### GASTROENTEROLOGY IN GERIATRIC PATIENTS (SITA CHOKHAVATIA)

# Constipation and Fecal Incontinence in the Elderly

Brototo Deb<sup>1</sup> • David O. Prichard<sup>1</sup> • Adil E. Bharucha<sup>1</sup>

Published online: 24 August 2020 © Springer Science+Business Media, LLC, part of Springer Nature 2020

### Abstract



**Purpose of Review** To review the epidemiology, pathogenesis, clinical features, and management of primary constipation and fecal incontinence in the elderly.

**Recent Findings** Among elderly people, 6.5%, 1.7%, and 1.1% have functional constipation, constipation-predominant IBS, and opioid-induced constipation. In elderly people, the number of colonic enteric neurons and smooth muscle functions is preserved; decreased cholinergic function with unopposed nitrergic relaxation may explain colonic motor dysfunction. Less physical activity or dietary fiber intake and postmenopausal hormonal therapy are risk factors for fecal incontinence in elderly people. Two thirds of patients with fecal incontinence respond to biofeedback therapy. Used in combination, loperamide and biofeedback therapy are more effective than placebo, education, and biofeedback therapy. Vaginal or anal insert devices are another option. **Summary** In the elderly, constipation and fecal incontinence are common and often distressing symptoms that can often be managed by addressing bowel disturbances. Selected diagnostic tests, prescription medications, and, infrequently, surgical options should be considered when necessary.

Keywords Defecatory disorder · Nursing home · Bowel leakage · Constipation · Elderly · Fecal incontinence

## Introduction

Constipation and fecal incontinence (FI) are common symptoms that may coexist and affect the quality of life in elderly people. This article reviews the burden, pathogenesis, and management of these symptoms with an emphasis on the elderly.

# Constipation

### Epidemiology

Constipation may be defined by self-report, symptom criteria, or based on laxative use. The Rome criteria categorize

Adil E. Bharucha bharucha.adil@mayo.edu constipated patients into four syndromes (i.e., functional constipation, constipation-predominant irritable bowel syndrome, defecatory disorders (DDs), and opioid-induced constipation (OIC) [1, 2]. Functional constipation (FC) and constipationpredominant IBS (IBS-C) are symptom-based diagnoses. DDs are defined by symptoms of FC or IBS-C and anorectal tests indicating impaired rectal evacuation. OIC is defined by worsening symptoms of constipation when initiating, changing, or increasing opioid therapy in patients who satisfy criteria for FC.

Among adults, the mean prevalence is approximately 14% with a range of 2–35% [3, 4]. In some but not all studies, chronic constipation was slightly more common in older people [4]. An internet-based survey from the USA, Canada, and the UK observed that constipation was *less* common in the 65+ group versus people aged 18–34 years [5•]. Whether this is explained by reduced internet access in the elderly is unknown. In that study, the prevalence of FC, IBS-C, and opioid-induced constipation diagnosed by Rome IV criteria in the 65+ age group was 6.5%, 1.7%, and 1.1% respectively. By contrast to these cross-sectional studies, the prevalence of self-reported constipation increased from 14 to 21% and laxative use increased from 6 to 15% over time among 239 elderly community residents in the Australian Longitudinal

This article is part of the Topical Collection on *Gastroenterology in Geriatric Patients* 

<sup>&</sup>lt;sup>1</sup> Division of Gastroenterology and Hepatology, Mayo Clinic, 200 1st Street, Rochester, MN 55905, USA

Study of Ageing [6]. By comparison to these communitybased studies, constipation is much common in institutionalized elderly adults with a prevalence ranging from 45 to 67% [7••].

The incidence of diagnosis of DD is associated with age [8]. In women, the incidence is greatest (57 per 100,000) between the age of 20–29 years, then declines with a second peak between the age of 80–89 years. In men, the incidence was less than or around 10 per 100,000 person-years until the age of 80–89 years, where it increased to 30 per 100,000 person-years.

Constipation is associated with an impaired quality of life (QOL) [9, 10]. The extent to which the impact on QOL is related to age and/or comorbid conditions (e.g., depression) is unknown.

### Pathophysiology

#### Effects of Age on the Enteric Nervous System

Phillips and Powley extensively reviewed the effects of aging on intrinsic and extrinsic (i.e., sympathetic and parasympathetic) innervation of the gastrointestinal tract [11]. In rodents, the age-associated loss of cholinergic neurons in the myenteric and submucosal plexus is more severe in the distal GI tract. By contrast, nitrergic neurons are spared. Aging is also associated with progressive accumulation of dystrophic axonal swellings and markedly dilated varicosities in the sympathetic, vagal, dorsal root, and enteric nitrergic innervation of the gut.

More recently, a landmark study compared enteric neuropathology and physiology, particularly the response to electrical field stimulation (EFS), in ex vivo colonic muscle strips obtained from 239 adult (35–60 years) or elderly ( $\geq$  70 years) individuals who had surgery for colon cancer [12..]. In older people, electrical stimulation was more likely to cause muscle relaxation rather than contraction in the ascending colon; the descending colon was not affected by age. By contrast to rodents, there was no loss of enteric cholinergic neurons and the density of intramuscular axon bundles, number of nitrergic neurons, and smooth muscle contraction and relaxation were also preserved. Hence, decreased cholinergic function with unopposed nitrergic relaxation probably explains the impaired contractile response to electrical stimulation. The number and volume of ICC networks also decline with age in the human stomach and colon [13]. Perhaps reduced cholinergicmediated contractile responses to stimulation predispose to acute colonic pseudo-obstruction in situations (e.g., after abdominal surgery) that are associated with sympathetic stimulation in older people.

In mice, aging is also associated with attenuated mechanosensation, more so for high-threshold than lowthreshold functions, and reduced chemosensory functions that are mediated by serotonin and transient receptor potential cation channel subfamily V member 1 (TrpV1) pathways [14••]. Perhaps reduced mechanosensation may explain why elderly people have a lower prevalence of painful diseases such as irritable bowel syndrome and have lesser awareness of gastrointestinal injury (e.g., with acute abdomen) [15].

### Effects of Aging on Colonic and Rectoanal Sensorimotor Functions

In small studies of patients with and without constipation, aging is not associated with slow colon transit [16, 17]. The effects of age on colon transit in *asymptomatic* healthy people are unclear. Increased age is associated with increased stiffness and reduced sensation in the colon and rectum, reduced anal resting, and to a lesser extent squeeze pressures, and increased perineal laxity [18, 19]. Increased age is also associated with longer and more polyphasic motor unit potentials, which may be markers of neurogenic injury, in nulliparous women [20]. Together, these changes may predispose to FI in elderly females [21••].

#### Pathophysiology

Colonic sensorimotor disturbances and pelvic floor dysfunction are the most widely recognized causes of chronic constipation; they may coexist [22...]. Other potential mechanisms are reduced caloric intake, disturbances of the microbiome, anatomical issues, and medications. Colonic motor dysfunction resulting from a loss of colonic nerves and/or interstitial cells of Cajal (ICC) is implicated to cause slow colonic transit constipation. However, normal and slow colonic transit may not necessarily reflect normal and impaired colonic motor function; indeed, intraluminal assessments of colonic motor function under fasting conditions, after a meal, and/or pharmacological stimuli suggest that some patients with normal transit have colonic motor dysfunctions and vice versa [23, 24]. Sensory disturbances, such as exaggerated perception of colonic distention, may explain abdominal pain and bloating, which are more common in patients who have normal transit constipation or IBS-C [25–27].

Germ-free mice colonized with fecal microbiome from constipated patients developed slow colonic transit suggesting that the colonic microbiome may also contribute to constipation [28, 29]. In humans, the colonic mucosal microbiome discriminated between constipated patients and controls with 94% accuracy even after adjusting for diet and colonic transit [30].

#### **Defecatory Disorders**

Defecation requires increased rectal pressure, which is generally secondary to increased abdominal pressure, coordinated with anal relaxation [31]. Defecatory disorders (DDs) result from abdomino-pelvic discoordination, which results in decreased rectal propulsive forces and/or increased resistance to evacuation [32]. Other disturbances (e.g., reduced rectal sensation [33, 34]) and anatomical abnormalities (e.g., large rectocele) may contribute. In the elderly, excessive straining can weaken the pelvic floor increasing the risk for excessive perineal descent [33], rectal intussusception [35], solitary ulcer syndrome [36], and pudendal neuropathy which in turn can increase the risk of FI [37].

### **Clinical Features**

Chronic constipation may be a primary symptom or secondary to another disease; the vast majority of patients have primary constipation. Similar to younger people, female sex, medications, especially opioids, lower socioeconomic status, less, selfreported activity, malnutrition, and depression are associated with idiopathic constipation in elderly [38–41]. While older people are more likely to have diseases that cause constipation, the proportion of older patients who have secondary constipation is unknown (Table 1). Of note, in Parkinson's disease, the constipation may precede motor symptoms by 20 years or longer [42].

The symptoms include infrequent stool passage (fewer than 3 bowel movements per week) and, more commonly, straining at stool, feeling of incomplete evacuation, need for digital assistance during defecation, bloating, and passing hard, lumpy stools [43]. The Rome IV defines functional constipation (FC) by the presence of constipation for at least 6 months and two or more symptoms for 25% of bowel movements for the last 3 months. By contrast, IBS-C is defined by the presence of abdominal pain that is associated with 2 of the

 Table 1
 Common medical

 conditions associated with
 constipation

features: altered stool form, altered frequency, or relief of abdominal pain with defecation.

Since patients with FC and IBS-C may have abdominal pain, it can be difficult to distinguish between them [44, 45]. Alternatively, constipated patients may be categorized as painless (or mild pain) and painful constipation. Compared to "mild pain" constipation, patients with "painful" constipation had more prominent bowel symptoms and were more likely to have upper gastrointestinal (e.g., dysphagia and dyspepsia) and anorectal symptoms, urinary and sexual symptoms, anxiety and depression, and slower rectosigmoid transit.

