# **Acute Diarrhea in Adults**

**Erica S. Meisenheimer, MD, MBA,** Fort Belvoir Community Hospital, Fort Belvoir, Virginia; Uniformed Services University of the Health Sciences, Bethesda, Maryland

Carly Epstein, DO, Fort Belvoir Community Hospital, Fort Belvoir, Virginia

**Derrick Thiel, MD, MPH,** Madigan Army Medical Center Family Medicine Residency, Joint Base Lewis-McChord, Washington; Uniformed Services University of the Health Sciences, Bethesda, Maryland

Acute diarrheal disease accounts for 179 million outpatient visits annually in the United States. Diarrhea can be categorized as inflammatory or noninflammatory, and both types have infectious and noninfectious causes. Infectious noninflammatory diarrhea is often viral in etiology and is the most common presentation; however, bacterial causes are also common and may be related to travel or foodborne illness. History for patients with acute diarrhea should include onset and frequency of symptoms, stool character, a focused review of systems including fever and other symptoms, and evaluation of exposures and risk factors. The physical examination should include evaluation for signs of dehydration, sepsis, or potential surgical processes. Most episodes of acute diarrhea in countries with adequate food and water sanitation are uncomplicated and self-limited, requiring only an initial evaluation and supportive treatment. Additional diagnostic evaluation and management may be warranted when diarrhea is bloody or mucoid or when risk factors are present, including immunocompromise or recent hospitalization. Unless an outbreak is suspected, molecular studies are preferred over traditional stool cultures. In all cases, management begins with replacing water, electrolytes, and nutrients. Oral rehydration is preferred; however, signs of severe dehydration or sepsis warrant intravenous rehydration. Antidiarrheal agents can be symptomatic therapy for acute watery diarrhea and can help decrease inappropriate antibiotic use. Empiric antibiotics are rarely warranted, except in sepsis and some cases of travelers' or inflammatory diarrhea. Targeted antibiotic therapy may be appropriate following microbiologic stool assessment. Hand hygiene, personal protective equipment, and food and water safety measures are integral to preventing infectious diarrheal illnesses. (Am Fam Physician. 2022;106(1):72-80. Copyright © 2022 American Academy of Family Physicians.)

**Acute diarrheal** disease is a common cause of clinic visits and hospitalizations. In the United States, diarrhea accounts for an estimated 179 million outpatient visits, 500,000 hospitalizations, and more than 5,000 deaths annually, resulting in approximately one episode of acute diarrheal illness per person per year.<sup>1-5</sup>

Diarrhea is the passage of three or more loose or watery stools in 24 hours.<sup>5-7</sup> Acute diarrhea is when symptoms typically last fewer than two weeks; however, some experts define acute as fewer than seven days, with seven to 14 days considered prolonged.<sup>2</sup> Diarrhea lasting 14 to 30 days is considered persistent, and chronic diarrhea lasts longer than one month.<sup>2,5,7,8</sup>

Diarrhea can be categorized as inflammatory or noninflammatory. Inflammatory diarrhea, or dysentery, typically presents with blood or mucus and is often caused by invasive pathogens

This clinical content conforms to AAFP criteria for CME. See CME Quiz on page 18.

Author disclosure: No relevant financial relationships.

or processes. Noninflammatory or watery diarrhea is caused by increased water secretion into the intestinal lumen or decreased water reabsorption.<sup>2</sup>

# **Differential Diagnosis**

Noninflammatory and inflammatory diarrhea may be subsequently categorized into infectious and noninfectious causes (*Table 1*).<sup>2,5,9,10</sup>

Infectious noninflammatory diarrhea, the most common presentation, is typically viral in etiology, but bacterial causes are also common and are usually related to travel or foodborne illness.<sup>9,11,12</sup>

Infectious inflammatory diarrhea is more severe and is typically caused by invasive or toxin-producing bacteria, although viral and parasitic causes exist.<sup>12</sup> Bloody stools, high fever, significant abdominal pain, or duration longer than three days suggests inflammatory infection. The most identified inflammatory pathogens in North America are *Salmonella*, *Campylobacter*, *Clostridioides difficile*, *Shigella*, and Shiga toxin-producing *Escherichia coli*.<sup>2,12</sup>

# **SORT: KEY RECOMMENDATIONS FOR PRACTICE**

Evidence rating	Comments
С	Consistent recommendation from practice guidelines with low-level evidence
Α	Consistent findings from systematic review
В	Single, high-quality randomized controlled trial
Α	Consistent findings from systematic review and meta-analysis
Α	Consistent recommendation from evidence-based practice guidelines with high-level evidence
	C A B

 $\mathbf{A}=$  consistent, good-quality patient-oriented evidence;  $\mathbf{B}=$  inconsistent or limited-quality patient-oriented evidence;  $\mathbf{C}=$  consensus, disease-oriented evidence, usual practice, expert opinion, or case series. For information about the SORT evidence rating system, go to https://www.aafp. org/afpsort.

