INTRODUCTION

Colonic diverticulosis is an acquired disease occurring by herniation of the mucosa through weak point in the colonic muscle wall. Approximately 60% of people older than 60 years in Westernized countries will develop asymptomatic or symptomatic diverticulosis. In most cases, the diverticulosis is asymptomatic, with only 10% to 25% of subjects with diverticulosis developing symptomatic diseases. Diverticular disease of the colon is defined as symptomatic and clinically significant diverticulosis which includes symptomatic uncomplicated diverticular disease, diverticulitis, diverticular bleeding, and segmental colitis associated with diverticular disease.

At present, diverticulosis is considered to be one of the most common GI disorders, and trends suggest an increasing incidence of associated complications and health care costs. Two features dominate the epidemiologic profile of diverticulosis:...
geography and age. Reports from countries such as Israel, Japan, Kenya, Singapore, and Uganda suggest a greater and growing prevalence of diverticulosis in these non-Western countries than previously realized.9,12 The incidence of colonic asymptomatic diverticulosis in Korea is also growing.13 This may be associated with a Westernized lifestyle, population aging, and increasing number of screening colonoscopy.14

Several risk factors are known to be associated with colonic diverticular disease. Lifestyle factors, including high dietary intake of red meat, low fiber intake, lack of vigorous physical activity, obesity, and smoking, have been associated with diverticular disease.15,16 However, except old age and male sex, other predisposing factors for asymptomatic diverticulosis are largely unknown. The role of low fiber consumption in the formation of diverticulosis is also controversial.14,16-18 In particular, data about this association in Asians are unknown. The purpose of our study was to assess the effect of vegetarianism on asymptomatic colonic diverticulosis in Korean Buddhist monks who are obligatory vegetarians for spiritual reasons compared with the general population.

**METHODS**

1. **Study Population**

This was a case-control, cross-sectional study. We enrolled all individuals who got colonoscopy during a health check-up at the Medical Healthcare Center of Dongguk University Ilsan Hospital (Goyang, Korea) from August 2005 to June 2018. A total of 12,602 subjects, including 809 Buddhist priests, underwent colonoscopy for screening purposes during this period. Exclusion criteria were incomplete study due to cecal intubation failure; poor bowel preparation; repeated colonoscopic examination during the study period; medical history of colectomy, colorectal cancer, or diverticular disease; and severe abdominal symptoms such as abdominal pain or dyspepsia.

Among the Buddhist monks, 151 subjects met 1 or more of the exclusion criteria; therefore, 658 Buddhist monks were included. To adjust for age and sex which are considered as important confounders in colonic diverticulosis, we formed a control group by randomly matching subjects from the general population according to age and sex (Fig. 1). The individuals who answered that they were vegetarian in the questionnaire were excluded from the general population group. Accordingly, 1,316 subjects were finally selected for this study. The Institutional Review Board of Dongguk University Hospital approved this study protocol (IRB No. 2019-02-017). The informed consent was waived.

2. **Clinical and Laboratory Evaluation**

All subjects were asked to complete questionnaires about abdominal symptoms (dyspepsia, abdominal pain, and hematochezia), smoking status, alcohol consumption, physical activity, and past medical history (Supplementary Material 1). This also included a question about the amount of weekly meat intake. All subjects’ height, weight, waist circumference, and blood pressure were measured by skilled personnel with a

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**Fig. 1.** Flow diagram of study population in this study.
standardized protocol. Venous blood samples were gathered after overnight fasting. Blood complete cell counts, total cholesterol, triglycerides, HDL cholesterol, LDL cholesterol, fasting glucose, and hemoglobin A1c (HbA1c) were measured.