### **Diagnostic Approach**

The history will identify alarm symptoms such as new onset rapidly progressive symptoms, rapid weight loss, hematochezia, or features of colonic obstruction [22, 41]. A digital rectal examination is essential to assess for a rectal mass, anal fissure, stool in the rectal vault, tenderness to palpation of the levator ani, and anal sphincter tone at rest, during squeeze, and evacuation. A complete blood count, age-appropriate screening for colon cancer, and, based on the clinical features, additional tests (e.g., thyroid functions, metabolic parameters) should be performed (Fig. 1). Thereafter, a therapeutic trial of fiber supplementation and/or osmotic or stimulant laxatives should be considered. Nonresponders who are candidates for pelvic floor biofeedback therapy should undergo anorectal manometry (ARM) and a rectal balloon expulsion tests (BET) to identify defecatory disorders followed by barium or MR defecography if necessary. The results of the colonic transit study classify patients into NTC and STC [22••].

Cause	Comments
Drug effects	
Mechanical obstruction: colon cancer, external compression from malignant lesion, strictures (diverticular or post ischemic), rectocele (if large), megacolon, anal fissure	Often associated with alarm clinical features or laboratory tests, apparent on digital rectal examination (fissure) or X-ray image of the abdomen, or preceded by the primary event (diverticulitis).
Metabolic conditions: diabetes mellitus, hypothyroidism, hypercalcemia, hypokalemia, hypomagnesemia, uremia, heavy metal poisoning, uremia, heavy metal poisoning	All are associated with/can be diagnosed by abnormal results from laboratory tests, which should be performed only when there is a high index of suspicion (such as in patients on diuretics).
Myopathies: amyloidosis, scleroderma	Typically associated with other clinical features of these conditions.
Neuropathies: Parkinson's disease, spinal cord injury or tumor, cerebrovascular disease, and multiple sclerosis	Constipation, either due to slow colon transit and/or DD, is common in patients with these disorders, which have many other features.
Other conditions: depression, degenerative joint disease, autonomic neuropathy, cognitive impairment, immobility, cardiac disease	The disorder and/or medications may contribute to constipation.

Reproduced with permission from [22..]



### **Medical Management**

A stepwise approach is recommended [22...]. Constipating medications should be discontinued where possible. Lifestyle modifications should be implemented diligently. Adequate exercise and fluid intake are generally beneficial but their effects on constipation are unclear [46]. When possible, dietary fiber intake, which is often deficient in elderly individuals, should be increased [47]. Fiber (e.g., fresh fruit and vegetables, legumes, whole grains, or a supplement) should be introduced at low doses and increased slowly to 25-30 g fiber day in an attempt to reduce the incidence of flatulence, bloating, and abdominal pain [48, 49]. In elderly patients, dietary fiber accelerates intestinal transit but the effects on symptoms are inconsistent [50, 51]. A prolonged trial is required because a response may take several weeks. At least in younger people, failure to respond to fiber may suggest the presence of slow transit constipation or a DD [52].

If lifestyle interventions are ineffective, medications are necessary; osmotic or stimulant laxatives are first-line therapy. These agents are at least as effective and less expensive than newer agents for treating chronic constipation in adults (Table 2) [22..]. Polyethylene glycol (PEG), lactulose, and sorbitol have been studied in elderly populations (Table 2). PEG, administered for 6 months to patients older than 65 years, is safe, not associated with malabsorption, electrolyte abnormalities, or micronutrient deficiencies, and more effective than lactulose [54]. Osmotic agents are associated with diarrhea, distension, flatulence, and bloating, which usually resolve with a lower dose [54, 55]. Senna products are more effective and as well tolerated as lactulose or placebo [58, 68]. Bisacodyl is effective but its effects in elderly patients have not been characterized [69]. A practical approach is to administer an osmotic laxative regularly, supplemented as required with a stimulant laxative suppository. The suppository should preferably be administered 15-30 min after the morning meal to coincide with the gastrocolonic reflex.

Among the newer drugs, the strongest data regarding efficacy in the elderly is for the 5-HT<sub>4</sub> receptor agonist prucalopride, which improved symptoms and quality of life in a 4-week study of 300 constipated patients over 65 years old [60]. The 1 and 2 mg doses were equally effective. The

	Number needed to treat in adult patients with CC and IBS-C	Cost per month (US\$, 2019)	Studies or analyses in patients over 65 years of age	Comments regarding patients over 65 years old
Bulking agents	CC: 2 (95% CI, 1–3) [53] IBS-C: 10 (95% CI, 6–33) [53]	\$8.34	Yes	Variable effect on symptoms of constipation [50, 51]. Failure to respond may suggest STC or DD.
Polyethylene glycol	CC: 3 (95% CI, 2–4) [2] IBS-C: NA	\$30.90	Yes	Effective at relieving symptoms of constipation in patients over 65 [54, 55]. PEG 400 is more palatable without than with electrolytes [56].
Lactulose and sorbitol	NA	\$11.20	Yes	Both may improve symptoms of constipation; no comparison with baseline bowel function is reported [57]. Nausea is more common with lactulose than sorbitol. Lactulose is inferior to PEG 4000 or senna with fiber [54, 58].
Bisacodyl	CC: 4 (95% CI, NA) [59]	\$5.17	No	No data.
Senna	1D5-C. NA	\$5.90	Yes	Long-stay elderly patients receiving a combination of senna with fiber experienced more frequent and softer bowel movements, passed with more ease, than patients receiving lactulose. However, drug effects were not compared with baseline [58].
Prucalopride	CC: 6 (95% CI, 5–9) [2] IBS-C: NA	\$395.67	Yes	Increases bowel motion frequency [60] without adverse cardiovascular events [60, 61].
Linaclotide	CC: 72 µg, 12 (95% CI, 6–29) 145 µg, 10 (95% CI, 6–19) [62] IBS-C: 290 µg, 6 (95% CI, 4–16)	\$395.41	No	No data.
Lubiprostone	CC: 24 µg, 4 (95% CI, 3–6) [53] IBS-C: 8 µg, 12 (95% CI, 8–25)	\$342.92	No	Abstracts suggest that lubiprostone is as effective and safe in patients older versus younger than 65 years [63, 64]. Patients over 65 are less likely to experience nausea [(65], [64)] or to discontinue lubiprostone [66]. Age is not associated with a higher likelihood of requiring diose adjustment [67].
Plecanatide	CC: 3 mg, 11 (95% CI, 8–19) 6 mg, 12 (95% CI, 8–23) [62] IBS-C: 3 mg, 9 (95% CI, 6–16); 6 mg, 9 (95% CI, 6–17)	\$384.36	No	No data.
Tenapanor	IBS-C: NA	NA	No	No data.

#### Table 2 Treatment of constipation treatment

NA not available

European Medicines Agency recommends a starting dose of 1 mg in patients over 65 years [70]. In contrast, the Food and Drug Administration in the USA recommends dose adjustment only in the setting of impaired renal function [71]. No cardiac side effects have been associated with prucalopride [60].

Abstracts suggest that safety and efficacy of lubiprostone are comparable in the elderly [63, 64]. Indeed, elderly patients were less likely to experience nausea [64, 65] or discontinue the medication [66]. There are no studies or analyses specifically describing the efficacy or safety of linaclotide, plecanatide, or tenapanor, a small-molecule inhibitor of the gastrointestinal sodium-hydrogen exchanger-3, in older populations. Two phase 3 trials of linaclotide in patients with constipation-predominant IBS, which were subsequently pooled for analyses, contained only 40 [72] and 45 [73] patients over 65 years old. In the largest single trial of linaclotide, comparing placebo, 72 µg linaclotide, and 145 µg linaclotide, there were 39 (10%), 36 (9%), and 43 (11%) patients over 65 years old in each cohort [74•]. Although not specifically reported, the phase 3 trials of plecanatide [75–77] and tenapanor [78•] appear to have few elderly patients. Thus, even pooled, retrospective analyses of this population are likely to be underpowered. The 5-HT<sub>4</sub> receptor agonist tegaserod is effective for treating constipation and constipationpredominant IBS and approved by the FDA for the latter [79, 80]. In comparative studies, tegaserod was inferior to PEG 3350 for constipation [81]. Efficacy trials enrolled 331 elderly patients (approximately 13% of participants) in studies regarding constipation, but no subgroup analyses were performed [82]. No dose adjustments are needed in the elderly or in patients with chronic kidney disease [83, 84]. The medication was withdrawn from market in the USA for many years due to cardiovascular events [53, 62]. After additional safety studies, it was reintroduced for treating adult women less than 65 years of age with irritable bowel syndrome with constipation [85].

Phosphate or tap water enemas or transanal irrigation are often useful on an as needed basis. Transanal irrigation, in which a device is used to irrigate the colon, is useful in retrospective studies [59, 86, 87]; prospective studies are underway. There is insufficient evidence to recommend the use of probiotics such as *Bifidobacterium longum* to manage constipation in the elderly [88].

### **Pelvic Floor Retraining**

DDs are managed with biofeedback-aided therapy that promotes relaxation of the pelvic floor muscles, coordinated with abdominal effort, during defecation [89]. Undertaken frequently and with motivation on the part of the patient and therapist, biofeedback therapy (BFT) can provide symptomatic benefit in the majority of adult patients with DD; five to six sessions, each 30–60-min long, at 2-week intervals, has demonstrated superiority to PEG [90], sham feedback [91], or diazepam [92]. A single small study suggests that BFT is effective in elderly patients with a DD [93•]. Irrespective of age, successful outcomes require adequate physical and mental capacity. Therefore, not all patients are suitable for BFT. BFT is underused because expertise is not widely available and the third-party coverage is suboptimal [94].

### **Surgical Management**

Patients with refractory symptoms despite discontinuation of constipating medications, a 4-week trial of pharmacologic therapy with each laxative and a 3-month trial of BFT if a DD was present, are considered for surgical therapy, i.e., a colectomy with ileorectal anastomosis [95, 96]. In patients with predominant pain or bloating, prior to a colectomy, a temporary loop ileostomy might be useful to determine if symptoms originate from the small intestine or colon; outcomes after colectomy are poorer in patients with generalized intestinal dysmotility [97].

Prior to surgery, rectoanal function and colonic transit, on laxatives, should be reevaluated. A DD, whether identified by manometry, BET, or defecography, is a contraindication for colectomy and warrants further BFT [22••]. Structural abnormalities identified on defecography may respond to targeted surgery if they obstruct defecation; we must strongly emphasize that pelvic organ prolapse is frequently encountered in health and rarely underlies constipation [98, 99].

