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Self-perceived Food Intolerances are Common and Associated with Clinical Severity in Childhood Irritable Bowel Syndrome

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Abstract

Background—Adults with irritable bowel syndrome (IBS) frequently identify foods as exacerbating their gastrointestinal (GI) symptoms. In children with IBS the prevalence of perceived food intolerances and their impact are unknown.

Objectives—To determine the prevalence of self-perceived food intolerances and the relationship of these intolerances to abdominal pain, psychosocial distress, and quality of life in children with IBS.

Design—Cross-sectional; questionnaire and prospective diary data were collected from 2008–2014 by trained research coordinators.

Participants/Setting—Children 7–18 years (pediatric Rome III IBS, n=154; age-sex matched healthy children (HC), n=32) in Houston, Texas.

Measures—Perceived food intolerances and avoided foods were captured using the Childhood Food and Symptom Association Questionnaire. IBS severity was assessed by a 7-day Pain Diary and validated psychosocial questionnaires assessing quality of life, somatization, functional disability, depression, and anxiety.

Statistical Analyses Performed—Descriptive, Spearman bivariate correlation, chi-square, Poisson log-linear generalized model with Wald chi-square statistics.

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CONFLICT OF INTEREST DISCLOSURE

Bruno Chumpitazi receives funding from the National Institutes of Health (NIH), QOL Medical Inc. and is a consultant for Mead Johnson Nutrition. Robert Shulman receives funding from the NIH, and Mead-Johnson Inc. and is a consultant for Nutrinia Inc. and Gerson Lehrman Group. The remaining authors do not conflicts of interest to disclose.

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Results—A greater proportion of children with IBS (143/154, 92.9%) vs HC (20/32, 62.5%) identified at least one self-perceived food intolerance ($\chi^2=22.5$, $P<0.001$). Children with IBS identified a greater number (median [25=75%]=4[2–6]) of perceived symptom-inducing foods than HC (median=2[0–4]; $\chi^2=28.6$, $P<0.001$). Children with IBS avoided more foods (median=2[1–4]) than HC (median=0 [0–2.75]; $\chi^2=20.8$, $P<0.001$). The number of self-perceived food intolerances was weakly associated (r-value range of –0.17 to 0.21) with pain frequency, pain severity, somatization, anxiety, functional disability, and decreased quality of life.

Conclusions—Children with IBS have a high prevalence of self-perceived food intolerances; the number of these intolerances is weakly associated with measures of IBS severity.

Keywords

Pediatric; Children; Diet; IBS; FODMAP

INTRODUCTION

Irritable bowel syndrome (IBS) is a prevalent functional gastrointestinal (GI) disorder affecting up to 20% of school aged children.¹ Factors which play a role in the clinical phenotype expressed include: visceral hyperalgesia, the gut microbiome, psychosocial distress (e.g., anxiety), and diet.^{2–4} Up to 84% of adults with IBS identify foods within their diet that exacerbate GI symptoms.^{5, 6} This is higher than the self-perceived food intolerance prevalence of up to 34% in the general population.^{7, 8}

The number of perceived food intolerances in adults with IBS is associated with more severe GI symptoms.^{6, 9} Self-reported food intolerances in adults also are associated with psychosocial distress such as somatization, anxiety, and decreased quality of life, though these associations have not been consistently identified.^{6, 10} Sex may also play a role, as women with IBS are more likely to identify food intolerances as compared to men.^{6, 10}

In children with IBS there is a dearth of information regarding self-perceived food intolerances and their clinical impact. A small mixed methods study in children with several functional GI disorders found that self-perceived food intolerances qualitatively negatively impacted several areas of quality of life.¹¹ Another study in Finnish 10 and 11-year-olds with functional GI disorders found that a majority of parents identified their child as having a food-induced GI symptom.¹² Neither study evaluated the relationship of intolerances to child-reported psychosocial distress and/or GI symptoms. Given the possibility that self-perceived food intolerances may have a large influence on the clinical severity of childhood IBS, further elucidating the prevalence and role of self-perceived food intolerances is clinically important.