3. Colonoscopy and Detection of Colonic Diverticulosis
All colonoscopies were carried out by 8 experienced endoscopists with endoscopy specialty board certification. Standard colonoscopes (CF-H260AL or CF-H290L; Olympus Medical Systems, Tokyo, Japan) were used for colonoscopy. Most subjects underwent conscious sedation colonoscopy using intravenous midazolam and pethidine. If diverticulosis were observed, their locations were recorded. Three endoscopy professionals also reviewed all photographs of colonoscopies to find cases, in which diverticulosis had been forgotten to record by mistake. In addition, diverticulum is generally located between fold and another fold. So, if the endoscopist do not take much time during withdrawal, missing rate may increase. So, we analyzed colonoscopy withdrawal time as a quality indicator in both groups. Withdrawal time was calculated by picture archiving and communication system, in which the endoscopists captured the colonoscopic image with the present time simultaneously. We calculated the withdrawal time by subtracting the time between cecum to anus only in colonoscopies with negative results. We defined the location of diverticula as follows: the right side of the colon consist of the cecum, ascending colon, or transverse colon; the left side of the colon consist of the splenic flexure, descending colon, sigmoid colon, or rectum.

4. Definitions
Smoking meant smoking more than 1 cigarette regularly during the previous 1 year. Drinking meant drinking >140 g of alcohol weekly. Hypertension meant ≥ 140/90 mmHg or taking antihypertensive drugs. Individuals were diagnosed as metabolic syndrome in case that they met over 3 of the next 5 criteria: (1) abdominal obesity (waist circumference > 90 cm in men and > 80 cm in women); (2) blood pressure ≥130/85 mmHg or antihypertensive medication; (3) fasting plasma glucose ≥ 100 mg/dL; (4) triglyceride levels ≥ 150 mg/dL; and (5) HDL cholesterol < 40 mg/dL for men and < 50 mg/dL for women.

5. Statistical Analyses
Statistical analysis was carried out by SPSS version 23.0 for Windows (IBM Corp., Armonk, NY, USA). Student t-test or the Mann-Whitney U-test for continuous factors and Pearson chi-square test (or Fisher exact test, when appropriate) for categorical factors were used to examine the association between colonic diverticulosis and the factors. ORs and 95% CIs were calculated using conditional logistic regression. Variables with P < 0.05 in the univariate analysis were added to the multivariate logistic regression analysis to detect independent risk factors for colonic diverticulosis. P-values < 0.05 were considered statistically significant.

RESULTS
1. Baseline Characteristics of the Study Population and Comparison between 2 Groups
A total of 1,316 individuals were enrolled in this study (Bud-
Vegetarianism and asymptomatic colonic diverticulosis

The overall prevalence of colonic diverticulosis was 8.7% (115/1,316). The mean age of the 2 groups was 52.6 ± 9.5 years, and 46.8% had male sex. Between the 2 groups, there were no statistically significant differences regarding hypertension, exercise ≥ 3 times/wk, AST, ALT, triglycerides, LDL, HDL levels and mean withdrawal time. Mean BMI (24.5 kg/m² vs. 23.7 kg/m², *P* < 0.001), HbA1c (5.71% vs. 5.57%, *P* = 0.002), and total cholesterol level (205.0 mg/dL vs. 198.2 mg/dL, *P* < 0.001) were higher and HDL level (53.1 mg/dL vs. 58.7 mg/dL, *P* < 0.001) was lower in the Buddhist monk group than in the general population group. Metabolic syndrome is more prevalent in the Buddhist monk group (23.7% vs. 17.8%, *P* = 0.008) and a lower prevalence of diverticulosis (6.7% vs. 10.8%, *P* = 0.008) than the control group. Both right-sided (6.1% vs. 9.9%, *P* = 0.010) and left-sided (0.9% vs. 2.7%, *P* = 0.006) diverticulosis were less prevalent in the Buddhist monk group (Table 1).

### 2. Risk Factors for Colonic Diverticulosis

It was investigated whether a vegetarian diet is associated with colonic diverticulosis using conditional regression analysis. By the univariate analysis, colonic diverticulosis was related with old age, male sex, BMI ≥ 25 kg/m², and a nonvegetarian diet. However, it was not associated with current smoking, alcohol consumption, or exercise (Table 2). Even in a conditional regression analysis model adjusted for age, male, BMI, and nonvegetarian diet, old age (age ≥ 60 years vs. ≤ 39 years: OR, 4.53; 95% CI, 1.36–15.12; *P* = 0.014), male sex (OR, 1.91; 95% CI, 1.28–2.85; *P* = 0.002), BMI ≥ 25 kg/m² (OR, 1.50; 95% CI, 1.01–2.23; *P* = 0.047), and nonvegetarian diet (OR, 1.82; 95% CI, 1.21–2.72; *P* = 0.004) were related with colonic diverticulosis (Table 3).

