Factor Analysis Defines Distinct Upper and Lower Gastrointestinal Symptom Groups Compatible With Rome IV Criteria in a Population-based Study

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BACKGROUND & AIMS:

The Rome IV criteria define functional gastrointestinal (GI) disorders by specific combinations of symptoms. It is possible to empirically evaluate these symptom combinations by factor analysis (a statistical procedure that groups variables that correlate). However, this analysis has not been performed for the Rome IV criteria, and factor analyses based on the previous versions of the Rome criteria did not use population-based data. We therefore investigated symptom grouping by the Rome IV questionnaire using factor analysis of a population-based sample.

METHODS:

The Rome IV questionnaire was completed online in English by 5931 respondents from the United Kingdom, United States, and Canada (49% female, age range, 18-92 years). We performed an exploratory factor analysis on the Rome IV questions. Next, we performed a confirmatory factor analysis to compare the exploratory factor result to that of the Rome IV criteria.

RESULTS:

The exploratory factor analysis identified 8 factors that accounted for 45% of the variance in response: constipation, diarrhea, irritable bowel syndrome, abdominal pain, heartburn, nausea or vomiting, globus, and other upper GI symptoms. Most factors corresponded to distinct functional GI disorders defined by the Rome IV criteria—exceptions included abdominal pain and upper GI symptoms. In confirmatory factor analysis, the exploratory model fitted slightly better than that based on the Rome IV criteria (root mean square error of approximation, 0.063 vs 0.077).

CONCLUSIONS:

We used factor analysis to identify distinct upper and lower GI symptom groups that are compatible with the Rome IV criteria. Our findings support the use of the Rome IV criteria in research and clinical practice as a basis for development of diagnostics and management of patients.

Keywords: IBS; FGID; validation; Internet survey.

F unctional gastrointestinal (GI) disorders (FGIDs) are syndromes defined by clusters of symptoms emanating from the GI tract, with population prevalences for many of these disorders ranging between 5% and 10%. More than 20 FGIDs are defined by the published Rome IV criteria, based on specific GI symptom combinations.

The distinction of each FGID as a separate entity may be corroborated by investigating how GI symptoms group together in the general population, and whether these symptom groupings are compatible with the Rome criteria. One method of doing so is factor analysis: a statistical procedure that groups variables with strong intercorrelations. Consistent agreement between factor analysis and the Rome symptom criteria would then suggest that the disorders represent distinct entities, which are identified by these symptom criteria. Previous

studies have performed factor analyses on GI symptoms and compared the resulting factors with the structures of the Rome I and II criteria, 3,4 and the Manning criteria for irritable bowel syndrome (IBS).⁵ For example, Whitehead et al⁶ found factors of IBS (ie, abdominal pain related to stool patterns), nausea, constipation (including straining and incomplete evacuation), and a heterogeneous group of other GI symptoms. A similar IBS factor was found by Taub et al⁷ and Talley et al.⁸ Whitehead et al⁹ performed a factor analysis on GI symptoms in gastroenterology clinic patients and reported 13 distinct factors, including IBS, constipation, diarrhea, and anorectal pain, but also several upper GI factors. Finally, Siah et al¹⁰ described among others a factor of bloating, associated with flatulence and belching. In conclusion, factor analyses can be used to substantiate the definitions of FGIDs.

The previously mentioned factor analyses assessed Rome I, II, and III diagnostic classifications, but such analyses have not been performed since the publication of Rome IV. Furthermore, most of the previous factor analyses did not use population-based samples. We therefore aimed to empirically substantiate the latest definitions of FGIDs in a large, population-based sample by factor analysis.

Methods

Rome IV Survey

We used an Internet-based Rome IV survey involving 6300 individuals, with methodology described previously. Briefly, participants were a nationally representative adult population sample from the United States, United Kingdom, and Canada, as selected by the survey vendor Qualtrics Inc. Quota-based sampling was used to ensure equal proportion of sex (50:50), age groups (40% aged 18-39 years, 40% aged 40-64 years, and 20% aged 65 years and older), and education level (30% maximum with more than 16 years of formal education) across the countries. In the US sample, the survey vendor also selected 20% African Americans and 20% Hispanic ethnicity respondents, but these demographic shifts were not done in the United Kingdom or Canada. The survey was presented as a health survey without specific mention of gastroenterology, to prevent enrichment of participants with GI problems. All respondents confirmed informed consent online at the start of the survey. To ensure reporting quality, 3 questions were presented twice, and there were 2 attention verification questions. The survey was completed by a total of 6300 respondents, (ie, 2100 for the United States, United Kingdom, and Canada, respectively). Before data collection started, the study was reviewed by the Institutional Review Board at the University of North Carolina and deemed Institutional Review Board-exempt because all study participants were anonymous to the investigators.

Data Preprocessing

Of the 6300 respondents, we included those who satisfied both attention verification questions and the response consistency checks with the 3 repeated symptom questions. This left 5931 respondents in the analysis sample. Of the 86 Rome IV questions, we included those with ordinal response scales, because these can be used to compute correlations. Excluded questions mostly related to the length of time the symptom had been present. No questions critical to a Rome IV FGID diagnosis were excluded. A total of 49 Rome IV questions were included in the analysis.