TABLE 1

### **Characteristics of Noninflammatory and Inflammatory Diarrhea**

Noninflammatory diarrhea	Inflammatory diarrhea
Infectious: often viral, but may be bacterial and, less likely, parasitic Noninfectious: dietary, psychosocial stressors	Infectious: frequently invasive or toxin- producing bacteria Noninfectious: Crohn disease, ulcerative colitis, radiation enteritis
Infectious: nausea, vomiting, abdominal discomfort  Noninfectious: nausea, abdominal discomfort, frequently without vomiting	Infectious: fever, abdominal pain, tenesmus, systemic signs and symptoms Noninfectious: abdominal pain, tenesmus, fatigue, weight loss
Infectious: often not performed; positive PCR or NAAT result Noninfectious: negative PCR or NAAT result, positive specific laboratory testing (e.g., antitissue transglutaminase antibody)	Infectious: positive PCR or NAAT result Noninfectious: negative PCR or NAAT result; positive for fecal calprotectin*
Bacterial: enterotoxigenic Escherichia coli, Clostridium perfringens, Bacillus cereus, Staphylococcus aureus, Vibrio cholerae Viral: Rotavirus, Norovirus Parasitic: Giardia, Cryptosporidium	Bacterial: Salmonella, Shigella, Campylo- bacter, Shiga toxin—producing E. coli, entero- invasive E. coli, Clostridioides difficile, Yersinia Parasitic: Entamoeba histolytica
	Infectious: often viral, but may be bacterial and, less likely, parasitic Noninfectious: dietary, psychosocial stressors  Infectious: nausea, vomiting, abdominal discomfort  Noninfectious: nausea, abdominal discomfort, frequently without vomiting  Infectious: often not performed; positive PCR or NAAT result Noninfectious: negative PCR or NAAT result, positive specific laboratory testing (e.g., antitissue transglutaminase antibody)  Bacterial: enterotoxigenic Escherichia coli, Clostridium perfringens, Bacillus cereus, Staphylococcus aureus, Vibrio cholerae Viral: Rotavirus, Norovirus

NAAT = nucleic acid amplification test; PCR = polymerase chain reaction.

Specific pathogenic causes differ based on location, travel or food exposures; work or living circumstances (e.g., health care, long-term care settings); and comorbidities, such as HIV or other immunocompromised states.

Noninfectious causes of diarrhea include dietary intolerance, adverse medication effects, abdominal surgical

processes such as appendicitis or mesenteric ischemia, thyroid or other endocrine abnormalities, and gastrointestinal diseases, such as Crohn disease or ulcerative colitis<sup>10,12</sup> (*Table* 2<sup>8,10,12</sup>). Although many of these processes are chronic, they can present acutely and should be included in the differential diagnosis of acute diarrhea.

<sup>\*—</sup>To be used if history is concerning for inflammatory bowel disease; otherwise, not routinely recommended. *Information from references 2, 5, 9, and 10.* 

### Initial Evaluation

In countries with adequate food and water sanitation, most episodes of acute diarrhea are uncomplicated and self-limited, requiring only an initial evaluation and supportive treatment.<sup>2,6,12</sup> The history and physical examination in patients with acute diarrhea should assess symptom severity, potential disease etiology, and personal

risk factors for complications and should guide additional workup and treatment decisions.2 Figure 1 provides an approach to the evaluation and management of acute diarrhea.2,5

### **HISTORY**

A history for patients with acute diarrhea should include symptom onset, severity (volume, frequency, duration of diarrhea), stool character, and a focused review of systems, including vomiting, fever, decreased urine output, and abdominal pain.<sup>9,12</sup>

Additional exposure history can help suggest potential etiology and the need for further evaluation. Physicians should ask about recent travel. food or water exposures, known contacts with sick people, occupations, residence, recent medical history (e.g., illnesses, hospitalizations, antibiotics, other medications), and sexual history.2,9,12

Assessing for medical comorbidities, including immunocompromised states or personal or family history of autoimmune conditions, can be useful in identifying patients at high risk of complicated diarrheal illnesses (Table 3<sup>2,5,9,12-14</sup>).

### PHYSICAL EXAMINATION

The initial physical examination should focus on disease severity and hydration status. Clinicians should take note of general appearance, fever, tachycardia, and hypotension. Peripheral pulses, skin turgor, and capillary refill may also assist in assessing volume status. The abdominal examination should evaluate for an acute abdomen requiring potential surgical intervention. 2,12,13

### **Additional Evaluation**

Further diagnostic evaluation is indicated for patients with inflammatory diarrhea or severe symptoms, including sepsis, or for those whose history includes risk factors for complications or potential outbreaks (e.g., patients who are immunocompromised, food handlers, health care or childcare workers, and community or nursing home residents).

