

# **IDSA Diarrhea Guidelines**

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Infectious Diseases Society of America





# IDSA

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## *Strategic Priorities...*



Establishing state of the art clinical guidelines



Advocating for ID prevention and public health funding



Leading the way in antimicrobial stewardship and addressing resistance



# IDSA

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## *Strategic Priorities...*



Promoting the value of ID Specialists



Bringing the best and brightest into our field



Putting ID and HIV research front, center, and into practice

# Disclosure

- I have no disclosures



# Objectives

- Highlight the structure of the 2017 IDSA Clinical Practice Guidelines for diagnosis and management of infectious diarrhea
- Replace IDSA guidelines published in 2001
- Discuss the impact of infectious diarrhea on clinical practice and public health
- Update diagnosis and treatment



# Promote Timely & Relevant **Guidelines**

- Implementing adherence to GRADE Methodology for all IDSA Practice Guidelines
- Have increased staff resources and technical expertise to make guideline development process more efficient



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ABSTRACT

RECOMMENDATIONS

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ADDITIONAL RESOURCES AND PRODUCTS

## "Clinical Practice Guidelines for Clostridium difficile Infection in Adults and Children: 2017 Update by the Infectious Diseases Society of America (IDSA) and Society for Healthcare Epidemiology of America (SHEA)

**CURRENT\***

Published: Clinical Infectious Diseases ; 2018 ; ; -

### Abstract

A panel of experts was convened by the Infectious Diseases Society of America (IDSA) and Society for Healthcare Epidemiology of America (SHEA) to update the 2010 clinical practice guideline on Clostridium difficile infection (CDI) in adults. The update, which has incorporated recommendations for children (following the adult recommendations for epidemiology, diagnosis, and treatment), includes significant changes in the management of this infection and reflects the evolving controversy over best methods for diagnosis. Clostridium difficile remains the most important cause of healthcare-associated diarrhea and has become the most commonly identified cause of healthcare-associated infection in adults in the United States. Moreover, C. difficile has established itself as an important community pathogen. Although the prevalence of the epidemic and virulent ribotype 027 strain has declined markedly along with overall CDI rates in parts of Europe, it remains one of the most commonly identified strains in the United States where it causes a sizable minority of CDIs, especially healthcare-associated CDIs. This guideline updates recommendations regarding epidemiology, diagnosis, treatment, infection prevention, and environmental management.

[Full text](#)

\*Every 12 to 10 months following publication, IDSA reviews its guidelines to determine whether an update is required. The guideline was published February of 2010 and is the most current version.

# Panel Composition for Diarrhea Guidelines

- Karen Kotloff
- Joseph Cantey
- Allen Chang
- John Crump
- Joanne Langley
- Rajal Mody
- Larry Pickering\*
- Andi Shane\*
- Theodore Steiner
- Christine Wanke
- Cirle Warren
- Phillip Tarr



# Practice Guidelines



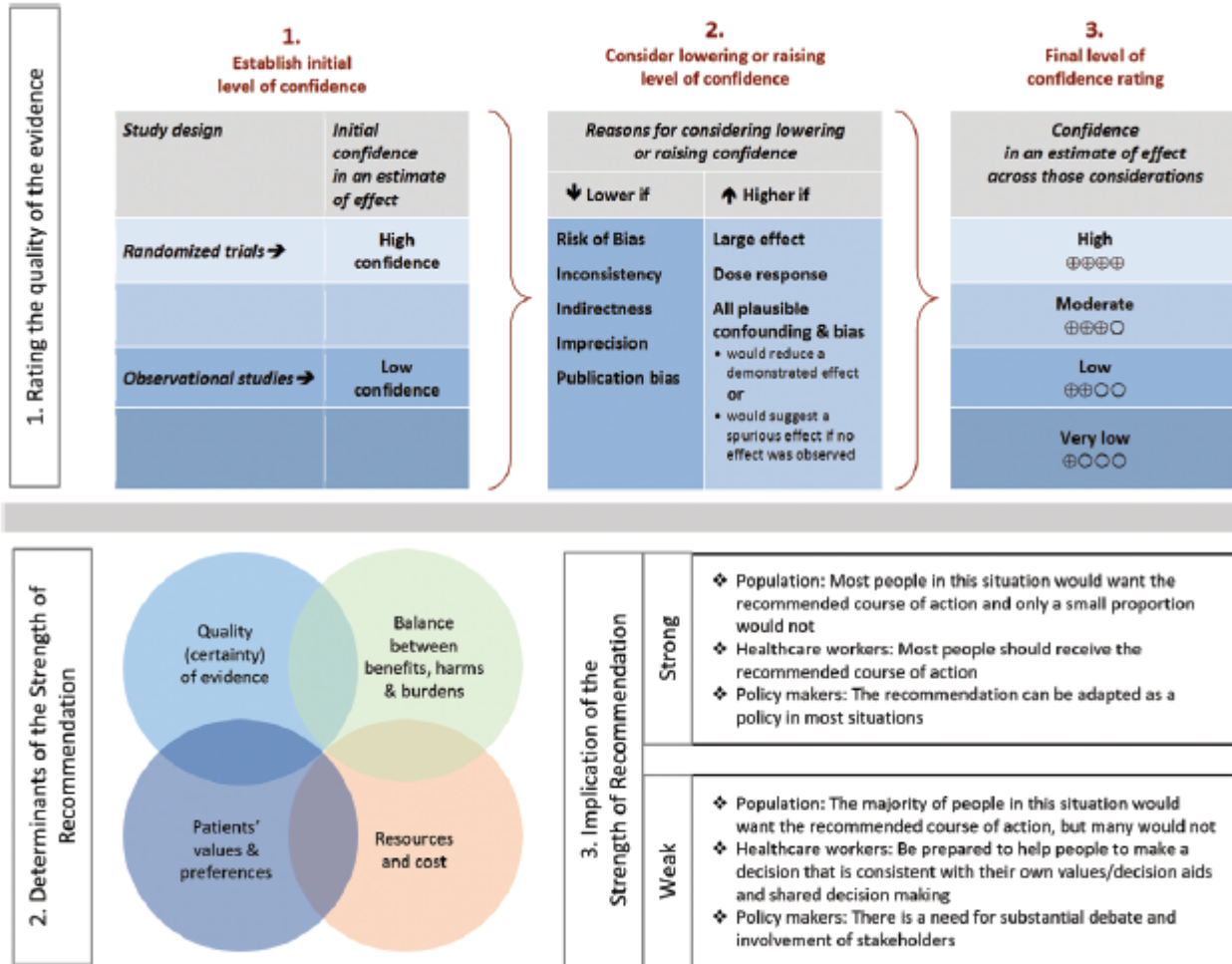
The Infectious Diseases Society of America (IDSA) represents physicians, scientists and other health care professionals who specialize in infectious diseases. IDSA's purpose is to improve the health of individuals, communities, and society by promoting excellence in patient care, education, research, public health, and prevention relating to infectious diseases.

