1.74) [‡]	1.64 (1.61~1.67) [‡]	<0.001 [§]
WBC count (10 ⁵)			6.48 (6.42~6.53) [‡]	6.90 (6.82~6.98) [‡]	<0.001 [§]		5.67 (5.63~5.72) [‡]	5.86 (5.78~5.94) [‡]	<0.001 [§]

* Weighted percent and 95% confidence interval

† Obtained from chi-square test

‡ Weighted mean and 95% confidence interval

§ Obtained from independent t-test

Table 2. Association between nutrients and periodontitis among males

		Total N	Healthy	Periodontitis	<i>P</i> -value	Total	25~44yr	45~64yr
			% (95% CI)	% (95% CI)		OR (95% CI)	OR (95% CI)	OR (95% CI)
Vit A	< EAR	2262	62.5 (59.9~65.1)	37.5 (34.9~40.1)	0.115	1.13 (0.97~1.32)	1.18 (0.93~1.49)	1.14 (0.94~1.38)
	< RI	1246	65.2 (62.0~68.2)	34.8 (31.8~38.0)		1.10 (0.93~1.30)	1.22 (0.96~1.56)	1.02 (0.81~1.27)
	≥ RI	3169	65.6 (63.4~67.8)	34.4 (32.2~36.6)		1.00 (reference)	1.00 (reference)	1.00 (reference)
Vit B ₁	< EAR	1692	60.6 (57.7~63.4)	39.4 (36.6~42.3)	0.002	1.14 (0.96~1.35)	1.27 (0.97~1.66)	1.11 (0.89~1.38)
	< RI	949	63.2 (59.4~66.8)	36.8 (33.2~40.6)		1.07 (0.88~1.30)	1.08 (0.80~1.46)	1.07 (0.85~1.36)
	≥ RI	4242	66.1 (64.1~68.1)	33.9 (31.9~35.9)		1.00 (reference)	1.00 (reference)	1.00 (reference)
Vit B ₂	< EAR	3504	61.0 (58.7~63.1)	39.0 (36.9~41.3)	<0.001	1.17 (0.99~1.37)	1.27 (1.00~1.59)	1.11 (0.91~1.35)
	< RI	780	69.0 (65.3~72.5)	31.0 (27.5~34.7)		0.90 (0.74~1.09)	0.86 (0.62~1.17)	0.93 (0.72~1.21)
	≥ RI	2599	67.5 (65.1~69.7)	32.5 (30.3~34.9)		1.00 (reference)	1.00 (reference)	1.00 (reference)
Vit C	< EAR	2511	64.1 (61.6~66.5)	35.9 (33.5~38.4)	0.204	1.11 (0.96~1.28)	1.23 (0.99~1.52)	1.09 (0.91~1.30)
	< RI	1141	62.0 (58.5~65.5)	38.0 (34.5~41.5)		1.18 (0.99~1.40)	1.16 (0.88~1.52)	1.18 (0.95~1.48)
	≥ RI	3231	65.5 (63.2~67.7)	34.5 (32.3~36.8)		1.00 (reference)	1.00 (reference)	1.00 (reference)
Fe	< EAR	598	62.3 (57.4~66.9)	37.7 (33.1~42.6)	0.180	1.21 (0.97~1.52)	1.45 (1.04~2.03)	1.10 (0.80~1.51)
	< RI	766	67.6 (63.4~71.5)	32.4 (28.5~36.6)		1.00 (0.81~1.23)	0.97 (0.71~1.31)	1.02 (0.76~1.37)
	≥ RI	5330	64.0 (62.1~65.9)	36.0 (34.1~37.9)		1.00 (reference)	1.00 (reference)	1.00 (reference)
Phos	< EAR	186	56.5 (48.0~64.7)	43.5 (35.3~52.0)	0.138	1.86 (1.24~2.80)	2.68 (1.55~4.63)	1.34 (0.79~2.27)
	< RI	224	62.9 (54.7~70.5)	37.1 (29.5~45.3)		1.08 (0.73~1.59)	0.84 (0.40~1.75)	1.27 (0.79~2.05)
	≥ RI	6430	64.7 (63.0~66.5)	35.3 (33.5~37.0)		1.00 (reference)	1.00 (reference)	1.00 (reference)
Ca	< EAR	4195	63.6 (61.6~65.7)	36.4 (34.3~38.4)	0.267	1.25 (1.06~1.47)	1.24 (0.97~1.59)	1.22 (0.99~1.49)
	< RI	1022	65.2 (61.7~68.5)	34.8 (31.5~38.3)		1.07 (0.88~1.31)	1.04 (0.75~1.43)	1.07 (0.82~1.39)
	≥ RI	1648	66.1 (63.2~69.0)	33.9 (31.0~36.8)		1.00 (reference)	1.00 (reference)	1.00 (reference)
Protein	< EAR	640	59.6 (54.9~64.1)	40.4 (35.9~45.1)	0.019	1.52 (1.20~1.93)	1.97 (1.41~2.74)	1.18 (0.86~1.62)
	< RI	662	61.3 (56.8~65.7)	38.7 (34.3~43.2)		1.14 (0.90~1.43)	1.20 (0.85~1.69)	1.07 (0.79~1.44)
	≥ RI	5581	65.3 (63.4~67.1)	34.7 (32.9~36.6)		1.00 (reference)	1.00 (reference)	1.00 (reference)

