

Nut, Corn, and Popcorn Consumption and the Incidence of Diverticular Disease

Lisa L. Strate, MD, MPH; Yan L. Liu, MS; Sapna Syngal, MD, MPH; Walid H. Aldoori, MD, MPA, ScD; Edward L. Giovannucci, MD, ScD JAMA. 2008;300(8):907-914.

Context Patients with diverticular disease are frequently advised to avoid eating nuts, corn, popcorn, and seeds to reduce the risk of complications. However, there is little evidence to support this recommendation.

Objective To determine whether nut, corn, or popcorn consumption is associated with diverticulitis and diverticular bleeding.

Design and Setting The Health Professionals Follow-up Study is a cohort of US men followed up prospectively from 1986 to 2004 via self-administered questionnaires about medical (biennial) and dietary (every 4 years) information. Men reporting newly diagnosed diverticulosis or diverticulitis were mailed supplemental questionnaires.

Participants The study included 47,228 men aged 40 to 75 years who at baseline were free of diverticulosis or its complications, cancer, and inflammatory bowel disease and returned a food-frequency questionnaire.

Main Outcome Measures Incident diverticulitis and diverticular bleeding.

Results During 18 years of follow-up, there were 801 incident cases of diverticulitis and 383 incident cases of diverticular bleeding. We found inverse associations between nut and popcorn consumption and the risk of diverticulitis. The multivariate hazard ratios for men with the highest intake of each food (at least twice per week) compared with men with the lowest intake (less than once per month) were 0.80 (95% confidence interval, 0.63-1.01; P for trend = .04) for nuts and 0.72 (95% confidence interval, 0.56-0.92; P for trend = .007) for popcorn. No associations were seen between corn consumption and diverticulitis or between nut, corn, or popcorn consumption and diverticular bleeding or uncomplicated diverticulosis.

Conclusions In this large, prospective study of men without known diverticular disease, nut, corn, and popcorn consumption did not increase the risk of diverticulosis or diverticular complications. The recommendation to avoid these foods to prevent diverticular complications should be reconsidered.

Diverticular disease is a common and costly digestive disorder in Western countries. One-third of the US population will develop diverticulosis by the age of 60 years and two-thirds by the age of 85 years.¹⁻³ Complications, including diverticulitis and diverticular bleeding, occur in an estimated 10% to 35% of persons with diverticulosis.⁴⁻⁵ Treatment of these complications frequently necessitates hospitalization and invasive procedures, including surgery. At least \$2.4 billion in direct health care costs and 3400 deaths are attributed to diverticular disease in the United States each year,⁶⁻⁷ and the medical and economic impact of this disorder is likely to increase substantially as the population ages.

Historically, physicians have advised individuals with diverticular disease to avoid nuts, seeds, popcorn, corn, and other high-residue foods.^{5, 8-9} This recommendation stems from the theory that luminal trauma is a causal mechanism for both diverticulitis and diverticular bleeding.¹⁰⁻¹² Stool may lodge within a diverticulum, obstruct the neck, or abrade the mucosa and precipitate inflammation or bleeding.^{1, 13} Nuts, seeds, popcorn, and corn are presumed to be particularly likely to abrade the mucosa or to lodge within small diverticula.⁵ However, the biological mechanisms responsible for diverticular complications remain poorly understood. Aside from luminal trauma, potential inciting factors include elevated colonic pressures, compromised colon wall integrity, and altered bacterial flora.¹⁴⁻²⁰ To our knowledge, there is no evidence to support consumption of nuts, corn, popcorn, or seeds as a risk factor.

Compelling evidence suggests that nuts are an important part of a healthy diet. Nuts are high in protein, unsaturated fats, fiber, vitamins, minerals, and other micronutrients. A diet high in nuts may protect against many common diseases, including cardiovascular disease, diabetes, cancer of the colon and prostate, and cholelithiasis.²¹⁻³²

The potential health benefits of nut consumption paired with the large number of individuals with diverticulosis makes it timely and important to study the long-held belief that eating nuts and other high-residue foods is associated with diverticular complications. We examined the association between nut, corn, and popcorn consumption and complicated diverticular disease in a large prospective cohort.

