Effects of dietary antioxidant vitamins on lung functions according to gender and smoking status in Korea: a population-based cross-sectional study

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ABSTRACT

Objective Cigarette smoke-induced oxidative stress plays an important role in the pathogenesis of chronic obstructive pulmonary disease (COPD). Dietary antioxidants are thought to prevent smoke-induced oxidative damage. The aim of this study was to investigate associations between lung function and the consumption of antioxidant vitamins in Korean adults.

Methods In total, 21,148 participants from the Korean National Health and Nutrition Examination Survey (2007–2014) were divided into four groups based on smoking history and gender. Multivariate regression models were used to evaluate associations between lung function and intake of dietary antioxidants.

Results Subjects in the highest intake quintile (Q5) of vitamin A, carotene and vitamin C intake had mean forced expiratory volume in 1 s (FEV1) measurements that were 30 mL, 32 mL and 36 mL higher than those of individuals in the lowest intake quintile (Q1), respectively (p for trend; p=0.008, p=0.010 and p<0.001, respectively). The risks of COPD for male smokers in Q1 increased 7.60-fold (95% CI 5.92 to 9.76), 7.16-fold (95% CI 5.58 to 9.19) and 7.79-fold (95% CI 6.12 to 9.92), for vitamin A, carotene and vitamin C, respectively, compared with those of female non-smokers in Q5. Among patients with COPD, men who smoked >20 pack-years had mean FEV1 measurements that were 192 mL, 149 mL and 177 mL higher than those of individuals in Q1 (p for trend; p=0.018, p=0.024 and p=0.043, for vitamin A, carotene and vitamin C, respectively).

Conclusions These findings indicate that the influence of antioxidant vitamins on lung function depends on gender and smoking status in the Korean COPD population.

INTRODUCTION

Chronic obstructive pulmonary disease (COPD) causes morbidity and mortality.1 Smoking is a primary risk factor for COPD; however, other factors also contribute as only 10%–20% of smokers develop airflow limitations.2 Dietary antioxidants protect against oxidative stress caused by smoking,3 and multiple studies have revealed associations between the intake of antioxidant vitamins or fibres and respiratory diseases.4–8 However, evidence supporting the benefits of vitamin supplementation is lacking.9 10

Because micronutrient status is affected by dietary intake and metabolic turnover, which are regulated by oxidative stress, the benefits of antioxidant vitamins may vary by gender and smoking status. Multiple studies have shown that different antioxidants exhibit different effects based on smoking status. Morabia et al reported an association between airway obstruction and vitamin A intake in smokers compared with former smokers, whereas Hu et al reported that carotene was less strongly associated with FEV1 in smokers compared with former smokers and non-smokers.11 12

This study used Korean National Health and Nutritional Examination Survey (KNHANES) data to investigate whether dietary antioxidant vitamins were independently associated with pulmonary function and COPD in the Korean population. This study also evaluated whether the effects of antioxidant vitamins on pulmonary function differed based on gender or smoking status.

PATIENTS AND METHODS

Study population

Participants were sampled from KNHANES (2007–2014) IV–VI, a nationwide survey
of antioxidant vitamins were calculated using the Korean Food Composition Table as the reference. Antioxidant vitamin consumption was adjusted for total energy intake.

**Potential confounders**

Data regarding demographic information, education level, household income, smoking status, smoking amount, alcohol intake, place of residence, body mass index (BMI) and comorbid diseases were obtained. Educational level was categorised as elementary school or lower, completion of middle school, completion of high school and college or higher. Household income was divided by quartile. Place of residence was divided into rural and urban.

Smokers were subjects who smoked more than 100 cigarettes in their lifetime. Participants were categorised in terms of smoking status as follows: smoker, ex-smoker or never smoked. Those who answered in the negative to the question ‘Do you currently smoke?’ were defined as ex-smokers.