### 3. Risk Factors for Right- or Left-Sided Colonic Diverticulosis

We also examined whether a vegetarian diet is associated with diverticulosis.

#### Table 2. Effect of Factors on Colonic Diverticulosis

| Factor                  | Control (n = 1,201) | Diverticulosis (n = 115) | OR (95% CI) | *P*-value |
|-------------------------|---------------------|--------------------------|-------------|-----------|
| Age (yr)                |                     |                          |             |           |
| ≤ 39                    | 103 (8.6)           | 3 (2.6)                  | 1 (reference) | < 0.001   |
| 40–49                   | 388 (32.3)          | 24 (20.9)                | 2.12 (0.63–7.19) | 0.226     |
| 50–59                   | 440 (36.6)          | 52 (45.2)                | 4.05 (1.24–13.25) | 0.020     |
| ≥ 60                    | 270 (22.5)          | 36 (31.3)                | 4.58 (1.38–15.19) | 0.013     |
| Male sex                | 544 (45.3)          | 72 (62.6)                | 2.02 (1.36–3.00) | < 0.001   |
| Current smoking         | 89 (7.4)            | 14 (12.2)                | 1.73 (0.95–3.15) | 0.069     |
| Alcohol consumption     | 96 (8.2)            | 19 (16.5)                | 1.59 (0.94–2.68) | 0.081     |
| Exercise (≥ 3 times/wk) | 271 (22.6)          | 24 (20.9)                | 0.91 (0.56–1.45) | 0.677     |
| BMI (kg/m²)             | 24.00 ± 3.23        | 25.01 ± 3.35             | 1 (reference) | 0.001     |
| < 25                    | 761 (92.7)          | 60 (7.3)                 | 1 (reference) |           |
| ≥ 25                    | 440 (88.9)          | 55 (11.1)                | 1.59 (1.08–2.33) | 0.019     |
| Hypertension (present)  | 228 (19.0)          | 28 (24.3)                | 1.37 (0.88–2.15) | 0.165     |
| Metabolic syndrome      | 243 (20.2)          | 30 (26.1)                | 1.39 (0.90–2.16) | 0.139     |
| HbA1c (%)               | 5.63 ± 0.84         | 5.82 ± 0.61              | -            | 0.090     |
| AST (IU/L)              | 31.17 ± 40.06       | 31.52 ± 17.28            | -            | 0.927     |
| ALT (IU/L)              | 29.68 ± 59.81       | 31.37 ± 27.12            | -            | 0.765     |
| Triglyceride (mg/dL)    | 120.40 ± 83.70      | 129.82 ± 83.87           | -            | 0.249     |
| LDL cholesterol (mg/dL) | 132.09 ± 35.12      | 134.53 ± 37.01           | -            | 0.479     |
| HDL cholesterol (mg/dL) | 56.00 ± 15.25       | 54.44 ± 15.05            | -            | 0.296     |
| Total cholesterol (mg/dL) | 201.60 ± 38.67    | 202.04 ± 42.04           | -            | 0.906     |
| Vegetarian diet         | 614 (51.1)          | 44 (38.3)                | 1.69 (1.14–2.50) | 0.008     |

Values are presented as mean ± SD or number (%).
HbA1c, hemoglobin A1c.
Differences were considered significant when *P*-values were < 0.05.
right-sided and left-sided diverticulosis. In univariate analysis, old age, male sex, higher BMI, and nonvegetarian diet were associated with right-sided colonic diverticulosis. In a multivariate analysis adjusted for age, male, BMI, and vegetarian diet, old age (age ≥ 60 years vs. ≤ 39 years: OR, 4.14; 95% CI, 1.24–13.87; P = 0.021), male sex (OR, 1.78; 95% CI, 1.17–2.69; P = 0.007), BMI ≥ 25 kg/m² (OR, 1.50; 95% CI, 1.01–2.27; P = 0.047), and nonvegetarian diet (OR, 1.82; 95% CI, 1.20–2.76; P = 0.005) were associated with an increased risk for right-sided colonic diverticulosis (Table 4).