Given this unexplored but clinically important area, the primary objective of this study was to determine the prevalence of self-perceived food intolerances in children with IBS. In addition, the study secondarily sought to determine whether there was a relationship between the number of self-perceived food intolerances and the severity of GI symptoms (e.g., abdominal pain), psychosocial distress (e.g., somatization), and quality of life.

MATERIALS AND METHODS

Participants

Participants included children ages 7–18 years meeting the Rome III diagnostic criteria for pediatric IBS¹ who were part of ongoing dietary intervention studies from 2008 – 2014. All children with IBS had completed a physician visit within the past year of enrollment and had not been identified with an organic etiology for their symptoms. Only baseline (prior to any intervention) information was used for this current report. Children with IBS and healthy children (HC) were recruited from the Texas Children’s Pediatric Associates general pediatrics practices and additional children with IBS from the Texas Children’s Hospital pediatric gastroenterology program as previously described.¹³ The screening process included investigator review of the subject’s medical record to ensure investigations of symptoms (e.g., lab testing) were negative.¹³

Healthy children were recruited based on absence of chronic illness and GI complaints and matched by age and sex in a 5:1 ratio to those with IBS. Though the differences between healthy children and those with IBS was unknown, the 5:1 ratio was initially chosen based on the much higher anticipated number of food intolerances in those with IBS vs. HC.

Demographic information was captured for all participants. Baylor College of Medicine Institutional Review Board approved the study protocol. Parents provided written informed consent, and children gave their assent.

Procedure

Study procedures were completed during a home visit or hospital visit for all children with IBS. Children completed all questionnaires in a room independent from their parent. For children who were 7–10 years of age, questionnaires were read by a research coordinator using a standardized protocol to ensure reading difficulties did not affect responses. Healthy children were interviewed over the phone in the same manner as would have occurred during a visit. Healthy children only completed the Childhood Food and Symptom Association Questionnaire. The comprehensive severity of IBS was evaluated through the use of the prospective Pain Diary as well as through the use of the questionnaires.

Determination of Food Intolerances and Avoided Foods

Participants completed the Childhood Food and Symptom Association Questionnaire assessing whether any of 97 foods (e.g., oranges) or food types (e.g., barbeque food) exacerbated their GI symptoms.¹¹ The food symptom association questionnaire was originally created and initially validated to assess symptoms in children with functional gastrointestinal disorders.¹¹ This questionnaire was read to the child by a coordinator outside the presence of a caregiver. If a food or food type was identified, the child was asked if he/she avoided eating the identified food. The questionnaire further assessed what type of symptom was elicited. Foods and food types were subsequently categorized into one of 8 groups for this study: Fruits, Vegetables, Dairy, Condiments, High Protein, Grains, Beverages, and Sweets (Table 1).

Pain Diary

Children with IBS were instructed on the completion of a daily Pain Diary for .7 days. Children rated abdominal pain for 3 intervals each day (morning, midday/afternoon, and evening/nighttime) using a numeric rating scale of 0 to 10, anchored with the phrases “no pain at all” and “the worst pain you can imagine”.¹⁴ This method of pain assessment has been validated in children and successfully used in children with IBS.^{15, 16} Parents were asked to remind children to complete the diaries daily and to allow children to independently rate abdominal pain without influencing their responses.¹⁷ Responses for each day were called in to a telephone data entry system. Research coordinators ensured compliance with completion of the Pain Diary by monitoring the phone call data entry and contacting families (at least once) when needed to ensure completion.

The pain severity scores were calculated as the mean pain rating over the course of the diary excluding ratings of “no pain” (i.e., zero). Pain frequency was calculated as the number of ratings ≥ 1 over the course of the diary.

Measures of Psychosocial Distress

The Children’s Somatization Inventory (CSI) includes 35 symptoms (e.g., “headache”) in which the child rates on a 5-point Likert scale (0=“not at all” to 4=“a whole lot”) how much the child is “bothered by each symptom” over the preceding two weeks.¹⁸ The CSI has been successfully used and validated in children with functional gastrointestinal disorders such as IBS.^{19, 20} Total scores for the CSI range between 0 and 140 with increasing scores indicating greater levels of somatization.

