

# (Toxigenic Vibrio cholerae O1, O139)

**Note:** Only toxigenic strains of *Vibrio cholerae* serogroups O1 and O139 cause epidemics and are reportable as cholera. This guidance is intended for management of patients with toxigenic strains of *V. cholerae* serogroups O1 and O139 and early management (i.e., before laboratory confirmation is available) of patients with cholera-like illness returning from regions where cholera activity has been reported (e.g., travelers returning from Haiti or the Dominican Republic).

For management of non-toxigenic strains of *V. cholerae* O1 and O139, toxigenic strains of other *V. cholerae* serogroups (e.g. O75 and O141), and other *Vibrio* species, refer to the guidance for vibriosis.

PR	OTOCOL CHECKLIST
	Enter available information into Merlin within 24 hours of notification Review background on disease, case definition, laboratory testing (section 2, 3, and 4)  Contact provider (section 5)
	<ul> <li>Confirm diagnosis</li> <li>Obtain available demographic and clinical information</li> <li>Determine what information was provided to the patient</li> </ul>
	Ensure collection and submission of appropriate specimens (section 4) Interview patient(s) (section 5)
_	Review disease facts (section 2)
	<ul><li>Description of illness</li><li>Modes of transmission</li></ul>
	<ul><li>☐ Ask about exposure to relevant risk factors (section 5)</li><li>☐ Travel to an area affected by cholera</li></ul>
	Exposure to untreated water sources
	<ul> <li>Exposure to raw shellfish or undercooked seafood</li> <li>Consumption of food imported from an area affected by cholera</li> <li>Pre-existing conditions</li> </ul>
	Identify any similar cases of illness among contacts
	<ul> <li>Determine whether case-patient or symptomatic contact is in a sensitive situation</li> <li>Provide recommendations for case management and management of contacts</li> </ul>
	☐ Infection control (section 6) ☐ Management of contacts (section 6)
	☐ Management of sensitive situations (section 7)
	<ul><li>☐ Provide education on transmission and prevention (section 6)</li><li>☐ Review cholera prevention messages</li></ul>
	Review proper hand washing
	Address case-patient's questions or concerns If associated with seafood exposure in the United States
	Determine source
	<ul> <li>Determine if an environmental investigation is warranted</li> <li>Provide education on transmission and prevention (section 6)</li> </ul>
	Avoid consumption of raw oysters and other raw shellfish
	<ul> <li>Avoid cross-contamination of food with raw seafood or juices from raw seafood</li> <li>Appropriate handling of shellfish</li> </ul>
	Follow-up on special situations, including outbreaks or case-patients in sensitive situations
	(section 7)

# Cholera (Toxigenic Vibrio cholerae O1, O139) Guide to Surveillance and Investigation Update Merlin case entry with any additional information Complete and submit the extended data in Merlin containing information from the Centers for Disease Control and Prevention's (CDC) Cholera and Other Vibrio Illness Surveillance (COVIS) form for all cases.

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#### 1. DISEASE REPORTING

# A. Purpose of reporting and surveillance

- 1. To identify exposures and sources of transmission and to take public health action to prevent additional cases.
- 2. To identify populations at risk and provide health education to reduce risk of exposure.
- 3. To monitor trends in the epidemiology of toxigenic *V. cholerae* O1 or O139.

#### B. Legal reporting requirements

Laboratories and physicians are required to report cholera to the county health department (CHD) immediately upon initial suspicion or laboratory test order.

#### C. County health department investigation responsibilities

- 1. Investigate all cases as soon as possible, but no later than one business day, after receiving a report from a provider or laboratory (Section 5).
- 2. Ensure appropriate control measures are taken to prevent the spread of cholera locally (Section 6). Instruct contacts what actions to take if they suspect they have cholera.
- 3. Ensure that laboratory specimens are submitted to the Bureau of Public Health Laboratories (BPHL) for confirmatory testing and additional characterization (Section 4).
- Report all cases by telephone to the Bureau of Epidemiology (BOE) Regional Environmental Epidemiologist who will help notify the CDC and other Department of Health (DOH) staff.
- 5. Enter cases in Merlin (Cholera [Vibrio cholerae Type O1] [Merlin disease code=00190]) within one business day of notification.
- 6. For all cases, complete the extended data in Merlin. The extended data in Merlin collects information for the COVIS Report form which can be accessed here:

  FloridaHealth.gov/diseases-and-conditions/disease-reporting-and-management/disease-reporting-and-surveillance/\_documents/crf-cholera.pdf

#### 2. THE DISEASE AND ITS EPIDEMIOLOGY

#### A. Etiologic agents

Cholera is an acute intestinal infection caused by toxigenic *Vibrio cholera* serogroups O1 and O139. Although other *V. cholerae* serogroups and non-toxigenic strains of *V. cholerae* O1 and O139 cause cholera-like illness, only toxigenic strains of serogroup O1 and O139 cause epidemics and are reportable as "cholera." *V. cholerae* O1 has two biotypes, Classical and El Tor. Each biotype has two distinct serotypes, Inaba and Ogawa. Cholera is a major cause of epidemic diarrhea in developing countries and causes an estimated 3–5 million cases and 100,000–120,000 deaths annually.