The smoking amount was determined in pack-years, which was calculated by multiplying the duration of smoking (years) by the number of packs of cigarettes smoked. Comorbid diseases included hypertension, stroke, cardiovascular disease, arthritis, tuberculosis, asthma, diabetes mellitus, thyroid disorders, renal failure, liver disease and malignancy.

**Statistical analysis**

A total of 21,148 subjects participated in this study (figure 1). The relationship between antioxidant vitamin intake and lung function was analysed using multiple linear regression analyses. We analysed the energy-adjusted antioxidant vitamin intake by quintiles. The adjustment factors were age, sex, BMI, educational level, household income, total energy intake, place of residence, number of comorbid diseases, smoking history, alcohol intake and pack-years. Assessments of linear trends across increasing antioxidant vitamin quintiles were also performed.

We estimated the ORs of COPD using multivariate logistic regression analyses of quintiles after adjusting for confounding factors. Participants were divided into four groups based on gender and smoking status (male smokers, male non-smokers, female smokers, female non-smokers) to determine whether the relationship between COPD risk and antioxidant vitamin intake is related to gender and smoking status. For combined analyses between the effects of antioxidant vitamin intake, gender and smoking status on the risk of COPD, interaction tests were performed. Patients with COPD and male patients with COPD were analysed separately. We attempted to determine whether the association of antioxidant vitamins and lung function varies with gender and smoking status in patients with COPD. Multiple linear regression analyses were performed after categorising patients with COPD by smoking status and amount.
Statistical analyses were performed using PASW Statistics V.20 (SPSS) and SAS V.9.4 (SAS Institute) software. P<0.05 was considered statistically significant.

RESULTS
The baseline characteristics of the 21148 participants are shown in table 1. All subjects were classified into four groups based on smoking history and gender. Of the 7986 smokers, 7178 were male (mean age, 57.8±11.0 years) and 808 were female (mean age, 57.4±12.5 years). Of the 13162 individuals who had never smoked, 1626 were male (mean age, 57.9±11.3 years) and 11536 were female (mean age, 57.1±10.8 years). Among all subjects, 3005 were diagnosed with COPD. The prevalence of COPD was highest in male smokers (26.4%) and lowest in female non-smokers (6.4%).

The four groups differed regarding age, BMI, educational level, household income and alcohol usage (p<0.001). Energy intake was significantly higher in men than women (men, 2256.5 kcal; women, 1648.3 kcal; p<0.001). The levels of vitamin A, carotene and vitamin C were highest in the male non-smoker group and lowest in the female smoker group. Korean male non-smokers are predisposed to COPD compared with female non-smokers (incidence rate of 15.7% vs 6.4%). Age and the percentage of alcohol intake were higher in Korean male non-smokers than female non-smokers.

Table 2 showed the association between lung function (FEV₁, FVC) and dietary antioxidant vitamin levels. Participants in the highest quintile (Q5) of vitamin A intake had 30 mL higher FEV₁ (p for trend across quintiles=0.008) and 33 mL higher FVC (p for trend across quintiles=0.007) compared with participants in the lowest quintile (Q1). Participants in Q5 for carotene intake had 32 mL higher FEV₁ (p for trend across quintiles=0.010) and 36 mL higher FVC (p for trend across quintiles=0.005) measurements compared with participants in Q1. Participants in Q5 of vitamin C intake had 36 mL higher FEV₁.
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(p for trend across quintiles<0.001) and 35mL higher FVC (p for trend across quintiles=0.014) measurements compared with participants in Q1. A statistically significant dose–response relationship was observed (all, p for trend across quintiles<0.005), but participants in Q3 of vitamin A and carotene had comparable lung function to those in Q5.