Adjusted for survey year, age and total energy

Bold denotes statistical significance at $P < 0.05$.

EAR= estimated average requirements, RI= recommended intake

		Male			Female		
		Model 1	Model 2	Model 3	Model 1	Model 2	Model 3
		OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)
Phosphorus	< EAR	2.19 (1.23~3.87)	2.31 (1.28~4.16)	2.68 (1.43~5.03)	1.30 (0.92~1.84)	1.19 (0.84~1.70)	1.18 (0.83~1.69)
	< RI	0.66 (0.32~1.36)	0.65 (0.32~1.35)	0.64 (0.29~1.40)	0.82 (0.58~1.16)	0.82 (0.58~1.17)	0.86 (0.61~1.24)
	≥ RI	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Ca	< EAR	1.23 (0.95~1.59)	1.20 (0.93~1.56)	1.20 (0.92~1.57)	1.08 (0.83~1.40)	1.10 (0.85~1.44)	1.10 (0.83~1.45)
	< RI	1.03 (0.74~1.43)	1.01 (0.72~1.41)	1.04 (0.75~1.46)	1.07 (0.77~1.48)	1.09 (0.78~1.54)	1.09 (0.76~1.54)
	≥ RI	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Protein	< EAR	1.67 (1.20~2.34)	1.63 (1.15~2.29)	1.68 (1.17~2.42)	1.30 (0.95~1.79)	1.28 (0.93~1.78)	1.29 (0.93~1.79)
	< RI	1.14 (0.80~1.63)	1.15 (0.80~1.65)	1.10 (0.76~1.59)	1.20 (0.90~1.60)	1.21 (0.90~1.63)	1.21 (0.90~1.63)
	≥ RI	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)

Model 1 was adjusted for survey year, age, education, occupation, income, marital status, residence, and total energy intake.

Model 2 was adjusted for survey year, age, education, occupation, income, marital status, residence, total energy intake, and WBC count.

Model 3 was adjusted for survey year, age, education, occupation, income, marital status, residence, diabetes, toothbrushing habit, regular oral check-up, smoking, and total energy intake.

Bold denotes statistical significance at $P < 0.05$.