Assessment of Nut, Corn, and Popcorn Consumption

Dietary information was ascertained from a semiquantitative food-frequency questionnaire that was validated in this cohort.³³ Participants were asked to indicate how frequently on average they consumed a standard portion of 131 specified food items during the previous year according to 9 response categories (never or less than once a month, 1-3 times per month, once per week, 2-4 times per week, 5-6 times per week, once a day, 2-3 times daily, 4-5 times daily, more than 6 times a day). Nut consumption was ascertained via questions regarding the average frequency of consumption of a 1-ounce serving of peanuts, walnuts, or other nuts. Walnuts were added as a distinct category of nuts in 1998. The food questionnaire also separately assessed consumption of fat-free or light popcorn and regular popcorn (1 standard unit was equivalent to 3 cups). Corn consumption was documented in a single question (a typical serving was equal to 1 ear or one-half cup of kernels). Simple updating was used to assess dietary information such that the food questionnaire immediately preceding the follow-up interval of interest was used in the analysis. The mean time interval between dietary assessment and study end point was 28.4 months. Total nut and popcorn consumption were calculated as the sum of the nut and popcorn items, respectively. Frequency categories for total nut, corn, and popcorn consumption were collapsed into 4 categories: less than once per month, 1 to 3 times per month, once per week, and 2 or more times per week. Categories were created for missing nut, corn, and popcorn data. Data on consumption was missing at baseline for 412 participants (0.9%) for nuts, 767 participants (1.6%) for corn, and 758 participants (1.6%) for popcorn.

Assessment of Other Potential Risk Factors

We also assessed a number of dietary, lifestyle, and medical factors that have been implicated as risk factors for complicated or symptomatic diverticular disease in this cohort and/or in other studies.^{20, 34-40} These included red meat and total dietary fiber and fat intake, which were assessed on the semiquantitative food-frequency questionnaires. Dietary fat and fiber amounts were calculated by multiplying the consumption frequency of a specified serving size by the corresponding nutrient content derived from food composition tables. Total fat and fiber consumption were adjusted for total energy intake using regression analysis.

We also considered physical activity as a potential risk factor. On biennial questionnaires, men were asked to indicate the average time spent each week engaged in any of 10 moderate to vigorous activities in addition to the number of flights of stairs climbed each day. Physical activity was summed and expressed in metabolic equivalent (MET) hours per week. (One MET is equivalent to 1 kcal of energy expended per kilogram of body weight per hour while at rest.) Current weight as reported on biennial questionnaires and height reported at baseline were used to calculate body mass index (weight in kilograms divided by height in meters squared).

Current, regular use of nonsteroidal anti-inflammatory drugs including aspirin and acetaminophen was also assessed and updated every 2 years, as was cigarette smoking.

Statistical Analysis

Person-years of follow-up accrued from the date of return of the 1986 baseline questionnaire to the month of the first diagnosis of diverticular disease (diverticulosis or a complication), death, or the end of follow-up (December 31, 2004). Men who reported a new diagnosis of diverticulitis, diverticular bleeding, uncomplicated diverticulosis, gastrointestinal cancer, or inflammatory bowel disease were censored at the date of diagnosis.

We identified 801 incident cases of diverticulitis and 383 incident cases of diverticular bleeding during 730 446 person-years of follow-up. Baseline characteristics of the cohort categorized by nut, corn, and popcorn consumption and standardized for age are shown in Table 1. Twenty-seven percent of participants reported eating nuts at least twice per week. Corn and popcorn each were consumed at least twice a week in 15%. Consumption of less than 1 serving per month of nuts, corn, and popcorn was noted in 25%, 12%, and 36% of men, respectively. Participants in the highest category of nut consumption tended to be older and to consume more alcohol but were slightly more physically active than those in the lowest category. Compared with men with the lowest intake of corn, men with the highest intake generally consumed more fiber and red meat but less alcohol. Men who ate popcorn at least twice per week were, in general, younger, more likely to use analgesics, and more likely to be nonsmokers than those who ate popcorn less than once per month.

Nut, corn, and popcorn consumption was not associated with an increased risk of complicated diverticular disease. Instead, we observed inverse relationships between nut and popcorn consumption and the risk of diverticulitis (Table 2). After adjustment for other known and potential risk factors for diverticular complications, the multivariate HRs for men with the highest intake of each food (at least twice per week) compared with men with the lowest intake (less than once per month) were 0.80 (95% CI, 0.63-1.01; P for trend = .04) for nuts and 0.72 (95% CI, 0.56-0.92; P for trend = .007) for popcorn. The population-attributable risks of developing incident diverticulitis associated with low consumption (less than 2 times per week compared with more than 2 times per week) were 6005 per 100 000 men for nuts and 6558 per 100 000 men for popcorn. Although the P for trends for the associations between increasing consumption of nuts and popcorn and the risk of diverticulitis were significant, the patterns of the HRs across consumption categories did not indicate dose-response relationships (Table 2). No association was seen between corn consumption and diverticulitis. For diverticular bleeding, there were no significant associations observed for nut, corn, or popcorn consumption (Table 3). Compared with men with the lowest intake, the multivariate HRs for men with the highest intake were 1.08 for nuts (95% CI, 0.77-1.49; P for trend = .89), 1.07 for corn (95% CI, 0.67-1.71; P for trend = .97), and 0.82 for popcorn (95% CI, 0.59-1.15; P for trend = .63).