### **BEST PRACTICES IN GASTROENTEROLOGY**

# **Recommendations From the Choosing Wisely Campaign**

Recommendation	organization
Do not order a comprehensive stool ova and parasite microscopic examination on patients presenting with diarrhea lasting less than seven days who have no immunodeficiency and no history of living in or traveling to endemic areas where gastrointestinal parasitic infections are prevalent. If symptoms of infectious diarrhea persist for seven days or more, start with molecular or antigen testing and next consider a full ova and parasite microscopic examination if other result is negative.	American Society for Clinical Laboratory Science

Source: For more information on the Choosing Wisely Campaign, see https://www. choosingwisely.org. For supporting citations and to search Choosing Wisely recommendations relevant to primary care, see https://www.aafp.org/pubs/afp/collections/choosing-wisely.html.

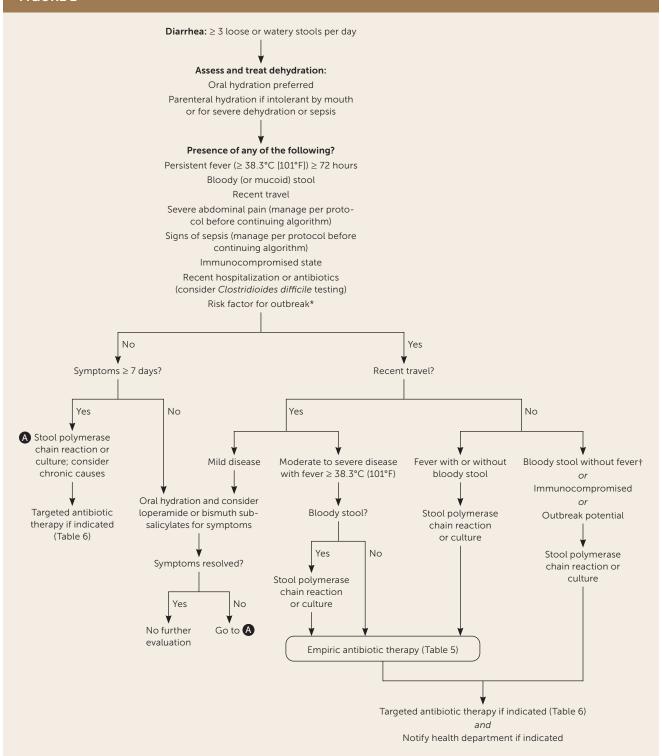
# TABLE 2

Potential causes

# **Common Causes of Noninfectious Diarrhea** Evamples

Potential causes	Examples
Abdominal processes	Appendicitis, ischemic colitis, malignancy
Dietary	Alcohol, caffeine, celiac disease (autoimmune), FODMAP malabsorption, lactose intolerance, nondigestible sugars
Endocrine abnormalities	Adrenal dysfunction, bile acid malabsorption, pancreatic exocrine insufficiency, thyroid dysfunction
Functional	Irritable bowel syndrome, mental health comorbidities, overflow diarrhea due to fecal impaction
Inflammatory	Crohn disease, microscopic colitis, radiation enteritis, ulcerative colitis
Medication effects	Antibiotics, antineoplastic drugs, behavioral health medications (e.g., selective serotonin reuptake inhibitors, serotonin-norepinephrine reuptake inhibitors, norepinephrine-dopamine reuptake inhibitors), hypoglycemic agents, magnesium-containing products

FODMAP = fermentable oligosaccharides, disaccharides, monosaccharides, and polyols. Information from references 8, 10, and 12.



<sup>\*—</sup>Including food handlers, childcare workers, staff or residents of nursing homes or other community dwellings, or exposure to a known or suspected outbreak pathogen.

# Evaluation and management of acute diarrhea.

Information from references 2 and 5.

<sup>†—</sup>Bloody stool without fever is concerning for Shiga toxin–producing *Escherichia coli*. Avoid antibiotics until stool microbiology confirms a pathogen.