EAR= estimated average requirements, RI= recommended intake

Table 5. Odds ratios of periodontitis by nutrient level among 45~64 years group

		Male			Female		
		Model 1	Model 2	Model 3	Model 1	Model 2	Model 3
		OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)
Vitamin A	< EAR	1.11 (0.92~1.35)	1.09 (0.90~1.32)	1.04 (0.85~1.27)	1.34 (1.13~1.59)	1.38 (1.16~1.64)	1.37 (1.15~1.63)
	< RI	1.05 (0.84~1.33)	1.05 (0.83~1.32)	1.03 (0.81~1.31)	1.16 (0.94~1.43)	1.18 (0.95~1.46)	1.16 (0.93~1.45)
	≥ RI	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Vitamin B ₁	< EAR	1.05 (0.84~1.32)	1.03 (0.82~1.30)	1.01 (0.80~1.27)	1.13 (0.93~1.37)	1.12 (0.92~1.37)	1.15 (0.94~1.40)
	< RI	1.01 (0.80~1.29)	1.04 (0.82~1.31)	1.00 (0.79~1.28)	1.03 (0.83~1.29)	1.02 (0.81~1.28)	1.04 (0.82~1.30)
	≥ RI	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Vitamin B ₂	< EAR	1.07 (0.87~1.32)	1.05 (0.85~1.30)	1.00 (0.81~1.24)	1.18 (0.97~1.43)	1.20 (0.99~1.46)	1.23 (1.01~1.49)
	< RI	0.93 (0.71~1.20)	0.93 (0.72~1.21)	0.91 (0.70~1.20)	0.91 (0.71~1.18)	0.95 (0.73~1.23)	0.95 (0.73~1.24)
	≥ RI	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Vitamin C	< EAR	1.09 (0.91~1.31)	1.07 (0.89~1.29)	0.99 (0.82~1.20)	1.13 (0.95~1.34)	1.15 (0.97~1.37)	1.15 (0.96~1.37)
	< RI	1.18 (0.94~1.49)	1.16 (0.92~1.47)	1.10 (0.86~1.39)	1.18 (0.95~1.46)	1.21 (0.97~1.50)	1.22 (0.98~1.51)
	≥ RI	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Iron	< EAR	1.05 (0.76~1.46)	1.01 (0.72~1.41)	0.92 (0.65~1.31)	1.19 (0.96~1.46)	1.19 (0.96~1.47)	1.17 (0.94~1.45)
	< RI	1.02 (0.75~1.38)	0.99 (0.72~1.34)	0.99 (0.72~1.36)	0.89 (0.71~1.11)	0.92 (0.73~1.16)	0.94 (0.74~1.18)
	≥ RI	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Phosphorus	< EAR	1.30 (0.74~2.29)	1.24 (0.69~2.22)	1.03 (0.58~1.85)	1.24 (0.96~1.59)	1.23 (0.94~1.59)	1.22 (0.94~1.58)
	< RI	1.25 (0.76~2.05)	1.21 (0.71~2.05)	1.08 (0.62~1.86)	1.01 (0.77~1.32)	0.98 (0.74~1.29)	0.95 (0.72~1.26)
	≥ RI	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Ca	< EAR	1.19 (0.97~1.47)	1.19 (0.96~1.48)	1.13 (0.91~1.40)	0.99 (0.81~1.22)	0.99 (0.81~1.22)	1.01 (0.82~1.24)
	< RI	1.05 (0.80~1.38)	1.06 (0.81~1.39)	1.02 (0.77~1.33)	0.89 (0.68~1.16)	0.89 (0.68~1.16)	0.87 (0.67~1.14)
	≥ RI	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)
Protein	< EAR	1.06 (0.77~1.47)	1.03 (0.74~1.43)	0.93 (0.66~1.30)	1.12 (0.90~1.39)	1.08 (0.86~1.35)	1.07 (0.85~1.34)
	< RI	1.00 (0.74~1.35)	0.96 (0.70~1.31)	0.94 (0.69~1.29)	1.07 (0.86~1.33)	1.07 (0.85~1.34)	1.10 (0.88~1.39)
	≥ RI	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)

Model 1 was adjusted for survey year, age, education, occupation, income, marital status, residence, and total energy intake.

Model 2 was adjusted for survey year, age, education, occupation, income, marital status, residence, total energy intake, and WBC count.

Model 3 was adjusted for survey year, age, education, occupation, income, marital status, residence, diabetes, toothbrushing habit, regular oral check-up, smoking, and total energy intake.

Bold denotes statistical significance at $P < 0.05$.

