Dietary habits of colorectal neoplasia patients in comparison to their first-degree relatives

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Abstract

AIM: To compare the dietary habits between colorectal neoplasia patients, their first-degree relatives, and unrelated controls.

METHODS: From July 2008 to April 2011, we collected epidemiological data relevant to colorectal cancer from patients with colorectal neoplasias, their first-degree relatives, and also from a control group consisting of people referred for colonoscopy with a negative family history of colorectal cancer and without evidence of neoplasia after colonoscopic examination. The first-degree relatives were divided into two groups following the colonoscopic examination: (1) patients with neoplasia or (2) patients without neoplasia. Dietary habits of all groups were compared. A χ² test was used to assess the association between two dichotomous categorical variables.

RESULTS: The study groups consisted of 242 patients with colorectal neoplasias (143 men, 99 women; mean age: 64 ± 12 years) and 160 first-degree relatives (66 men, 94 women; mean age: 48 ± 11 years). Fifty-five of the first-degree relatives were found to have a neoplastic lesion upon colonoscopy, while the remaining 105 were without neoplasia. The control group contained 123 individuals with a negative family history for neoplastic lesions (66 men, 57 women; mean age: 54 ± 12 years). Two hypotheses were tested. In the first, the dietary habits of first-degree relatives with neoplasia were more similar to those of patients with neoplasia, while the dietary habits of first-degree relatives without neoplasia were similar to those of the control group. In the second, no sex-related differences in dietary habits were expected between the particular groups. Indeed, no significant differences were observed in the dietary habits between the groups of patients, controls and first-degree relatives with/without neoplastic lesions. Nevertheless, statistically significant sex-related differences were observed in all groups, wherein women had healthier dietary habits than men.

CONCLUSION: In all groups examined, women had healthier dietary habits than men. Modification of screening guidelines according to sex may improve the efficiency of screening programs.

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Key words: Colorectal neoplasms; Family; Food habits; Risk factors; Mass screening

Core tip: We compared the dietary habits of patients with neoplasia (patients and their first-degree relatives with neoplasia) and without neoplasia (first-degree relatives without neoplasia and an unrelated control group). We did not identify significant differences in dietary habits between the groups; however, we did identify statistically significant differences between the...
dietary habits of men and women in all groups. In all
groups, women had healthier dietary habits. Modification
of screening guidelines according to sex may im-
prove the efficiency of screening programs, although
further studies are needed to support this hypothesis.

INTRODUCTION

Colorectal cancer is the second leading cause of cancer-
related death in developed countries. The Czech Republic
has the highest prevalence of colorectal cancer in the
world. In 2008, the incidence of colorectal cancer in the
Czech Republic was 94.2/100000 men and 61.8/100000
women[1]. It is well established that colonoscopic screen-
ing reduces both the occurrence and mortality of colo-
rectal cancer[2]. In 2000, the Czech Republic intro-
duced a nationwide cancer-screening program that in-
cluded fecal occult blood testing of people over 50 years
of age. The program was then updated in 2009 to include
the possibility of a primary colonoscopy screening for
those over 55 years of age[3,4].

Colorectal neoplasias (CRN) are associated with non-
ereditary as well as hereditary risks. Colorectal cancer
is the most common familial form of cancer. More than
30% of cases can be attributed to hereditary causes, of
which only 5% are due to hereditary cancer syndromes
such as familial adenomatous polyposis syndrome and
hereditary non-polyposis colorectal cancer[5]. First-degree
relatives (FDR) of patients with CRN (either colorectal
cancer or advanced adenomas) show up to a 4-fold in-
creased risk for CRN when compared with the general
population and are at increased risk for advanced or mul-
tiple adenomas[6-9].

Non-hereditary risk factors for colon cancer in-
clude advanced age, male sex, alcohol consumption and
smoking[10-12]. Dietary factors, such as elevated red meat
consumption and low intake of fruit, vegetables, dairy
products and dietary fiber, have been associated with an
increased risk for CRN[13]. Obesity, sedentary lifestyle,
inflammatory bowel diseases and several other conditions
such as acromegaly, diabetes mellitus and ischemic heart
disease have also been shown to increase risk for colon
cancer[14-17].