### **Significant Historical Factors for Acute Diarrhea**

	61 10	
Finding	Significance	Further measures indicated
Associated signs and symp Bloody stool Mucoid stool Persistent fever	Inflammatory process	Stool studies (culture or multiplex polymerase chain reaction with reflex culture)
Severe abdominal pain		Shiga toxin, if not included in polymerase chain reaction*
Confusion Decreased urine output Dizziness Syncope	Dehydration Possible sepsis	Restore volume status If septic, obtain blood cultures and stool studies
Symptoms lasting ≥ 7 days	Less reassuring for acute, self-limited process	Consider stool studies
Vomiting	Reassuring for likely viral etiology	None, unless otherwise indi- cated below
Personal risk factors		
Age ≥ 65 years	Increased risk of severe disease	Keep higher concern for dehydration or invasive disease and evaluate as indicated
Childcare workers Community or nurs- ing home workers or residents Food handlers	Increased risk of potential outbreak	Public health reporting as indicated Stool studies
History of pelvic radiation	Radiation enteritis	Consider specialty consultatio
Immunocompromised	Increased risk of severe or invasive disease	Stool studies†
Known autoimmune disorders	Concern for flare-up or new diagnosis of inflam- matory bowel disease	Additional measures based on suspicion for inflammatory bowel disease; consider specialty consultation
Recent hospitalization or antibiotic use	Concern for Clos- tridioides difficile	C. difficile testing per guidelines‡
Recent travel	Increased concern for enteric fever or other location- specific pathogens	No additional measures unles indicated by symptoms above persistent diarrhea, or associated with recent antibiotic use
*—If Escherichia coli O157 or S	Shiga toxin-producing E.	coli, monitor complete blood cour

Escherichia coli O157 or Shiga toxin-producing *E. coli*, monitor complete blood count and basic metabolic panel for hemolytic uremic syndrome; these pathogens are more commonly found with bloody diarrhea without fever.

Information from references 2, 5, 9, and 12-14.

### STOOL TESTING

Most patients who are immunocompetent with acute, nonbloody diarrhea without systemic symptoms do not require stool testing.2,5,6,12,13 Stool studies should be obtained in patients with acute bloody diarrhea, fever, severe abdominal pain, signs of sepsis, or duration of seven days or greater, and patients with occupational or residential risk factors for an outbreak. Molecular, or culture-independent studies, also known as multiplex polymerase chain reaction or nucleic acid amplification tests, are generally preferred over traditional stool cultures.<sup>2,5,15</sup> These methods allow for rapid detection of bacterial, viral, and parasitic pathogens concurrently. Multiplex testing in inflammatory diarrhea should include Salmonella, Shigella, Campylobacter, Yersinia, C. difficile, and Shiga toxin. Specific exposures and travel history may dictate the need to test for additional pathogens. For example, patients who report swimming in brackish water or consuming undercooked shellfish warrant evaluation for Vibrio species.2

A limitation of molecular testing is that it identifies the presence of an organism but does not indicate clinical significance. Molecular methods are also of limited use in patients whose symptoms are related to an outbreak. If a clinician suspects an outbreak requiring a public health investigation, separate stool cultures are required to assist with reporting.<sup>2,5</sup> Nationally reportable causes of death include specific pathogens (i.e., Campylobacter, giardiasis, Listeria monocytogenes, and others), foodborne outbreaks, waterborne outbreaks, and postdiarrheal hemolytic uremic syndrome.2

# C. DIFFICILE

Adult patients who were recently hospitalized for more than three days or who used antibiotics within the past 12 weeks should undergo nucleic acid

<sup>†-</sup>In patients with AIDS, perform additional testing for Cryptosporidium, Cyclospora, Cystoisospora, microsporidia, Mycobacterium avium complex, and cytomegalovirus.

<sup>‡—</sup>See algorithm for the diagnosis of *C. difficile* infection (https://www.aafp.org/pubs/afp/ issues/2020/0201/p168.html#afp20200201p168-f1).14

amplification testing or multistep testing for *C. difficile*. Testing requires a fresh, nonsolid sample. Evaluation and management guidelines for *C. difficile* can be found in separate updates. <sup>14,16</sup>

### **BLOOD TESTING**

Blood testing is not routinely recommended for patients with acute diarrhea.<sup>12</sup> A chemistry panel evaluating electrolytes and kidney function is appropriate for patients with dehydration. Complete blood count with differential and blood cultures should be obtained in patients with signs of sepsis. Patients with confirmed or suspected *E. coli* O157 or other Shiga toxin infections are at risk of hemolytic uremic syndrome. A blood cell count and chemistry panels should be obtained in these patients to monitor for changes in hemoglobin, platelets, electrolytes, and kidney function.<sup>2</sup>

### NOT RECOMMENDED

Fecal leukocytes, lactoferrin, and calprotectin are of limited utility and are not recommended in the evaluation of acute infectious diarrhea.<sup>2,5,12</sup> Fecal leukocytes break down easily in samples, reducing sensitivity. Fecal lactoferrin is more sensitive and can be associated with severe dehydration and bacterial etiology, but it lacks the specificity to be clinically useful.<sup>2,15,17</sup> Fecal calprotectin, a marker of inflammatory bowel disease, has shown associations with bacterial causes for acute diarrhea, including *C. difficile*, but results lack consistency.<sup>2,17</sup> Fecal occult blood testing generally should not be used in the evaluation of diarrhea.<sup>18</sup>

### **ENDOSCOPY**

Patients with acute diarrhea do not require endoscopic evaluation unless symptoms persist for more than one month and laboratory results are nondiagnostic.<sup>5,19</sup>

Patients with HIV, AIDS, and other immunocompromised conditions are at greater risk of invasive or opportunistic infections. An endoscopic examination should be considered in patients who are immunocompromised and whose initial stool testing is nondiagnostic or whose diarrhea is functionally disabling.<sup>2,20,21</sup>

### **Management**

Treatment of acute diarrhea depends on the suspected or determined cause. Supportive therapy with early rehydration and refeeding is recommended regardless of etiology.