Among adult patients undergoing colectomy with ileorectal anastomosis in order to manage constipation, approximately 88% report being "satisfied" or "very satisfied" with the outcome [95]. Perioperative complications, recurrent small bowel obstruction, and mortality occur in 20-30%, 10-20%, and 0.4% of patients. Other functional complications including diarrhea and incontinence, abdominal pain, recurrent constipation, and bloating remain or recur in 5-15%, 30-50%, 10-30%, and 10-40% of patients respectively [95]. Outcomes in elderly patients have not been described. However, prospective studies of oncologic colorectal surgery suggest that age is associated with higher complication rates and is independently associated with a higher 1-year overall and cardiovascular-specific mortality [100, 101]. Thus, especially in the elderly, surgical management should not be considered lightly. Patients should be counseled appropriately with regards postoperative morbidity and the risk of persistent symptoms.

### **Fecal Incontinence**

## Epidemiology

FI is the involuntary passage of solid or liquid feces; anal incontinence also includes the loss of flatus. The prevalence in the community increases with age, for example, from 2.6% in the third decade to 15% in the eighth decade [102, 103]. Among elderly individuals who do not reside at home, the prevalence is higher, i.e., 18-33% in hospitals, 38% in home health, and 50-70% in nursing homes [104–106]. In a retrospective study of 15,432 patients served by a hospice agency mostly 75 years or older, most patients with FI complained of incontinence at the time of the hospice admission [107•]. There is mixed evidence as to whether FI contributes to institutionalization independent of cognitive impairment in the elderly [108]. Twenty percent of initially continent individuals develop FI within 10 months of admission to a nursing perhaps due to comorbid conditions and low functional status [109].

FI can profoundly impair the quality of life in several domains, especially those that involve eating or leaving home, and can be associated with considerable psychosocial distress [110]. In a retrospective analysis of 41,932 communitydwelling older adults aged 65 and older who required some assistance, FI was associated with an increased likelihood of death (i.e., a hazard ratio 1.28) [111]. In this context, FI probably represents a marker of underlying frailty rather than the cause of death.

#### Associated Factors

1 The risk factors and associated conditions are listed in Table 3 [103, 112, 113, 117, 119–122]. While obstetric anal sphincter injury can cause FI, most community women develop FI in the seventh decade. Among women in the community, rectal urgency and diarrhea are the strongest and independent risk factors for FI [113, 123, 124]; obstetric anal sphincter injury is not independently significant after adjusting for other risk factors [113, 121]. Rectal urgency was also associated with severity of FI [125]. Risk factors that are more germane to older individuals include dementia [107, 109, 115, 126], impaired mobility [127, 128], comorbid conditions [109, 114], polypharmacy [116], depression symptoms, and poor self-related health [129]. Conversely, in the Nurses' Health Study, more physical activity and increased long-term dietary fiber intake were associated with a modest reduction in FI [130•].

#### Pathophysiology

Fecal continence is maintained by the internal and external anal sphincters and the puborectalis muscle of the levator ani complex, rectal compliance, and rectoanal sensation [125]. Bowel disturbances typically diarrhea, anal sphincter weakness due to obstetric or iatrogenic trauma, or neurogenic causes like pudendal neuropathy, reduced rectal compliance, and/or altered rectal sensation cause FI [131, 132]; many patients have more than one disturbance [133]. The structural injury is documented by imaging [134]. Anal weakness is assessed by measuring the anal

Table 3 Fee	cal incontinence-	-associated	conditions	and risk	factors

In population-based and/or case-control studies

- Old age \* [103, 107, 109, 112]
- Diarrhea \* [103, 113, 114]
- Rectal urgency
- Multiple chronic comorbidities \* [103, 112, 114]
- Urinary incontinence \* [103, 109, 112-114]
- Poor self-related health \* [103, 114–116]
- Major depression \* [112, 114, 116, 117]
- Obesity [113]
- Current smoking [113]
- Stroke and other neurological diseases \* [107, 109, 114, 117]
- Dementia, cognitive disorders, and functional limitation with poor mobility \* [107, 109, 115]
- Prostate diseases with prostatectomy and radiation for prostate cancer \* [114, 115]
- Hysterectomy with oophorectomy\* [114]
- Not in population-based and/or case-control studies
- Type 2 diabetes mellitus\* [117]
  Chronic kidney disease \* [115]
- Multinomity \* [117]
- Multiparity \* [117]Polypharmacy \* [116]
- Menopausal hormonal therapy \* [118]

resting and squeeze pressure, reflecting IAS and EAS function with a digital rectal examination [135] or with manometry (HR-ARM) [136, 137]. During manometry, rectal sensation [138], which can be increased, reduced, or normal in FI, can also be assessed [139, 140]. When decreased, it leads to a weakened reflex for the sphincters and the pelvic muscles to contract when the rectum is distended, leading to FI. Conversely, rectal hypersensitivity can be secondary to a reduced rectal compliance or capacity and/or capacity, partly explaining the symptom of urgency [141, 142]. IBS is associated with rectal hypersensitivity [143]. Compared to women, anal sphincter dysfunction is less common whereas impaired rectal sensation and compliance were more common in men [144]. In conclusion, old age affects all the mechanisms which maintain continence, lending support to the observation that aging is an important risk factor of FI (Table 4).

#### **Clinical Features**

A substantial proportion of FI patients, approximately 50% in some studies, do not share the symptom with a physician, indeed not even with friends or close relatives [102, 145]. Hence, older people should always be asked if they have FI. Among four characteristics are necessary to assess the severity of FI: frequency, type of stool, urgency, and amount of stool [21, 110]. So characterized, the severity of FI is strongly correlated with the QOL-weighted symptom severity score [110]. The severity of FI is however not related to age.

### Management

The goals are to reduce the physical and social consequences of FI [146, 147]. The critical step is to identify FI through direct enquiry. Thereafter, management progresses in a stepwise manner (Fig. 2).

### Lifestyle Management

Toileting advice, dietary triggers, and pads and undergarments should be thoroughly discussed with patients (Fig. 2) [148]; their efficacy should be emphasized. Elderly women acknowledge the association between diet and FI and are willing to modify their personal triggers [149, 150]. Smokers should be counseled to quit. Adequate fiber consumption should be promoted for prevention [151, 152] and treatment [153, 154]. Therapeutically, psyllium, but not carboxymethylcellulose, guar gum, or placebo, reduced the frequency of FI episodes but did not affect quality of life [153••]. In a placebo-controlled crossover study (68% male), psyllium was as effective as loperamide for reducing FI episodes but was associated

<sup>\*</sup> Risk factors apply to studies done in an elderly population

### Table 4 Components of a comprehensive history in fecal incontinence

Question	Rationale			
Elucidate whether a patient has incontinence	Patients may not volunteer the symptom spontaneously			
Onset, natural history, and risk factors	<ul> <li>Relationship of symptom onset/deterioration to other risk factors may suggest etiology</li> <li>Natural history, e.g., recent symptomatic deterioration may reveal reason for seeking medical attention</li> </ul>			
Bowel habits/type of leakage	<ul> <li>Disordered bowel habits are critical to pathogenesis of incontinence</li> <li>Incontinence for solid stool suggests more severe sphincter weakness than for liquid stool</li> <li>Management should be tailored to specific bowel disturbance</li> </ul>			
Degree of warning before incontinence	<ul> <li>Urge and passive incontinence are associated with more severe weakness of the external and internal anal sphincter respectively</li> <li>These symptoms may also reflect rectoanal sensory disturbances, potentially amenable to biofeedback therapy</li> </ul>			
Diurnal variation in incontinence	<ul> <li>Nocturnal incontinence occurs uncommonly in idiopathic fecal incontinence and is most frequently encountered in diabetes and scleroderma</li> </ul>			
Impact of FI on quality of life	Critical to ascertain severity of incontinence			
Urinary incontinence-presence and type	<ul><li>Association between urinary and FI</li><li>Same therapy may be effective for both conditions</li></ul>			
Evaluate possible causes for incontinence	<ul> <li>Most conditions listed in Table 2 are associated with other, i.e., non-anorectal manifestations</li> <li>The obstetric history must inquire specifically for known risk factors for pelvic trauma, e.g., forceps delivery, episiotomy, and prolonged second stage of labor</li> <li>Medications, including laxatives and artificial stool softeners, may cause or exacerbate incontinence</li> </ul>			

Reproduced with permission from [119]

with fewer side effects [154]. However, the effect was maintained during the washout phase in both groups

**Fig. 2** Algorithm for managing fecal incontinence

suggesting an effect from factors unrelated to the study interventions.

1. Identify patients with fecal incontinence		
2. Identify,	investigate and treat	: FI risk factors (e.g. diarrhea, urgency)
	History:	Adjust medications
	Investigation:	Laboratory, stool tests, endoscopy and biopsy, etc
3. Exclude of	defecatory disorder /	constipation with overflow
	History Examination: Investigation:	Pelvic floor dynamics and stool in rectum Anorectal manometry and possibly abdominal X-Ray
3. Educatio	n and lifestyle modif	ications
Simple dietary interventions (e.g., caffeine and trigger-food avoidance, fiber) Use pads for protection Squeeze before you sneeze, laugh, or lift Schedule times during the day to have a bowel movement Clean, squeeze, reclean technique Walk, don't run, to the toilet Practice delaying your bowel movements Squeeze exercises		
5. Initiate targeted therapy for FI		
	Medications to manag Pelvic floor biofeedbac Anal insert devices	e bowel disturbances :k therapy
6. Consider	6. Consider advanced interventions	

#### **Treatment of Constipation**

This is summarized in the section on constipation. Between 13 and 36% of patients of FI have a DD, which should be managed with BFT [144]. When BFT is not possible, medical therapy may be helpful. In a cohort of elderly constipated patients with cognitive and physical impairment and impaired rectal evaluation, neither lactulose nor the combination of lactulose, a daily glycerin suppository, and weekly tap water enemas improved FI [155]. However, FI improved in the subgroup of patients with adequate rectal clearance (i.e., no stool in the rectal vault on repeated examinations). In contrast, compared to placebo, lactulose reduced the number of days with FI in elderly patients with uncharacterized constipation [156].