Exploratory Factor Analysis

We performed an exploratory factor analysis using the "factanal" function in R version 3.3.3. This is a statistical procedure that groups variables with strong intercorrelations, and as such identifies the key variable groupings. A correlation matrix of the Rome IV questions was computed, and orthogonal factors with an eigenvalue larger than 1 were retained. Factor analysis was performed with maximum likelihood estimation and varimax rotation. Loadings of 0.40 and larger were deemed relevant.

Confirmatory Factor Analysis

We used confirmatory factor analysis to confirm that Rome IV successfully captures the GI symptoms into distinct FGID entities. This is done by fitting a prespecified model of the Rome IV structure, and comparing this model with the data-driven structure resulting from the exploratory factor analysis. In case the models are similar of composition and statistical fit, this further reinforces the Rome IV criteria.

The factor structure of the Rome IV criteria was prespecified as follows (see Table 1 for reference). Globus was defined as a lump in the throat occurring between meals (items 1 and 3). Functional heartburn was defined by burning sensations in the chest (items 7 and 9). Functional dysphagia was defined by foods sticking in the esophagus¹² (items 8 and 12). Postprandial distress syndrome was defined by postprandial fullness and early satiety (items 14 and 16). Chronic nausea and vomiting syndrome was defined by nausea and vomiting¹³ (items 21 and 23). IBS was defined by pain anywhere in the abdomen related to defecation (items 40, 41, 42, and 43). Functional constipation was defined by stools that are hard, infrequent, and hard to pass¹⁴ (items 49, 51, 52, 53, 54, and 55). Biliary pain was defined by pain in the right upper quadrant 15 (items 68, 69, 70, 71, and 72). Other FGIDs were defined by only 1 eligible Rome IV question, and were not in the confirmatory model to be consistent with the exploratory model. The exploratory and Rome IV-based

Table 1. Factor Loadings After Varimax Rotation

Item	Topic	Constipation	Upper Gl	Abdominal pain	IBS	Nausea/ vomiting	Diarrhea	Heartburn	Globus
1	Lump in throat	.15	.36	.03	.02	.12	.08	.08	.78
3	Lump in throat when not eating	.09	.05	.19	.14	.08	.04	.19	.48
4	Pain with swallowing	.12	.17	.23	.08	.22	.03	.30	.07
5	Chest pain	.20	.49	.10	01	.18	.08	.22	.18
7	Chest pain as burning sensation	.07	.12	.19	.13	.05	.05	.62	.07
8	Chest pain and food sticking after swallowing	.09	.10	.08	.12	.18	.05	.45	.09
9	Heartburn	.15	.41	.07	.05	.11	.12	.49	.08
12	Foods stuck in chest	.18	.43	.05	.03	.20	.11	.33	.22
14	Postprandial fullness	.18	.41	.18	.04	.25	.13	.21	.08
16	Early satiety	.22	.36	.17	.05	.30	.13	.15	.09
18	Epigastric pain/burning	.15	.55	.29	.03	.25	.09	.26	.04
19	Epigastric pain/burning improved by defecation	.12	.03	05	.45	02	.00	.21	.10
21	Nausea	.16	.38	.29	.08	.54	.11	.11	.07
23	Vomiting	.11	.27	.15	.05	.66	.07	.10	.05
25	Self-induced vomiting	.06	.00	01	.00	.17	.09	.18	.07
32	Food coming back up after swallowing	.16	.44	.07	.08	.37	.12	.21	.12
34	Retching	.14	.14	.16	.07	.58	.14	.15	.09
35	Vomiting when food comes back up	.20	.15	.21	.09	.63	.13	.09	.01
37	Food coming back up is recognizable	.04	.10	.11	.12	.18	01	.05	.06
38	Belching	.09	.40	.15	.04	.27	.10	.19	.05
40	Pain anywhere in the abdomen	.25	.63	.33	.11	.13	.18	.02	.05
41	Abdominal pain related to defecation	.14	.03	.06	.73	01	.13	.03	.01
42	Abdominal pain related to stool consistency	.20	.07	.11	.85	.08	.19	.05	.04
43	Abdominal pain related to stool frequency	.22	.08	.16	.75	.13	.23	.10	.03
45	Abdominal pain related to a meal	.09	.13	.33	.38	.13	.25	.22	.04
46	Abdominal pain limits usual activities	.17	.14	.54	.27	.24	.16	.10	.05
49	Hard stools	.61	.13	.10	.14	.08	03	.12	.07
51	Infrequent stools	.55	.08	.18	.08	.14	.03	.04	.06
52	Straining during bowel movements	.80	.16	.09	.18	.08	.04	.11	.06
53	Incomplete bowel emptying	.72	.19	.13	.20	.12	.20	.09	.06
54	Anorectal obstruction / blockage	.81	.20	.14	.14	.10	.10	.09	.06
55	Manual maneuvers to facilitate defecation	.52	.14	.13	.05	.11	.04	.10	01
59	Loose stools	.07	.21	.09	.21	.15	.73	.08	.03
61	Loose stools Loose stools following a meal	.03	.05	.17	.21	.09	.64	.11	.06
63	<u>-</u>	.20	.25	.17	.20	.12	.64	.10	.03
65	Urgency Bloating/abdominal distention	.20 .37	.23 .42	.20	.15	.03	. 04 .27	.10	.03
	•		.42 .63	.38					
68	Right upper quadrant pain	.25	. 03 .18	.36 .71	.09 .08	.14	.17	.01	.03 .07
69	Long-lasting right upper quadrant pain	.18				.12	.09	.10	
70	Right upper quadrant pain of steady severe level	.19	.17	.74	.09	.14	.14	.07	.05
71	Right upper quadrant pain at different intervals	07	12	12	.11	10	.01	.03	02
72	Right upper quadrant pain limits usual activities	.12	.19	.63	.07	.26	.07	.07	.04
73	Right upper quadrant pain related to defecation	.17	.07	.13	.48	.05	.14	.01	.03
74	Right upper quadrant pain related to posture	.10	.02	.08	.13	.09	.06	.15	.00
75	Right upper quadrant pain improved by medicine	.05	.08	.04	.01	.05	.05	.42	01
76	Right upper quadrant pain related to vomiting	.11	.16	.37	.05	.57	.10	.17	.04
77	Right upper quadrant pain related to back pain	.18	.21	.40	.07	.14	.07	.17	.05
78	Right upper quadrant pain waking one up	.13	.25	.51	.04	.26	.15	.15	.00
80	Fecal incontinence	.11	.28	.07	.09	.13	.36	.05	.01
83	Anorectal pain	.38	.38	.13	.12	.11	.17	.02	.04