The effects of gender, smoking and dietary antioxidant vitamins on the risk of COPD are summarised in table 3. The risk of COPD for male smokers in Q1 for vitamin A, carotene and vitamin C intake increased by 3.26-fold (95% CI 2.24 to 4.75), 3.35-fold (95% CI 2.31 to 4.86) and 3.28-fold (95% CI 2.27 to 4.73), respectively, compared with female non-smokers in Q5 for antioxidant vitamin intake. The risk of COPD for male non-smokers in Q1 for vitamin A, carotene and vitamin C intake increased by 2.80-fold (95% CI 1.90 to 4.12), 3.25-fold (95% CI 2.21 to 4.78) and 3.17-fold (95% CI 2.04 to 4.91), respectively, compared with female non-smokers in Q1 for antioxidant vitamin intake. These results suggest that men may have other causes of COPD as well as smoking, compared with women who took similar amounts of antioxidant vitamins.

The interaction exists between the antioxidant vitamin intake and gender/smoking status on the risk of COPD.

Table 1  Study population characteristics

|                     | Total (n=21 148) | Male smokers (n=7178) | Male non-smokers (n=1 626) | Female smokers (n=808) | Female non-smokers (n=11 536) | P values |
|---------------------|------------------|----------------------|---------------------------|------------------------|-------------------------------|----------|
| **Age**             | 57.4 (10.9)      | 57.8 (11.0)          | 57.9 (11.3)               | 57.4 (12.5)            | 57.1 (10.8)                   | <0.001   |
| 40–49               | 6048 (28.6)      | 1998 (27.8)          | 464 (28.5)                | 273 (33.8)             | 3313 (28.7)                   | <0.001   |
| 50–59               | 6131 (29.0)      | 1981 (27.6)          | 431 (26.5)                | 199 (24.6)             | 3520 (30.5)                   |          |
| 60–69               | 5387 (25.5)      | 1913 (26.7)          | 430 (26.4)                | 158 (19.6)             | 2866 (25.0)                   |          |
| 70–                 | 3582 (16.9)      | 1286 (17.9)          | 301 (18.5)                | 178 (22.0)             | 1817 (15.8)                   |          |
| BMI*                | 24.2 (3.0)       | 24.2 (2.8)           | 24.3 (2.8)                | 23.8 (3.6)             | 24.2 (3.2)                    | 0.007    |
| Education           | <0.001           |                      |                           |                        |                               |          |
| Elementary          | 7229 (34.2)      | 1763 (24.6)          | 321 (19.7)                | 381 (47.2)             | 4764 (41.3)                   |          |
| Middle school       | 3315 (15.7)      | 1216 (16.9)          | 267 (16.4)                | 112 (13.9)             | 1720 (14.9)                   |          |
| High school         | 6427 (30.4)      | 2366 (33.0)          | 458 (28.2)                | 228 (28.2)             | 3375 (29.3)                   |          |
| More than college   | 4169 (19.7)      | 1831 (25.5)          | 580 (35.7)                | 87 (10.8)              | 1671 (14.5)                   |          |
| Household income    | <0.001           |                      |                           |                        |                               |          |
| First quartile      | 4763 (22.5)      | 1440 (20.1)          | 289 (17.8)                | 315 (39.0)             | 2719 (23.6)                   |          |
| Second quartile     | 5427 (25.7)      | 1874 (26.1)          | 391 (24.1)                | 223 (27.6)             | 2939 (25.5)                   |          |
| Third quartile      | 5162 (24.4)      | 1869 (26.1)          | 414 (25.5)                | 145 (17.9)             | 2734 (23.7)                   |          |
| Fourth quartile     | 5780 (27.3)      | 1988 (27.7)          | 530 (32.6)                | 125 (15.5)             | 3137 (27.2)                   |          |
| Comorbidity*        | 0.9 (1.1)        | 0.9 (1.0)            | 0.8 (0.9)                 | 1.0 (1.2)              | 1.0 (1.1)                     | <0.001   |
| Pack-years*         | 4.7 (13.6)       | 13.3 (20.3)          | 0.2 (2.2)                 | 3.3 (11.9)             | 0.0 (0.0)                     | <0.001   |
| Alcohol             | 17 554 (83.0)    | 6877 (95.8)          | 1399 (86.0)               | 714 (88.4)             | 8564 (74.2)                   | <0.001   |
| Energy intake (kcal/ day)* | 1901.5 (797.8) | 2266.5 (869.7)       | 2212.6 (855.9)            | 1538.2 (653.4)         | 1656.0 (630.0)                | <0.001   |
| Vitamin A (μg RE/day)* | 822.5 (1118.5) | 881.9 (1067.5)       | 925.5 (1095.2)            | 600.3 (644.2)          | 786.6 (1173.9)                | <0.001   |
| Carotene (μg/day)*  | 4337.3 (6206.0)  | 4596.2 (5557.8)      | 4803.8 (5506.6)          | 3143.5 (3862.6)        | 4194.1 (6780.6)               | <0.001   |
| Vitamin C (mg/day)* | 111.9 (107.6)    | 111.8 (97.9)         | 128.8 (107.1)            | 84.8 (96.9)            | 111.5 (113.5)                 | <0.001   |
| FEV1 (mL)*          | 2.60 (0.67)      | 3.02 (0.68)          | 3.09 (0.66)               | 2.23 (0.56)            | 2.30 (0.46)                   | <0.001   |
| FVC (mL)*           | 3.38 (0.84)      | 4.07 (0.72)          | 4.04 (0.73)               | 2.88 (0.62)            | 2.89 (0.51)                   | <0.001   |
| FEV1/FVC (%)*       | 77.3 (7.9)       | 73.9 (9.1)           | 76.6 (7.9)                | 77.2 (8.0)             | 79.5 (6.1)                    | <0.001   |
| COPD                | 3005 (14.2)      | 1893 (26.4)          | 256 (15.7)                | 119 (14.7)             | 737 (6.4)                     | <0.001   |