### **Medical Management**

Osmotic laxatives should be discontinued if loose stool or urgency is present. Loperamide, an opioid receptor agonist, reduced urgency and increased stool consistency and anal resting pressure in patients with both normal [157–159] and surgically altered anatomy [160]. Diphenoxylate plus atropine reduced stool weight and frequency versus placebo; however, the duration of treatment (i.e., 3 days each with drug and placebo) was insufficient to assess the effects on fecal continence [161]. Loperamide was equivalent to codeine and superior to diphenoxylate plus atropine for reducing urgency and stool frequency in patients with diarrhea [162, 163]. However, the effects in patients with FI have not been studied. More recent studies suggest that loperamide is equivalent to psyllium [154] or placebo, education, and BFT [164•] for the treatment of FI. Colesevelam, cholestyramine, and amitriptyline improved fecal continence in small uncontrolled studies [165-167]. Clonidine did not improve continence in women with urge predominant FI; however, clonidine reduced diarrhea and tended to improve fecal continence in patients with diarrhea [168]. The main side effect of these therapies is constipation. This can usually be overcome by use of a stimulant laxative or enema if no bowel motion has been passed in 48-72 h. The average age of patients in the majority of these studies is approximately 60 years.

### Biofeedback

Conceptually, biofeedback therapy (BFT) seeks to improve pelvic floor muscle contraction in response to rectal distension, maintain sustained contraction, and address rectal sensory abnormalities [89]. BFT was not superior to education [169] or pelvic floor exercises taught by digital rectal examination [170, 171]. However, it was superior to exercises taught verbally in patients who had failed medical therapy [172]. Follow-up studies suggest that two thirds of patients respond to BFT and half maintain a durable response [173•]. Age is associated with a greater likelihood of completing BFT but whether older age is associated with response to BFT is unclear [174, 175]. Therapy provided remotely or in secondary care has shown equal efficacy to care provided in tertiary care institutions and could facilitate greater access in the future [176, 177]. Used in combination, loperamide and BFT were more effective than placebo, education, and BFT [178•]; loperamide, fiber, and BFT were more successful than the individual therapies [179]; and cholestyramine and BFT were more beneficial than BFT alone [166]. However, in a randomized study comparing (1) placebo and education; (2) placebo, education, and BFT; (3) loperamide and education; and (4) loperamide, education, and BFT, no intervention group demonstrated superiority [164•].

### **Barrier Devices**

In a multicenter, prospective, open-label study of the Renew anal insert, 62% achieved a  $\geq$  50% reduction in FI frequency in an intention-to-treat analysis [180]; 78% of patients who completed that study were very or extremely satisfied with the device and no serious adverse events occurred. In an openlabel study of a vaginal insert and pressure-regulated pump, 61 of 110 participants successfully completed the fitting period. Of these, 79% achieved treatment success, defined as greater than a 50% reduction of incontinent episodes at 1 month [181, 182]. These devices may be an effective treatment option for patients who fail standard conservative or surgical therapy.

### **Perianal Injection of Bulking Agents**

When compared to sham injections in 206 patients with moderately severe FI, patients receiving dextranomer were more likely to experience a 50% reduction in FI episodes (odds ratio 2.36, 95% CI 1.24–4.47) [183]. The beneficial effects on symptoms and quality of life were observed at 36 months [184]. However, anal bulking agents were not superior to anorectal BFT in FI patients [185].

### Surgery

The most widely used option, sacral nerve stimulation, reduced the frequency of FI and improved quality of life in open-label studies [186, 187]. However, the mechanisms underlying these benefits are unclear [188]. The largest study of 120 implanted patients demonstrated that 83% of subjects had a > 50% reduction in FI episodes and that 41% achieved complete continence [189]. Efficacy persisted after 5 years in the 76 patients (63%) evaluated

[190]. Complications were frequently; among the 120 patients who initially underwent implantation, 33% experienced implant site pain, 19% experienced paresthesia, 12% experienced a "change in sensation of stimulation," 10% experienced implant site infection, and 8% experienced urinary incontinence [190]. Furthermore, 10 patients (8%) underwent 10 device revisions, 29 (24%) patients underwent 40 device replacements (12 of which were for battery depletion), and 22 patients (18%) had the device removed (11 of which were for lack of efficacy) [190]. Because sham stimulation is associated with clinically and statistically meaningful improvement, more controlled studies are necessary [187...]. Age did not substantially influence the outcomes after SNS for FI [191, 192]. Percutaneous tibial nerve stimulation (PTNS) was not better than sham electrical stimulation [193..] and was comparable to SNS in a small study [194]. The effects of anal sphincteroplasty are not durable [109]. The artificial anal sphincter has been abandoned due to high complication rates [195...]. A colostomy is the last resort for patients with FI [196].

### **Compliance with Ethical Standards**

**Conflict of Interest** Dr. Bharucha has a patented anorectal manometry device with royalties paid to Medspira Inc, a patented anorectal catheter fixation clip pending to Medtronic, and a patented anal insert device for fecal incontinence pending to Minnesota Medical Technologies. The other authors declare that they have no competing interests.

Human and Animal Rights and Informed Consent This article does not contain any studies with human or animal subject performed by any of the authors.

### References

Papers of particular interest, published recently, have been highlighted as:

- Of importance
- •• Of major importance
  - 1. Mearin F, Lacy BE, Chang L, Chey WD, Lembo AJ, Simren M, et al. Bowel disorders. Gastroenterology. 2016;18:18.
  - Rao S, Bharucha AE, Chiarioni G, Felt-Bersma RJ, Knowles CH, Malcolm A, et al. Functional anorectal disorders. Gastroenterology. 2016;150(6):1430–42.
  - Mugie SM, Benninga MA, Di Lorenzo C. Epidemiology of constipation in children and adults: a systematic review. Best Pract Res Clin Gastroenterol. 2011;25(1):3–18.
  - Suares NC, Ford AC. Prevalence of, and risk factors for, chronic idiopathic constipation in the community: systematic review and meta-analysis. Am J Gastroenterol. 2011;106(9):1582–91 quiz 1, 92.

- 5•. Palsson OS, Whitehead W, Tornblom H, Sperber AD, Simren M. Prevalence of Rome IV functional bowel disorders among adults in the United States, Canada, and the United Kingdom. Gastroenterology. 2020. Recent multinational study on epidemiology of functional bowel disorders.
- 6. Werth BL, Williams KA, Pont LG. A longitudinal study of constipation and laxative use in a community-dwelling elderly population. Arch Gerontol Geriatr. 2015;60(3):418–24.
- 7••. Lamas K, Karlsson S, Nolen A, Lovheim H, Sandman PO. Prevalence of constipation among persons living in institutional geriatric-care settings—a cross-sectional study. Scand J Caring Sci. 2017;31(1):157–63. Prevalence of constipation in elderly people in geriatric-care.
- Noelting J, Eaton J, Choung RS, Zinsmeister AR, Locke GR 3rd, Bharucha AE. The incidence rate and characteristics of clinically diagnosed defecatory disorders in the community. Neurogastroenterol Motil. 2016;28(11):1690–7.
- Dennison C, Prasad M, Lloyd A, Bhattacharyya SK, Dhawan R, Coyne K. The health-related quality of life and economic burden of constipation. Pharmacoeconomics. 2005;23(5):461–76.
- Belsey J, Greenfield S, Candy D, Geraint M. Systematic review: impact of constipation on quality of life in adults and children. Aliment Pharmacol Ther. 2010;31(9):938–49.
- Phillips RJ, Powley TL. Innervation of the gastrointestinal tract: patterns of aging. Auto Neuroscience-BasiC Clin. 2007;136(1–2): 1–19.
- 12••. Broad J, Kung VWS, Palmer A, Elahi S, Karami A, Darreh-Shori T, et al. Changes in neuromuscular structure and functions of human colon during ageing are region-dependent. Gut. 2019;68(7):1210–23. The most recent and comprehensive study on the effects of aging in the human colon.
- Gomez-Pinilla PJ, Gibbons SJ, Sarr MG, Kendrick ML, Shen KR, Cima RR, et al. Changes in interstitial cells of Cajal with age in the human stomach and colon. Neurogastroenterol Motil. 2011;23(1): 36–44.
- Keating C, Nocchi L, Yu Y, Donovan J, Grundy D. Ageing and gastrointestinal sensory function: altered colonic mechanosensory and chemosensory function in the aged mouse. J Physiol Lond. 2016;594(16):4549–64.
- Lovell RM, Ford AC. Global prevalence of and risk factors for irritable bowel syndrome: a meta-analysis. Clin Gastroenterol Hepatol. 2012;10(7):712–21.e4.
- Melkersson M, Andersson H, Bosaeus I, Falkheden T. Intestinal transit time in constipated and non-constipated geriatric patients. Scand J Gastroenterol. 1983;18(5):593–7.
- Merkel IS, Locher J, Burgio K, Towers A, Wald A. Physiologic and psychologic characteristics of an elderly population with chronic constipation. Am J Gastroenterol. 1993;88(11):1854–9.
- Fox JC, Fletcher JG, Zinsmeister AR, Seide B, Riederer SJ, Bharucha AE. Effect of aging on anorectal and pelvic floor functions in females. Dis Colon Rectum. 2006;49(11):1726–35.
- Odunsi ST, Camilleri M, Bharucha AE, Papathanasopoulos A, Busciglio I, Burton D, et al. Reproducibility and performance characteristics of colonic compliance, tone, and sensory tests in healthy humans. Dig Dis Sci. 2010;55(3):709–15.
- Bharucha AE, Daube J, Litchy W, Traue J, Edge J, Enck P, et al. Anal sphincteric neurogenic injury in asymptomatic nulliparous women and fecal incontinence. Am J Physiol Gastrointest Liver Physiol. 2012;303(2):G256–62.
- 21••. Bharucha AE, Dunivan G, Goode PS, Lukacz ES, Markland AD, Matthews CA, et al. Epidemiology, pathophysiology, and classification of fecal incontinence: state of the science summary for the National Institute of Diabetes and Digestive and Kidney Diseases (NIDDK) workshop. Am J Gastroenterol. 2015;110(1):127–36. Systematic review on epidemiology, pathophysiology and classification of fecal incontinence.