models were compared by root mean square error of approximation.

Results

There were 5931 respondents in the analysis sample. The demographics approximated those from recent census reports^{16–18} by age and sex categories. The age ranged from 18 to 92 years old and 49% was female (Figure 1). Common FGIDs were functional dyspepsia, postprandial distress syndrome (7.7%); functional constipation (6.3%); proctalgia fugax (5.9%); IBS (5.7%); functional abdominal bloating/distention (5.6%); and functional diarrhea (5.4%).

Exploratory Factor Analysis

Eight distinct factors emerged from the factor analysis and were characterized as constipation, diarrhea, IBS, abdominal pain, heartburn, nausea/vomiting, globus, and other upper GI symptoms (Table 1). Taken together, they explained 45% of the response variance. The constipation factor held all items that define functional constipation in Rome IV. Diarrhea was described by loose stools, including those after a meal, and urgency, and was thus compatible with functional diarrhea in Rome IV. The IBS factor held the Rome IV defining questions for IBS (abdominal pain related to defecation, stool frequency, and stool consistency) with large loadings ($r \ge 0.73$), but also pain in the epigastric or right upper quadrant regions related to defecation, albeit with smaller loadings (r = 0.45 and 0.48). In the factor of abdominal pain, pain was not associated with defecation

but did interfere with usual activities (ie, symptoms partially covering both biliary pain¹⁵ and centrally mediated abdominal pain syndrome in Rome IV). 19 Heartburn and globus overlapped with the Rome IV definitions for functional heartburn and globus. The factor nausea/vomiting contained the items used to define chronic nausea and vomiting syndrome in Rome IV, plus retching and vomiting related to right upper quadrant pain (r = 0.58 and 0.63). Finally, there was a factor of upper GI symptoms, containing 2 symptoms of functional dyspepsia in Rome IV (epigastric pain and postprandial fullness), but also bloating, belching, abdominal pain, and some esophageal symptoms. Some items stood in isolation, particularly pain with swallowing, early satiety, abdominal pain related to a meal, fecal incontinence, and anorectal pain (r < 0.40). All items and their rotated factor loadings are shown in Table 1. Results separate for the 3 country samples are available in Supplementary Tables 1-3.

Of the 5931 respondents, 62% met the Rome IV symptom threshold for at least 1 single symptom that is part of the definition of FGIDs (Figure 2). Symptoms were sometimes confined to 1 factor (19%), but often covered items from 2 (11%) or more (32%), indicating a degree of overlap among the various GI symptoms. The most common symptom was abdominal pain related to defecation (21%) followed by bloating/abdominal distention and loose stools (19% each), whereas no esophageal or gastroduodenal symptom was particularly common (maximum 10%).

As can be appreciated in Figure 2, the IBS factor had the strongest within-factor loadings and correlations: of the 30% who met the Rome IV symptom threshold for at least 1 of the items in the IBS factor, 77% met the

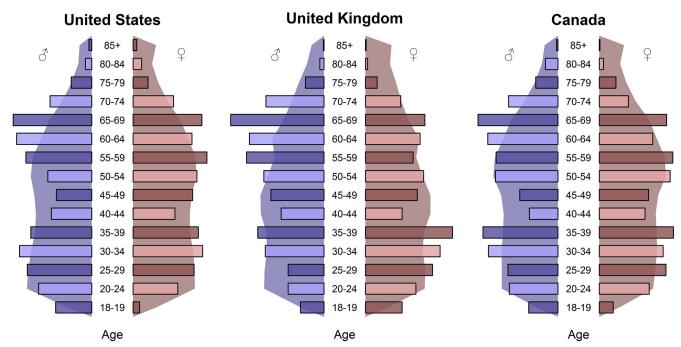


Figure 1. Age and sex of the respondents in the present study (bars) compared with those of recent census reports (shade).