Furthermore, old age, male sex, and nonvegetarian diet were associated with left-sided colonic diverticulosis in univariate analysis. Even in a conditional regression model, old age (OR, 3.50; 95% CI, 1.51–8.11; P = 0.004), male sex (OR, 5.55; 95% CI, 1.86–16.5; P = 0.002), and nonvegetarian diet (OR, 3.90; 95% CI, 1.43–10.67; P = 0.008) continued to be associated with increased risk for left-sided colonic diverticulosis (Table 5).

**DISCUSSION**

This study showed that a nonvegetarian diet was significantly associated with the risk of asymptomatic colonic diverticulo-
Vegetarianism was found to be a protective factor for asymptomatic colonic diverticulosis even after adjusting for several confounding factors. Regardless of the location of diverticulosis, vegetarianism showed a protective effect. Old age, male sex, and high BMI were independently associated with the risk of asymptomatic colonic diverticulosis. The prevalence of diverticulosis in this study was 10.8% in the general population group which is similar with that of previous reports.

In our study, the risk factors for diverticulosis were similar between right-sided and left-sided diverticulosis. In Asians, colonic diverticulosis occurs mainly in the right-sided colon, whereas the sigmoid colon is more commonly involved in Westerners. Most Korean studies, including this study, have also reported that colonic diverticula are more common in the right side than in the left side (about 80%).14,20 The pathogenesis of right-sided and left-sided diverticulosis may differ. However, a Japanese study reported that the etiology of right-sided diverticulosis in Asians may be similar to that of left-sided diverticulosis in Western countries. The authors concluded that the differences in location may stand for a difference in the morphology of colon between Asians and Western people, rather than differences in environment.21

In addition to race or genetic influences, many investigators have showed that dietary pattern, colonic abnormal motility, and alterations in colonic structure are the causes of diverticulosis development.21,22 Painter and Burkitt17 first suggested that a fiber deficiency is an etiology of diverticular disease. This hypothesis is that the deficiency of dietary fiber intake causes reduced intestinal contents to make bowel lumen smaller. Subsequently, pressure from muscular contractions is applied to the colonic wall more than to the luminal contents. High pressure on the intestinal wall causes the development of colonic diverticulosis at the weak points of wall.17 In 1985, a lifespan study using 1,800 Wistar rats showed a significant inverse association between the incidence of diverticulosis and the concentration of fiber in their diets. This study offers strong support for the hypothesis of Painter and Burkitt that human diverticular disease is caused by fiber deficiency, provided the association can be extrapolated from rats to humans.22

In contrast, some studies showed no relationship between fiber consumption and asymptomatic diverticulosis. A recent study of over 2,000 patients who underwent colonoscopy in the United States revealed that high intake of fiber did not reduce the prevalence of diverticulosis. Dietary information was gathered using questionnaire, a food frequency questionnaire with over 100 food items. This study found that dose-dependent higher fiber diets were actually associated with a higher prevalence of diverticulosis. In a Korean study of 848 subjects who underwent colonoscopy, the authors concluded that higher fiber diets were actually associated with a higher prevalence of diverticulosis. In a Korean study of 848 subjects who underwent colonoscopy, no correlation between a high-fiber diet and colonic diverticula was observed. Information about subjects’ dietary styles was obtained through a questionnaire using the mini dietary assessment (MDA) index. This index scores the frequencies of vegetable, fruit, and kimchi intakes. The mean fiber MDA score of subjects with diverticulosis was lower than that of subjects without diverticulosis.24