### REHYDRATION AND REFEEDING

Supportive management begins with replacing water, electrolytes, and nutrients. Oral rehydration is the preferred method for fluid replacement. Severe dehydration associated

### **TABLE 4**

# **Home Recipes for Oral Rehydration Solution**

Base beverage	Recipe
Chicken broth	2 cups liquid broth (not low sodium) 2 cups water 2 tablespoons sugar
Gatorade	32 oz Gatorade G2 3/4 teaspoon table salt
Water	1 quart water 1/4 teaspoon table salt 2 tablespoons sugar

Information from reference 23.

### TABLE 5

# **Empiric Antibiotics for Travelers' Diarrhea**

Antibiotic	(orally)	Three-day dose (orally)
Azithromycin (Zithromax)	1,000 mg	500 mg per day*
Ciprofloxacin	750 mg	500 mg per day
Levofloxacin (Levaquin)	500 mg	500 mg per day
Ofloxacin	400 mg	400 mg per day
Rifaximin (Xifaxan)†	None	200 mg three times per day

<sup>\*—</sup>Preferred empiric regimen for febrile illness with or without bloody stool or in areas where fluoroquinolone-resistant *Campylobacter* is common.

Adapted with permission from Riddle MS, DuPont HL, Connor BA. ACG clinical guideline: diagnosis, treatment, and prevention of acute diarrheal infections in adults. Am J Gastroenterol. 2016;111(5):610.

with shock, sepsis, or altered mental status warrants rehydration with isotonic intravenous fluid.<sup>2,5</sup>

Oral rehydration solution has decreased the worldwide mortality of acute diarrhea, particularly in children and older people. Oral rehydration has been shown to decrease total stool output, episodes of emesis, and the need for intravenous rehydration, by slowly releasing glucose into the gut to improve salt and water reabsorption.<sup>22</sup> Prepackaged oral rehydration solution is available commercially, but similar recipes can be made at home with readily available ingredients (*Table 4*).<sup>23</sup>

<sup>†—</sup>Avoid when invasive pathogens are suspected.

### TABLE 6

### Targeted Antibiotic Therapy for Infectious Diarrhea With an Identified Pathogen

Organism	Indication for treatment	First choice	Alternative
Campylobacter	All identified cases	Azithromycin (Zithromax)	Ciprofloxacin
Clostridioides difficile	All identified cases	Fidaxomicin (Dificid)*	Oral vancomycin
Giardia	All identified cases	Tinidazole (Tindamax), nitazoxanide (Alinia)	Metronidazole (Flagyl)
Nontyphoidal Salmo- nella enterica	Consider in patients older than 50 years with immunosuppression or significant joint or cardiac disease	Ceftriaxone, cipro- floxacin, TMP/SMX, amoxicillin	Not applicable
Non-Vibrio cholerae	Invasive disease only, including bloody diarrhea or sepsis	Ceftriaxone plus doxycycline	TMP/SMX plus aminoglycoside
Salmonella enterica typhi or paratyphi	All identified cases	Ceftriaxone or ciprofloxacin	Ampicillin, TMP/SMX, azithromycin
Shigella spp.	All identified cases	Azithromycin, ciproflox- acin, or ceftriaxone	TMP/SMX, ampicillin if susceptible
Vibrio cholerae	All identified cases	Doxycycline	Ciprofloxacin, azithromycin, ceftriaxone
Yersinia enterocolitica	All identified cases	TMP/SMX	Cefotaxime (Claforan), ciprofloxacin

TMP/SMX = trimethoprim/sulfamethoxazole.

Adapted with permission from Shane AL, Mody RK, Crump JA, et al. 2017 Infectious Diseases Society of America clinical practice guidelines for the diagnosis and management of infectious diarrhea. Clin Infect Dis. 2017;65(12):e66, with additional information from reference 16.