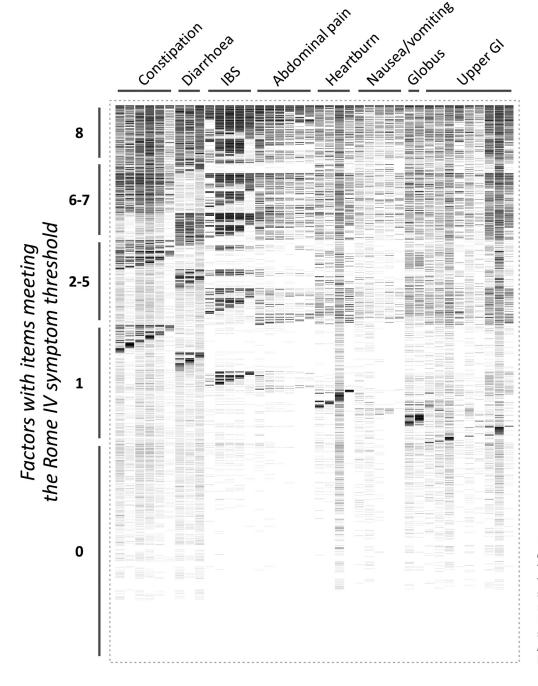


Figure 2. Raw data. Columns are Rome IV questions sorted numerically by factor. Rows are respondents, sorted by the number of factors with items meeting the Rome IV symptom threshold. Darker colors represent higher (worse) responses.

criteria for at least 2 items (note the contrasting black and white color patterns for IBS in Figure 2). By contrast, of the 33% who met the Rome IV symptom threshold for at least 1 upper GI item, only 51% met the criteria for at least 2. Many upper GI items thus stood in isolation, possibly because many upper GI FGIDs are based on a single key symptom rather than a combination of symptoms, unlike most functional bowel disorders.

Confirmatory Factor Analysis

The exploratory factor model had a slightly better fit than that of the Rome IV model (root mean square error of approximation, 0.063 vs 0.077; chi-square test, 17500 vs 8932; df, 712 vs 247). However, the difference in fit was small, also given that the most factors and FGIDs were represented in both the exploratory and Rome IV model, with near-identical compositions (Figure 3). As can be seen in Figure 3, the confirmatory model had coefficients of a magnitude comparable with those of the exploratory model (these are the values on the diagonal lines: compare left with right). One item had a negative coefficient in the confirmatory model: right upper quadrant pain at different intervals (item 71, -0.16, Figure 3), indicating that it did not match well with the factor for biliary pain as hypothesized based on Rome IV.

Exploratory model

Confirmatory model

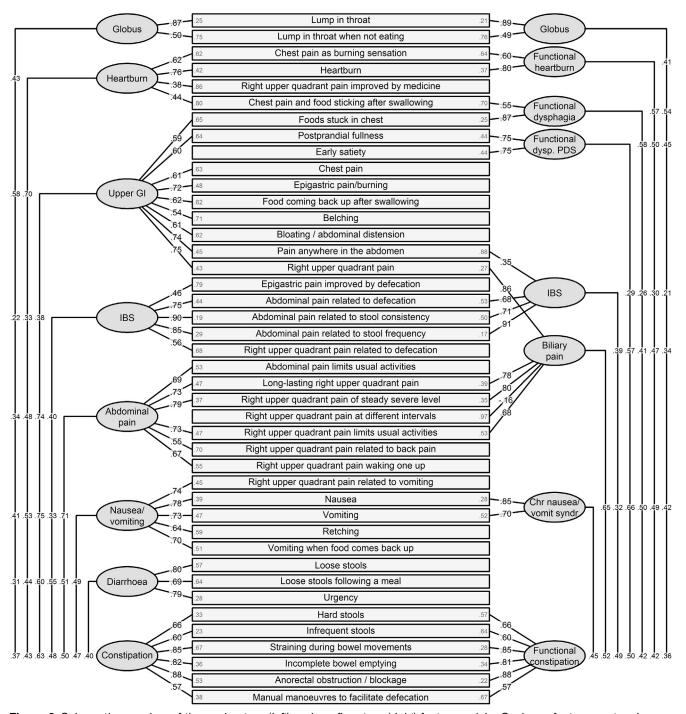


Figure 3. Schematic overview of the exploratory (*left*) and confirmatory (*right*) factor models. *Ovals* are factors, *rectangles* are Rome IV items. Values on *diagonal lines* are standardized coefficients (factor loadings) and indicate how strongly each item loads on the factor that it defines. Values on *vertical lines* denote covariances of factors. Values in *boxes* are error terms.

Discussion

The present study empirically evaluated the symptom combinations in the Rome IV criteria, and found 8 distinct symptom groupings (constipation, diarrhea, IBS, abdominal pain, heartburn, nausea/vomiting, globus, and other upper GI symptoms). All in all, these corresponded well with the Rome IV structure.²

The factors for IBS, constipation, diarrhea, globus, heartburn, and nausea/vomiting closely matched the Rome IV criteria for IBS, functional constipation, functional diarrhoea, 14 globus, functional heartburn, 12 and chronic nausea and vomiting syndrome, 13 respectively, thereby reinforcing the Rome IV criteria for these FGIDs. Moreover, these factors were quite similar to those reported in factor analyses of earlier Rome

editions.⁶⁻¹⁰ The IBS factor was the most distinct, with respondents scoring either very high or very low on all defining questions. Although constipation and diarrhea are often seen in IBS patients, IBS is a heterogeneous condition, and as such it is not unexpected to see factors for the core element of IBS (pain related to stool patterns) and for the separate stool patterns.