The Behavioral Assessment System for Children, Second Edition (BASC-2) is a questionnaire assessing a wide array of behavior and emotional dimensions of children, and has extensive support for reliability, content and construct validity (including in children with functional gastrointestinal disorders).^{21–23} The child self-report questionnaires are available for children ages 6 to 21 years. The Anxiety and Depression T scores were used for this study. The child self-report scales are derived from established norms with higher scorers having more anxiety or depression characteristics.²¹

The Functional Disability Index is a 15-item questionnaire which measures the degree to which children have difficulty in physical and psychosocial functioning.²⁴ It has been successfully used and validated in children with functional gastrointestinal disorders.^{24, 25} The total scores range from 0–60 with higher scores reflecting higher levels of perceived disability.^{24, 26}

The Pediatric Health-Related Quality of Life 4.0 Generic Core Scales is a 23-item, extensively validated questionnaire assessing health-related quality of life in healthy children and in those with medical conditions (including IBS).^{27, 28} Pediatric Health-Related Quality of Life total scores range from 0–100 with higher scores reflecting better quality of life.²⁷

Statistical Analyses

All statistical analyses were conducted using IBS® SPSS® Statistics (version 22, 2013, IBM Corporation). Chi square (for proportions) or a generalized linear model with Poisson log-

linear distribution (for data using counts) was used to compare data between groups. A binary Spearman's rho correlation coefficient was calculated between the number of self-perceived food intolerances and pain-related variables, psychosocial questionnaire scores, and quality of life scores. A generalized linear model with Poisson log-linear distribution using factors found to significantly correlate with the total number of perceived foods was used to evaluate the main effects and interactions of these factors on the total number of self-perceived food intolerances. All factors (main effects and their potential interactions) were initially included; thereafter, statistically redundant interactions were removed one at a time until all remaining factors and interactions in the final model were statistically significant.

Unless otherwise indicated, values are presented as mean \pm standard deviation or median [25% quartile, 75% quartile]. Two tailed P values <0.05 were considered statistically significant.

RESULTS

Comparison of Frequency and Type of Self-Perceived Food Intolerances in Children with IBS versus Healthy Children

A total of 154 children with IBS [89 (57.8%) girls] and 32 HC [19 (59.4%) girls] were enrolled in the study. Children in both groups were of similar age (12.5 ± 2.9 (SD) vs. 12.4 ± 3.6 years, IBS vs. HC, respectively). Children with IBS differed from HC with respect to race/ethnicity composition [97 (62.9%) vs. 10 (31.3%) White, 25 (16.2%) vs. 9 (28.1%) Hispanic, 22 (14.3%) vs. 9 (28.1%) Black, 5 (3.2%) vs. 4 (12.5%) Asian, 1 (0.6%) vs. 0 American Indian, respectively, $\chi^2=15.5$, $P<0.01$].

A greater proportion of children with IBS (143/154, 92.9%) vs. HC (20/32, 62.5%) identified at least one food as being able to exacerbate a GI symptom ($\chi^2=22.5$, $P<0.001$). Children with IBS identified a greater number (median = 4 [2–6]; Range=35) of symptom-inducing foods than HC (median = 2 [0–4]; Range=11; $\chi^2=28.6$, $P<0.001$). In parallel, children with IBS avoided a greater median number (2 [1–4]; Range=16) of foods than HC (median=0 [0–2.75]; $\chi^2=20.8$, Range=9; $P<0.001$). The number of self-perceived food intolerances and avoided foods correlated in both those with IBS ($r=0.73$; $P<0.0001$) and HC ($r=0.58$; $P=0.001$). Children with IBS vs. HC identified a wide range of self-perceived food intolerances (Figure 1) and avoided foods (Figure 2).

Children with IBS vs. HC were more likely to have GI symptom inducing foods within the Dairy, Beverages, and Grains groups but not foods in the other assessed categories (Table 2). The ten most commonly identified GI symptom inducing foods in children with IBS in descending order of frequency were cow's milk, fast food, cheese, ice cream, spicy food, beans, pizza, sodas, chocolate, and fried foods (Table 3).