Table 5. Effect of Vegetarian Diet on Left-Sided Colonic Diverticulosis by Univariate and Multivariate Analysis

| Risk factor            | Control (n = 1,201) | Left-sided diverticulosis (n = 23) | Univariate analysis | Multivariate analysis* |
|------------------------|---------------------|-----------------------------------|---------------------|------------------------|
|                        | No. of person (%)   | OR (95% CI) P-value               | OR (95% CI) P-value |
| Current smoking        | 89 (7.4)            | 4 (17.4)                          | 2.63 (0.88–7.90)   | 0.091                  |
| Alcohol consumption    | 133 (11.1)          | 5 (21.7)                          | 2.23 (0.82–6.11)   | 0.170                  |
| Hypertension           | 84 (7.9)            | 21 (8.8)                          | 1.13 (0.69–1.87)   | 0.692                  |
| Metabolic syndrome    | 243 (20.2)          | 2 (8.7)                           | 0.38 (0.09–1.61)   | 0.289                  |
| Exercise (≥ 3 times/wk)| 271 (22.6)          | 5 (21.7)                          | 0.95 (0.35–2.59)   | 1.000                  |
| Age (yr)               |                     |                                   |                     |                        |
| ≤ 59                   | 931 (77.5)          | 11 (47.8)                         | 3.76 (1.64–8.62)   | 0.002                  |
| ≥ 60                   | 270 (22.5)          | 12 (52.2)                         | 5.74 (1.94–16.96)  | < 0.001                |
| Male sex               | 544 (45.3)          | 19 (82.6)                         | 0.92 (0.39–2.19)   | 0.855                  |
| BMI (≥ 25 kg/m²)       | 440 (36.6)          | 8 (34.8)                          | 3.77 (1.39–10.21)  | 0.006                  |
| Nonvegetarian diet     | 18 (3.0)            | 5 (21.7)                          | 3.90 (1.43–10.67)  | 0.008                  |

*Adjusted for age, male and vegetarian diet.
sis was not significantly different with that of those without. Reproducibility problems may result from the use of repeated 24-hour or quantitative dietary recalls; therefore, a study of confirmed vegetarians, such as Buddhist monks, might solve this problem.

In Buddhism, food is considered to be the only means of maintaining life; pleasure is excluded.23 Buddhist monks consume a vegetarian diet according to specific rules.24,25 In general, the Korean Buddhist diet avoid food of any animal origin except milk, several stinky vegetables including garlic and Welsh onion, alcohol, and excessive quantity of processed food.25,26 Unlike the previous studies that investigated vegetarians using interviews or questionnaires, our study yielded statistically significant results of the prevalence of diverticulosis in Buddhist monks who are strict vegetarians compared with that in the general population.

In this study, the prevalence of diverticulosis was age-dependent, increasing with increasing age. This indicated a dose-dependent relationship between age and the risk of colonic diverticulosis which was compatible with many previous studies.22,27-29 The prevalence of diverticulosis ranges from less than 10% in subjects younger than 40 years to an about 70% in those older than 80 years in Western countries.5,30-32 Although an equal distribution or a female preponderance was suggested in a Western study,1 a male preponderance was reported in previous Asian studies.13,20,33-35 The ratio of men to women in our study was 2.02 which is similar to that of previous studies. A future study should investigate the cause of this male preponderance.

The prevalence of obesity has grown considerably through the years, and this has been associated with several GI diseases, including diverticular disease. In a recent systematic review, it was suggested that even moderate increases in BMI may increase not only the risk of diverticular disease, but also diverticular disease complications.36 Notably, the BMI of Buddhist monks was higher than that of the general population; this was consistent with other previous studies of Korean Buddhist vegetarians.26,37-41 The reason for this is that Buddhist monks’ calorie- and carbohydrate intakes were significantly higher, and their preference for sweet or salty food was high. It may also be caused by a lack of exercise as less than 20% of Buddhist monks exercise 2 or more times a week.38,42 As shown in Table 1, although Buddhist monks had a higher prevalence of obesity and metabolic syndrome, which were risk factors for diverticulosis in several studies, than the general population, it could be speculated that a vegetarian diet had a strong protective effect against colonic diverticulosis.

This study had several strong points. Firstly, we enrolled Buddhist monks as a pure vegetarian group. They are obligatory vegetarians for the spiritual reason. This ensured an accurate assessment of the effect of vegetarian diet on diverticulosis. Secondly, the collected data had high quality because all the measurements were taken by skilled personnel with standardized protocol. Third, we analyzed the risk factors according to the location of the diverticula.