EAR= estimated average requirements, RI= recommended intake

4. Discussion

Using an FFQ to estimate daily nutrient intake in a representative Korean sample, we found that, in their daily diets, those who ingested lower quantities of nutrients (calcium in males, vitamin-A, -B₁, -B₂, -C, and iron in females, phosphorus and protein in both genders) associated with having periodontitis in survey year-, age-, and total energy intake-adjusted model. This association modestly attenuated with further adjustment of socioeconomic status (education, occupation, income, marital status, and residence) but remain statistically significant.

Vitamin A has been shown to increase the peroxisome proliferator-activity binding to the peroxisome proliferator-response element that participates in the induction of the Superoxide Dismutase gene and to increase the activity of catalase, Superoxide Dismutase, and glutathione reductase, suggesting that vitamin A may improve the antioxidant defense system¹⁴. All these findings indicate that vitamin A deficiency related inflammation and oxidative stress impairs activities of antioxidant enzymes in vasculature and play a critical role in vascular injury¹⁵ and this might be related to the periodontal tissue destruction. Although there has been a few studies about the association between vitamin A and periodontitis¹⁶, the result of this study partly justified the previous relationship in females.

Thiamin, also known as vitamin B₁, is required for the normal function of muscles and nerves because it converts glucose to energy¹⁷. Riboflavin, also known as vitamin B₂, have an antioxidant action independently or as a component of the glutathione redox cycle and can protect the body against oxidative stress, especially lipid peroxidation and reperfusion oxidative injury. The mechanisms by which riboflavin protects the body against oxidative stress may be attributed to the glutathione redox cycle and also to other possible

mechanisms such as the conversion of reduced riboflavin to the oxidised form¹⁸. One previous study demonstrated that nutrient supplements, in particular vitamin-B complex (Vit-B), can positively influence periodontal wound healing processes¹⁹. The importance of ascorbic acid, better known as vitamin C, for periodontal health has long been known⁸. During the last decades there is growing evidence for the importance of vitamin C as antioxidant against reactive oxygen species (ROS) and in leucocyte function in healthy as well as periodontally diseased subjects²⁰. In a rat periodontitis model, increases in plasma vitamin C levels after vitamin C intake was beneficial in improving periodontitis-induced oxidative stress by down-regulating inflammatory gene expression²¹. Our results did not support the previous study for vitamin B₁, B₂, and C in males but in females only in a survey year-, age adjusted model.

On the other hand, phosphorus and protein intakes were associated with periodontitis among males in our study. From a nutritional standpoint, phosphorus and calcium are essential for the formation of hydroxyapatite, the main crystalline component of bone²². There is evidence that low calcium and high phosphorus intake may have a negative effect on bone health²³. Utilizing data from the third National Health and Nutrition Examination Survey, it has been shown that low dietary intake of calcium results in more severe periodontal disease²³. Most population-based studies¹⁷ suggest that greater dietary protein intake is associated with higher bone mineral density (BMD) in middle-aged and older adults²⁴. Some results showed positive association between protein intake and higher BMD among women²⁵. However, our results showed inconsistency with previous reports. The positive associations between phosphorus/protein and periodontitis were seen in younger male population.

Restriction of dietary intake of phosphorus generally requires some reduction in the allowable protein intake²⁶. The similar associations of phosphorus and protein in this study might be due to the characteristics of phosphorus and protein intake.

Previous studies found no significant difference of serum iron level between periodontitis patients and healthy controls²⁷. Although there was a weak association between iron intake and periodontitis among females in our study, no association was found in a sociodemographic factor-, somatic problem-, and health behavior-adjusted model.

Gender differences have been reported for dietary intakes and eating behaviours²⁸. Previous studies show that women tend to have better overall diet quality than men²⁹. Furthermore, studies also show that dietary intakes of low energy density are associated with a better quality of dietary intakes and a lower BMI³⁰ and evidence of the beneficial influence of healthy eating habits on health status is well established in the literature³¹. In our gender stratified analyses in Table 4 and 5, women had no significant association of nutrients with periodontitis except vitamin A in 45~64 years group. This might be due to overall better diet quality of women prohibit the impact of low nutrient intake to the periodontal tissue destruction. Moreover, low nutrient intake in younger men especially for phosphorus and protein showed strong association with periodontitis. Our results suggested that periodontal bone health might be vulnerable to the low intake of phosphorus and protein among younger population.