Association of Self-Perceived Food Intolerances with Sex, Race/Ethnicity, and Age

In children with IBS, girls identified a greater number (median = 4 [2–7]; Range=35) of self-perceived food intolerances than boys (median = 3 [1.5–5]; Range=15; $\chi^2=1337.8$, $P<0.001$). In addition, girls avoided a higher number (median = 2 [1–5]; range: 0–16) of foods than boys (median = 1 [1–3]; Range=10; $\chi^2=378.8$, $P<0.001$) within the IBS cohort.

No differences were found in HC between boys and girls in the total number of food intolerances or avoided foods (data not shown). We did not identify an association between either the total number food intolerances or avoided foods with race/ethnicity (data not shown). In children with IBS there is a weak correlation between age and the number of self-perceived food intolerances ($r=0.16$, $P=0.05$). In contrast, there is a weak negative correlation in HC between age and the number of self-perceived food intolerances ($r=-0.38$, $P<0.05$).

Association of Self-Perceived Food Intolerances with Abdominal Pain Characteristics, Psychosocial Distress, and Quality of Life in Children with IBS

In children with IBS, the number of self-perceived food intolerances was weakly related (r -value range, -0.17 to 0.21 ; Table 4) with abdominal pain frequency, pain severity, somatization, anxiety, functional disability, and lower quality of life scores. Depression was not associated with the number of identified food intolerances.

DISCUSSION

To date the self-perceived role of diet in IBS symptom generation in children has not been comprehensively assessed. This study suggests that children with IBS, particularly girls, frequently perceive diet as playing an important role in exacerbating their IBS symptoms. Presumably due to these self-perceived food intolerances, children with IBS avoid a high number of foods. The number of food intolerances was weakly associated with more frequent and severe abdominal pain, as well as increased functional disability, increased somatization and anxiety, and decreased quality of life. Overall, these findings in children with IBS suggest a high prevalence of self-perceived food intolerances in children but a weak correlation with clinical phenotype.

The prevalence of reported food intolerance in the children with IBS in this study is higher than that (65–84%) reported in adult IBS studies evaluating postprandial symptoms.^{5, 6, 10, 29} Whether this suggests differences in children with IBS vs. adults with IBS with respect to food responses (e.g., immunological) needs to be evaluated in future studies. Given the high prevalence of perceived food intolerances in children with IBS, future dietary studies in this population may consider taking into account the number of self-perceived food intolerances. In addition, intervention studies (both dietary and non-dietary) may need to formally address dietary regimens consumed during the intervention as they may influence GI symptom expression.

Though not the primary aim of this study, this study reports for the first time the number of food intolerances in a small population of HC. A group of HC was used in this initial assessment to provide context to the findings in those children with IBS. HC were matched for age/sex and not by race/ethnicity to those with IBS to reflect better Houston (where the study took place) demographics. The healthy children participants had a wide range and overall higher prevalence of self-perceived food intolerances as compared to primarily adult-based population studies which identified perceived food reactions in 20–34%, though some of the reported symptoms were not specifically GI related.^{7, 8} This may be related in part either to differences in biological factors related to food perception such as food neophobia

(which is more pronounced in children vs. adults) or methodological differences through this study's use of a more comprehensive questionnaire assessing more foods and/or food types. Young et al. only evaluated 8 foods, while Zuberbier et al. mentioned "more than 40" assessed foods and/or food additives.^{7, 8} Healthy children (and those with IBS) may have incorrectly associated dislike of the food due to taste or other reasons with a food intolerance. Future studies of perceived food intolerances in HC of various races/ethnicities and of different geographic locations with different cuisines are needed.

This study found that avoidance of foods that may induce symptoms was common in children with IBS as has been reported in adults with IBS.¹⁰ Avoidance of a culprit food would be an expected coping strategy that children with IBS may employ to better control their symptoms. Though this study did not address whether nutritional deficiencies were present due to food avoidance, studies in adults with IBS have found nutritionally inadequate diets in a small number of subjects.⁵ Families and clinicians caring for children with IBS with a large number of self-perceived food intolerances and subsequent avoidance of these foods may consider formally assessing the child's nutrient intake.