To account for the possibility that symptoms of undiagnosed diverticular disease may have led to a reduction in nut, corn, or popcorn consumption, we performed a 2-year time lag analysis with respect to ascertainment of diet and diagnosis of diverticular complications (eg, data from the 1986 food-frequency questionnaire were used for the 1990-1992 disease follow-up). The inverse association between frequent nut consumption and the risk of diverticulitis was strengthened (multivariate HR, 0.76; 95% CI, 0.61-0.96; P for trend = .03 in a high/low comparison), and the association for popcorn was attenuated (multivariate HR, 0.89; 95% CI, 0.70-1.13; P for trend = .21). No significant associations were observed between nut, corn, or popcorn consumption and diverticular bleeding in the time lag analysis.

In addition, we performed several analyses to explore the possibility that nut, corn, and popcorn consumption influences the development of diverticulosis. We examined the relationship between consumption of these foods and uncomplicated diverticulosis (restricting the analysis to men who had undergone prior lower endoscopy to minimize detection bias) and found no significant associations. The multivariate HR for nuts was 0.91 (95% CI, 0.78-1.07; P for trend = .26), for corn was 0.98 (95% CI, 0.78-1.23; P for trend = .83), and for popcorn was 0.93 (95% CI, 0.78-1.10; P for trend = .38). To better isolate the independent effect of very recent diet from the long-term effect of past diet, we added baseline consumption of nuts, corn, and popcorn to the multivariate models. The associations between recent intake of these foods and diverticulitis and diverticular bleeding remained unchanged. No significant associations were seen for baseline consumption of nuts, corn, or popcorn and diverticulitis or diverticular bleeding. For diverticulitis, the multivariate HRs for baseline consumption were 0.95 for nuts (95% CI, 0.77-1.16; P for trend = .52), 1.02 for corn (95% CI, 0.75-1.38; P for trend = .44), and 0.92 for popcorn (95% CI, 0.73-1.15; P for trend = .40) in high to low comparisons. The multivariate HRs for baseline consumption and diverticular bleeding were 0.85 for nuts (95% CI, 0.61-1.18; P for trend = .18), 0.97 for corn (95% CI, 0.62-1.52; P for trend = .64), and 1.30 for popcorn (95% CI, 0.89-1.92; P for trend = .73).

Finally, although we were unable to assess total seed intake, we did examine the relationship between combined strawberry and blueberry consumption and diverticular complications and found no significant association. (The small seeds found in berries have been implicated in diverticular complications.) The multivariate HRs for consumption of berries at least twice per week compared with less than once a month were 0.87 for diverticulitis (95% CI, 0.65-1.16; P for trend = .37) and 0.86 for diverticular bleeding (95% CI, 0.57-1.30; P for trend = .54).

Data from this large prospective cohort of men followed up for 18 years indicate that frequent nut, corn, and popcorn consumption is not associated with an increased risk of diverticular complications. Indeed, nut and popcorn consumption appears to be inversely associated with the risk of diverticulitis independent of other known or potential risk factors, including age; body mass index; dietary fat, fiber, and red meat consumption; physical activity; cigarette smoking; and the use of nonsteroidal anti-inflammatory drugs and acetaminophen. No associations were found between nut, corn, or popcorn consumption and diverticular bleeding.

To our knowledge, no prior study has examined the association between nut, corn, or popcorn consumption and diverticular complications. Nonetheless, in a recent survey of colorectal surgeons, 47% felt that patients with diverticular disease should avoid these foods.⁸ The commonly held belief that these foods can incite complications of diverticulosis and the recommendation that individuals with this condition avoid them apparently evolved from the theory that luminal trauma is a causal mechanism.⁵ In this model, inflammation or bleeding is precipitated when a fecalith or particulate fecal material lodges within a diverticulum and erodes the mucosa.^{1, 13} Foods with poorly digested particles, such as nuts, corn, and seeds, are presumed to be particularly abrasive or apt to lodge within diverticula. Although fecal matter is commonly found within wide-necked diverticula, the relationship between the ingestion of a particular food and subsequent trauma to a diverticulum is largely speculative, and the exact mechanisms leading to diverticular complications are not known.

The differences in our results with respect to diverticulitis and diverticular bleeding may be explained by potentially distinct mechanisms underlying these complications. A histological study of bleeding colonic diverticula found an absence of mucosal inflammation or evidence of diverticulitis at the bleeding sites.⁴²⁻⁴³ Abnormalities were confined to the vasa rectum and included

eccentric intimal thickening near the site of bleeding and asymmetric rupture toward the lumen, findings consistent with a nonspecific vascular response to injury.