There were also several weak points in this study. Firstly, because it was a retrospective, cross-sectional study, the temporal association and causality between vegetarianism and asymptomatic diverticulosis could not be evaluated. Secondly, although a multivariate analysis was performed, no subject in the vegetarian group smoked or drank alcohol, and these were risk factors in several studies.11,44 This may have limited the ability to adjust for multiple variables. Thirdly, we did not evaluate medications, including NSAIDs, statins, or opiates, which were reported to be risk factors for diverticular disease45 due to missing data. However, no association between these medications and asymptomatic diverticulosis has been reported. Forth, we did not evaluate the association between dietary fiber intake and symptomatic colonic diverticulosis. Among Buddhist monks who underwent colonoscopy, only 1 person replied that he had history of diverticulitis. This may result from the fact that questionnaire included open question about past medical history, not specific question about diverticular disease.

In conclusion, old age, male sex, higher BMI, and a nonvegetarian diet were independently associated with colonic diverticulosis. Therefore, our study suggested that a vegetarian diet is inversely associated with asymptomatic colonic diverticulosis irrespective of the location of the diverticulosis in Asians.

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**CONFLICT OF INTEREST**

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

**AUTHOR CONTRIBUTION**

Conception and design: Kang HW. Data collection: Bong J. Cho
H. Nam JH, Jang DK. Writing: Bong J, Kang HW, Jang DK. Review and manuscript supervision: Kim JH, Lee JK, Lim YJ, Koh MS, Lee JH. Approval of final manuscript: all authors.

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SUPPLEMENTARY MATERIAL

Supplementary materials are available at the Intestinal Research website (https://www.irjournal.org).

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See “Vegetarianism as a protective factor for asymptomatic colonic diverticulosis in Asians: a retrospective cross-sectional and case-control study” on page 121-129.

Supplementary Material 1. Questionnaire

### Past Medical History

Please give details of any important illness, operation or medications:

| Current or Past Medical Problems | Dates of Diagnosis |
|----------------------------------|--------------------|
|                                  |                    |
|                                  |                    |
|                                  |                    |

| Surgeries | Dates | Reason |
|-----------|-------|--------|
|           |       |        |
|           |       |        |
|           |       |        |

| Medications | Dosage/Frequency | Reason |
|-------------|------------------|--------|
|             |                  |        |
|             |                  |        |
|             |                  |        |

### Life Style

- Check one of the following about smoking within recent 1 year
  - □ Never smoked  □ Former smoked  □ Smoke some days  □ Smoke every day
  - If you smoke or used to smoke, how many packs do/did you smoke per day?
  - How many years did you smoke/have you smoked?
  - If you quit, when did you quit?

- How often do you drink alcohol on average?
  - □ Never  □ < 4/month  □ 1-3/week  □ 4-6/week  □ daily
  - If you drink alcohol, please indicate the quantity per week of each
    - Soju glass (40cc) of Soju _____________
- Beer glasses (250cc) of beer _____________
- Wine glass (85cc) of wine _____________
- Shot (25cc) of hard liquor (such as whisky) _____________

• Do you take regular exercise?  ☐ no    ☐ yes
  If yes, what kind of exercise do you take? _____________

How often have you ate meat
  ☐ < 1/week,  ☐ 1-2/week,  ☐ 3-6/week,  ☐ daily

How long do you exercise on average when you take exercise? ________ minutes

Symptoms

• Abdominal pain  ☐ none    ☐ mild    ☐ moderate    ☐ severe
• Dyspepsia  ☐ none    ☐ mild    ☐ moderate    ☐ severe
• Nausea/vomiting  ☐ none    ☐ yes
• Diarrhea  ☐ none    ☐ yes (defecation /day)
• Constipation  ☐ none    ☐ yes (defecation /week)
• Jaundice  ☐ none    ☐ yes
• Black stool  ☐ none    ☐ yes
• Blood in stool  ☐ none    ☐ yes
• Difficulty in swallowing  ☐ none    ☐ yes

Diet

• How often have you eat red meat or poultry?
  ☐ Never (vegetarian),  ☐ ≤ 1/week,  ☐ 2-3/week,  ☐ ≥ 4/week