Girls with IBS have increased visceral hypersensitivity, more frequent autonomic abnormalities, and more psychological distress (e.g., depression) as compared to boys with IBS.^{30, 31} The etiology for these sex differences has not been fully established but is also seen in adult women versus men.³² Paralleling these sex differences, girls with IBS in the study had a higher number of self-perceived food intolerances than boys. These differences (particularly increased visceral hypersensitivity) are believed to play a significant role in food-induced symptoms. Alternatively, increased psychological distress influencing perception may account for the sex findings. Another possibility may be differences in self-awareness between boys and girls. Given that these sex differences may extend into adulthood, future studies are needed to determine the true sex-based pathophysiologic connections with food intolerance (including differences in self-awareness).^{6, 10}

The primary limitation of the study is that it is cross-sectional. Given the number of factors/relationships evaluated, the number of children studied was modest. Although this approach allows for insight into the evaluated relationships to food intolerance, causality will need to be determined in future studies. Another limitation in the study is that food intolerances were not documented by blind challenges, prospective means (e.g., electronic diaries) or corroboration with healthcare providers and/or parents.³³ However, while the role of healthcare provider opinion is unknown, significant differences between parental perception and child complaints have been reported in children with functional GI disorders.^{34, 35} Another limitation (given mode of administration of a questionnaire may influence results) is having HC interviewed via phone rather than in person.³⁶ However differences were minimized by having the same research coordinators administer the food symptom association questionnaire using a standardized protocol.

Areas where future studies may be better able to address self-perceived food intolerances in children with IBS include the role of self-awareness, the influence of ongoing treatments, and parental influences (e.g., parental IBS status). Double-blind placebo controlled food challenge trials (often employed in food allergy research) may help determine causality of

foods and IBS symptoms.³⁷ In addition these trials may be able to be paired with non-invasive mechanistic evaluations (such as breath testing for assessment of colonic fermentation) to understand the underlying pathophysiology.¹⁵ Information from future trials will help healthcare providers and families in determining how best to identify (e.g., self-perception vs. formal challenges) and address potential foods that induce symptoms.

There are several strengths of this study. First, all IBS children were defined as having IBS per Rome III criteria. Second, only validated and previously published measures were used, increasing the validity of the study. Third, children themselves completed the measures with the assistance of research coordinators independent of parents rather than having parents fill out the measures as surrogates. Fourth, abdominal pain characteristics were captured prospectively via using the Pain Diary which has been shown to be more reliable than relying on recall alone in children with IBS.³⁸ Finally, children with IBS were recruited from both primary care and tertiary care settings, lending greater generalizability to the results.

CONCLUSIONS

Children with IBS have a high prevalence of self-perceived food intolerances and avoid foods due to this perception. The number of self-perceived food intolerances in children with IBS is weakly associated with a more severe IBS clinical phenotype characterized by increased GI symptoms, increased psychosocial distress, and decreased quality of life. These study findings indicate a need for clinicians to address self-perceived food intolerance in children with IBS. Future studies related to self-perceived food intolerance may lead to identification of successful strategies to address this important and common experience of children with IBS.

Abbreviations

FODMAP	fermentable oligosaccharides disaccharides monosaccharides and polyols
GI	gastroenterology
IBS	irritable bowel syndrome