#### **B.** Description of illness

The clinical picture for cholera ranges from asymptomatic infection to severe diarrheal illness. Approximately 75% of persons infected with *V. cholerae* are asymptomatic. Most persons with symptoms develop a mild to moderate diarrheal illness that is difficult to differentiate from other causes of diarrheal disease. Five to ten percent of infections cause severe disease. Signs generally appear 24–48 hours after inoculation. Symptoms

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generally begin with the sudden onset of painless watery diarrhea that is often followed by vomiting often before becoming voluminous. Patients also may experience accompanying abdominal cramps. Severe cholera is characterized by acute profuse watery diarrhea, sometimes described as "rice-water stools," that can lead to rapid fluid loss and hypovolemic shock. Additional symptoms of severe illness may include vomiting, tachycardia, loss of skin turgor, muscle cramps, dry mucous membranes, hypotension, vascular collapse, dehydration, and thirst. Without treatment, death can occur within hours.

#### C. Reservoirs

The main reservoir is humans, although environmental reservoirs also exist. Toxigenic *V. cholerae* O1 and O139 are naturally occurring in fresh and brackish waters where reservoirs include shellfish and plankton. In clusters without definite human transport, ballast water from maritime shipping has been implicated.

#### D. Modes of transmission

Cholera transmission occurs by ingestion of contaminated water or food. In the outbreak setting, fecal contamination of water and food supplies by infected individuals contributes to the widespread transmission of cholera. Transmission can also occur through consumption of raw or undercooked shellfish from waters where natural reservoirs of *V. cholerae* exist. Person-to-person transmission is rarely documented.

#### E. Incubation period

The incubation period for cholera ranges from a few hours to five days (usually 2-3 days).

# F. Period of communicability

*V. cholerae* is communicable for as long as stool cultures are positive. Stools typically remain positive only a few days following resolution of symptoms. A long-term carrier state, as long as several months, is uncommon and not thought to play an important role in the transmission of disease.

#### G. Treatment

Prompt and adequate rehydration is the cornerstone of treatment. Immediate administration of oral rehydration solution (ORS) or intravenous fluids can dramatically reduce case-fatality rates. Antimicrobial therapy is indicated for patients with severe diarrhea and has been shown to reduce fluid requirements and shorten the duration of diarrhea and *Vibrio* excretion. Antibiotics commonly used to treat cholera include doxycycline, azithromycin, erythromycin, and ciprofloxacin. When possible and in the setting of an outbreak, antibiotic choice should be based on antimicrobial susceptibility testing. Additional details about rehydration and antibiotic treatment are available at <a href="https://www.cdc.gov/cholera/treatment/index.html">www.cdc.gov/cholera/treatment/index.html</a>.

# H. Prophylaxis

None indicated in the United States.

#### I. Vibrio cholerae in Florida

Cholera in Florida is almost exclusively due to travel outside the United States. Florida experienced an increased number of cases of imported cholera during the Latin American cholera epidemic in the early 1990s. No cases of illness due to toxigenic *V. cholerae* were reported from 1997–2009. Cases associated with travel to Hispaniola were reported following identification of the outbreak in October 2010. Since 2013, cases of cholera have been reported in travelers returning from Cuba and Haiti.

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#### 3. CASE DEFINITIONS

#### A. Clinical criteria for case classification

An infection of variable severity characterized by diarrhea and/or vomiting; severity is variable.

#### B. Laboratory criteria for case classification

Either of the following:

- Isolation of toxigenic (i.e., cholera toxin-producing) Vibrio cholerae O1 or O139 from stool or vomitus or
- Serologic evidence of recent infection (testing performed at the CDC).

#### C. Epidemiological criteria for case classification

Not applicable.

#### D. Case classification

Confirmed: a clinically compatible illness in a person with laboratory evidence.

# E. Criteria to distinguish a new case from previous reports

Not applicable.

#### F. Comments

Illnesses caused by strains of *V. cholerae* other than toxigenic *V. cholerae* O1 or O139 should not be reported as cases of cholera.