- **Practice guidelines are systematically developed statements to assist practitioners and patients in making decisions about appropriate health care for specific clinical circumstances.**
- **Attributes of good guidelines include validity, reliability, reproducibility, clinical applicability, clinical flexibility, clarity, multidisciplinary process, review of evidence, and documentation.**

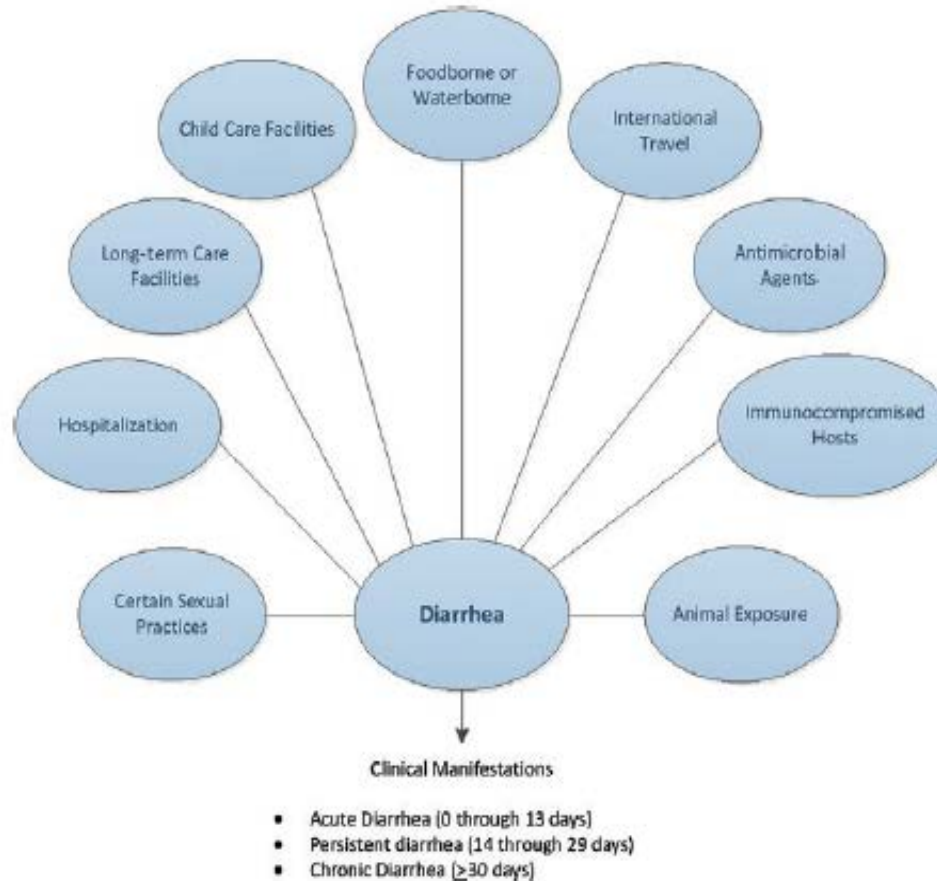
**[Institute of Medicine Committee to Advise the Public Health Service on Clinical Practice Guidelines, 1990]**



# GRADE Approach



# Considerations When Evaluating People With Infectious Diarrhea



**Figure 1.** Considerations when evaluating people with infectious diarrhea. Modified from Long SS, Pickering LK, Pober CG, eds. Principles and Practice of Pediatric Infectious Diseases, 4th ed. New York: Elsevier Saunders, 2012.

# Structure of the Diarrhea Guidelines

- ❑ **Clinical, demographic and epidemiologic features (10 headings with 29 recommendations)**
- ❑ **Empiric management of infectious diarrhea (2 headings with 8 recommendations)**
- ❑ **Directed management of infectious diarrhea (1 heading with 1 recommendation)**
- ❑ **Supportive treatment (2 headings with 7 recommendations)**
- ❑ **Ancillary management (3 headings with 6 recommendations)**
- ❑ **Prevention (3 headings with 9 recommendations)**



# Burden of diarrheal diseases in the U.S.

- ❑ 179 million outpatient visits
- ❑ 500,000 hospitalizations
- ❑ 5,000 deaths
- ❑ Norovirus and *Salmonella enterica* subspecies were the leading pathogens among the 24 gastrointestinal tract pathogens transmitted by food
- ❑ Missing organisms: rotavirus (vaccine) and norovirus (not reportable)



# Summary of Top 10 Nationally Notifiable Diseases in the United States, 2015

- Three cause gastroenteritis (Salmonella, Shigella, Giardia) (missing are rotavirus (vaccine) and norovirus (not reportable))
- Four are sexually transmitted
- Two are vaccine-preventable
- One is caused by a tickborne spirochete
- All 10 involve children, adolescents, and adults



# Tables included in guidelines

1. Modes of acquisition of enteric organisms and sources of guidelines
2. Exposure or condition associated with pathogens causing diarrhea
3. Clinical presentations suggestive of infectious diarrhea etiologies
4. Post-infectious manifestations associated with enteric pathogens
5. Laboratory diagnostics for organisms associated with infectious diarrhea
6. Recommended antimicrobial agents by pathogen
7. Fluid and nutritional management of diarrhea