Reintroducing food as early as tolerated decreases the intestinal permeability caused by infection, reducing illness duration and improving nutritional outcomes. The evidence for a particular refeeding diet is insufficient; therefore, an age-appropriate regular diet is recommended.6,24

### ANTIDIARRHEAL MEDICATIONS

The antisecretory properties of bismuth subsalicylates and the antimotility properties of loperamide (Imodium) make these two drugs the most useful symptomatic therapies for acute watery diarrhea and can help decrease inappropriate antibiotic use. A loperamide and simethicone combination has demonstrated faster and more complete relief of acute watery diarrhea and gas-related abdominal discomfort compared with either medication alone.<sup>25</sup>

In cases of travelers' diarrhea (TD), adjunct loperamide therapy with antibiotics has been shown to increase the likelihood of a clinical cure at 24 and 48 hours.<sup>26</sup>

Loperamide and other antimotility agents should be avoided in patients with bloody or suspected inflammatory diarrhea because of the potential for toxic megacolon and prolonged illness. Bismuth subsalicylates are a safe alternative in patients with fever and inflammatory diarrhea.2,5

### **ANTIBIOTIC THERAPY**

Most cases of acute watery diarrhea are self-limited; therefore, antibiotics are not routinely recommended. To avoid overuse of antibiotics and related complications, empiric antibiotics should be used in specific instances, including moderate to severe TD, bloody stool with fever, sepsis, and immunocompromised states.

A fluoroquinolone or macrolide is an appropriate firstline option when empirically treating TD, depending on local resistance patterns and recent travel history. Antibiotics can decrease the duration of TD to just over 24 hours after dosing. 27,28 A single dose of antibiotics is usually sufficient, but three-day dosing is also an option (Table 5).5

Once a pathogen has been identified, targeted antibiotic therapy may be appropriate depending on the organism (Table 6<sup>2,16</sup>).

# **STANDBY ANTIBIOTICS**

Although previous guidelines recommended providing standby antibiotics to most travelers, studies have shown no significant illness reduction with self-treatment and have raised concerns about effects on the gut microbiota and antibiotic resistance. The decision to provide standby antibiotics should be individualized based on the patient's risk of developing moderate to severe TD.5,29

<sup>\*—</sup>Fidaxomicin is more expensive than oral vancomycin but shows less resistance and requires fewer daily doses.

### **PROBIOTICS**

There are no formal recommendations for or against probiotics to treat acute diarrheal illness due to a lack of consistent evidence. One Cochrane review found that probiotics decrease the mean duration of acute diarrhea by one day and decrease the risk of diarrhea lasting more than four days. Another large meta-analysis found a 15% relative decrease in the risk of TD with prophylactic probiotic use. <sup>30,31</sup> Other studies have shown inconclusive or insufficient evidence. The decision to use probiotics should be individualized for the patient and situation. <sup>30,31</sup>

### **Prevention**

### **HYGIENE**

Adequate handwashing, proper use of personal protective equipment, and food and water safety practices are mainstays for preventing the transmission of diarrheal illness. Proper handwashing with soap and water for at least 20 seconds before rinsing shows the most benefit in reducing transmission of the more contagious pathogens (e.g., Norovirus, Shigella) and should be promoted in settings of community or institutional outbreaks. Patients who are hospitalized, including those with *C. difficile* colitis, may require contact precautions with appropriate personal protective equipment, including gloves and a gown.

Implementation of guidelines from the Centers for Disease Control and Prevention and the U.S. Food and Drug Administration on food and water safety decreases the risk of foodborne diarrheal illness in the United States; however, the risk increases with travel. Pretravel counseling about high-risk food and beverages can be used to help prevent TD.<sup>5,9</sup> A previous article in *American Family Physician* discussed foodborne illness in more detail.<sup>9</sup>

### **VACCINATION**

Typhoid and cholera vaccines are recommended for people traveling to endemic areas or with known personal or occupational exposures.<sup>2</sup> Vaccines for other diarrhea-causing agents continue to be investigated.

### Sequelae

Postinfectious irritable bowel syndrome is a common sequela of acute gastroenteritis, occurring in 4% to 36% of patients depending on the infective pathogen, with a pooled prevalence of 11% in one meta-analysis. 32,33 Presenting as unexplained persistent diarrhea in the absence of an infectious agent, postinfectious irritable bowel syndrome is thought to be caused by an inability to downregulate intestinal inflammation following acute bacterial infection. Symptoms may last for months or years and are most common following *Campylobacter*,

### **TABLE 7**

### **Potential Postinfectious Diarrhea Sequelae**

Erythema nodosum

Glomerulonephritis

Guillain-Barré syndrome

Hemolytic anemia

Hemolytic uremic syndrome

Immunoglobulin A nephropathy

Intestinal perforation

Lactose intolerance

Meningitis

Postinfectious irritable bowel syndrome

Reactive arthritis

Information from reference 2.