The etiologic agent of a case of cholera should be reported as either *V. cholerae* O1 or *V. cholerae* O139.

Infections due to V. cholerae non-O1 should be reported as vibriosis (Vibrio cholerae Type Non-O1) (Merlin reporting code = 00198).

Any available isolates of the organism must be sent to BPHL for confirmation and serotyping. Toxigenic production for *V. cholerae* O1 or O139 must be performed by CDC. This condition has been identified as a potential bioterrorism agent by the CDC.

#### 4. LABORATORY TESTING

#### A. Criteria for diagnosis

Isolation of toxigenic (i.e., cholera toxin-producing) *Vibrio cholerae* O1 or O139 from stool or vomitus OR serologic evidence of recent infection.

Isolation of *Vibrio cholerae* from stool is widely available at most hospital and commercial laboratories. The test request must be specific for *Vibrio cholerae* because identification of the organism by culture requires special techniques that are not routinely performed.

Any available isolates of the organism must be sent to BPHL for confirmation and serotyping.

Confirmation of toxin production must be performed by the CDC. Serologic testing is only available at CDC.

#### B. Specimens for laboratory testing

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- 1. Stool: Isolation of *Vibrio cholerae* from stool is the preferred method of diagnosis for cholera.
- 2. Serum: Please consult with BPHL prior to the collection and submission of sera for testing. Serology may be performed by CDC under select circumstances. For optimal results, three serum samples (S1-S3) should be collected for each case according to the following time schedule:
  - a. Serum 1 (S1): Collect as soon as possible following onset of symptoms: ideally within the first three days.
  - b. Serum 2 (S2): Collect 10-14 days after illness onset, but no later than three weeks after illness onset.
  - c. Serum 3 (S3): Collect about two weeks after S2, but no later than six weeks after illness onset.

If the optimal time for collection of S1 has passed, collect S2 and S3 specimens. If more than three weeks has elapsed from illness onset, then collect S3 only.

#### C. Testing requests

- 1. Submitting specimens/isolates to BPHL
  - a. All submissions should be accompanied by Clinical Lab Submission Form 1847 FloridaHealth.gov/diseases-and-conditions/disease-reporting-and-management/disease-reporting-and-surveillance/surveillance-and-investigation-guidance/\_documents/dh1847clinicallabsubmissionform.pdf.
  - b. Electronic Laboratory Ordering (ELO) may also be used by entering a request into the Health Management System (HMS) State Laboratory System, placing a bar coded label on the Cary-Blair vial, and writing the date collected on the vial.
- 2. Specimen collection and storage
  - a. Stool or rectal swabs should be transported in Cary-Blair medium at room temperature (not refrigerated or frozen). Cary-Blair medium should be at room temperature before stool is place in the media.
  - b. Isolates should be submitted in a screw-top agar slant that will support growth (e.g., blood agar).
  - c. Storage of stool specimens and isolates prior to shipping should be at ambient temperatures. Prolonged storage in the refrigerator may adversely affect the laboratory's ability to culture the organism.
  - d. Serum should be separated from the clot or packed cells in a serum separator tube within one hour of collection and transferred to a separate sterile tube for storage and shipment. Storage should be at 4 degrees Celsius or frozen if stored for more than two weeks.
- 3. Packaging and shipping
  - a. Contact BPHL prior to shipment.
  - b. Stool specimens and isolates should be shipped in a cooler without cool-packs.
  - c. Serum: If refrigerated, specimens should be shipped in a cooler with cool-packs. If frozen, specimens should be shipped using dry ice. Specimens should be shipped to the state laboratory in Jacksonville, Florida, for transfer to CDC for testing. Specify the type of testing requested and that the specimen should be forwarded to CDC for testing on the submission form.
  - d. Contact BPHL for packaging and shipping training dates. BPHL conducts approximately 20 face-to-face trainings per year all over Florida, free of charge. DOH employees must register for the classes in the DOH online training system, TRAIN.

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For shipping guidance, contact BPHL. Additional shipping trainings are also available commercially through vendors.