# Table 1: Modes of Acquisition of Enteric Organisms and Sources of Guidelines

**Table 1. Modes of Acquisition of Enteric Organisms and Sources of Guidelines**

Mode	Title	URL	Author/Issuing Agency
International travel	Expert Review of the Evidence Base for Prevention of Travelers' Diarrhea	<a href="http://www.ncbi.nlm.nih.gov/pubmed/19538575">http://www.ncbi.nlm.nih.gov/pubmed/19538575</a>	DuPont et al [113]
	Medical Considerations Before International Travel	<a href="http://www.ncbi.nlm.nih.gov/pubmed/27468061">http://www.ncbi.nlm.nih.gov/pubmed/27468061</a>	Freedman et al [207]
	The Yellow Book	<a href="http://wwwnc.cdc.gov/travel/page/yellowbook-home-2014">http://wwwnc.cdc.gov/travel/page/yellowbook-home-2014</a>	CDC
	Travelers Health	<a href="http://wwwnc.cdc.gov/travel">http://wwwnc.cdc.gov/travel</a>	CDC
Immunocompromised hosts	Guidelines for the Prevention and Treatment of Opportunistic Infections in HIV-Infected Adults and Adolescents	<a href="http://aidsinfo.nih.gov/contentfiles/vguidelines/adult_ol.pdf">http://aidsinfo.nih.gov/contentfiles/vguidelines/adult_ol.pdf</a>	CDC/NIH/HIVMA/IDSA
	Guidelines for the Prevention and Treatment of Opportunistic Infections in HIV-Exposed and HIV-Infected Children	<a href="http://aidsinfo.nih.gov/contentfiles/vguidelines/ol_guidelines_pediatrics.pdf">http://aidsinfo.nih.gov/contentfiles/vguidelines/ol_guidelines_pediatrics.pdf</a>	CDC/NIH/HIVMA/IDSA
Foodborne and waterborne	Surveillance for Foodborne Disease Outbreaks—United States, 2009–2010	<a href="http://www.cdc.gov/mmwr/preview/mmwrhtml/mm6203a1.htm?s_cid=mm6203a1_w">http://www.cdc.gov/mmwr/preview/mmwrhtml/mm6203a1.htm?s_cid=mm6203a1_w</a>	CDC
	Food Safety	<a href="http://www.cdc.gov/foodsafety/">http://www.cdc.gov/foodsafety/</a>	CDC
	Healthy Water	<a href="http://wwwnc.cdc.gov/healthywater">http://wwwnc.cdc.gov/healthywater</a>	CDC
Antimicrobial-associated ( <i>C. difficile</i> )	Clinical Practice Guidelines for <i>Clostridium difficile</i> Infection in Adults and Children 2017 Update (In press)	<a href="http://www.jstor.org/stable/10.1086/651706">http://www.jstor.org/stable/10.1086/651706</a>	IDSA/SHEA
	2010 Clinical Practice Guidelines for <i>Clostridium difficile</i> Infection in Adults	<a href="https://www.idsociety.org/Organ_System/#Clostridiumdifficile">https://www.idsociety.org/Organ_System/#Clostridiumdifficile</a>	IDSA/SHEA
Healthcare-associated	Healthcare-Associated Infections	<a href="http://www.cdc.gov/hai/">http://www.cdc.gov/hai/</a>	CDC
Child care settings	Caring for Our Children: National Health and Safety Performance Standards; Guidelines for Early Care and Education Programs	<a href="http://nrckids.org">http://nrckids.org</a>	AAP, APHA, NRC
	Recommendations for Care of Children in Special Circumstances—Children in Out-of-Home Child Care (pp 132–51)	<a href="http://redbook.solutions.aap.org/redbook.aspx">http://redbook.solutions.aap.org/redbook.aspx</a>	AAP
	Managing Infectious Diseases in Child Care and Schools	<a href="http://ebooks.aapublications.org/content/managing-infectious-diseases-in-child-care-and-schools-3rd-edition">http://ebooks.aapublications.org/content/managing-infectious-diseases-in-child-care-and-schools-3rd-edition</a>	AAP
Long-term care settings	Nursing Homes and Assisted Living (Long-term Care Facilities)	<a href="http://www.cdc.gov/longtermcare/">http://www.cdc.gov/longtermcare/</a>	CDC
	Infection Prevention and Control in the Long-term Care Facility	<a href="http://www.shea-online.org/assets/files/position-papers/ic-itc97.pdf">http://www.shea-online.org/assets/files/position-papers/ic-itc97.pdf</a>	SHEA/APIC
Zoonoses	Compendium of Measures to Prevent Disease Associated With Animals in Public Settings	<a href="http://www.cdc.gov/mmwr/preview/mmwrhtml/r6004a1.htm?s_cid=r6004a1_w">http://www.cdc.gov/mmwr/preview/mmwrhtml/r6004a1.htm?s_cid=r6004a1_w</a>	CDC
	Exposure to Nontraditional Pets at Home and to Animals in Public Settings: Risks to Children	<a href="http://pediatrics.aappublications.org/content/122/4/876">http://pediatrics.aappublications.org/content/122/4/876</a>	Pickering et al [51]
	Review of Institute of Medicine and National Research Council Recommendations for One Health Initiative	<a href="http://wwwnc.cdc.gov/eid/article/19/12/12-1659_article.htm">http://wwwnc.cdc.gov/eid/article/19/12/12-1659_article.htm</a>	Rubin et al [208]

Abbreviations: AAP, American Academy of Pediatrics; APHA, American Public Health Association; APIC, Association for Professionals in Infection Control and Epidemiology; CDC, Centers for Disease Control and Prevention; HIV, human immunodeficiency virus; HIVMA, HIV Medicine Association; IDSA, Infectious Diseases Society of America; NIH, National Institutes of Health; NRC, National Resource Center for Health and Safety in Child Care and Early Education; SHEA, Society for Healthcare Epidemiology of America.



# Modes of Acquisition of Enteric Organisms

- International travel |
- Immunocompromised hosts
- Foodborne and waterborne
- Antimicrobial associated (C.difficile)
- Healthcare associated
- Child care settings
- Long term care settings
- Zoonoses



# Diseases Associated with Raw Milk

<b>Before milk pasteurized in 1920s</b>	Tuberculosis Diphtheria Severe streptococcal infections Typhoid fever
<b>Current diseases</b>	Campylobacter 55% Salmonella 22% STEC 14% Listeria/Brucella/Shigella each 3%





## POLICY STATEMENT

# Consumption of Raw or Unpasteurized Milk and Milk Products by Pregnant Women and Children

COMMITTEE ON INFECTIOUS DISEASES and COMMITTEE ON NUTRITION

### KEY WORDS

raw milk/milk products, unpasteurized milk/milk products, pregnant women, children

### ABBREVIATIONS

AAP—American Academy of Pediatrics

FDA—Food and Drug Administration

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## abstract

FREE

Sales of raw or unpasteurized milk and milk products are still legal in at least 30 states in the United States. Raw milk and milk products from cows, goats, and sheep continue to be a source of bacterial infections attributable to a number of virulent pathogens, including *Listeria monocytogenes*, *Campylobacter jejuni*, *Salmonella* species, *Brucella* species, and *Escherichia coli* O157. These infections can occur in both healthy and immunocompromised individuals, including older adults, infants, young children, and pregnant women and their unborn fetuses, in whom life-threatening infections and fetal miscarriage can occur. Efforts to limit the sale of raw milk products have met with opposition from those who are proponents of the purported health benefits of consuming raw milk products, which contain nat-

# Sporadic *Salmonella* Infections

- 1.4 million nontyphoidal human *Salmonella* infections occur annually in the U.S.
- 15,000 hospitalizations
- 400 deaths
- Reptile associated salmonellosis represents 6% of salmonellosis and 11% in people < 21 years of age
- Reptile or amphibian associated Salmonellosis: 74,000 per year