There are several plausible mechanisms by which nut intake may protect against diverticulitis, although our data did not enable us to make causal inferences. Nuts are rich in nutrients with anti-inflammatory properties such as vitamin E, -linolenic acid and other unsaturated fatty acids, and phytochemicals. Nut consumption is inversely correlated with levels of inflammatory markers including C-reactive protein and IL-6^{22, 27-28} and is protective against certain inflammatory disease states. The high mineral content in nuts, including zinc and magnesium, may also favorably influence the colon. Alterations in the structure and content of collagen in the colon have been noted in diverticular disease, which may in part be due to decreased expression of zinc-containing endopeptidases.⁴⁴⁻⁴⁵ Magnesium may reduce oxidative stress in the colon, a mechanism that is also postulated for the inverse association seen between magnesium intake and colon cancer.⁴⁶⁻⁴⁷ Popcorn is less nutrient dense than nuts but does contain lutein, a micronutrient with anti-inflammatory and chemoprotective properties that is also found in nuts, as well as modest amounts of magnesium.

The high fiber content of nuts and popcorn does not appear to provide adequate explanation for the observed inverse associations between nuts and popcorn and diverticulitis. The multivariate analyses adjusted for total dietary fiber intake, and the consistency between the age-adjusted and multivariate models does not imply significant confounding by fiber or the other covariates. Nor do nuts and popcorn appear to be markers for generally healthy habits because the data on other lifestyle and dietary factors such as physical activity and red meat consumption do not suggest a health-conscious pattern in men with frequent nut and popcorn consumption. Notably, the majority of our data were acquired during a time period when nuts were not perceived as a healthy food. Nonetheless, given the observational nature of the study, we cannot exclude the possibility of residual confounding by a lifestyle or other unmeasured factor.

We were unable to specifically study the effect of nut, corn, and popcorn consumption in men with established diverticulosis because the presence of diverticulosis was not known at baseline in all individuals. Men with known asymptomatic diverticulosis likely represent a biased subset of men who have undergone lower endoscopy or colon imaging, and men with known diverticulosis are likely to alter their behaviors, including decreasing the consumption of nuts, corn, and popcorn. However, the prevalence of undiagnosed asymptomatic diverticulosis was presumably quite high in this cohort of middle-aged and elderly men, and therefore, our findings likely apply to patients with established diverticular disease.

A long-term influence of nut and popcorn consumption on the development of diverticulosis does not appear to account for the inverse associations with diverticulitis observed in this study. We did not find significant associations between nut and popcorn consumption and incident uncomplicated diverticulosis. Furthermore, our results remained unchanged after adjustment for baseline consumption. The later analysis aims to distinguish the independent effect of very recent diet (relevant to the risk of diverticular complications) from past diet (relevant to the development of diverticulosis).

In the 2-year time lag analysis, the inverse relationship between nuts and diverticulitis was strengthened, but the results for popcorn were attenuated. Some men may have reduced their consumption of popcorn in response to symptoms of undiagnosed diverticulitis under the assumption that popcorn is harmful. However, if recent popcorn consumption is protective, their subsequent risk of diverticulitis would increase. This would enhance the inverse relationship seen between popcorn consumption and diverticulitis in the standard analysis and attenuate it in

the time lag analysis. Overall, there was no evidence from the time lag analysis that consumption of nuts, popcorn, or corn was positively associated with diverticular complications. Lastly, we had not hypothesized that these foods would decrease the incidence of diverticular complications, and it is possible that this finding is the result of chance alone.

Our study has certain limitations. As noted earlier, it is possible that the associations found between nuts and popcorn were the result of residual confounding. In addition, diverticular complications and dietary intake were based on self-report. However, several factors limit the possibility of misclassification bias. Study participants were health professionals, review of 179 medical records endorsed the validity of self-report, and secondary analyses supported our findings. Food consumption was assessed prior to the development of diverticular complications using a validated food-frequency questionnaire. On the whole, any misclassification bias would likely be random and thus would weaken any true association. The generalizability of our results to younger patients and women may be limited. However, diverticulosis is rare before the age of 40 years,¹ and there are no clear sex differences in diverticular disease.^{1, 48} Lastly, we were unable to directly examine the relationship between seed consumption and diverticular complications. However, consumption of strawberries and blueberries (fruits with small seeds) was not associated with diverticular complications.

In conclusion, our results suggest that nut, corn, and popcorn consumption is not associated with an increased risk of incident diverticulitis or diverticular bleeding and may be protective for the former. These findings refute the pervasive but unproven belief that these foods are associated with diverticular complications and suggest that the recommendation to avoid these foods in diverticular disease should be reconsidered.