The goal of this study was to compare the dietary
habits of patients with CRN and a control group with
the dietary habits of FDR with regard to the findings
obtained after a colonoscopy screening. The first tested
hypothesis was that dietary habits of FDR with neoplasia
are similar to those of patients with CRN and that the
dietary habits of FDR without neoplasia are similar to
those of the control group. The second tested hypothesis
was that there are no sex-related differences of dietary
habits between the particular groups.

MATERIALS AND METHODS

Study subjects and clinical data

From July 2008 to April 2011, we collected epidemiolo-
gical data relevant to colorectal cancer, both from patients
with CRN and their FDR as well as from a control group.
Epidemiological data, including smoking status (current/
former/never), fat intake (low vs high), body mass index
(BMI; < 30 vs ≥ 30 kg/m²), beer consumption (daily/
occasionally/never), consumption of dairy products,
fruits, vegetables and red meat (daily/less/frequent) and
education attainment (primary/secondary/tertiary), were collected from the patients with CRN, FDR and
controls by a medical doctor. A single specialist in
gastroenterology and nutrition performed the interview
about the respondent’s dietary habits (the amounts of red
meat, fat, dairy products, etc.) and made a categorization
according to the answers (high intake/low intake in each
category). Collection of epidemiological data was part of
The Family Project, a unique direct medical counseling
project targeting FDR that took place at a single center
(non-university), Hospital Frydek-Mistek. The goals of
the project were to promote proper colonoscopic surve-
illance of FDR and to identify FDR at highest risk for
CRN. The project was approved by the local ethics
committee. All participants signed an informed consent.
Simultaneously, an informative campaign was launched in
the local media to promote and support public awareness
of the project.

FDR were referred to colonoscopic examinations
and, dependent on the findings, were divided into FDR
with or FDR without neoplasia. The control group con-
tained people with a negative family history that had been
referred for colonoscopy and were confirmed to be with-
out neoplasia according to the findings from the colono-
scopic examination.

Statistical analysis

Ages are presented as mean ± SD. The dietary habits of
all groups (patients with CRN, FDR with neoplasia, FDR
without neoplasia, and control group) were compared. A
χ² or Fisher’s exact test was used to assess the association
between two dichotomous categorical variables. Because
of a heterogeneous representation of men and women in
the FDR without neoplasia group, the men and women in
all groups were compared separately.

RESULTS

The study groups consisted of 242 patients with CRN
(143 men, 99 women; 64 ± 12 years) and 160 FDR (66
men, 94 women; 48 ± 11 years). Fifty-five patients in the
FDR group were found to have neoplastic lesions upon
colonoscopy, while 105 patients had no evidence of neo-

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plasia. The control group consisted of 123 individuals with a negative family history of colon cancer and without neoplastic lesion following colonoscopic examination (66 men, 57 women; 54 ± 12 years). Characteristics of all groups are presented in Table 1.

We first tested the hypothesis that dietary habits of FDR with neoplasia are similar to those of patients with CRN and that dietary habits of FDR without neoplasia are similar to those of the control group. We next tested the hypothesis that there are no sex-related differences in the dietary habits between the particular groups. Comparisons of the groups are presented in Tables 2 and 3. The comparison between men and women in all groups is shown in Table 4.

In summary, both of our hypotheses were disproven. There were no significant differences in the dietary habits between the groups of patients, controls and FDR with/without neoplasia. In all groups, however, there were statistically significant differences in the dietary habits between men and women, despite no differences in education attainment among them.