# Table 2: Exposure or Condition Associated With Specific Pathogens Causing Diarrhea

Exposure or Condition	Pathogen(s)
<b>Foodborne</b>	
Foodborne outbreaks in hotels, cruise ships, resorts, restaurants, catered events	Norovirus, nontyphoidal <i>Salmonella</i> , <i>Clostridium perfringens</i> , <i>Bacillus cereus</i> , <i>Staphylococcus aureus</i> , <i>Campylobacter</i> spp, ETEC, STEC, <i>Listeria</i> , <i>Shigella</i> , <i>Cyclospora cayetanensis</i> , <i>Cryptosporidium</i> spp
Consumption of unpasteurized milk or dairy products	<i>Salmonella</i> , <i>Campylobacter</i> , <i>Yersinia enterocolitica</i> , <i>S. aureus</i> toxin, <i>Cryptosporidium</i> , and STEC. <i>Listeria</i> is infrequently associated with diarrhea, <i>Brucella</i> (goat milk cheese), <i>Mycobacterium bovis</i> , <i>Coxiella burnetii</i>
Consumption of raw or undercooked meat or poultry	STEC (beef), <i>C. perfringens</i> (beef, poultry), <i>Salmonella</i> (poultry), <i>Campylobacter</i> (poultry), <i>Yersinia</i> (pork, chitterlings), <i>S. aureus</i> (poultry), and <i>Trichinella</i> spp (pork, wild game meat)
Consumption of fruits or unpasteurized fruit juices, vegetables, leafy greens, and sprouts	STEC, nontyphoidal <i>Salmonella</i> , <i>Cyclospora</i> , <i>Cryptosporidium</i> , norovirus, hepatitis A, and <i>Listeria monocytogenes</i>
Consumption of undercooked eggs	<i>Salmonella</i> , <i>Shigella</i> (egg salad)
Consumption of raw shellfish	<i>Vibrio</i> species, norovirus, hepatitis A, <i>Plesiomonas</i>
<b>Exposure or contact</b>	
Swimming in or drinking untreated fresh water	<i>Campylobacter</i> , <i>Cryptosporidium</i> , <i>Giardia</i> , <i>Shigella</i> , <i>Salmonella</i> , STEC, <i>Plesiomonas shigelloides</i>
Swimming in recreational water facility with treated water	<i>Cryptosporidium</i> and other potentially waterborne pathogens when disinfectant concentrations are inadequately maintained
Healthcare, long-term care, prison exposure, or employment	Norovirus, <i>Clostridium difficile</i> , <i>Shigella</i> , <i>Cryptosporidium</i> , <i>Giardia</i> , STEC, rotavirus
Child care center attendance or employment	Rotavirus, <i>Cryptosporidium</i> , <i>Giardia</i> , <i>Shigella</i> , STEC
Recent antimicrobial therapy	<i>C. difficile</i> , multidrug-resistant <i>Salmonella</i>
Travel to resource-challenged countries	<i>Escherichia coli</i> (enteroaggregative, enterotoxigenic, enteroinvasive), <i>Shigella</i> , Typhi and nontyphoidal <i>Salmonella</i> , <i>Campylobacter</i> , <i>Vibrio cholerae</i> , <i>Entamoeba histolytica</i> , <i>Giardia</i> , <i>Blastocystis</i> , <i>Cyclospora</i> , <i>Cystoisospora</i> , <i>Cryptosporidium</i>
Exposure to house pets with diarrhea	<i>Campylobacter</i> , <i>Yersinia</i>
Exposure to pig feces in certain parts of the world	<i>Balantidium coli</i>
Contact with young poultry or reptiles	Nontyphoidal <i>Salmonella</i>
Visiting a farm or petting zoo	STEC, <i>Cryptosporidium</i> , <i>Campylobacter</i>
<b>Exposure or condition</b>	
Age group	Rotavirus (6–18 months of age), nontyphoidal <i>Salmonella</i> (infants from birth to 3 months of age and adults >50 years with a history of atherosclerosis), <i>Shigella</i> (1–7 years of age), <i>Campylobacter</i> (young adults)
Underlying immunocompromising condition	Nontyphoidal <i>Salmonella</i> , <i>Cryptosporidium</i> , <i>Campylobacter</i> , <i>Shigella</i> , <i>Yersinia</i>
Hemochromatosis or hemoglobinopathy	<i>Y. enterocolitica</i> , <i>Salmonella</i>
AIDS, immunosuppressive therapies	<i>Cryptosporidium</i> , <i>Cyclospora</i> , <i>Cystoisospora</i> , microsporidia, <i>Mycobacterium avium</i> –intercellulare complex, cytomegalovirus
Anal-genital, oral-anal, or digital-anal contact	<i>Shigella</i> , <i>Salmonella</i> , <i>Campylobacter</i> , <i>E. histolytica</i> , <i>Giardia lamblia</i> , <i>Cryptosporidium</i> as well as sexually transmitted infections

Abbreviations: ETEC, enterotoxigenic *Escherichia coli*; STEC, Shiga toxin-producing *Escherichia coli*.



# Exposure or condition associated with pathogens

- Foodborne outbreaks
- Unpasteurized milk or dairy products
- Raw shellfish
- Raw or undercooked meat or poultry
  
- Swimming
- Childcare center
- Travel
- Farm or petting zoo



# Known Causes of Foodborne Illness Outbreaks, U.S., 2006–2010

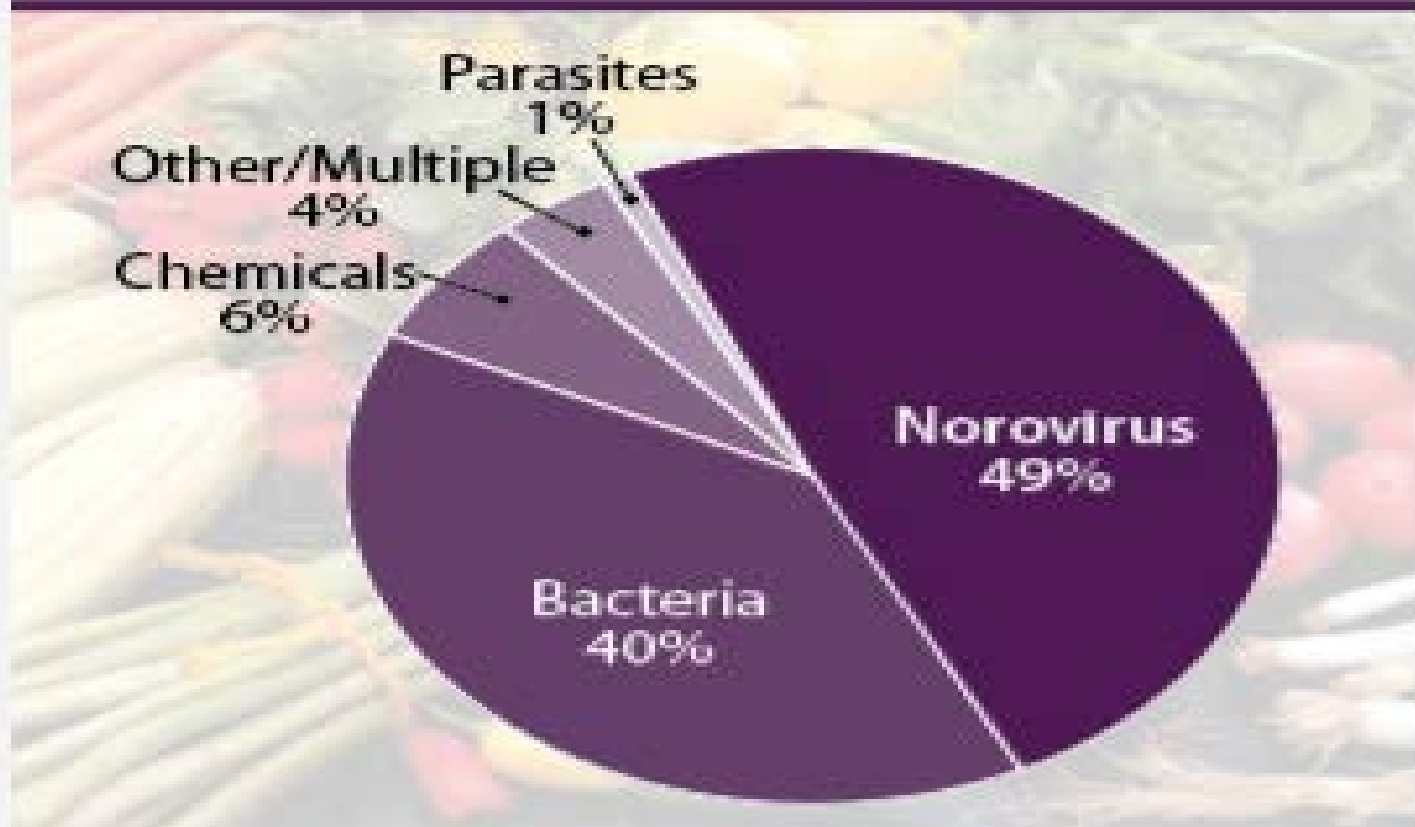


Figure 3: Known Causes of Foodborne Illness Outbreaks, U.S., 2006-2010. Norovirus 49%; Bacteria 40%; Chemicals 6%; Parasites 1%; Other/Multiple 4%.