- 22••. Bharucha AE, Lacy BE. Chronic constipation: mechanisms, evaluation and management. Gastroenterology. 2020;158(5):1232–49. Comprehensive updated review on chronic constipation.
- Bassotti G, de Roberto G, Chistolini F, Sietchiping-Nzepa F, Morelli O, Morelli A. Twenty-four-hour manometric study of colonic propulsive activity in patients with diarrhea due to inflammatory (ulcerative colitis) and non-inflammatory (irritable bowel syndrome) conditions. Int J Color Dis. 2004;19(5):493–7.
- Ravi K, Bharucha AE, Camilleri M, Rhoten D, Bakken T, Zinsmeister AR. Phenotypic variation of colonic motor functions in chronic constipation. Gastroenterology. 2010;138(1):89–97.
- Mertz H, Naliboff B, Mayer EA. Symptoms and physiology in severe chronic constipation. Am J Gastroenterol. 1999;94(1):131–8.
- Posserud I, Syrous A, Lindstrom L, Tack J, Abrahamsson H, Simren M. Altered rectal perception in irritable bowel syndrome is associated with symptom severity. Gastroenterology. 2007;133(4):1113–23.
- Manabe N, Wong BS, Camilleri M, Burton D, McKinzie S, Zinsmeister AR. Lower functional gastrointestinal disorders: evidence of abnormal colonic transit in a 287 patient cohort. Neurogastroenterol Motil. 2010;22(3):293–e82.
- Cao H, Liu X, An Y, Zhou G, Liu Y, Xu M, et al. Dysbiosis contributes to chronic constipation development via regulation of serotonin transporter in the intestine. Sci Rep. 2017;7(1):10322.
- 29. Ge X, Zhao W, Ding C, Tian H, Xu L, Wang H, et al. Potential role of fecal microbiota from patients with slow transit constipation in the regulation of gastrointestinal motility. Sci Rep. 2017;7(1):441.
- Parthasarathy G, Chen J, Chen X, Chia N, O'Connor HM, Wolf PG, et al. Relationship between colonic vs fecal microbiota and symptoms, colonic transit, and methane production in female patients with chronic constipation. Gastroenterology. 2016;150(2):367–79.e1.
- Palit S, Bhan C, Lunniss PJ, Boyle DJ, Gladman MA, Knowles CH, et al. Evacuation proctography: a reappraisal of normal variability. Color Dis. 2014;16(7):538–46.
- Rao SS, Bharucha AE, Chiarioni G, Felt-Bersma R, Knowles C, Malcolm A, et al. Functional anorectal disorders. Gastroenterology. 2016;25:25.
- Henry MM, Parks AG, Swash M. The pelvic floor musculature in the descending perineum syndrome. Br J Surg. 1982;69(8):470–2.
- Bharucha AE, Fletcher JG, Seide B, Riederer SJ, Zinsmeister AR. Phenotypic variation in functional disorders of defecation. Gastroenterology. 2005;128:1199–210.
- Parks AG, Swash M, Urich H. Sphincter denervation in anorectal incontinence and rectal prolapse. Gut. 1977;18(8):656–65.
- Schweiger M, Alexander-Williams J. Solitary-ulcer syndrome of the rectum its association with occult rectal prolapse. Lancet. 1977;309(8004):170–1.
- Bartram CI, Turnbull GK, Lennard-Jones JE. Evacuation proctography: an investigation of rectal expulsion in 20 subjects without defecatory disturbance. Gastrointest Radiol. 1988;13(1):72–80.
- Talley NJ, Fleming KC, Evans JM, O'Keefe EA, Weaver AL, Zinsmeister AR, et al. Constipation in an elderly community: a study of prevalence and potential risk factors. Am J Gastroenterol. 1996;91(1):19–25.
- 39•. Fragakis A, Zhou J, Mannan H, Ho V. Association between drug usage and constipation in the elderly population of Greater Western Sydney Australia. Int J Environ Res Public Health. 2018;15(2):29.
- Dore MP, Pes GM, Bibbo S, Tedde P, Bassotti G. Constipation in the elderly from Northern Sardinia is positively associated with depression, malnutrition and female gender. Scand J Gastroenterol. 2018;53(7):797–802.
- Bharucha AE, Wald A. Chronic constipation. Mayo Clin Proc. 2019;94(11):2340–57.

- Savica R, Carlin JM, Grossardt BR, Bower JH, Ahlskog JE, Maraganore DM, et al. Medical records documentation of constipation preceding Parkinson disease. A case-control study. 2009;73(21):1752–8.
- Palsson OS, Whitehead WE, van Tilburg MA, Chang L, Chey W, Crowell MD, et al. Rome IV diagnostic questionnaires and tables for investigators and clinicians. In: Gastroenterology; 2016.
- 44•. Bharucha AE, Sharma M. Painful and painless constipation: all roads lead to (a change in) Rome. Dig Dis Sci. 2018;21:21.
   Review of studies highlighting an alternate classification of constipation i.e., as painless and painful constipation.
- 45. Bouchoucha M, Devroede G, Mary F, Bon C, Bejou B, Benamouzig R. Painful or mild-pain constipation? A clinically useful alternative to classification as irritable bowel syndrome with constipation versus functional constipation. Dig Dis Sci. 2018;63(7):1763–73.
- Bouaziz W, Vogel T, Schmitt E, Kaltenbach G, Geny B, Lang PO. Health benefits of aerobic training programs in adults aged 70 and over: a systematic review. Arch Gerontol Geriatr. 2017;69:110– 27.
- King DE, Mainous AG, Lambourne CA. Trends in dietary fiber intake in the United States, 1999-2008. J Acad Nutr Diet. 2012;112(5):642–8.
- Eswaran S, Muir J, Chey WD. Fiber and functional gastrointestinal disorders. Am J Gastroenterol. 2013;108(5):718–27.
- Ford AC, Moayyedi P, Lacy BE, Lembo AJ, Saito YA, Schiller LR, et al. American College of Gastroenterology monograph on the management of irritable bowel syndrome and chronic idiopathic constipation. Am J Gastroenterol. 2014;109(Suppl 1):S2– 26 quiz S7.
- Anderson H, Bosaeus I, Falkheden T, Melkersson M. Transit time in constipated geriatric patients during treatment with a bulk laxative and bran: a comparison. Scand J Gastroenterol. 1979;14(7): 821–6.
- Nour-Eldein H, Salama H, Abdulmajeed A, Heissam K. The effect of lifestyle modification on severity of constipation and quality of life of elders in nursing homes at Ismailia city, Egypt. J Fam Community Med. 2014;21(2):100–6.
- Voderholzer WA, Schatke W, Muhldorfer BE, Klauser AG, Birkner B, Muller-Lissner SA. Clinical response to dietary fiber treatment of chronic constipation. Am J Gastroenterol. 1997;92(1):95–8.
- Morganroth J, Rüegg PC, Dunger-Baldauf C, Appel-Dingemanse S, Bliesath H, Lefkowitz M. Tegaserod, a 5-hydroxytryptamine type 4 receptor partial agonist, is devoid of electrocardiographic effects. Am J Gastroenterol. 2002;97(9):2321–7.
- 54. Chassagne P, Ducrotte P, Garnier P, Mathiex-Fortunet H. Tolerance and long-term efficacy of polyethylene glycol 4000 (Forlax(R)) compared to lactulose in elderly patients with chronic constipation. J Nutr Health Aging. 2017;21(4):429–39.
- Dipalma JA, Cleveland MV, McGowan J, Herrera JL. A randomized, multicenter, placebo-controlled trial of polyethylene glycol laxative for chronic treatment of chronic constipation. Am J Gastroenterol. 2007;102(7):1436–41.
- 56. Seinela L, Sairanen U, Laine T, Kurl S, Pettersson T, Happonen P. Comparison of polyethylene glycol with and without electrolytes in the treatment of constipation in elderly institutionalized patients: a randomized, double-blind, parallel-group study. Drugs Aging. 2009;26(8):703–13.
- Lederle FA, Busch DL, Mattox KM, West MJ, Aske DM. Costeffective treatment of constipation in the elderly: a randomized double-blind comparison of sorbitol and lactulose. Am J Med. 1990;89(5):597–601.
- Passmore AP, Wilson-Davies K, Stoker C, Scott ME. Chronic constipation in long stay elderly patients: a comparison of

lactulose and a senna-fibre combination. BMJ. 1993;307(6907): 769-71.

- Emmett CD, Close HJ, Yiannakou Y, Mason JM. Trans-anal irrigation therapy to treat adult chronic functional constipation: systematic review and meta-analysis. BMC Gastroenterol. 2015;15(139).
- Muller-Lissner S, Rykx A, Kerstens R, Vandeplassche L. A double-blind, placebo-controlled study of prucalopride in elderly patients with chronic constipation. Neurogastroenterol Motil. 2010;22(9):991–8 e255.
- Camilleri M, Beyens G, Kerstens R, Robinson P, Vandeplassche L. Safety assessment of prucalopride in elderly patients with constipation: a double-blind, placebo-controlled study. Neurogastroenterol Motil. 2009;21(12):1256–e117.
- Fidelholtz J, Smith W, Rawls J, Shi Y, Zack A, Rüegg P, et al. Safety and tolerability of tegaserod in patients with irritable bowel syndrome and diarrhea symptoms. Am J Gastroenterol. 2002;97(5):1176–81.
- Ryuji Ueno TRJ, Wahle A, Zhu Y, Holland PC. Efficacy and safety of lubiprostone for the treatment of chronic constipation in elderly vs non-elderly subjects. Gastroenterol. 2006;130(4): A–189.
- Ryuji Ueno RP, Wahle A, Zhu Y, Holland PC. Long-term safety and efficacy of lubiprostone for the treatment of chronic constipation in elderly subjects. Gastroenterol. 2006;130(4):A–188.
- Cryer B, Drossman DA, Chey WD, Webster L, Habibi S, Wang M. Analysis of nausea in clinical studies of lubiprostone for the treatment of constipation disorders. Dig Dis Sci. 2017;62(12): 3568–78.
- 66. Chey WD, Drossman DA, Johanson JF, Scott C, Panas RM, Ueno R. Safety and patient outcomes with lubiprostone for up to 52 weeks in patients with irritable bowel syndrome with constipation. Aliment Pharmacol Ther. 2012;35(5):587–99.
- Lembo AJ, Johanson JF, Parkman HP, Rao SS, Miner PB, Ueno R. Long-term safety and effectiveness of lubiprostone, a chloride channel (ClC-2) activator, in patients with chronic idiopathic constipation. Dig Dis Sci. 2011;56(9):2639–45.
- Bub S, Brinckmann J, Cicconetti G, Valentine B. Efficacy of an herbal dietary supplement (smooth move) in the management of constipation in nursing home residents: a randomized, doubleblind, placebo-controlled study. J Am Med Dir Assoc. 2006;7(9):556–61.
- Kamm MA, Mueller-Lissner S, Wald A, Richter E, Swallow R, Gessner U. Oral bisacodyl is effective and well-tolerated in patients with chronic constipation. Clin Gastroenterol Hepatol. 2011;9(7):577–83.
- 70. Agency. EM. Resolor 2019 [Available from: https://www.ema. europa.eu/en/medicines/human/EPAR/resolor.
- Administration. FaD. Motegrity 2018 [Available from: https:// www.accessdata.fda.gov/drugsatfda\_docs/label/2018/ 210166s000lbl.pdf.
- Chey WD, Lembo AJ, Lavins BJ, Shiff SJ, Kurtz CB, Currie MG, et al. Linaclotide for irritable bowel syndrome with constipation: a 26-week, randomized, double-blind, placebo-controlled trial to evaluate efficacy and safety. Am J Gastroenterol. 2012;107(11): 1702–12.
- 73. Rao S, Lembo AJ, Shiff SJ, Lavins BJ, Currie MG, Jia XD, et al. A 12-week, randomized, controlled trial with a 4-week randomized withdrawal period to evaluate the efficacy and safety of linaclotide in irritable bowel syndrome with constipation. Am J Gastroenterol. 2012;107(11):1714–24 quiz p.25.
- 74•. Schoenfeld P, Lacy BE, Chey WD, Lembo AJ, Kurtz CB, Reasner DS, et al. Low-dose linaclotide (72 mug) for chronic idiopathic constipation: a 12-week, randomized, double-blind. Placebo-Controlled Trial Am J Gastroenterol. 2018;113(1):105–14.