Some limitations should be considered in interpreting our findings. First is the selection bias during enrollment, in that more malnourished patients were less likely to enroll. However, selection bias in this direction would lead to bias toward the null; therefore, without this bias, our results may have been even

stronger. Second is the recall bias during the FFQs. FFQs may underestimate the amount of daily nutrient intake. However, our findings are given on the basis of EAR and RI rather than absolute amounts of dietary intakes. Finally, the cross-sectional design prohibited us from inferring causal relationships. Further well designed prospective investigations are required to determine the causality between nutrients intake and periodontitis to reduce the above mentioned limitations. The Food and Nutrition Board cautions that FFQ data may not be accurate enough to use to assess the adequacy of dietary intakes of individuals or small groups of people. FFQs can provide estimates of the long-term, usual dietary intake of certain foods and nutrients in populations of subjects³². Because the FFQ is a reliable tool for ranking individuals according to their dietary intake, assessment of dietary nutrients by FFQ was a suitable choice for this study.

There are several strengths to this study, including the relatively large representative sample of Koreans, the comprehensive dietary evaluations and the detailed evaluation of the clinical and comorbid states of the patients by study physicians. Our study has been extensively characterized for markers of inflammation and nutritional status, and energy intake. A unique feature of this study is its novelty in assessing various nutrients intake measured by FFQ, which is a validated method to estimate long-term usual dietary intake at a population basis and almost certainly a more valid tool for estimating overall nutrients burden.

5. Conclusions

There is an association between low intake of nutrients and periodontitis. The increased risk of periodontitis apparent in some low-level nutrient

intake suggests that people with periodontal disease should be vigilant about undernourishment. Screening programmes for undernourished people would have yielded substantial help of treatable periodontal risks. We need more prospective studies with large representative populations, good descriptions of the nutritional status, long follow-ups, and reliable measurements of health outcomes. The knowledge generated from such studies could be used to develop mechanism-specific interventions aimed at improving the oral health of adults with malnutrition.

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Conflicts

No potential conflict of interest relevant to this article was reported.

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ABSTRACT

Lower Nutrient Intake and Periodontitis: Findings from the Korea National Health and Nutrition Examination Surveys

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Background: Limited information is available regarding the associations of various nutrients and periodontitis in Korea. Furthermore, these associations have been controversial in previous studies. Therefore, the present study aimed to evaluate the associations between several nutrients and periodontitis.

Methods: Pooled data from the 2007~2010 and 2012 Korea National Health and Nutrition Examination Surveys (KNHANES) were used for the present study. Periodontitis was defined using the World Health Organization (WHO) Community Periodontal Index. All participants completed the Food Frequency Questionnaire (FFQ), which was analyzed using CAN-Pro 4.0. Vitamins A, B₁, B₂, and C, iron, phosphorus, calcium, and protein were selected for analysis. Multiple logistic regression analyses were used to estimate the odds ratios (ORs) for periodontitis.

Results: Lower intake of phosphorus (adjusted odds ratio [AOR] = 1.86), calcium (AOR = 1.25), and protein (AOR = 1.52) were associated with periodontitis in males, and lower intake of vitamins A (AOR = 1.32), B₁ (AOR = 1.21), B₂ (AOR = 1.24), and C (AOR = 1.21), iron (AOR = 1.25), phosphorus (AOR = 1.33), and protein (AOR = 1.27) showed significant relationships with periodontitis in females. In the sociodemographic, somatic, and health behavior-adjusted models, lower phosphorus (AOR = 2.68) and protein (AOR = 1.68) intake in younger males, and lower vitamin A intake (AOR = 1.37) in middle-aged females were significantly associated with periodontitis.

Conclusions: The results of the present study demonstrated a significant association between periodontitis and nutrient intake. To avoid an insufficient nutrient supply for a patient with periodontal disease, the patient's diet should be closely monitored.

Keywords: Diet, nutrients; nutrition; oxidative stress; periodontitis