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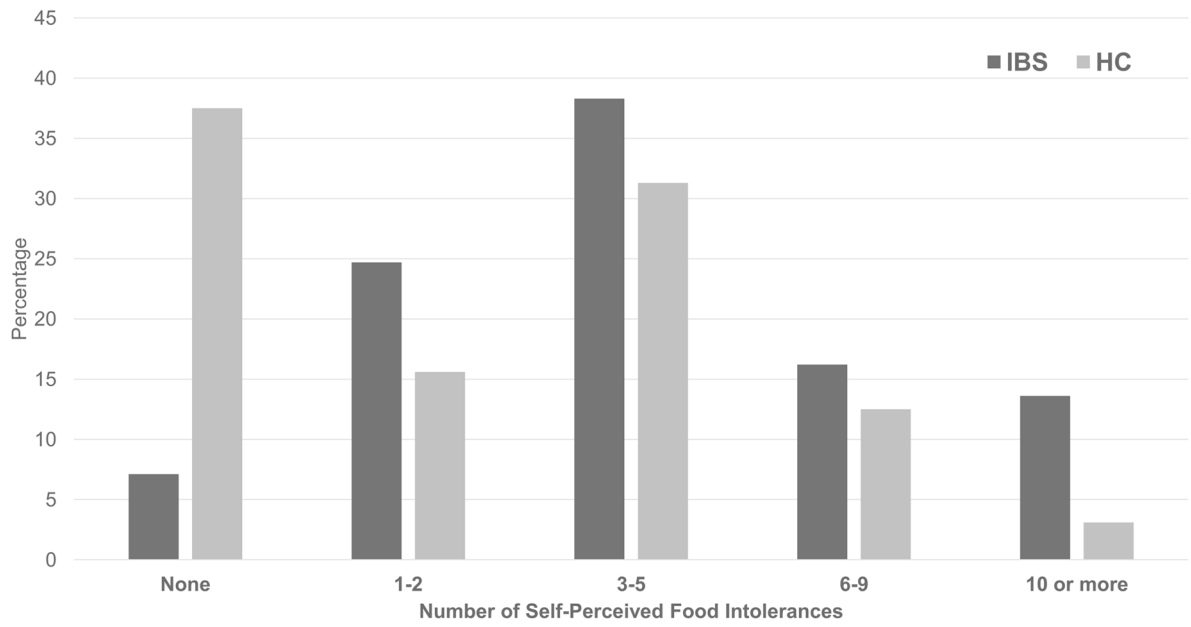


Figure 1. Number of self-perceived food intolerances in children with irritable bowel syndrome (n=154) versus healthy children (n=32). IBS = children with irritable bowel syndrome; HC = healthy children

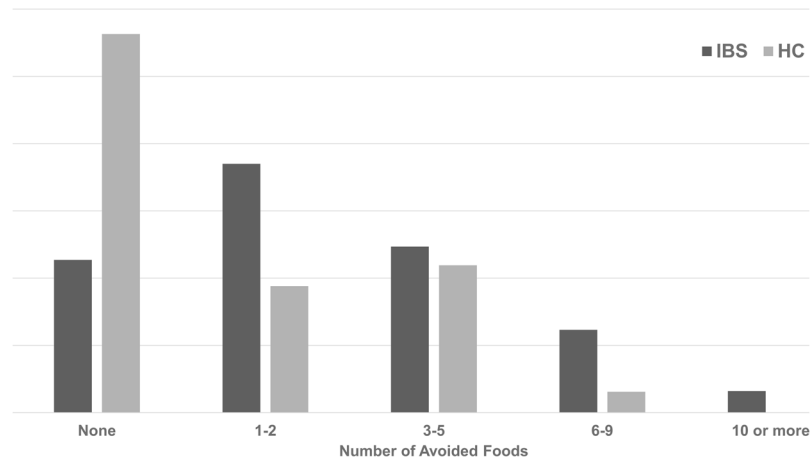


Figure 2. Number of avoided foods in children with irritable bowel syndrome (n=154) versus healthy health controls (n=32). IBS = children with irritable bowel syndrome; HC = healthy children

Table 1

Food and Food Type Assignments into Food Categories from Items in the Childhood Food and Symptom Association Questionnaire¹¹

Food Category	Food and/or Food Type
Fruits	Apples, Pears, Mangoes, Watermelon, Melons, Oranges, Grapefruit, Lemons, Peaches, Plums, Cherries, Nectarines, Bananas, Apricots, Grapes, Raisins, Berries, Fruit juices, Dried fruits, Other fruits
Vegetables	Onions, Broccoli, Cauliflower, Asparagus, Cabbage, Beans, Peas, Tomatoes, Corn, Potatoes, Brussels sprout, Green or Red peppers, Cooked vegetables, Raw vegetables, Other vegetables
Dairy	Milk (cow's), Soy milk, Ice cream, Cheese, Creams, Yogurt, Butter, Margarine, Coconut milk/cream
Condiments	BBQ sauce, Ketchup, Mayonnaise, Salad dressing, Mustard, Pancake syrup, Honey, Jellies/Jam, Peanut butter, Sweet and sour sauce, Gravy, Salsa
Meats/Protein	Pork, Sausage, Ham, Beef, Turkey, Hot dogs, Chicken, Fish, Eggs, Peanuts, Other nuts
Grains	Pasta, Pasta sauce, Wheat bread, Rye bread, White bread, Bagels, Sourdough bread, Cold breakfast cereal, Cookies, Crackers, Cakes, Pizza, Rice
Beverages/Other	Pickles, Sodas, Diet sodas, Sports drinks, Coffee, Tea, Alcohol, Smoked foods, Deep fried foods, Fried foods, Spicy foods, Fast food
Sweets	Pastries, Chocolate, Non-chocolate candy, Sugar-free gum, Sugar gum