*Numbers represent mean percentages (SD).
BMI, body mass index; COPD, chronic obstructive pulmonary disease; FVC, forced vital capacity; FEV1, forced expiratory volume in 1 s; RE, retinol equivalent.
Table 2  Mean values of adjusted lung function measurements across quintiles of vitamin A, carotene and vitamin C intake

|                      | Q1     | Q2     | Q3     | Q4     | Q5     | Difference between Q5 and Q1 (95% CI) | P values for trend |
|----------------------|--------|--------|--------|--------|--------|---------------------------------------|--------------------|
| **Vitamin A**        |        |        |        |        |        |                                       |                    |
| Mean intake (μg RE)  | 151.2  | 353.6  | 573.1  | 893.9  | 2140.8 |                                        |                    |
| FEV₁ (mL)            | 2379   | 2389   | 2410   | 2397   | 2409   | 30 (10 to 50)                         | 0.008              |
| FVC (mL)             | 3119   | 3136   | 3158   | 3148   | 3152   | 33 (10 to 57)                         | 0.007              |
| Predicted FEV₁ (%)   | 91.37  | 91.44  | 91.91  | 91.45  | 91.94  | 0.57 (-0.08 to 1.22)                  | 0.185              |
| Predicted FVC (%)    | 90.93  | 91.06  | 91.45  | 91.22  | 91.48  | 0.55 (0.00 to 1.10)                   | 0.195              |
| **Carotene**         |        |        |        |        |        |                                       |                    |
| Mean intake (μg)     | 691.1  | 1747.4 | 2938.9 | 4736.1 | 11574.1|                                        |                    |
| FEV₁ (mL)            | 2347   | 2363   | 2376   | 2370   | 2379   | 32 (12 to 52)                         | 0.010              |
| FVC (mL)             | 3088   | 3117   | 3127   | 3119   | 3124   | 36 (13.59)                            | 0.005              |
| Predicted FEV₁ (%)   | 91.55  | 92.03  | 92.26  | 91.85  | 92.31  | 0.76 (0.12 to 1.39)                   | 0.096              |
| Predicted FVC (%)    | 91.02  | 91.68  | 91.78  | 91.41  | 91.82  | 0.80 (0.26 to 1.33)                   | 0.015              |
| **Vitamin C**        |        |        |        |        |        |                                       |                    |
| Mean intake (mg)     | 24.2   | 53.6   | 84.2   | 128.8  | 268.9  |                                        |                    |
| FEV₁ (mL)            | 2411   | 2423   | 2436   | 2441   | 2453   | 36 (16.56)                            | <0.001             |
| FVC (mL)             | 3117   | 3122   | 3132   | 3140   | 3154   | 35 (12.58)                            | 0.014              |
| Predicted FEV₁ (%)   | 91.3   | 91.5   | 91.9   | 91.99  | 92.21  | 0.91 (0.27 to 1.55)                   | 0.050              |
| Predicted FVC (%)    | 91.29  | 91.33  | 91.58  | 91.77  | 92.0   | 0.71 (0.17 to 1.26)                   | 0.118              |