# Table 3: Clinical Presentations Suggestive of Infectious Diarrhea Etiologies

**Table 3. Clinical Presentations Suggestive of Infectious Diarrhea Etiologies**

Finding	Likely Pathogens
Persistent or chronic diarrhea	<i>Cryptosporidium</i> spp., <i>Giardia lamblia</i> , <i>Cyclospora cayentanensis</i> , <i>Cystoisospora belli</i> , and <i>Entamoeba histolytica</i>
Visible blood in stool	STEC, <i>Shigella</i> , <i>Salmonella</i> , <i>Campylobacter</i> , <i>Entamoeba histolytica</i> , noncholera <i>Vibrio</i> species, <i>Yersinia</i> , <i>Balantidium coli</i> , <i>Plesiomonas</i>
Fever	Not highly discriminatory—viral, bacterial, and parasitic infections can cause fever. In general, higher temperatures are suggestive of bacterial etiology or <i>E. histolytica</i> . Patients infected with STEC usually are not febrile at time of presentation
Abdominal pain	STEC, <i>Salmonella</i> , <i>Shigella</i> , <i>Campylobacter</i> , <i>Yersinia</i> , noncholera <i>Vibrio</i> species, <i>Clostridium difficile</i>
Severe abdominal pain, often grossly bloody stools (occasionally nonbloody), and minimal or no fever	STEC, <i>Salmonella</i> , <i>Shigella</i> , <i>Campylobacter</i> , and <i>Yersinia enterocolitica</i>
Persistent abdominal pain and fever	<i>Y. enterocolitica</i> and <i>Y. pseudotuberculosis</i> ; may mimic appendicitis
Nausea and vomiting lasting <24 hours	Ingestion of <i>Staphylococcus aureus</i> enterotoxin or <i>Bacillus cereus</i> (short-incubation emetic syndrome)
Diarrhea and abdominal cramping lasting 1–2 days	Ingestion of <i>Clostridium perfringens</i> or <i>B. cereus</i> (long-incubation emetic syndrome)
Vomiting and nonbloody diarrhea lasting 2–3 days or less	Norovirus (low-grade fever usually present during the first 24 hours in 40% if infections)
Chronic watery diarrhea, often lasting a year or more	Brainerd diarrhea (etiologic agent has not been identified); postinfectious irritable bowel syndrome

Abbreviation: STEC, Shiga toxin-producing *Escherichia coli*.



# Clinical presentations suggestive of etiologies

- Persistent or chronic: parasite
- Visible blood: STEC, shigella, campylobacter, *Balantidium coli*
- Severe abdominal pain: similar to above
- Fever: not highly discriminatory
- Persistent abdominal pain and fever: *Yersinia enterocolitica* and *Y. pseudotuberculosis*: may mimic appendicitis
- Vomiting and nonbloody diarrhea for 2-3 days: norovirus



# Table 4: Postinfectious Manifestations Associated With Enteric Infections

**Table 4. Postinfectious Manifestations Associated With Enteric Pathogens**

Manifestation	Organism(s)
Erythema nodosum	<i>Yersinia</i> , <i>Campylobacter</i> , <i>Salmonella</i> , <i>Shigella</i>
Glomerulonephritis	<i>Shigella</i> , <i>Campylobacter</i> , <i>Yersinia</i>
Guillain-Barré syndrome	<i>Campylobacter</i>
Hemolytic anemia	<i>Campylobacter</i> , <i>Yersinia</i>
Hemolytic uremic syndrome	STEC, <i>Shigella dysenteriae</i> serotype 1
Immunoglobulin A nephropathy	<i>Campylobacter</i>
Reactive arthritis <sup>a</sup>	<i>Salmonella</i> , <i>Shigella</i> , <i>Campylobacter</i> , <i>Yersinia</i> , rarely <i>Giardia</i> , and <i>Cyclospora cayentanensis</i>
Postinfectious irritable bowel syndrome	<i>Campylobacter</i> , <i>Salmonella</i> , <i>Shigella</i> , STEC, <i>Giardia</i>
Meningitis	<i>Listeria</i> , <i>Salmonella</i> (infants <3 months of age are at high risk)
Intestinal perforation	<i>Salmonella</i> including <i>Salmonella</i> Typhi, <i>Shigella</i> , <i>Campylobacter</i> , <i>Yersinia</i> , <i>Entamoeba histolytica</i>
Ekiri syndrome (lethal, toxic encephalopathy) and/or seizure	<i>Shigella</i>
Aortitis, osteomyelitis, extravascular deep tissue focus	<i>Salmonella</i> , <i>Yersinia</i>

Abbreviation: STEC, Shiga toxin-producing *Escherichia coli*.

<sup>a</sup>Includes Reiter syndrome.



# Postinfectious Manifestations

- Guillain-Barre syndrome: *Campylobacter*
- Hemolytic uremic syndrome: STEC, *S. dysenteriae*
- Meningitis: *Listeria*, *Salmonella* (infants < 3 months)
- Ekiri syndrome (lethal, toxic encephalopathy, seizure): *Shigella*
- Aortitis, osteo, deep tissue: *Salmonella*, *Yersinia*