Linaclotide is better than placebo for chronic idiopathic constipation; 10% of patients were older than 65 years.

- DeMicco M, Barrow L, Hickey B, Shailubhai K, Griffin P. Randomized clinical trial: efficacy and safety of plecanatide in the treatment of chronic idiopathic constipation. Ther Adv Gastroenterol. 2017;10(11):837–51.
- 76. Miner PB Jr, Koltun WD, Wiener GJ, De La Portilla M, Prieto B, Shailubhai K, et al. A randomized phase III clinical trial of plecanatide, a uroguanylin analog, in patients with chronic idiopathic constipation. Am J Gastroenterol. 2017;112(4):613–21.
- 77•. Brenner DM, Fogel R, Dorn SD, Krause R, Eng P, Kirshoff R, et al. Efficacy, safety, and tolerability of plecanatide in patients with irritable bowel syndrome with constipation: results of two phase 3 randomized clinical trials. Am J Gastroenterol. 2018;113(5):735–45. Phase 3 trial of plecanatide for IBS-C; proportion of elderly patients is not reported.
- 78•. Chey WD, Lembo AJ, Rosenbaum DP. Efficacy of tenapanor in treating patients with irritable bowel syndrome with constipation: a 12-week, placebo-controlled phase 3 trial (T3MPO-1). Am J Gastroenterol. 2020;115(2):281–93. Large phase 3 trial of tenapanor for IBS-C.
- Müller-Lissner SA, Fumagalli I, Bardhan KD, Pace F, Pecher E, Nault B, et al. Tegaserod, a 5-HT(4) receptor partial agonist, relieves symptoms in irritable bowel syndrome patients with abdominal pain, bloating and constipation. Aliment Pharmacol Ther. 2001;15(10):1655–66.
- Johanson JF, Wald A, Tougas G, Chey WD, Novick JS, Lembo AJ, et al. Effect of tegaserod in chronic constipation: a randomized, double-blind, controlled trial. Clin Gastroenterol Hepatol. 2004;2(9):796–805.
- Di Palma JA, Cleveland MV, McGowan J, Herrera JL. A randomized, multicenter comparison of polyethylene glycol laxative and tegaserod in treatment of patients with chronic constipation. Am J Gastroenterol. 2007;102(9):1964–71.
- 82. Baun RF, Levy HB. Tegaserod for treating chronic constipation in elderly patients. Ann Pharmacother. 2007;41(2):309–13.
- Appel-Dingemanse S, Horowitz A, Campestrini J, Osborne S, McLeod J. The pharmacokinetics of the novel promotile drug, tegaserod, are similar in healthy subjects-male and female, elderly and young. Aliment Pharmacol Ther. 2001;15(7):937–44.
- Swan SK, Zhou H, Horowitz A, Alladina L, Hubert M, Appel-Dingemanse S, et al. Tegaserod pharmacokinetics are similar in patients with severe renal insufficiency and in healthy subjects. J Clin Pharmacol. 2003;43(4):359–64.
- Administration. FaD. Zelnorm—highlights of prescribing information 2019 [Available from: https://www.accessdata.fda.gov/ drugsatfda\_docs/label/2019/021200Orig1s015lbl.pdf.
- Etherson KJ, Minty I, Bain IM, Cundall J, Yiannakou Y. Transanal irrigation for refractory chronic idiopathic constipation: patients perceive a safe and effective therapy. Gastroenterol Res Pract. 2017;2017:3826087.
- Emmett C, Close H, Mason J, Taheri S, Stevens N, Eldridge S, et al. Low-volume versus high-volume initiated trans-anal irrigation therapy in adults with chronic constipation: study protocol for a randomised controlled trial. Trials. 2017;18(1):151.
- Martinez-Martinez MI, Calabuig-Tolsa R, Cauli O. The effect of probiotics as a treatment for constipation in elderly people: a systematic review. Arch Gerontol Geriatr. 2017;71:142–9.
- Narayanan SP, Bharucha AE. A practical guide to biofeedback therapy for pelvic floor disorders. Curr Gastroenterol Rep. 2019;21(5):21.
- Chiarioni G, Whitehead WE, Pezza V, Morelli A, Bassotti G. Biofeedback is superior to laxatives for normal transit constipation due to pelvic floor dyssynergia. [see comment]. Gastroenterology. 2006;130(3):657–64.

- Rao SSC, Seaton K, Miller M, Brown K, Nygaard I, Stumbo P, et al. Randomized controlled trial of biofeedback, sham feedback, and standard therapy for dyssynergic defecation. Clin Gastroenterol Hepatol. 2007;5(3):331–8.
- 92. Heymen S, Scarlett Y, Jones K, Ringel Y, Drossman D, Whitehead WE. Randomized, controlled trial shows biofeedback to be superior to alternative treatments for patients with pelvic floor dyssynergia-type constipation. Dis Colon Rectum. 2007;50(4):428–41.
- Simon MA, Bueno AM, Otero P, Vazquez FL, Blanco V. A randomized controlled trial on the effects of electromyographic biofeedback on quality of life and bowel symptoms in elderly women with dyssynergic defecation. Int J Environ Res Public Health. 2019;16(18).
- Rao SS, Benninga MA, Bharucha AE, Chiarioni G, Di Lorenzo C, Whitehead WE. ANMS-ESNM position paper and consensus guidelines on biofeedback therapy for anorectal disorders. Neurogastroenterol Motil. 2015;27(5):594–609.
- Knowles CH, Grossi U, Chapman M, Mason J. Surgery for constipation: systematic review and practice recommendations: results I: colonic resection. Color Dis. 2017;19(Suppl 3):17–36.
- 96•. Wilkinson-Smith V, Bharucha AE, Emmanuel A, Knowles C, Yiannakou YA, Corsetti M. When all seems lost: management of refractory constipation—surgery, rectal irrigation, percutaneous endoscopic colostomy, and more. Neurogastroenterol Motil. 2018;30(5):e13352. Review of non-medical approaches to manage chronic constipation.
- Redmond JM, Smith GW, Barofsky I, Ratych RE, Goldsborough DC, Schuster MM. Physiological tests to predict long-term outcome of total abdominal colectomy for intractable constipation. Am J Gastroenterol. 1995;90(5):748–53.
- Shorvon PJ, McHugh S, Diamant NE, Somers S, Stevenson GW. Defecography in normal volunteers: results and implications. Gut. 1989;30(12):1737–49.
- Tirumanisetty P, Prichard D, Fletcher JG, Chakraborty S, Zinsmeister AR, Bharucha AE. Normal values for assessment of anal sphincter morphology, anorectal motion, and pelvic organ prolapse with MRI in healthy women. Neurogastroenterol Motil. 2018;30(7):e13314.
- Aquina CT, Mohile SG, Tejani MA, Becerra AZ, Xu Z, Hensley BJ, et al. The impact of age on complications, survival, and cause of death following colon cancer surgery. Br J Cancer. 2017;116(3):389–97.
- Fagard K, Casaer J, Wolthuis A, Flamaing J, Milisen K, Lobelle J-P, et al. Postoperative complications in individuals aged 70 and over undergoing elective surgery for colorectal cancer. Color Dis. 2017;19(9):O329–O38.
- Bharucha AE, Zinsmeister AR, Locke GR, Seide B, McKeon K, Schleck CD, et al. Prevalence and burden of fecal incontinence: a population based study in women. Gastroenterology. 2005;129: 42–9.
- 103. Whitehead WE, Borrud L, Goode PS, Meikle S, Mueller ER, Tuteja A, et al. Fecal incontinence in US adults: epidemiology and risk factors. Gastroenterology. 2009;137(2):512–7.
- Nelson R, Furner S, Jesudason V. Fecal incontinence in Wisconsin nursing homes: prevalence and associations. Dis Colon Rectum. 1998;41(10):1226–9.
- Nelson RL. Epidemiology of fecal incontinence. Gastroenterology. 2004;126(1 Suppl 1):S3–7.
- Bliss DZ, Harms S, Garrard JM, Cunanan K, Savik K, Gurvich O, et al. Prevalence of incontinence by race and ethnicity of older people admitted to nursing homes. J Am Med Dir Assoc. 2013;14(6):451.e1–7.
- 107•. Chughtai B, Thomas D, Russell D, Phongtankuel V, Bowles K, Prigerson H. Prevalence and risk factors for fecal incontinence in

home hospice. Am J Hosp Palliat Med. 2019;36(1):33–7. Fecal incontinence in home hospice care.