We found a separate factor for abdominal pain (ie, of a different nature than IBS), which also has been described in earlier studies. 9 It covered both items from biliary pain (right upper quadrant pain)¹⁵ and centrally mediated abdominal pain syndrome (pain restricting usual activities). 19 This factor does not cleanly represent biliary pain, because the item of the pain taking place at different intervals loaded negatively on this factor, while being a defining item in Rome IV. Instead it may represent centrally mediated abdominal pain syndrome, in which the pain is nearly continuously present. Nevertheless, it is difficult for respondents to sense exactly where the pain is located (possibly because of diffuse activation of nociceptive systems in visceral organs),²⁰ explaining the overlap between pains in the various abdominal subregions. Moreover, irrespective of factor structure, it is possible that symptoms overlap between different disorders because of common pathophysiological mechanisms.

Upper GI symptoms factored into heartburn, globus, and nausea/vomiting. There was also a factor with a combination of upper GI symptoms including functional dyspepsia and overlapping symptoms, such as belching and bloating, which was also found by Siah et al. 10 Whitehead et al⁹ reported separate factors of heartburn, dysphagia, globus, rumination syndrome, dyspepsia, capturing the structure proposed in Rome I. Others saw factors of nausea and gas/belching/acid^{6,7} or reflux.8 Factor analyses have also been performed on different questionnaires, such as the reflux disease questionnaire, where factors were found for regurgitation (eg, acid taste), heartburn, and dyspepsia, although upper GI symptoms overlapped considerably, 21 in line with the present study. Siah et al¹⁰ further discussed a meal-related factor. Meal-related abdominal pain in the present study did not load on any factor, with the largest loadings being for IBS (r = 0.38) and abdominal pain (r = 0.33). The existence of a broad factor of upper GI symptoms may imply a degree of common pathophysiology between these items. Common mechanisms may be peripheral or central. For example, Fischler et al²² found a factor of bloating and postprandial fullness to be associated with delayed gastric emptying, a factor of belching associated with gastric hypersensitivity, and a factor of epigastric pain associated with both gastric hypersensitivity and several psychosocial factors. In summary, the factor structure of upper GI symptoms found in the present study is quite consistent with those of prior studies, and partially reinforces the Rome IV criteria, although some upper GI symptoms were difficult to separate out using the Rome IV questionnaire.

Finally, confirmatory factor analysis showed that Rome IV groups symptoms in a manner that captures the variance in the population, close to the factors found through computerized means (ie, the exploratory factor analysis). The item "right upper quadrant pain at different intervals," defining biliary pain in Rome IV, was the only odd one out to have a negative coefficient for this FGID. It may be considered to rephrase this item in future Rome editions.

Our study has 2 strengths. First, although factor analytical studies have contributed to amendments of Rome I, we present a factor analysis for the published Rome IV questionnaire. The results reinforce the Rome IV criteria, which can be used in research and clinical care for diagnostics and specific management of FGID patients. Second, we did so in a large sample in 3 countries, with demographics close to those of the general population. Generalizability of our findings to the population should therefore be high. It remains to be seen whether other languages, countries, or continents (eg, Asia) would render similar findings, 23 although the recent factor analysis by Siah et al¹⁰ of Rome III shows that any differences are likely to be subtle. A limitation is the absence of information on how many potential respondents chose to ignore the invitation to partake in the survey, despite the close similarity between sample and population demographics. Furthermore, some FGIDs are described by only 1 key symptom in Rome IV, so that these are not supported by factor analysis. However, it is useful to include these symptoms into the exploratory analysis to determine whether there are unpredicted symptom associations that should be recognized.

To conclude, upper and lower GI symptom groupings compatible with Rome IV diagnoses exist in the general population and can be identified using the English Rome IV questionnaire in a large population-based sample from 3 English-speaking countries. Our results provide further basis for practicing clinicians to use the Rome IV criteria throughout the process going from symptoms to diagnosis and management. Although our results reinforce the Rome IV criteria, some upper GI symptoms were difficult to separate out, and more work is needed to reach a distinct classification.

Supplementary Material

Note: To access the supplementary material accompanying this article, visit the online version of *Clinical Gastroenterology and Hepatology* at www.cghjournal.org, and at https://doi.org/10.1016/j.cgh.2018.02.042.

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Reprint requests

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Conflicts of interest

These authors disclose the following: William E. Whitehead received research grants from Ironwood Pharmaceuticals, Takeda Pharmaceuticals, Salix Pharmaceuticals, and the Rome Foundation; and is a consultant to Ono Pharma USA, Olafur S, Palsson received research grants from Ironwood Pharmaceuticals. Takeda Pharmaceuticals. Salix Pharmaceuticals. and the Rome Foundation. Hans Törnblom has served as consultant/advisory board member for Almirall and Allergan; and as a speaker for Tillotts, Takeda, Shire, and Almirall. Lukas Van Oudenhove has received grant support from Abide Therapeutics and Nestlé; and has given scientific advice to Grünenthal. Jan Tack has given scientific advice to Abide Therapeutics, AlfaWassermann, Allergan, Christian Hansen, Danone, Genfit, Ironwood, Janssen, Kiowa Kirin, Menarini, Mylan, Novartis, Nutricia, Ono Pharma, Rhythm, Shionogi, Shire, SK Life Sciences, Takeda, Theravance, Tsumura, Yuhan, Zealand, and Zeria pharmaceuticals; has received research grant or support from Abide Therapeutics, Shire, and Zeria; and has served on the speaker bureau for Abbott, Allergan, AstraZeneca, Janssen, Kiowa Kirin, Menarini, Mylan, Novartis, Shire, Takeda, and Zeria. Magnus Simrén has received unrestricted research grants from Danone and Ferring Pharmaceuticals; served as a consultant/advisory board member for AstraZeneca, Danone, Nestlé, Menarini, Almirall, Allergan, Albireo, Glycom, and Shire; and as a speaker for Tillotts, Takeda, Menarini, Allergan Shire, and Almirall. The remaining authors disclose no conflicts.