Table 2

Comparison of Self-Perceived Food Intolerances by Food Category in Children with Irritable Bowel Syndrome versus Healthy Children using the Food and Symptom Association Questionnaire¹¹

Food category ^a	Children with Irritable Bowel Syndrome (n=154)	Healthy Children (n=32)	P-value
Fruit	61 (39.6%)	11 (34.4%)	N.S. ^b
Vegetables	59 (38.3%)	10 (31.3%)	N.S.
Dairy	82 (53.2%)	8 (25%)	<0.005
Condiments	44 (28.6%)	6 (18.8%)	N.S.
Meat/High Protein	39 (25.3%)	3 (9.4%)	N.S.
Grains	48 (31.2%)	4 (12.5%)	<0.05
Beverages/Other	84 (54.5%)	8 (25%)	<0.005
Sweets	29 (18.8%)	3 (9.4%)	N.S.
Item From Any Food Category	143 (92.9%)	20 (62.5%)	<0.001

^aPlease see Table 1 for foods and food types included in each category.

^bN.S. = Not Significant

Table 3

Top Ten Most Commonly Identified Foods or Food Types^a Causing Self-Perceived Food Intolerance in Children with Irritable Bowel Syndrome vs. Healthy Children Using the Food and Symptom Association Questionnaire¹¹

Food or Food Type ^a	Children with Irritable Bowel Syndrome (n=154)	Healthy Children (n=32)
Cow's milk	51 (33.1%)	3 (9.4%)
Fast food	37 (24%)	3 (9.4%)
Cheese	35 (22.7%)	2 (6.3%)
Ice cream	34 (22.1%)	3 (9.4%)
Spicy Food	32 (20.8%)	4 (12.5%)
Beans	23 (14.9%)	3 (9.4%)
Pizza	23 (14.9%)	1 (3.1%)
Sodas	21 (13.6%)	1 (3.1%)
Chocolate	20 (13%)	3 (9.4%)
Fried foods	18 (11.7%)	2 (6.3%)

^aThe food or food type is from the Food and Symptom Association Questionnaire¹¹

Table 4

In children with Irritable Bowel Syndrome (n=154), the Number of Self-perceived Food Intolerances Correlates with Somatization, Anxiety, Functional Disability, Number of Abdominal Pain Episodes and Median Pain Severity. The Number of Food Intolerances Negatively Correlates with Quality of Life but is not Correlated to Depression.

Characteristic	Median Value [25–75% quartile]	Spearman Correlation r-value	P value
Number of daily pain episodes ^a	1 [0.5–1.6]	0.17	<0.05
Median pain severity ^a	3.25 [2.3–4.0]	0.2	<0.05
Somatization ^b	24 [15–35.7]	0.22	<0.01
Anxiety (BASC-2 T-score) ^c	51 [43–62]	0.21	0.01
Functional disability ^d	9 [4–19]	0.16	<0.05
Quality of life ^e	81.8 [70.7–89.1]	– 0.17	<0.05
Depression (BASC-2 T- score) ^c	45 [41–51]	0.1	0.2

^aAbdominal pain characteristics were captured by a two week Pain Diary¹³

^bDetermined using Children's Somatization Inventory¹⁸

^cBASC-2= Behavioral Assessment for Children, Second Edition²¹

^dDetermined using Functional Disability Inventory²⁴

^eDetermined using PedsQL®.²⁷