- Grover M, Whitehead WE. Is fecal incontinence a risk factor for institutionalization in the elderly? Am J Gastroenterol. 2011;106(2):366-7 author reply 7.
- Chassagne P, Landrin I, Neveu C, Czernichow P, Bouaniche M, Doucet J, et al. Fecal incontinence in the institutionalized elderly: incidence, risk factors, and prognosis. Am J Med. 1999;106(2): 185–90.
- Bharucha AE, Zinsmeister AR, Locke GR, Schleck C, McKeon K, Melton LJ. Symptoms and quality of life in community women with fecal incontinence. Clin Gastroenterol Hepatol. 2006;4(8): 1004–9.
- 111. Jamieson HA, Schluter PJ, Pyun J, Arnold T, Scrase R, Nisbet-Abey R, et al. Fecal incontinence is associated with mortality among older adults with complex needs: an observational cohort study. Am J Gastroenterol. 2017;112(9):1431–7.
- Melville JL, Fan MY, Newton K, Fenner D. Fecal incontinence in US women: a population-based study. Am J Obstet Gynecol. 2005;193(6):2071–6.
- 113. Bharucha AE, Zinsmeister AR, Schleck CD, Melton LJ 3rd. Bowel disturbances are the most important risk factors for late onset fecal incontinence: a population-based case-control study in women. Gastroenterology. 2010;139(5):1559–66.
- Goode PS, Burgio KL, Halli AD, Jones RW, Richter HE, Redden DT, et al. Prevalence and correlates of fecal incontinence in community-dwelling older adults. J Am Geriatr Soc. 2005;53(4): 629–35.
- Shamliyan TA, Bliss DZ, Du J, Ping R, Wilt TJ, Kane RL. Prevalence and risk factors of fecal incontinence in communitydwelling men. Rev Gastroenterol Disord. 2009;9(4):E97–110.
- Demir N, Yuruyen M, Atay K, Yavuzer H, Hatemi I, Doventas A, et al. Prevalence of fecal incontinence and associated risk factors in elderly outpatients: a cross-sectional study. Aging-Clin Exper Res. 2017;29(6):1165–71.
- 117. Matthews CA, Whitehead WE, Townsend MK, Grodstein F. Risk factors for urinary, fecal, or dual incontinence in the Nurses' Health Study. Obstet Gynecol. 2013;122(3):539–45.
- Staller K, Townsend MK, Khalili H, Mehta R, Grodstein F, Whitehead WE, et al. Menopausal hormone therapy is associated with increased risk of fecal incontinence in women after menopause. Gastroenterology. 2017;152(8):1915–21.e1.
- Bharucha AE. Fecal incontinence. Gastroenterology. 2003;124(6): 1672–85.
- Varma MG, Brown JS, Creasman JM, Thom DH, Van Den Eeden SK, Beattie MS, et al. Fecal incontinence in females older than aged 40 years: who is at risk? Dis Colon Rectum. 2006;49(6):841– 51.
- 121. Bharucha AE, Fletcher JG, Melton LJ 3rd, Zinsmeister AR. Obstetric trauma, pelvic floor injury and fecal incontinence: a population-based case-control study. Am J Gastroenterol. 2012;107(6):902–11.
- 122. Markland AD, Dunivan GC, Vaughan CP, Rogers RG. Anal intercourse and fecal incontinence: evidence from the 2009-2010 National Health and Nutrition Examination Survey. Am J Gastroenterol. 2016;111(2):269–74.
- 123. Bharucha AE, Zinsmeister AR, Locke GR, Seide BM, McKeon K, Schleck CD, et al. Risk factors for fecal incontinence: a population-based study in women. Am J Gastroenterol. 2006;101(6):1305–12.
- 124. Bharucha AE, Seide BM, Zinsmeister AR, Melton LJ 3rd. Relation of bowel habits to fecal incontinence in women. Am J Gastroenterol. 2008;103(6):1470–5.
- 125. Bharucha AE. Pelvic floor: anatomy and function. Neurogastroenterol Motil. 2006;18(7):507–19.

- 126. Johanson JF, Lafferty J. Epidemiology of fecal incontinence: the silent affliction. Am J Gastroenterol. 1996;91(1):33–6.
- 127. Akpan A, Gosney MA, Barret J. Factors contributing to fecal incontinence in older people and outcome of routine management in home, hospital and nursing home settings. Clin Interv Aging. 2007;2(1):139–45.
- Schnelle JF, Simmons SF, Beuscher L, Peterson EN, Habermann R, Leung F. Prevalence of constipation symptoms in fecally incontinent nursing home residents. J Am Geriatr Soc. 2009;57(4): 647–52.
- Andy UU, Vaughan CP, Burgio KL, Alli FM, Goode PS, Markland AD. Shared risk factors for constipation, fecal incontinence, and combined symptoms in older U.S. adults. J Am Geriatr Soc. 2016;64(11):e183–e8.
- 130. Staller K, Song M, Grodstein F, Matthews CA, Whitehead WE, Kuo B, et al. Physical activity, BMI, and risk of fecal incontinence in the Nurses' Health Study. Clin Transl Gastroenterol. 2018;9(10):200.
- 131. Rao SS. Pathophysiology of adult fecal incontinence. Gastroenterology. 2004;126:S14–22.
- Andrews CN, Bharucha AE. The etiology, assessment, and treatment of fecal incontinence. Nat Clin Prac Gastroenterol Hepatol. 2005;2(11):516–25.
- 133. Sun WM, Donnelly TC, Read NW. Utility of a combined test of anorectal manometry, electromyography, and sensation in determining the mechanism of 'idiopathic' faecal incontinence. Gut 1992;33:807–13.
- 134. Ledgerwood-Lee M, Zifan A, Kunkel DC, Sah R, Mittal RK. High-frequency ultrasound imaging of the anal sphincter muscles in normal subjects and patients with fecal incontinence. Neurogastroenterol Motil. 31(4):e13537.
- Tantiphlachiva K, Rao P, Attaluri A, Rao SSC. Digital rectal examination is a useful tool for identifying patients with dyssynergia. Clin Gastroenterol Hepatol. 2010;8(11):955–60.
- 136. Carrington EV, Knowles CH, Grossi U, Scott SM. High-resolution anorectal manometry measures are more accurate than conventional measures in detecting anal hypocontractility in women with fecal incontinence. Clin Gastroenterol Hepatol. 2019;17(3):477–85.e9.
- 137•. Carrington EV, Heinrich H, Knowles CH, Fox M, Rao S, Altomare DF, et al. The international anorectal physiology working group (IAPWG) recommendations: standardized testing protocol and the London classification for disorders of anorectal function. Neurogastroenterol Motil. 2020;32(1):e13679. Recent guidelines for conducting anorectal manometry.
- Lee TH, Bharucha AE. How to perform and interpret a highresolution anorectal manometry test. J Neurogastroenterol Motil. 2016;22(1):46–59.
- Bharucha AE, Fletcher JG, Harper CM, Hough D, Daube JR, Stevens C, et al. Relationship between symptoms and disordered continence mechanisms in women with idiopathic fecal incontinence. Gut. 2005;54:546–55.
- Andrews CN, Seide B, Zinsmeister AR, Bharucha AE. Effect of distention rate on rectal sensory thresholds in health and fecal incontinence. Gastroenterology. 2005;128(4):A-264.
- 141. Siproudhis L, El Abkari M, El Alaoui M, Juguet F, Bretagne JF. Low rectal volumes in patients suffering from fecal incontinence: what does it mean? Aliment Pharmacol Ther. 2005;22(10):989–96.
- 142. Deutekom M, Dobben AC, Terra MP, Engel AF, Stoker J, Bossuyt PM, et al. Clinical presentation of fecal incontinence and anorectal function: what is the relationship? Am J Gastroenterol. 2007;102(2):351–61.
- Azpiroz F, Enck P, Whitehead WE. Anorectal functional testing: review of collective experience. Am J Gastroenterol. 2002;97(2): 232–40.
- 144. Townsend DC, Carrington EV, Grossi U, Burgell RE, Wong JY, Knowles CH, et al. Pathophysiology of fecal incontinence differs

between men and women: a case-matched study in 200 patients. Neurogastroenterol Motil. 2016;28(10):1580–8.

- Leigh RJ, Turnberg LA. Faecal incontinence: the unvoiced symptom. Lancet. 1982;1(8285):1349–51.
- Miner PB Jr. Economic and personal impact of fecal and urinary incontinence. Gastroenterology. 2004;126(1 Suppl 1):S8–13.
- Bliss DZ, Funk T, Jacobson M, Savik K. Incidence and characteristics of incontinence-associated dermatitis in communitydwelling persons with fecal incontinence. J Wound Ostomy Continence Nurs. 2015;42(5):525–30.
- Fader M, Cottenden A, Getliffe K, Gage H, Clarke-O'Neill S, Jamieson K, et al. Absorbent products for urinary/faecal incontinence: a comparative evaluation of key product designs. Health Technol Assess. 2008;12(29):iii–v ix-185.
- 149. Andy UU, Ejike N, Khanijow KD, Flick LC, Markland AD, Arya LA, et al. Diet modifications in older women with fecal incontinence: a qualitative study. Female Pelvic Med Reconstruct Surg. 2020;26(4):239–43.
- Colavita K, Andy UU. Role of diet in fecal incontinence: a systematic review of the literature. Int Urogynecol J. 2016;27(12): 1805–10.
- 151. Markland AD, Richter HE, Burgio KL, Bragg C, Hernandez AL, Subak LL. Fecal incontinence in obese women with urinary incontinence: prevalence and role of dietary fiber intake. Am J Obstet Gynecol. 2009;200(5):566.e1–6.
- 152•. Staller K, Song M, Grodstein F, Whitehead WE, Matthews CA, Kuo B, et al. Increased long-term dietary fiber intake is associated with a decreased risk of fecal incontinence in older women. Gastroenterology. 2018;155(3):661–7.e1. Key study which highlights the long-term benefits of dietary fiber intake for reducing the risk of FI.
- 153••. Bliss DZ, Savik K, Jung H-JG, Whitebird R, Lowry A, Sheng X. Dietary fiber supplementation for fecal incontinence: a randomized clinical trial. Res Nurs Health. 2014;37(5):367–78. Dietary fiber supplementation benefits some patients with FI.
- 154. Markland AD, Burgio KL, Whitehead WE, Richter HE, Wilcox CM, Redden DT, et al. Loperamide versus psyllium fiber for treatment of fecal incontinence: the Fecal Incontinence Prescription (Rx) Management (FIRM) randomized clinical trial. Dis Colon Rectum. 2015;58(10):983–93.
- Chassagne P, Jego A, Gloc P, Capet C, Trivalle C, Doucet J, et al. Does treatment of constipation improve faecal incontinence in institutionalized elderly patients? Age Ageing. 2000;29(2):159–64.
- Ryan D, Wilson A, Muir TS, Judge TG. The reduction of faecal incontinence by the use of "Duphalac" in geriatric patients. Curr Med Res Opin. 1974;2(6):329–33.
- 157. Sun WM, Read NW, Verlinden M. Effects of loperamide oxide on gastrointestinal transit time and anorectal function in patients with chronic diarrhoea and faecal incontinence. Scand J Gastroenterol. 1997;32(1):34–8.
- 158. Read M, Read NW, Barber DC, Duthie HL. Effects of loperamide on anal sphincter function in patients complaining of chronic diarrhea with fecal incontinence and urgency. Dig Dis Sci. 1982;27(9):807–14.
- Fox M, Stutz B, Menne D, Fried M, Schwizer W, Thumshim M. The effects of loperamide on continence problems and anorectal function in obese subjects taking orlistat. Dig Dis Sci. 2005;50(9):1576–83.
- Hallgren T, Fasth S, Delbro DS, Nordgren S, Oresland T, Hulten L. Loperamide improves anal sphincter function and continence after restorative proctocolectomy. Dig Dis Sci. 1994;39(12): 2612–8.
- Harford WV, Krejs GJ, Santa Ana CA, Fordtran JS. Acute effect of diphenoxylate with atropine (Lomotil) in patients with chronic diarrhea and fecal incontinence. Gastroenterology. 1980;78(3): 440–3.