Supplementary Material

This supporting document presents exploratory factor analysis results separate for the 3 countries. Methods are identical to those described in the main text. Results are presented in Supplementary Tables 1–3. Briefly,

factors were highly similar across the individual countries. There emerged factors for IBS, constipation, diarrhea, abdominal pain, nausea and/or vomiting, and upper GI symptoms in all models. Heartburn and globus were part of the upper GI symptom factor in the US sample.

Supplementary Table 1. Factor Loadings in US Sample After Varimax Rotation

Item	Topic	Upper GI	Abdominal pain	Constipation	Upper GI II	Nausea/ vomiting	IBS	Diarrhea
1	Lump in throat	.54	.11	.19	.17	.11	.00	.08
3	Lump in throat when not eating	.33	.20	.14	.49	.10	.07	02
4	Pain with swallowing	.34	.33	.15	.27	.20	.07	.02
5	Chest pain	.59	.21	.22	.05	.13	.01	.12
7	Chest pain as burning sensation	.49	.17	.12	.28	.07	.22	.06
8	Chest pain and food sticking after swallowing	.40	.08	.11	.39	.18	.17	.00
9	Heartburn	.61	.09	.15	.12	.08	.13	.11
12	Foods stuck in chest	.63	.14	.16	.19	.16	.04	.10
14	Postprandial fullness	.46	.26	.22	.05	.23	02	.16
16	Early satiety	.40	.24	.27	.02	.28	.04	.17
18	Epigastric pain/burning	.60	.37	.17	.02	.24	.03	.13
19	Epigastric pain/burning improved by defecation	.08	.01	.13	.59	04	.37	.03
21	Nausea	.42	.31	.21	.03	.58	.08	.15
23	Vomiting	.42	.18	.13	.07	.66	.05	.08
25	Self-induced vomiting	.20	.05	.10	.54	.26	.02	.04
32	Food coming back up after swallowing	.59	.11	.14	.17	.33	.09	.14
34	Retching	.30	.24	.12	.33	.55	.12	.16
35	Vomiting when food comes back up	.22	.33	.21	.11	.62	.17	.10
37	Food coming back up is recognizable	.13	.19	.05	.50	.24	.11	.01
38	Belching	.49	.19	.13	.22	.25	.01	.12
40	Pain anywhere in the abdomen	.56	.44	.26	10	.13	.12	.20
41	Abdominal pain related to defecation	.02	.04	.18	.33	.00	.68	.16
42	Abdominal pain related to stool consistency	.08	.09	.23	.16	.11	.85	.15
43	Abdominal pain related to stool consistency Abdominal pain related to stool frequency	.13	.19	.25 .25	.14	.11	.74	.20
45 45	Abdominal pain related to a meal	.13	.35	.25 .15	.14	.12	.36	.20
46		.23	.59	.23	.00	.28	.24	.19
46 49	Abdominal pain limits usual activities Hard stools	.14 .21	. 59 .15	.∠ა .56	.00 .14	.∠o .11	.12	06
			.13					06 .04
51	Infrequent stools	.13		.55 .80	.12	.12	.12	
52	Straining during bowel movements	.22	.09		.09	.05	.23	.07
53	Incomplete bowel emptying	.25	.14	.72	.05	.11	.24	.21
54	Anorectal obstruction/blockage	.23	.20	.79	.07	.12	.14	.11
55	Manual maneuvers to facilitate defecation	.20	.23	.46	.22	.15	.02	.02
59	Loose stools	.29	.08	.08	.10	.15	.23	.75
61	Loose stools following a meal	.16	.20	.03	.22	.11	.20	.64
63	Urgency	.32	.19	.22	.15	.10	.18	.62
65	Bloating/abdominal distention	.39	.25	.41	03	.04	.14	.29
68	Right upper quadrant pain	.55	.49	.24	11	.15	.14	.20
69 70	Long-lasting right upper quadrant pain Right upper quadrant pain of steady severe	.28 .23	.67 .69	.24 .24	.03 .10	.12 .13	.10 .10	.08 .15
71	level Right upper quadrant pain at different intervals	09	09	06	.32	06	.04	.08
72	Right upper quadrant pain limits usual activities	.17	.68	.12	.10	.26	.06	.03
73	Right upper quadrant pain related to defecation	.05	.14	.24	.42	.06	.33	.18
74	Right upper quadrant pain related to posture	.06	.08	.08	.29	.06	.06	.09
75	Right upper quadrant pain improved by medicine	.25	.05	.07	.36	01	.06	.06
76	Right upper quadrant pain related to vomiting	.30	.36	.14	.25	.59	.04	.11
77	Right upper quadrant pain related to back pain	.31	.44	.20	.28	.08	.02	.03
78	Right upper quadrant pain waking one up	.33	.51	.10	.20	.20	.05	.14
80	Fecal incontinence	.38	.10	.13	.19	.15	.09	.30
83	Anorectal pain	.40	.17	.36	.17	.19	.12	.15
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NOTE. Loadings larger than 0.4 are shown in boldface. Gl, gastrointestinal; IBS, inflammatory bowel syndrome.