Data were adjusted for age, sex, body mass index, energy intake, number of comorbid diseases, alcohol consumption, place of residence smoking history, pack-years (smoking amount), household income and education level. P values were determined using tests for linear trends across increasing quintiles (means) of antioxidant vitamin intake.

FVC, forced vital capacity; FEV₁, forced expiratory volume in 1 s; RE, retinol equivalent; Q1, lowest quintile; Q5, highest quintile.

Table 3  Association between vitamin A, carotene and vitamin C intake and COPD according to gender and smoking status

|                      | Intake | COPD | OR     | P interaction |
|----------------------|--------|------|--------|---------------|
|                      | Q5     | Q1   | Q5     | Q1            |
| **Vitamin A**        |        |      |        |               |
| Female non-smokers   | 2096   | 2564 | 105    | 242           | Ref           | 1.16 (0.89, 1.49) | <0.001 |
| Female smokers       | 109    | 264  | 16     | 53            | 3.90 (2.12, 7.17) | 2.42 (1.63, 3.58) |      |
| Male non-smokers     | 394    | 225  | 53     | 47            | 3.26 (2.24, 4.75) | 3.15 (2.10, 4.72) |      |
| Male smokers         | 1630   | 1176 | 320    | 444           | 5.54 (4.28, 7.16)* | 7.60 (5.92, 9.76)* |      |
| **Carotene**         |        |      |        |               |
| Female non-smokers   | 2118   | 2529 | 108    | 226           | Ref           | 1.10 (0.85, 1.42) | <0.001 |
| Female smokers       | 104    | 268  | 15     | 49            | 3.47 (1.86, 6.47) | 2.16 (1.45, 3.23) |      |
| Male non-smokers     | 397    | 243  | 55     | 50            | 3.35 (2.31, 4.86) | 3.24 (2.18, 4.82) |      |
| Male smokers         | 1610   | 1189 | 321    | 425           | 5.83 (4.51, 7.53)* | 7.16 (5.58, 9.19)* |      |
| **Vitamin C**        |        |      |        |               |
| Female non-smokers   | 2303   | 2466 | 112    | 465           | Ref           | 1.00 (0.77, 1.30) | <0.001 |
| Female smokers       | 107    | 294  | 12     | 35            | 2.37 (1.20, 4.71) | 2.27 (1.55, 3.34) |      |
| Male non-smokers     | 401    | 191  | 55     | 55            | 3.28 (2.27, 4.73) | 3.24 (2.07, 5.06) |      |
| Male smokers         | 1419   | 1278 | 317    | 204           | 6.20 (4.82, 7.98)* | 7.79 (6.12, 9.92)* |      |

OR was determined following adjustment for age, body mass index, energy intake, number of comorbid diseases, alcohol consumption, place of residence, household income and education level.