Shigella, or Salmonella infection. Effective prevention and treatment of acute bacterial illness decrease the chances of developing postinfectious irritable bowel syndrome. <sup>32,33</sup> Potential postinfectious diarrhea sequelae are listed in *Table 7.*<sup>2</sup>

### COVID-19

Gastrointestinal adverse effects are common in patients with SARS-CoV-2. Symptoms vary among patients with COVID-19, and diarrhea may be the only presenting complaint for some. COVID-19–associated diarrhea is self-limited but may be severe in some patients. The overall clinical significance of COVID-19–associated diarrhea is unclear but causes concern about potential fecal-oral transmission of the virus.<sup>34</sup>

This article updates a previous article on this topic by Barr and Smith.<sup>12</sup>

**Data Sources:** A search was completed using PubMed, EBSCOhost, DynaMed, Essential Evidence Plus, and the Cochrane database. Key terms searched were acute diarrhea, travelers' diarrhea, acute diarrhea evaluation, acute diarrhea testing, acute diarrhea treatment, diarrhea antibiotics, acute diarrhea prevention, postinfectious diarrhea, and COVID diarrhea. Also searched were the Centers for Disease Control and Prevention and World Health Organization websites. Search dates: August 2021 to January 2022, and April 2022.

The opinions and assertions contained herein are the private views of the authors and are not to be construed as official or as reflecting the views of the Uniformed Services University of the Health Sciences, the U.S. Army Medical Department, or the U.S. Army at large.

### **ACUTE DIARRHEA**

### The Authors

**ERICA S. MEISENHEIMER, MD, MBA, FAAFP,** is a faculty physician at the National Capital Consortium Family Medicine Residency at Fort Belvoir (Va.) Community Hospital, and an assistant professor in the Department of Family Medicine at the Uniformed Services University of the Health Sciences, Bethesda, Md.

**CARLY EPSTEIN, DO,** is a chief resident in the Department of Family Medicine at Fort Belvoir Community Hospital.

**DERRICK THIEL, MD, MPH,** is an adjunct faculty member at the Madigan Army Medical Center Family Medicine Residency, Joint Base Lewis-McChord, Wash., and an assistant professor at the Uniformed Services University of the Health Sciences.

Address correspondence to Erica S. Meisenheimer, MD, MBA, 9300 Dewitt Loop, Fort Belvoir, VA 22060 (email: erica.j. meisenheimer.mil@mail.mil). Reprints are not available from the authors.

### References

- 1. Scallan E, Griffin PM, Angulo FJ, et al. Foodborne illness acquired in the United States—unspecified agents. *Emerg Infect Dis.* 2011;17(1):16-22.
- Shane AL, Mody RK, Crump JA, et al. 2017 Infectious Diseases Society of America clinical practice guidelines for the diagnosis and management of infectious diarrhea. Clin Infect Dis. 2017;65(12):e45-e80.
- 3. Imhoff B, Morse D, Shiferaw B, et al.; Emerging Infections Program FoodNet Working Group. Burden of self-reported acute diarrheal illness in FoodNet surveillance areas, 1998-1999. *Clin Infect Dis.* 2004; 38(suppl 3):S219-S226.
- 4. Jones TF, McMillian MB, Scallan E, et al. A population-based estimate of the substantial burden of diarrhoeal disease in the United States; Food-Net, 1996-2003. *Epidemiol Infect*. 2007;135(2):293-301.
- Riddle MS, DuPont HL, Connor BA. ACG clinical guideline: diagnosis, treatment, and prevention of acute diarrheal infections in adults. Am J Gastroenterol. 2016;111(5):602-622.
- Guerrant RL, Van Gilder T, Steiner TS, et al.; Infectious Diseases Society of America. Practice guidelines for the management of infectious diarrhea. Clin Infect Dis. 2001;32(3):331-351.
- 7. World Health Organization. Diarrhoeal disease. 2017. Accessed September 27, 2021. https://www.who.int/news-room/fact-sheets/detail/diarrhoeal-disease
- 8. Burgers K, Lindberg B, Bevis ZJ. Chronic diarrhea in adults: evaluation and differential diagnosis. *Am Fam Physician*. 2020;101(8):472-480.
- 9. Switaj TL, Winter KJ, Christensen SR. Diagnosis and management of foodborne illness. *Am Fam Physician*. 2015;92(5):358-365.
- Arasaradnam RP, Brown S, Forbes A, et al. Guidelines for the investigation of chronic diarrhoea in adults: British Society of Gastroenterology, 3rd edition. Gut. 2018;67(8):1380-1399.
- Jones TF, Bulens SN, Gettner S, et al. Use of stool collection kits delivered to patients can improve confirmation of etiology in foodborne disease outbreaks. Clin Infect Dis. 2004;39(10):1454-1459.
- 12. Barr W, Smith A. Acute diarrhea. Am Fam Physician. 2014;89(3):180-189.
- DuPont HL. Acute infectious diarrhea in immunocompetent adults. N Engl J Med. 2014;370(16):1532-1540.
- Mounsey A, Lacy Smith K, Reddy VC, et al. Clostridioides difficile infection: update on management. Am Fam Physician. 2020;101(3): 168-175