Supplementary Table 2. Factor Loadings in Canadian Sample After Varimax Rotation

Item	Topic	Upper GI	Abdominal pain	Constipation	IBS	Diarrhea	Nausea/ vomiting	Globus
1	Lump in throat	.41	03	.13	.04	.06	.07	.43
3	Lump in throat when not eating	.05	.09	.07	.15	.10	.03	.64
4	Pain with swallowing	.20	.12	.09	.12	.03	.05	.11
5	Chest pain	.54	.06	.17	02	.07	.09	.21
7	Chest pain as burning sensation	.11	.18	.07	.06	.06	.02	.16
8	Chest pain and food sticking after swallowing	.21	.00	.08	.11	.00	.08	08
9	Heartburn	.49	.10	.14	.05	.11	.02	.05
12	Foods stuck in chest	.54	.01	.16	.04	.11	.06	.04
14	Postprandial fullness	.56	.16	.14	.03	.11	.04	06
16	Early satiety	.53	.10	.17	.06	.12	.11	02
18	Epigastric pain/burning	.58	.34	.14	.05	.07	.03	.04
19	Epigastric pain/burning improved by defecation	.01	12	.12	.43	01	.06	.17
21	Nausea	.50	.34	.11	.08	.10	.32	.07
23	Vomiting	.33	.26	.10	.04	.04	.48	.06
25	Self-induced vomiting	08	.05	.02	09	.02	.08	.11
32	Food coming back up after swallowing	00 .53	.08	.17	.09	.10	.26	.09
34	Retching	. 33 .21	.00	.07	.03	.03	.49	.03
35	Vomiting when food comes back up	.17	.16	.18	.02	.03	. 49 .76	05
37	Food coming back up is recognizable	.17	.01	.01	.02	02	.05	03 .12
38	Belching	.12 .46	.10	.07	.03	02 .06	.03	05
40	Pain anywhere in the abdomen	. 40 .52	.10 .41	.07 .24	.03	.19	.03	05 .15
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41	Abdominal pain related to defecation	.01	.09	.14	.73	.16	.02	.08
42	Abdominal pain related to stool consistency	.10	.17	.19	.88	.19	.02	.11
43	Abdominal pain related to stool frequency	.13	.22	.21	.66	.25	.04	.02
45	Abdominal pain related to a meal	.18	.34	.09	.30	.28	05	.08
46	Abdominal pain limits usual activities	.23	.53	.18	.17	.17	.10	.05
49	Hard stools	.18	.06	.64	.12	.00	03	.09
51	Infrequent stools	.08	.21	.53	.01	.04	.14	.09
52	Straining during bowel movements	.21	.11	.80	.12	.04	.05	.10
53	Incomplete bowel emptying	.22	.22	.71	.16	.17	.11	.02
54	Anorectal obstruction/blockage	.23	.12	.82	.12	.09	.08	.05
55	Manual maneuvers to facilitate defecation	.11	.14	.56	.07	.01	.04	10
59	Loose stools	.19	.15	.04	.12	.74	.08	.07
61	Loose stools following a meal	.06	.11	.00	.16	.66	.00	.14
63	Urgency	.23	.20	.18	.14	.68	.09	02
65	Bloating/abdominal distention	.39	.20	.34	.14	.33	03	03
68	Right upper quadrant pain	.48	.40	.26	.05	.19	.04	.11
69	Long-lasting right upper quadrant pain	.06	.71	.15	.09	.14	.04	.10
70	Right upper quadrant pain of steady severe level	.12	.75	.17	.13	.17	.01	.02
71	Right upper quadrant pain at different intervals	12	14	08	.07	02	05	.02
72	Right upper quadrant pain limits usual activities	.14	.69	.13	.09	.08	.13	.04
73	Right upper quadrant pain related to defecation	.06	.14	.16	.58	.16	.06	.02
74	Right upper quadrant pain related to posture	.12	.07	01	.10	.01	.08	03
75	Right upper quadrant pain improved by medicine	.15	.06	.05	.02	.02	.07	06
76	Right upper quadrant pain related to vomiting	.16	.49	.05	.04	.05	.42	.06
77	Right upper quadrant pain related to back pain	.20	.46	.17	.06	.07	.09	04
78	Right upper quadrant pain waking one up	.16	.58	.16	.05	.09	.15	03
80	Fecal incontinence	.16	.10	.09	.14	.36	.10	03
83	Anorectal pain	.26	.15	.37	.13	.20	.05	.00

NOTE. Loadings larger than 0.4 are shown in boldface. Gl, gastrointestinal; IBS, inflammatory bowel syndrome.