*COPD, chronic obstructive pulmonary disease; Q1, lowest quintile; Q5, highest quintile.
The effect of the antioxidant vitamin intake depends on the gender/smoking status. When assessing the risk of COPD following reduction of antioxidant intake from Q5 to Q1, only male smokers showed significant difference in risk of COPD, but other three groups did not. Figure 2 shows that the risk of COPD was influenced by dietary antioxidant vitamin levels in male smokers in detail. In male smokers, the risk of COPD in subjects in Q5 for antioxidant vitamins intake was significantly lower than that for subjects in Q1 (vitamin A, OR=0.77, 95% CI 0.63 to 0.94, p=0.009; carotene, OR=0.81, 95% CI 0.67 to 0.99, p=0.041; vitamin C, OR=0.74, 95% CI 0.61 to 0.91, p=0.004). The dose–dependent effect of vitamin C was observed between COPD risk and dietary antioxidant vitamin levels, but it was not for vitamin A and carotene. Although not significant, Q3 group of carotene had increased risk to develop COPD than Q1 group of carotene.

The prevalence of COPD did not increase significantly as the intake of dietary antioxidant vitamins increased in male non-smokers, female smokers or female non-smokers. No significant interaction between the effects of antioxidant vitamins on COPD and smoking status was observed. The correlation between the risk of COPD and antioxidant vitamin intake was stronger in male smokers who smoked less than 20 pack-years (not shown).

We investigated the association between dietary antioxidant vitamin intake and lung function after limiting the analyses to individuals with COPD. The changes in FEV₁ were not statistically significant based on the levels of dietary antioxidant vitamins in subjects with COPD. Similar to the previous results, only male smokers in subjects with COPD, exhibited a beneficial association between dietary antioxidant vitamin intake and FEV₁ (figure 3). Male smokers with COPD in Q5 for vitamin A intake had a 71 mL higher FEV₁ (p for trend across quintiles=0.019) compared with those in Q1. Male smokers with COPD in Q5 of carotene and vitamin C intake had 71 mL higher FEV₁ (p for trend across quintiles=0.037) and a 109 mL higher FEV₁ (p for trend across quintiles=0.026), respectively, compared with individuals in Q1.

Additional analyses were performed to determine if lung function was reduced by smoking amount or smoking status in male smoker patients with COPD. Male patients with COPD who had smoked ≥20 pack-years exhibited a beneficial association between dietary antioxidant vitamin intake and FEV₁ (figure 4). Male patients with COPD in Q5 of vitamin A intake who had smoked ≥20 pack-years had a 192 mL higher FEV₁ (p for trend across quintiles=0.018) compared with individuals in Q1. Patients with COPD in Q5 of carotene intake who had smoked ≥20 pack-years had a 149 mL higher FEV₁ (p for trend across quintiles=0.024) compared with patients in Q1. Patients with COPD in Q5 of vitamin C intake who had smoked >20 pack-years had a 177 mL higher FEV₁ (p for trend across quintiles=0.043) compared with patients in Q1.

**DISCUSSION**

This study examined the association between the intake of antioxidant vitamins and lung functions in the Korean population.

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Figure 2 OR for the association between antioxidant vitamin intake and COPD among (A) male and (B) female smokers and non-smokers. OR were adjusted for age, body mass index, energy intake, number of comorbid diseases, alcohol consumption, place of residence, household income and education level.
population. Previous studies showed that antioxidant vitamins, including vitamin C, were protective of the human lung, whereas high levels of vitamin A and carotene were also associated with increased lung functions in multiple studies. In a randomised controlled trial, Keranis et al reported that increasing the intake of antioxidants improved lung function.

Cigarette smoking is the primary cause of COPD as it increases oxidative stress in the lungs and activates inflammatory responses. Notably, one inhalation from a cigarette generates more than $10^{15}$ free radicals and other oxidants.