# Table 5: Laboratory Diagnostics for Organisms Associated With Infectious Diarrhea

**Table 5. Laboratory Diagnostics for Organisms Associated With Infectious Diarrhea**

Etiologic Agent	Diagnostic Procedures	Optimal Specimen
<i>Clostridium difficile</i>	NAAT GDH antigen with or without toxin detection followed by cytotoxin or <i>Clostridium difficile</i> toxin or toxigenic <i>C. difficile</i> strain	Stool
<i>Salmonella enterica</i> , <i>Shigella</i> spp, <i>Campylobacter</i> spp	Routine stool enteric pathogen culture <sup>a</sup> or NAAT	Stool
<i>Salmonella enterica</i> serovars Typhi and Paratyphi (enteric fever)	Routine culture	Stool, blood, bone marrow, and duodenal fluid
Shiga toxin-producing <i>Escherichia coli</i>	Culture for <i>E. coli</i> O157:H7 <sup>b</sup> and Shiga toxin immunoassay or NAAT for Shiga toxin genes	Stool
<i>Yersinia</i> spp, <i>Plesiomonas</i> spp, <i>Edwardsiella tarda</i> , <i>Staphylococcus aureus</i> , <i>E. coli</i> (enterotoxigenic, enteroinvasive, enteropathogenic, enteroaggregative)	Specialized stool culture or molecular assays <sup>c</sup> or NAAT	Stool
<i>Clostridium perfringens</i>	Specialized procedure for toxin detection <sup>d</sup>	Stool
<i>Bacillus cereus</i> , <i>S. aureus</i>	Specialized procedure for toxin detection <sup>d</sup>	Food
<i>Clostridium botulinum</i>	Mouse lethality assay (performed at a state public health laboratory, or CDC) <sup>e,f,g</sup>	Serum, stool, gastric contents, vomitus
<i>Entamoeba histolytica</i> ; <i>Blastocystis hominis</i> <sup>h</sup> ; <i>Dientamoeba fragilis</i> <sup>h</sup> ; <i>Balantidium coli</i> ; <i>Giardia lamblia</i> ; nematodes (generally not associated with diarrhea) including <i>Ascaris lumbricoides</i> , <i>Strongyloides stercoralis</i> , <i>Trichuris trichiura</i> , hookworms; cestodes (tapeworms); trematodes (flukes)	Ova and parasite examination including permanent stained smear <sup>i</sup> or NAAT	Stool Duodenal fluid for <i>Giardia</i> and <i>Strongyloides</i>
<i>E. histolytica</i>	<i>E. histolytica</i> species-specific immunoassay or NAAT	Stool
<i>G. lamblia</i>	EIA or NAAT	Stool
<i>Cryptosporidium</i> spp [121] <sup>j</sup>	Direct fluorescent immunoassay, EIA, or NAAT	Stool
<i>Cyclospora cayentanensis</i> , <i>Cystoisospora belli</i> <sup>k</sup>	Modified acid-fast stain <sup>k</sup> performed on concentrated specimen, ultraviolet fluorescence microscopy, or NAAT	Stool
Microsporidia (now classified as a fungus)	Modified trichrome stain <sup>k</sup> performed on concentrated specimen Histologic examination with electron microscopic confirmation	Stool Small bowel biopsy
Calicivirus (norovirus, sapovirus) <sup>k</sup> ; enteric adenovirus; enterovirus/parechovirus <sup>k</sup> ; rotavirus	NAAT	Stool
Rotavirus, enteric adenovirus	EIA	Stool
Enteric adenovirus <sup>l</sup> ; enterovirus/parechovirus	Viral culture	Stool
Cytomegalovirus	Histopathological examination Cytomegalovirus culture	Biopsy Biopsy



# Indications for laboratory investigation

- Epidemiologic considerations (child care center, nursing home)
- Immunocompromised host
- Extraintestinal manifestations
- Potential to impact management
- Bloody diarrhea
- Suspicion of an outbreak
- Travel
- Suspect Shiga toxin producing organism (HUS)



# Laboratory diagnosis by etiologic agent

- Optimal stool specimen
- Diagnostic procedures
  - Routine or specialized stool specimen
  - Nucleic acid amplification test
  - Toxin detection assay
  - Specific immunoassay
  - Microscopy
  - Histology



# Multiplex gastrointestinal tract panels

- Facilitate appropriate treatment
- Avoid unnecessary antimicrobial treatment
- Expedite recognition of foodborne, waterborne and other outbreaks



# Culture-independent diagnostic tests (CIDTs)

- Multiplex GI panels rapidly detect a wide range of bacterial, viral and parasitic organisms
- Guide management in specific areas: child care, foodborne, nursing homes
- Facilitate public health surveillance
- Detect pathogens unsuspected by clinicians
- Reflex directed cultures may be needed for susceptibility testing



# Multiplex GI Panels: Bacteria

- *Campylobacter* spp.
- *C. difficile*
- *Plesiomonas*
- *Salmonella* spp.
- *Vibrio* spp.
- EAEC, EPEC, ETEC, STEC (*E. coli* 0157)
- *Shigella*, EIEC

.....

- **bioMerieux Film Array**



# Multiplex GI Panels: Viruses and Parasites Detected

- Viruses: Adenovirus 40/41
- Astrovirus
- Norovirus
- Rotavirus
- Sapovirus
- Parasites: Cryptosporidium
- *Cyclospora cayetanensis*
- *Entamoeba histolytica*
- *Giardia*

**bioMerieux Film Array**



## Other diagnostic techniques to establish a diagnosis

- Fecal leukocyte and stool lactoferrin: NO
- Serologic tests: NO, but exception is post-diarrheal HUS
- White blood cell count: NO but may be useful)
- Endoscopy or proctoscopy: persistent diarrhea and AIDS, underlying medical conditions, clinical colitis and proctitis, anal intercourse,
- Duodenal aspirate for giardia, strongyloides, microsporidia, cystoisospora
- Imaging: occasionally using CT, MRI, ultrasound



# Serial stool specimens

- Generally not recommended except for public health reasons and for culture dependent methods for susceptibility testing
- Public health reasons: return to child care, work, or group social activities and other areas where transmission is a consideration



# Table 6: Recommended Antimicrobial Agents by Pathogen

Indication	First Choice	Alternative	Comments/Considerations
<b>Bacteria<sup>a</sup></b>			
<i>Campylobacter</i> <i>Clostridium difficile</i>	Azithromycin Oral vancomycin	Ciprofloxacin Fidaxomicin	Fidaxomicin not currently recommended for people <18 years of age. Metronidazole is still acceptable treatment for nonsevere CDI in children and as a second-line agent for adults with nonsevere CDI (eg, who cannot obtain vancomycin or fidaxomicin at a reasonable cost).
Nontyphoidal <i>Salmonella enterica</i> <sup>b</sup>	Usually not indicated for uncomplicated infection	NA	Antimicrobial therapy should be considered for groups at increased risk for invasive infection: neonates (up to 3 months old), persons >50 years old with suspected atherosclerosis, persons with immunosuppression, cardiac disease (valvular or endovascular), or significant joint disease. If susceptible, treatment with ceftriaxone, ciprofloxacin, TMP-SMX, or amoxicillin.
<i>Salmonella enterica</i> Typhi or Paratyphi <sup>b</sup> Shigella <sup>b</sup>	Ceftriaxone or ciprofloxacin  Azithromycin <sup>c</sup> or ciprofloxacin <sup>c</sup> , or ceftriaxone	Ampicillin or TMP-SMX or azithromycin  TMP-SMX or ampicillin if susceptible	Clinicians treating people with shigellosis for whom antibiotic treatment is indicated should avoid prescribing fluoroquinolones if the ciprofloxacin MIC is 0.12 µg/mL or higher even if the laboratory report identifies the isolate as susceptible. See <a href="https://www.emergency.cdc.gov/henhan00401.asp">https://www.emergency.cdc.gov/henhan00401.asp</a>
<i>Vibrio cholerae</i>	Doxycycline <sup>d</sup>	Ciprofloxacin, azithromycin, or ceftriaxone	
Non- <i>Vibrio cholerae</i> <sup>d</sup>	Usually not indicated for noninvasive disease. Single-agent therapy for noninvasive disease if treated.  Invasive disease: ceftriaxone plus doxycycline	Usually not indicated for noninvasive disease. Single-agent therapy for noninvasive disease if treated.  Invasive disease: TMP-SMX plus an aminoglycoside	
<i>Yersinia enterocolitica</i>	TMP-SMX	Cefotaxime or ciprofloxacin	
<b>Parasites</b>			
<i>Cryptosporidium</i> spp	Nitazoxanide (HIV-uninfected, HIV-infected in combination with effective cART):	Effective cART: Immune reconstitution may lead to microbiologic and clinical response [154, 200, 210]	NA
<i>Cyclospora cayentensis</i>	TMP-SMX	Nitazoxanide (limited data)	Patients with HIV infection may require higher doses or longer durations of TMP-SMX treatment
<i>Giardia lamblia</i>	• Tinidazole Note: Based on data from HIV-uninfected children • Nitazoxanide	Metronidazole Note: Based on data from HIV-uninfected children	• Tinidazole is approved in the United States for children aged >3 years. It is available in tablets that can be crushed. • Metronidazole has high frequency of gastrointestinal side effects. A pediatric suspension of metronidazole is not commercially available but can be compounded from tablets. Metronidazole is not FDA approved for the treatment of giardiasis.
<i>Cystoisospora belli</i>	TMP-SMX	Pyrimethamine Potential second-line alternatives: • Ciprofloxacin • Nitazoxanide	
<i>Trichinella</i> spp	Albendazole	Alternative: mebendazole	• Therapy less effective in late stage of infection, when larvae encapsulate in muscle
<b>Fungus</b>			
Microsporidia	For disseminated (not ocular) and intestinal infection attributed to microsporidia other than <i>Enterocytozoon bieneusi</i> or <i>Vitiforma corneae</i> : • Albendazole after initiation of cART and resolution of signs and symptoms For <i>E. bieneusi</i> or <i>V. corneae</i> infections: • Fumagillin recommended for treatment of infections due to <i>E. bieneusi</i> in HIV-infected adults	NA	Effective cART therapy: • Immune reconstitution may lead to microbiologic and clinical response • Fumagillin for systemic use is unavailable in the United States and data on dosing in children are unavailable. • Consultation with an expert is recommended.