Supplementary Table 3. Factor Loadings in UK Sample After Varimax Rotation

Item	Topic	Constipation		Abdominal pain		Diarrhea	Vomiting	Nausea	Heartburn	Abdomina pain II
1	Lump in throat	.18	.56	.03	.01	.10	04	02	02	.02
3	Lump in throat when not eating	.08	.14	.12	.07	.00	.03	.02	.11	.01
4	Pain with swallowing	.08	.35	.14	.05	.03	.18	.09	.20	.03
5	Chest pain	.21	.49	.05	04	.09	.14	.06	.07	.16
7	Chest pain as burning sensation	.03	.11	.15	.10	.03	.00	.00	.97	.01
8	Chest pain and food sticking after swallowing	.02	.44	.11	.17	.05	.05	.00	.27	16
9	Heartburn	.19	.40	.02	01	.14	.04	.07	.42	.15
12	Foods stuck in chest	.20	.68	.00	.05	.10	.05	.02	.11	.00
14	Postprandial fullness	.17	.52	.20	.08	.12	.18	.10	.02	.09
16	Early satiety	.20	.42	.22	.03	.13	.20	.13	.03	.07
18	Epigastric pain/burning	.15	.57	.22	.04	.10	.06	.22	.12	.26
19	Epigastric pain/burning improved by defecation	.11	.07	04	.44	01	.07	.05	.22	04
21	Nausea	.18	.40	.26	.02	.12	.19	.51	.01	.16
23	Vomiting	.10	.31	.11	.10	.08	.40	.39	.01	.10
25	Self-induced vomiting	.03	.03	10	.09	.09	.10	05	.07	.03
32	Food coming back up after swallowing	.16	.45	.04	.10	.13	.22	.25	.07	.08
34	Retching	.15	.19	.13	.10	.12	.68	.09	01	.00
35	Vomiting when food comes back up	.13	.19	.17	.07	.14	.79	.15	02	.06
37	Food coming back up is recognizable	.04	.10	.09	.13	.00	.13	.09	.01	.04
38	Belching	.07	.40	.20	.07	.15	.08	.30	.02	.11
40	Pain anywhere in the abdomen	.28	.29	.23	.09	.24	.09	.03	.01	.68
41	Abdominal pain related to defecation	.12	01	.05	.75	.11	.02	04	03	.03
42	Abdominal pain related to stool consistency	.20	.07	.07	.80	.25	.05	03	01	.09
43	Abdominal pain related to stool frequency	.23	.08	.08	.73	.29	.09	.06	.03	.08
45	Abdominal pain related to a meal	.04	.16	.32	.36	.27	.13	.14	.08	.07
46	Abdominal pain limits usual activities	.11	.17	.54	.29	.17	.10	.21	.05	.12
49	Hard stools	.66	.11	.10	.14	02	.06	.05	.06	.02
51	Infrequent stools	.57	.10	.11	.08	01	.14	.05	01	.07
52	Straining during bowel movements	.82	.13	.08	.14	.01	.07	.06	.04	.09
53	Incomplete bowel emptying	.73	.09	.06	.14	.24	.10	.04	.08	.14
54	Anorectal obstruction/blockage	.83	.17	.13	.14	.14	.06	.03	.04	.07
55	Manual maneuvers to facilitate defecation	.56	.17	.02	.05	.11	.00	.06	01	04
59	Loose stools	.10	.13	.07	.22	.70	.14	.02	.03	.11
61	Loose stools following a meal	.07	.08	.15	.22	.64	.04	.01	.06	04
63	Urgency	.19	.18	.16	.26	.62	.12	.08	01	.09
65	Bloating/abdominal distention	.39	.27	.19	.08	.27	08	.10	.05	.33
68	Right upper quadrant pain	.28	.32	.30	.07	.21	.11	.07	.07	.53
69	Long-lasting right upper quadrant pain	.15	.11	.74	.02	.02	.04	.08	.08	.14
70	Right upper quadrant pain of steady severe level	.15	.11	.83	.02	.12	.13	.05	.05	.00
71	Right upper quadrant pain at different intervals	06	21	16	.16	04	13	.03	01	02
72	Right upper quadrant pain limits usual activities	.11	.30	.58	.03	.14	.06	.29	04	.00
73	Right upper quadrant pain related to defecation	.15	.00	.08	.54	.13	.01	.08	05	02
74	Right upper quadrant pain related to posture	.22	.05	.08	.17	.06	02	.23	.08	10

Supplementary Table 3. Continued

Item	Topic	Constipation	- 1- 1	Abdominal pain		Diarrhea	Vomiting	Nausea	Heartburn	Abdominal pain II
75	Right upper quadrant pain improved by medicine	.02	.18	.01	05	.02	06	.10	.25	.00
76	Right upper quadrant pain related to vomiting	.14	.17	.27	.00	.11	.21	.62	.11	.00
77	Right upper quadrant pain related to back pain	.15	.23	.19	.10	.08	.09	.13	.02	.02
78	Right upper quadrant pain waking one up	.12	.26	.39	.04	.22	.15	.29	.00	.14
80	Fecal incontinence	.12	.15	02	.03	.44	.02	.15	.03	.14
83	Anorectal pain	.42	.24	.08	.09	.22	.00	.10	02	.21

NOTE. Loadings larger than 0.4 are shown in boldface.
Gl, gastrointestinal; IBS, inflammatory bowel syndrome. Loadings larger than 0.4 are shown in bold style.