 Contact the regional laboratory with questions: FloridaHealth.gov/programs-andservices/public-health-laboratories/index.html

# 5. CASE INVESTIGATION

#### A. Contact the physician or hospital

- 1. Confirm diagnosis of a Vibrio cholerae infection.
- 2. Obtain and review available medical records and note the following:
  - a. Date of onset
  - b. Signs and symptoms
  - c. Predisposing conditions (e.g., immunosuppression)
  - d. Request available test results
  - e. Treatment (especially antimicrobials)
  - f. Current clinical status
- 3. Ensure collection and submission of required clinical specimens for the confirmation of cholera.
- 4. Determine that the patient is aware of the diagnosis and ascertain what information has been provided.
- 5. Obtain demographic information, including contact information (home, cellular, pager, and work numbers). Determine current location of the patient (i.e., hospital or home) and the preferred method of contact.
- 6. Notify the physician that you will be contacting the patient as DOH follows up on all cases of cholera to assess risk factors, to better characterize the occurrence of cholera infections in Florida, and to identify potential means for preventing further illness. It may also be appropriate at this point to determine if the physician has any concerns about the health department contacting the patient.

#### B. Interview the patient

- 1. Contact the patient by telephone, home visit, or visit to the hospital. Interviews should be completed as soon as possible after being reported to optimize recall.
- 2. The extended data in Merlin must be completed for all cases or you can use the COVIS form to capture the information before entering into Merlin. This form can be found at FloridaHealth.gov/diseases-and-conditions/disease-reporting-and-management/disease-reporting-and-surveillance/\_documents/crf-cholera.pdf. Ensure that the relevant data for this form is collected during the interview.
- 3. For interviews, try to develop a timeline for household and close contacts, shared meals, and recreational activities.
- 4. Items to cover during interview include:
  - a. Briefly review general information for this disease.
  - b. Exposures during exposure period (seven days prior to onset):
    - i. Travel outside Florida or the United States. Determine dates of travel.
    - ii. Exposure to untreated sources of water.
    - iii. Handling of raw seafood.
    - iv. Consumption of seafood, particularly raw or undercooked seafood. If a seafood vehicle is identified, interview others who ate the same item.
    - v. Consumption of food items imported from areas affected by cholera; interview others who ate the same item and provide information on actions to take if they become ill.

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- vi. Restaurant meals: Obtain the name(s) of the restaurant(s), date(s), and location(s) of the meal(s).
- c. Determine if others (e.g., family, friends, travel companions, coworkers, etc.) are known or thought to be ill with similar symptoms. If so, inquire about possible common source exposures. Obtain the name, phone number or address, and clinical information of the ill person and follow up, if appropriate.
- d. Determine if the patient or any of their symptomatic household or other close contacts are associated with sensitive situations (e.g., an attendee or employee of a daycare/childcare setting, a food handler, or an employee in a health care setting with direct patient care). Determine the dates and times he/she worked to assess the risk of transmission to others. See Section 7 for recommended exclusions for symptomatic patients or contacts in sensitive situations.
- e. Provide basic instruction to patients and contacts regarding control measures for preventing further spread of the disease (Section 6).

#### C. Environmental evaluation

An environmental assessment is indicated if a commercial food service facility, child care center, or public drinking water supply is suspected as the source of infection. If illness was associated with shellfish, determine what shellfish was consumed, and information on preparation and handling. Obtain the shellfish labeling tags and collect information on the supplier and harvest site.

# D. Merlin data entry

Create a case in Merlin under disease code Cholera (Vibrio cholerae Type O1) (Merlin disease code=00190) within one business day of notification. Enter the data collected into Merlin, being sure to include all required fields on the Basic Data screen and complete the Case Symptoms screen, Extended Data screen, Travel History screen, and associate all relevant labs. Please associate all labs received via electronic laboratory reporting (ELR) to the case and include serogroup, biotype, and toxigenicity information as available.

#### 6. CONTROLLING FURTHER SPREAD

#### A. Infection Control Recommendations

Standard precautions are recommended for hospitalized patients. Contact precautions should be used for diapered or incontinent persons for the duration of the illness or to control institutional outbreaks.

#### B. Patient/ household education on prevention recommendations

Education for the patient and caregivers should include:

- 1. The importance of effective hand washing, particularly after using the toilet or changing diapers and before preparing or eating food.
- 2. Persons with diarrhea should not be involved in food preparation.
- 3. Importation of food items by travelers from cholera-affected areas is not recommended.
- 4. Avoid consumption of raw oysters or other raw shellfish, particularly if a predisposing factor for infection exists. Persons at increased risk for infection and more severe disease include persons with immunocompromising conditions, persons with chronic liver disease, and persons using antacids, histamine receptor blockers, and proton pump inhibitors.
- 5. Shellfish should be obtained from approved sources and should be adequately boiled or steamed (at least 10 minutes) prior to consumption.

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- 6. Avoid cross-contamination of food with raw seafood or juices from raw seafood.
- 7. Ensure proper temperature control of shellfish prior to preparation and consumption.
- 8. Eat shellfish promptly after cooking and immediately refrigerate leftovers.