- 15. Lee HM, Lee S, Lee BI, et al. Clinical significance of fecal lactoferrin and multiplex polymerase chain reaction in patients with acute diarrhea. *Gut Liver.* 2015;9(5):636-640.
- Johnson S, Lavergne V, Skinner AM, et al. Clinical practice guideline by the Infectious Diseases Society of America and Society for Healthcare Epidemiology of America: 2021 focused update guidelines on management of Clostridioides difficile infection in adults. Clin Infect Dis. 2021; 73(5):755-757.
- 17. Weh J, Antoni C, Weiß C, et al. Discriminatory potential of C-reactive protein, cytokines, and fecal markers in infectious gastroenteritis in adults. *Diagn Microbiol Infect Dis*. 2013;77(1):79-84.
- 18. Lee MW, Pourmorady JS, Laine L. Use of fecal occult blood testing as a diagnostic tool for clinical indications: a systematic review and meta-analysis. *Am J Gastroenterol*. 2020;115(5):662-670.
- 19. Wahnschaffe U, Ignatius R, Loddenkemper C, et al. Diagnostic value of endoscopy for the diagnosis of giardiasis and other intestinal diseases in patients with persistent diarrhea from tropical or subtropical areas. *Scand J Gastroenterol*. 2007;42(3):391-396.
- Cello JP, Day LW. Idiopathic AIDS enteropathy and treatment of gastrointestinal opportunistic pathogens [published correction appears in *Gastroenterology*. 2009;137(1):393]. *Gastroenterology*. 2009;136(6): 1952-1965.
- 21. Krones E, Högenauer C. Diarrhea in the immunocompromised patient. *Gastroenterol Clin North Am.* 2012;41(3):677-701.
- Gregorio GV, Gonzales MLM, Dans LF, et al. Polymer-based oral rehydration solution for treating acute watery diarrhoea. Cochrane Database Syst Rev. 2016;(12):CD006519.
- 23. Parrish CR, DiBaise JK. Hydrating adult patient with short bowel syndrome. *Pract Gastroenterol*. 2015;39(2):10-18.
- 24. Duggan C, Nurko S. Feeding the gut: the scientific basis for continued enteral nutrition during acute diarrhea. *J Pediatr.* 1997;131(6):801-808.
- Hanauer SB, DuPont HL, Cooper KM, et al. Randomized, double-blind, placebo-controlled clinical trial of loperamide plus simethicone versus loperamide alone and simethicone alone in the treatment of acute diarrhea with gas-related abdominal discomfort. Curr Med Res Opin. 2007; 23(5):1033-1043.
- 26. Riddle MS, Arnold S, Tribble DR. Effect of adjunctive loperamide in combination with antibiotics on treatment outcomes in traveler's diarrhea: a systematic review and meta-analysis. *Clin Infect Dis.* 2008;47(8): 1007-1014.
- 27. De Bruyn G, Hahn S, Borwick A. Antibiotic treatment for travellers' diarrhoea. *Cochrane Database Syst Rev.* 2000;(3):CD002242.
- Adachi JA, Ericsson CD, Jiang ZD, et al. Azithromycin found to be comparable to levofloxacin for the treatment of US travelers with acute diarrhea acquired in Mexico. Clin Infect Dis. 2003;37(9):1165-1171.
- 29. Riddle MS, Ericsson CD, Gutierrez RL, et al. Stand-by antibiotics for travellers' diarrhoea: risks, benefits and research needs. *J Travel Med*. 2018; 25(1):1-3.
- 30. Wilkins T, Sequoia J. Probiotics for gastrointestinal conditions: A summary of the evidence. *Am Fam Physician*. 2017;96(3):170-178.
- Collinson S, Deans A, Padua-Zamora A, et al. Probiotics for treating acute infectious diarrhoea. Cochrane Database Syst Rev. 2020;(12): CD003048.
- 32. Connor BA, Riddle MS. Post-infectious sequelae of travelers' diarrhea. *J Travel Med*. 2013;20(5):303-312.
- Klem F, Wadhwa A, Prokop LJ, et al. Prevalence, risk factors, and outcomes of irritable bowel syndrome after infectious enteritis: a systematic review and meta-analysis. Gastroenterology. 2017;152(5):1042-1054.e1.
- 34. Megyeri K, Dernovics Á, Al-Luhaibi ZII, et al. COVID-19-associated diarrhea. World J Gastroenterol. 2021;27(23):3208-3222.