Antioxidants protect against the damage caused by smoking in multiple ways. For example, as it is...
water soluble, vitamin C scavenges free radicals in the cytoplasm. Koike et al reported that vitamin C diminished smoke-induced oxidative stress and corrected emphysematous lungs in vivo.27

Carotenoids quench singlet oxygen and inhibit lipid peroxidation.3 In an animal study, the respiratory epithelial of retinol-deficient animals had atrophied ciliated cells and modified lipid contents.28 The pathological features of the retinol-deficient animals were similar to those of human smokers.29

Smokers exhibit nicotine-induced reductions in intestinal absorption and elevated metabolic turnover.20 The metabolism or destruction of antioxidant vitamins increases in inflammatory environments31–34 which suggests that smokers with COPD require larger amounts of antioxidant vitamins to achieve the same blood levels as non-smokers. A study by Sargeant and Jaeckel found that vitamin C may modify the adverse effects of smoking and the risk of COPD in the European population.35 Additionally, Shin et al reported that Korean smokers with adequate vitamin C intake had acceptable pulmonary functions36 and Park et al showed that dietary vitamin C provides protection against COPD.37 Additionally, Morabia et al identified that airway obstruction was reduced by vitamin A in smokers.38

One notable finding in the current study was that the effects of antioxidant vitamin intake on lung function were stronger among male smokers. Additionally, the association between the risk of COPD and antioxidant vitamin intake was clear for male but not female smokers. Male smokers with lower antioxidant vitamin intakes had increased ORs of COPD compared with female smokers. Although the dose-dependent effect on COPD risk was not obvious in vitamin A and carotene, contrary to vitamin C (figure 2), male smokers with Q5 intake showed a clearly reduced risk to develop COPD than male smokers with Q1 intake in all three antioxidant vitamins.

After limiting the analysis to subjects with COPD, a significant association between antioxidant vitamin intake and FEV₁ was observed in male smokers but not in other groups. This finding was similar to that of Joshi et al, where changes in COPD risk and dietary vitamin C and vitamin E intake differed between men and women.38

It is not known how gender differences impact pulmonary functions based on antioxidant vitamin intake; however, animal studies have revealed gender differences in antioxidant vitamin requirements. Al-Rejaie et al reported gender-related differences in the protective roles of ascorbic acid against oxidative stress,39 whereas Jiao et al revealed gender differences in the regulation and expression of oxidative genes in mice.40

Studies detailing the effects of antioxidant vitamins on lung function in smokers and non-smokers are lacking.5,23 In the US population, Britton et al revealed that the relationship between vitamin C intake and FEV₁ was stronger in ex-smokers than non-smokers or current smokers.35 Shahar et al reported a relationship between individuals in Q1 of vitamin A intake and airway obstruction among individuals who smoked >41 pack-years.25 Among male patients with COPD, those smoking ≥20 pack-years had improved lung functions as antioxidant vitamin intake increased. These results support that associations between antioxidants and lung function may differ according to smoking status in patients with COPD patients. However, it is unknown what causes such differences. One hypothesis is that the efficacy of antioxidant vitamins is proportional to the level of oxidant burden in COPD. Additional studies are required to determine whether the benefits of antioxidant vitamins depend on the smoking duration or dose in patients with COPD.

This study has several limitations that should be noted. As we used a cross-sectional design, the data cannot be used to answer questions regarding causation. Additionally, because data on nutritional intake were obtained by 24 hours recall, inaccurate responses may have been offered. This study used the prebronchodilator FEV₁ for determining COPD; however, the definition of COPD is based on postbronchodilator FEV₁.15 This study failed to obtain data regarding air pollution or occupational exposure and, therefore, could not associate these variables with lung function; however, the strength of this study is that these data represent the Korean population.

**CONCLUSION**

This study supports that antioxidant vitamins have beneficial effects on pulmonary function in the Korean population. The data indicate that there is a stronger association between antioxidant vitamin intake and the risk of COPD in male smokers. The beneficial effects of antioxidant vitamins in patients with COPD differed by gender and smoking status, and future investigations should determine the roles of dietary antioxidant vitamins in specific groups.

**Contributors** JYH and YSK equally contributed to the conception and design of the research; YSK contributed to the design of the research; CYL contributed to the acquisition and analysis of the data; MGL and YSK contributed to the interpretation of the data and JYH drafted the manuscript. All authors critically revised the manuscript, agreed to be fully accountable for ensuring the integrity and accuracy of the work and read and approved the final manuscript.

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