**DISCUSSION**

Our study was based on epidemiological data relevant to colorectal cancer that was obtained from patients with CRN, their FDR with neoplasia, FDR without neoplasia, and from a control group.

| Characteristics | Patients | FDR with neoplasia | FDR without neoplasia | Controls | P value (χ²) |
|-----------------|----------|--------------------|-----------------------|----------|-------------|
| Male sex        | 143/242 (59) | 30/55 (56)       | 36/105 (34)        | 66/123 (54) | 0.001 |
| Obesity         | 68/242 (28)   | 15/55 (27)       | 23/105 (22)        | 27/123 (22) | 0.478 |
| Smoking, current/former | 123/242 (51)   | 28/55 (51)       | 32/105 (30)        | 48/123 (39) | 0.006 |
| High fat intake | 102/242 (42)   | 28/55 (51)       | 35/105 (33)        | 52/123 (42) | 0.175 |
| High red meat consumption | 171/242 (71)  | 37/55 (67)       | 65/105 (62)        | 62/123 (50) | 0.002 |
| Beer consumption | 155/242 (64)  | 35/55 (64)       | 54/105 (51)        | 83/123 (67) | 0.070 |
| Low intake of dairy products | 81/242 (33)    | 22/55 (40)       | 27/105 (26)        | 45/123 (37) | 0.219 |
| Low fruit and vegetable consumption | 72/242 (30)  | 14/55 (25)       | 25/105 (24)        | 46/123 (37) | 0.128 |
| Primary education attainment | 134/242 (55) | 15/55 (27)       | 27/105 (26)        | 47/123 (38) | 0.001 |

FDR: First-degree relatives.

| Comparison | Male patients vs FDR with neoplasia | Male patients vs FDR without neoplasia | Female patients vs FDR with neoplasia | Female patients vs FDR without neoplasia |
|------------|-------------------------------------|----------------------------------------|--------------------------------------|----------------------------------------|
| Obesity    | 0.274                               | 0.101                                  | 0.207                                | 0.642                                  |
| Smoking    | 0.975                               | 0.001                                  | 0.727                                | 0.645                                  |
| High fat intake | 0.247                                | 0.451                                  | 0.456                                | 0.460                                  |
| High red meat consumption | 0.621                                | 0.956                                  | 0.474                                | 0.474                                  |
| Beer consumption | 0.674                                | 0.558                                  | 0.316                                | 0.316                                  |
| Low intake of dairy products | 0.932                                | 0.143                                  | 0.328                                | 0.328                                  |
| Low fruit and vegetable consumption | 0.353                                | 1.000                                  | 0.707                                | 0.707                                  |
| Education attainment | 0.002                               | 0.260                                  | 0.001                                | 0.001                                  |

1Higher education attainment in first-degree relatives (FDR).

| Comparison | Male controls vs FDR with neoplasia | Male controls vs FDR without neoplasia | Female controls vs FDR with neoplasia | Female controls vs FDR without neoplasia |
|------------|-------------------------------------|----------------------------------------|--------------------------------------|----------------------------------------|
| Obesity    | 0.816                               | 0.833                                  | 0.379                                | 0.959                                  |
| Smoking    | 0.281                               | 0.078                                  | 0.289                                | 0.578                                  |
| High fat intake | 0.281                                | 0.090                                  | 0.375                                | 0.685                                  |
| High red meat consumption | 0.284                                | 0.187                                  | 0.052                                | 0.041¹                                |
| Beer consumption | 0.045¹                              | 0.749                                  | 0.456                                | 0.535                                  |
| Low intake of dairy products | 0.618                                | 0.215                                  | 0.315                                | 0.315                                  |
| Low fruit and vegetable consumption | 0.006¹                              | 1.000                                  | 0.794                                | 0.794                                  |
| Education attainment | 0.095                               | 0.444                                  | 0.199                                | 0.199                                  |

¹Male controls have higher beer consumption and lower consumption of fruits and vegetables; ²Female controls have higher red meat consumption. FDR: First-degree relatives.
It is well established that risks for colorectal cancer can be either hereditary or non-hereditary. Non-hereditary risks are well described, as mentioned in the Introduction. There is also an association of colorectal cancer with the gut microbiome. Intestinal microbiota can transform food compounds into genotoxic agents, activate proto-oncogenes, or inactivate tumor suppressor genes\(^\text{[18-20]}\).