- 162. Pelemans W, Vantrappen F. A double blind crossover comparison of loperamide with diphenoxylate in the symptomatic treatment of chronic diarrhea. Gastroenterology. 1976;70(6):1030–4.
- Palmer KR, Corbett CL, Holdsworth CD. Double-blind cross-over study comparing loperamide, codeine and diphenoxylate in the treatment of chronic diarrhea. Gastroenterology. 1980;79(6): 1272–5.
- 164•. Jelovsek JE, Markland AD, Whitehead WE, Barber MD, Newman DK, Rogers RG, et al. Controlling faecal incontinence in women by performing anal exercises with biofeedback or loperamide: a randomised clinical trial. \Lancet Gastroenterol Hepatol. 2019;4(9):698–710.
- Santoro GA, Eitan BZ, Pryde A, Bartolo DC. Open study of lowdose amitriptyline in the treatment of patients with idiopathic fecal incontinence. Dis Colon Rectum. 2000;43:1676–81.
- Remes-Troche JM, Ozturk R, Philips C, Stessman M, Rao SSC. Cholestyramine—a useful adjunct for the treatment of patients with fecal incontinence. Int J Color Dis. 2008;23(2):189–94.
- 167. Wedlake L, Thomas K, Lalji A, Anagnostopoulos C, Andreyev HJ. Effectiveness and tolerability of colesevelam hydrochloride for bile-acid malabsorption in patients with cancer: a retrospective chart review and patient questionnaire. Clin Ther. 2009;31(11): 2549–58.
- Bharucha AE, Fletcher JG, Camilleri M, Edge J, Carlson P, Zinsmeister AR. Effects of clonidine in women with fecal incontinence. Clin Gastroenterol Hepatol. 2014;12(5):843–51 e2 quiz e44.
- 169. Ilnyckyj A, Fachnie E, Tougas G. A randomized-controlled trial comparing an educational intervention alone vs education and biofeedback in the management of faecal incontinence in women. Neurogastroenterol Motil. 2005;17(1):58–63.
- Norton C, Chelvanayagam S, Wilson-Barnett J, Redfern S, Kamm MA. Randomized controlled trial of biofeedback for fecal incontinence. Gastroenterology. 2003;125:1320–9.
- 171. Solomon MJ, Pager CK, Rex J, Roberts R, Manning J. Randomized, controlled trial of biofeedback with anal manometry, transanal ultrasound, or pelvic floor retraining with digital guidance alone in the treatment of mild to moderate fecal incontinence. Dis Colon Rectum. 2003;46(6):703–10.
- 172. Heymen S, Scarlett Y, Jones K, Ringel Y, Drossman DA, Whitehead W. Randomized controlled trial shows biofeedback to be superior to alternative treatments for fecal incontinence. Dis Colon Rectum. 2009;52:1730–7.
- 173•. Mazor Y, Ejova A, Andrews A, Jones M, Kellow J, Malcolm A. Long-term outcome of anorectal biofeedback for treatment of fecal incontinence. Neurogastroenterol Motil. 2018:e13389.
   Highlights the long-term outcomes after anorectal biofeedback for fecal incontinence.
- 174. Byrne CM, Solomon MJ, Young JM, Rex J, Merlino CL. Biofeedback for fecal incontinence: short-term outcomes of 513 consecutive patients and predictors of successful treatment. Dis Colon Rectum. 2007;50(4):417–27.
- 175•. Mazor Y, Prott G, Jones M, Ejova A, Kellow J. Malcolm A. Factors associated with response to anorectal biofeedback therapy in patients with fecal incontinence. Clin Gastroenterol Hepatol. 2020;(20)30427–4. https://doi.org/10.1016/j.cgh.2020.03.050. Approximately two-thirds of patients had a response to biofeedback therapy.
- 176. Vasant DH, Solanki K, Balakrishnan S, Radhakrishnan NV. Integrated low-intensity biofeedback therapy in fecal incontinence: evidence that "good" in-home anal sphincter exercise practice makes perfect. Neurogastroenterol Motil. 2017;29(1): e12912.
- 177. Young CJ, Zahid A, Koh CE, Young JM, Byrne CM, Solomon MJ, et al. A randomized controlled trial of four different regimes

of biofeedback programme in the treatment of faecal incontinence. Color Dis. 2018;20(4):312–20.

- 178. Richter HE, Jelovsek JE, Iyer P, Rogers RG, Meyer I, Newman DK, et al. Characteristics associated with clinically important treatment responses in women undergoing nonsurgical therapy for fecal incontinence. Am J Gastroenterol. 2020;115(1):115–27.
- 179. Sjödahl J, Walter SA, Johansson E, Ingemansson A, Ryn AK, Hallböök O. Combination therapy with biofeedback, loperamide, and stool-bulking agents is effective for the treatment of fecal incontinence in women—a randomized controlled trial. Scand J Gastroenterol. 2015;50(8):965–74.
- Lukacz ES, Segall MM, Wexner SD. Evaluation of an anal insert device for the conservative management of fecal incontinence. Dis Colon Rectum. 2015;58(9):892–8.
- 181. Richter HE, Matthews CA, Muir T, Takase-Sanchez MM, Hale DS, Van Drie D, et al. A vaginal bowel-control system for the treatment of fecal incontinence. Obstet Gynecol. 2015;125(3): 540–7.
- 182••. Bharucha AE, SSC R, Shin AS. Surgical interventions and the use of device-aided therapy for the treatment of fecal incontinence and defecatory disorders. Clin Gastroenterol Hepatol. 2017;15(12): 1844–54. Systematic review of surgical procedures and other devices for fecal incontinence.
- 183. Graf W, Mellgren A, Matzel KE, Hull T, Johansson C, Bernstein M. Efficacy of dextranomer in stabilised hyaluronic acid for treatment of faecal incontinence: a randomised, sham-controlled trial. Lancet. 2011;377(9770):997–1003.
- Mellgren A, Matzel KE, Pollack J, Hull T, Bernstein M, Graf W, et al. Long-term efficacy of NASHA Dx injection therapy for treatment of fecal incontinence. Neurogastroenterol Motil. 2014;26(8):1087–94.
- 185. Dehli T, Stordahl A, Vatten LJ, Romundstad PR, Mevik K, Sahlin Y, et al. Sphincter training or anal injections of dextranomer for treatment of anal incontinence: a randomized trial. Scand J Gastroenterol. 2013;48(3):302–10.
- 186. Simillis C, Lal N, Qiu S, Kontovounisios C, Rasheed S, Tan E, et al. Sacral nerve stimulation versus percutaneous tibial nerve stimulation for faecal incontinence: a systematic review and meta-analysis. Int J Color Dis. 2018;33(5):645–8.
- 187. Tan K, Wells CI, Dinning P, Bissett IP, O'Grady G. Placebo response rates in electrical nerve stimulation trials for fecal incontinence and constipation: a systematic review and meta-analysis. Neuromodulation. 2019.
- Carrington EV, Evers J, Grossi U, Dinning PG, Scott SM, O'Connell PR, et al. A systematic review of sacral nerve stimulation mechanisms in the treatment of fecal incontinence and constipation. Neurogastroenterol Motil. 2014;26(9):1222–37.
- 189. Wexner SD, Coller JA, Devroede G, Hull T, McCallum R, Chan M, et al. Sacral nerve stimulation for fecal incontinence: results of a 120-patient prospective multicenter study. Ann Surg. 2010;251(3):441–9.
- Hull T, Giese C, Wexner SD, Mellgren A, Devroede G, Madoff RD, et al. Long-term durability of sacral nerve stimulation therapy for chronic fecal incontinence. Dis Colon Rectum. 2013;56(2): 234–45.
- 191. Gallas S, Michot F, Faucheron JL, Meurette G, Lehur PA, Barth X, et al. Predictive factors for successful sacral nerve stimulation in the treatment of faecal incontinence: results of trial stimulation in 200 patients. Color Dis. 2011;13(6):689–96.
- 192. Le Fouler A, Duchalais E, Loong TH, et al. Long-term outcome following implanted pulse generator change in patients treated with sacral nerve modulation for fecal incontinence. Neuromodulation. 2018;21(7):694–99. https://doi.org/10.1111/ner.12806.

- 193. Knowles CH, Horrocks EJ, Bremner SA, Stevens N, Norton C, O'Connell PR, et al. Percutaneous tibial nerve stimulation versus sham electrical stimulation for the treatment of faecal incontinence in adults (CONFIDENT): a double-blind, multicentre, pragmatic, parallel-group, randomised controlled trial. Lancet. 2015;386(10004):1640–8.
- 194. Thin NN, Taylor SJC, Bremner SA, Emmanuel AV, Hounsome N, Williams NS, et al. Randomized clinical trial of sacral versus percutaneous tibial nerve stimulation in patients with faecal incontinence. Br J Surg. 2015;102(4):349–58.
- 195••. Forte ML, Andrade KE, Lowry AC, Butler M, Bliss DZ, Kane RL. Systematic review of surgical treatments for fecal incontinence. Dis Colon Rectum. 2016;59(5):443–69. Systematic review of surgical treatments for fecal incontinence.
- Norton C, Burch J, Kamm MA. Patients' views of a colostomy for fecal incontinence. Dis Colon Rectum. 2005;48(5):1062–9.

**Publisher's Note** Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.