# Antimicrobial agents by pathogen

- Campylobacter: azithromycin/ciprofloxacin
- *C. difficile*: oral vancomycin/fidaxomicin
- Nontyphoidal Salmonella: none
- Typhoid or paratyphoid fever: ceftriaxone/cipro
- Shigella: azithromycin or cipro or ceftriaxone
- *Vibrio cholerae*: doxycycline/cipro or azithromycin or ceftriaxone
- *Yersinia enterocolitica*: TMX-SMX



# No Clinical Benefit of Empirical Antimicrobial Therapy for Pediatric Diarrhea in a High-Use, High-Resistance Setting.

Duong VT<sup>1,2</sup>, Tuyen HT<sup>1</sup>, Van Minh P<sup>1</sup>, Campbell JI<sup>1</sup>, Phuc HL<sup>2</sup>, Nhu TDH<sup>1</sup>, Tu LTP<sup>1,3</sup>, Chau TTH<sup>1</sup>, Nhi LTQ<sup>1,4</sup>, Hung NT<sup>2</sup>, Ngoc NM<sup>5</sup>, Huong NTT<sup>5</sup>, Thompson CN<sup>7</sup>, Thwaites GE<sup>1,7</sup>, de Alwis R<sup>1,7</sup>, Baker S<sup>1,7,8</sup>.

## + Author information

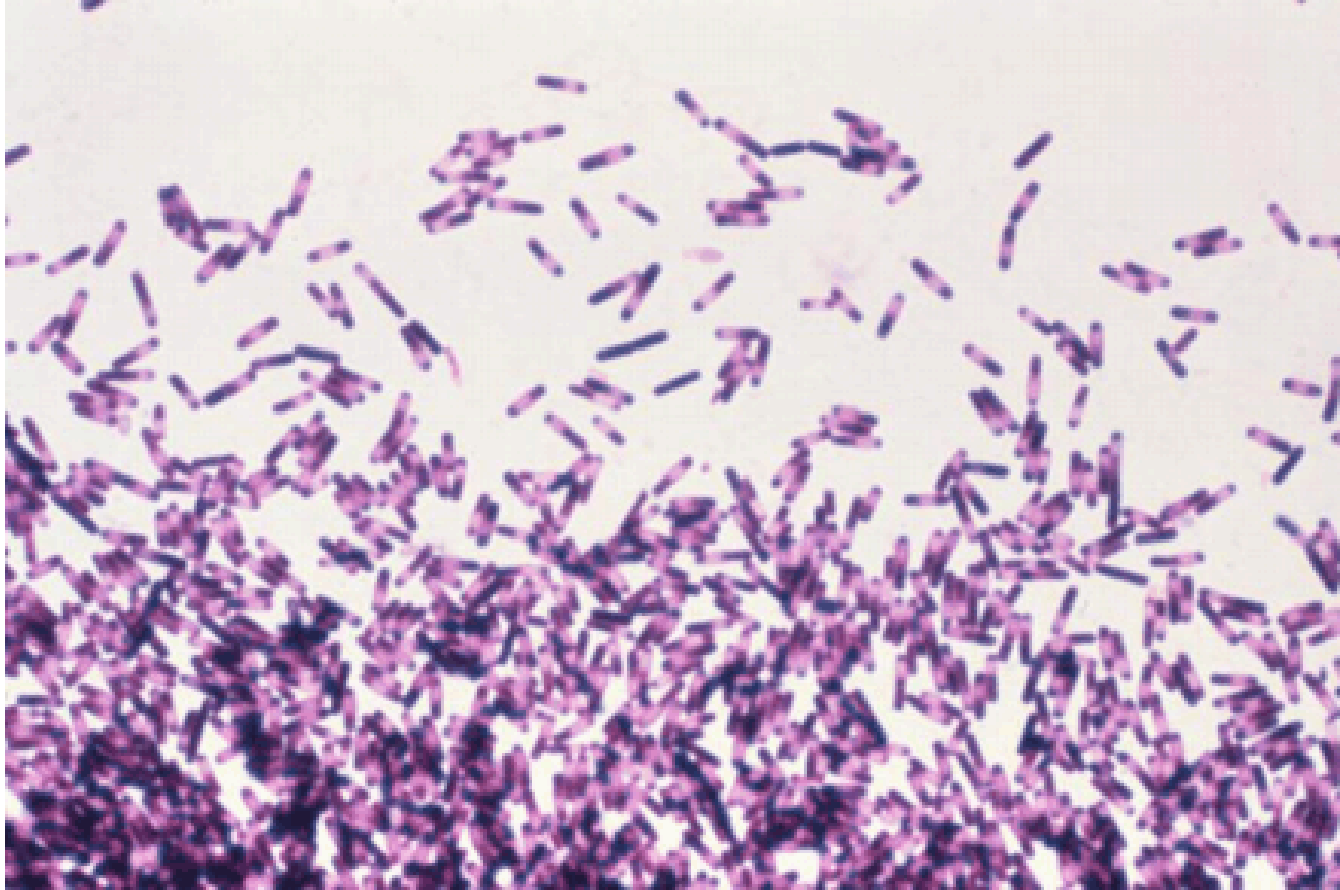
### Abstract

**BACKGROUND:** Pediatric diarrheal disease presents a major public health burden in low- to middle-income countries. The clinical utility of empirical antimicrobial treatment for diarrhea are unclear in settings that lack reliable diagnostics and have high antimicrobial resistance (AMR).