Genetic factors associated with an increased risk for CRN include low-penetrant susceptibility loci and specific polymorphisms. Certain genetic variants and polymorphisms in a number of genes have been associated with increased colon cancer risk; APC-I1307K, HRAS1-VNTR and MTHFR variants represent the strongest candidates for low penetrance susceptibility alleles\(^\text{[21,22]}\). In genome-wide association studies, as many as 170 common but separate genetic variations have been implicated in CRN susceptibility\(^\text{[23]}\). Based on current data, there are three main pathways of colorectal carcinogenesis: chromosomal instability, microsatellite instability, and hypermethylation\(^\text{[31,32]}\). One important question, however, is how hereditary risks may be confounded by familial similarities in diet, physical activity level, or other environmental exposures.

Our first tested hypothesis was that the dietary habits of FDR with neoplasia are similar to those of CRN patients, while the dietary habits of FDR without neoplasia are different and more similar to those of the control group. We hypothesized that both the controls and FDR without neoplasia have a healthier lifestyle, while patients with CRN and FDR with neoplasia have worse, shared dietary habits. Because of the heterogeneous representation of men and women FDR without neoplasia, men and women in all groups were compared separately.

To our surprise, all groups had very similar dietary habits. We only observed a difference in the male CRN patients, where there were significantly more smokers than in the group of FDR males without neoplasia. It has been shown that smoking can increase risk of colorectal cancer by up to 18%\(^\text{[21]}\). Paradoxically, male controls consumed more beer and lower amounts of fruits and vegetables than FDR males with neoplasia. Female controls consumed more red meat than FDR females without neoplasia. It is surprising that we did not observe any association between poor dietary habits and occurrence of neoplasia in patients with CRN and their FDR with neoplasia, despite all the proven non-hereditary risk factors.

The second tested hypothesis was that there would be no sex-related differences between the particular groups. Regardless of the colonoscopic findings in all groups, however, males had worse dietary habits than females, despite no difference in education attainment between the men and women. It is well known that women gain more health resources in their screening programs. This fact, together with a known higher incidence of CRN in men, places men at a disadvantage. Thus, we can assume that the one-third higher incidence of colorectal cancer in men could be, in part, attributed to their less healthy lifestyle. Media campaigns should, therefore, be targeted to the male population, since there is a great need for improvement of their lifestyle and dietary habits.

This study has several limitations. The sample size of each group was relatively small and made up of individuals stemming from a population with the highest prevalence of colorectal cancer in the world. The results, therefore, are specific and may only apply to the Czech population surveyed. Diabetes mellitus was not observed throughout all groups (only in the CRN group of patients), so we cannot evaluate obesity and dietary habits with respect to diabetes mellitus. The mean ages across the groups examined were different and represent another weakness of the study.

In conclusion, we did not find significant differences between patients and their FDR with/without neoplastic lesions, although we did identify statistically significant differences between the habits of men and women in all groups. Women in all groups had healthier dietary habits. We propose that media campaigns should be targeted to the male population, due to a need to improve their lifestyle. Modification of screening guidelines according to sex may improve the efficiency of screening programs but further studies are needed to support this hypothesis.

### COMMENTS

**Background**

Colorectal neoplasias are associated with hereditary and non-hereditary risks. Colorectal cancer is the most common familial form of cancer. First-degree relatives of patients with colorectal neoplasia, both colorectal cancer and advanced
adennomas, show up to a 4-fold increased risk for colorectal neoplasias when compared to the general population.

Research frontiers
It is important to understand how hereditary risks may be confounded by familial similarities in diet, physical activity level or other environmental exposures and whether it is possible to modify screening programs according to different risk groups to achieve higher efficiency in reduction of colorectal neoplasia.

Innovations and breakthroughs
The authors did not find significant differences between healthy controls, patients and their first-degree relatives with/without neoplastic lesions. The authors identified statistically significant differences between the dietary habits of men and women in all groups. In all groups examined, women had healthier dietary habits.

Applications
The authors propose a media campaign to target the male population and promote ways to improve the health-related aspects of their lifestyle. Modification of screening guidelines according to sex may improve the efficiency of screening programs, but further studies are needed to support this hypothesis.

Terminology
First-degree relatives: a family member who shares approximately 50% of their genes with a particular individual in a family; first-degree relatives include parents, offspring and siblings.

Peer review
This is an important epidemiological study comparing the dietary habits of persons with and without colorectal neoplasia. This is a well-designed study and has clinical applications for understanding the risks of colorectal cancer.

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