**METHODS:** We conducted a prospective multicenter cross-sectional study of pediatric patients hospitalized with diarrhea containing blood and/or mucus in Ho Chi Minh City, Vietnam. Clinical parameters, including disease outcome and treatment, were measured. Shigella, nontyphoidal Salmonella (NTS), and Campylobacter were isolated from fecal samples, and their antimicrobial susceptibility profiles were determined. Statistical analyses, comprising log-rank tests and accelerated failure time models, were performed to assess the effect of empirical antimicrobials on disease outcome.



# Diarrheal Disease



*Red Book Online Visual Library. Clostridium difficile*



# Directed management of infectious diarrhea

- Rehydration
- Human milk
- Diet
- Antimotility drugs (generally not recommended)
- Antinausea and antiemetic drugs (occasionally)
- Probiotic preparations
- Oral zinc supplementation



# Vaccines that prevent diarrhea

- Rotavirus (routine recommendation for infants)
- Typhoid (two vaccines, one oral and one injectable, are available in the U.S. but are not routinely recommended)
- Cholera (live attenuated vaccine available as a single dose oral vaccine in the U.S. for adults 18-64 years of age who travel to cholera-affected areas)
- See <https://www.cdc.gov/vaccines/hcp/acip-recs/index.html>



## Trends in Rate of Seizure-Associated Hospitalizations Among Children <5 Years Old Before and After Rotavirus Vaccine Introduction in the United States, 2000-2013.

[Pringle KD](#)<sup>1,2</sup>, [Burke RM](#)<sup>1,2</sup>, [Steiner CA](#)<sup>3</sup>, [Parashar UD](#)<sup>1</sup>, [Tate JE](#)<sup>1</sup>.

### + Author information

### Abstract

**BACKGROUND:** Rotavirus is a common cause of acute gastroenteritis and has also been associated with generalized tonic-clonic afebrile seizures. Since rotavirus vaccine introduction, hospitalizations for treatment of acute gastroenteritis have decreased. We assess whether there has been an associated decrease in seizure-associated hospitalizations.

**METHODS:** We used discharge codes to abstract data on seizure hospitalizations among children <5 years old from the State Inpatient Databases of the Healthcare Cost and Utilization Project. We compared seizure hospitalization rates before and after vaccine introduction, using Poisson regression, stratifying by age and by month and year of admission. We performed a time-series analysis with negative binomial models, constructed using prevaccine data from 2000 to 2006 and controlling for admission month and year.

**RESULTS:** We examined 962899 seizure hospitalizations among children <5 years old during 2000-2013. Seizure rates after vaccine introduction were lower than those before vaccine introduction by 1%-8%, and rate ratios decreased over time. Time-series analyses demonstrated a decrease in the number of seizure-coded hospitalizations in 2012 and 2013, with notable decreases in children 12-17 months and 18-23 months.

# Vaccine Recommendations and Guidelines of the ACIP

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[Smallpox \(Vaccinia\)](#)

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# Table 7: Fluid and Nutritional Management of Diarrhea

**Table 7. Fluid and Nutritional Management of Diarrhea**

Degree of Dehydration <sup>a</sup>	Rehydration Therapy	Replacement of Losses During Maintenance <sup>c</sup>
Mild to moderate dehydration	Infants <sup>b</sup> and children: ORS, 50–100 mL/kg over 3–4 hours Adolescents and adults (>30 kg): ORS, 2–4 L	Infants and children: <10 kg body weight: 60–120 mL ORS for each diarrheal stool or vomiting episode, up to ~500 mL/day >10 kg body weight: 120–240 mL ORS for each diarrheal stool or vomiting episode; up to ~1 L/day Adolescents and adults: Ad libitum, up to ~2 L/day Replace losses as above as long as diarrhea or vomiting continues
Severe dehydration	Infants: Malnourished infants may benefit from smaller-volume, frequent boluses of 10 mL/kg body weight due to reduced capacity to increase cardiac output with larger volume resuscitation. Children, adolescents, and adults: Intravenous isotonic crystalloid boluses, per current fluid resuscitation guidelines, until pulse, perfusion, and mental status return to normal. Adjust electrolytes and administer dextrose based on chemistry values. Administer up to 20 mL/kg body weight until pulse, perfusion, and mental status return to normal.	Infants and children: <10 kg body weight: 60–120 mL ORS for each diarrheal stool or vomiting episode, up to ~500 mL/day >10 kg body weight: 120–240 mL ORS for each diarrheal stool or vomiting episode; up to ~1 L/day Adolescents and adults: Ad libitum, up to ~2 L/day Replace losses as above as long as diarrhea or vomiting continues. If unable to drink, administer either through a nasogastric tube or give 5% dextrose 0.25 normal saline solution with 20 mEq/L potassium chloride intravenously.

Adapted from Centers for Disease Control and Prevention. Managing acute gastroenteritis among children: oral rehydration, maintenance, and nutritional therapy. MMWR Recomm Rep 2003; 52(RR-16):1–16 and World Health Organization. The treatment of diarrhoea: a manual for physicians and other senior health workers (<https://www.cdc.gov/mmwr/preview/mmwrhtml/rr5216a1.htm>).

Low-osmolarity ORS can be given to all age groups, with any cause of diarrhea. It is safe in the presence of hyponatremia as well as hyponatremia (except when edema is present). Some commercially available formulations that can be used as ORS include Pedialyte Liters (Abbott Nutrition), CeraLyte (Cero Products), and Enfalac Lytren (Mead Johnson). Popular beverages that should not be used for rehydration include apple juice, Gatorade, and commercial soft drinks.

Abbreviation: ORS, oral rehydration solution.

<sup>a</sup>A variety of scales are available to grade the severity of dehydration in young children but no single, standard, validated method exists. Note that signs of dehydration may be masked when a child is hypernatremic.

<sup>b</sup>Breastfed infants should continue nursing throughout the illness.

<sup>c</sup>After rehydration is complete, maintenance fluids should be resumed along with an age-appropriate normal diet offered every 3–4 hours. Children previously receiving a lactose-containing formula can tolerate the same product in most instances. Diluted formula does not appear to confer any benefit.



# Prevention

- Hand hygiene
- Infection control
- Food safety practices
- Direct educational efforts
- What people with diarrhea should avoid
- Report nationally notifiable diseases
- Vaccines
  - Rotavirus (infants)
  - Typhoid (children and adults)
  - Cholera (adults)



# Unusual Features of O104:H4 Strains from German

- Large percentage of case patients had HUS
- HUS occurred in adults
- Frequent development of neurologic symptoms in patients when clinical and laboratory markers of HUS were improving



# So, what was the causative food?

- A. Peppers
- B. Tomatoes
- C. Lettuce
- D. Sprouts
- E. Spinach



# Food Safety



USDA Meat and Poultry Hotline  
1-888-MPHOTLINE  
(1-888-674-6854)

E-mail: [mphotlinea.fsis@usda.gov](mailto:mphotlinea.fsis@usda.gov)  
Web site: [fsis.outreach@usda.gov](mailto:fsis.outreach@usda.gov)

For People with  
**HIV/AIDS**

*A need-to-know guide  
for those who have been  
diagnosed with*