#### C. Isolation of patients

Patients in non-sensitive situations should be counseled regarding disease transmission, food preparation, and hand washing practices. Follow-up or release from isolation based on stool culture results is not required. See Section 7 for recommended exclusions for symptomatic patients in sensitive situations (i.e., an attendee or employee of a daycare/childcare setting, a food handler, or an employee in a health care setting with direct patient care).

# D. Management of contacts

Although person-to-person transmission is not likely, CHD epidemiology staff should identify persons at risk of acquiring cholera infection from identified patients and those persons with similar exposures. Household and other close contacts should be educated about the importance of hand washing after caring for ill persons, changing diapers, defecating, and before preparing food. These contacts should be advised to seek immediate medical attention and to alert their health care provider of their exposure to a cholera case if they should become ill. If contacts with acute watery diarrhea are identified, CHD staff should ensure that these persons receive appropriate medical care and diagnostic testing for cholera. These persons should be managed as probable cases until proven otherwise. This is especially important when secondary transmission to persons in sensitive situations is suspected. Stool culture or serology can be used to investigate possible infection among persons exposed to a common source. Hospital infection control of affected facilities should be aware of the case and evaluate risk of transmission within in their facility.

#### E. Food or water is implicated as the source of an outbreak

Contact your Regional Environmental Epidemiologist for investigation guidance FloridaHealth.gov/diseases-and-conditions/disease-reporting-and-management/disease-reporting-and-surveillance/surveillance-and-investigation-guidance/\_documents/environmental-epi-map.pdf

#### F. Immunization recommendations

There is one oral cholera vaccine currently available on the market that is Food and Drug Administration- (FDA) approved for use in the United States <a href="https://www.cdc.gov/cholera/vaccines.html">www.cdc.gov/cholera/vaccines.html</a>. However, CDC does not recommend cholera vaccines for most travelers because protection is incomplete and of short duration (3–6 months).

# 7. MANAGING SENSITIVE SITUATIONS

Guidance for management of cholera cases in sensitive situations is not specifically addressed in Chapter 64D-3 of the Florida Administrative Code. Current DOH recommendations for sensitive situations are addressed below.

#### A. Patient is a food handler

The patient should be instructed not to work as a food handler until symptoms resolve completely. Consult with the Regional Environmental Epidemiologist to determine if further follow-up is warranted.

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#### B. Patient works at a health care or residential care facility

The patient should be instructed not to work until complete resolution of symptoms. Investigate if the patient had contact with any persons with a similar illness and determine if there has been an unusual incidence of diarrheal illness at the facility within the past week. Consult with the case reviewer to determine if further investigation is necessary.

#### C. Patient attends or works at a child care facility

The patient should be instructed not to return to the day care facility until complete resolution of symptoms. Determine if there has been an unusual incidence of diarrheal illnesses at the facility within the past week. Consult with the case reviewer to determine if further investigation is warranted.

# 8. IMPORTANT LINKS

# A. Cholera Case Report Form

FloridaHealth.gov/diseases-and-conditions/disease-reporting-and-management/disease-reporting-and-surveillance/\_documents/crf-cholera.pdf

#### B. Cholera FAQs

www.cdc.gov/cholera/index.html

#### C. Surveillance Case Definitions

FloridaHealth.gov/diseases-and-conditions/disease-reporting-and-management/disease-reporting-and-surveillance/index.html

# D. Food and Waterborne Disease Program - Contact List

FloridaHealth.gov/diseases-and-conditions/disease-reporting-and-management/disease-reporting-and-surveillance/surveillance-and-investigation-guidance/\_documents/environmental-epi-map.pdf

#### **E. County Health Department Contact Information**

FloridaHealth.gov/public-health-in-your-life/county-health-departments/CHDlisting.html

#### F. FDOH Bureau of Laboratory Contact Information.

Jacksonville 904-791-1500 Miami 305-324-2432 Tampa 813-974-8000

#### 9. REFERENCES

- A. David L. Heymann (ed.), *Control of Communicable Diseases Manual*, 20th ed., American Public Health Association Press, Washington, District of Columbia, 2015.
- B. WHO Cholera. Retrieved from www.who.int/topics/cholera/en/.
- C. CDC Cholera. Retrieved from www.cdc.gov/cholera/index.html www.cdc.gov/cholera/treatment/index.html

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www.cdc.gov/cholera/prevention.html.

D. Annual Summaries of *Vibrio* Illnesses. Retrieved from www.cdc.gov/cholera/usa/surveillance.html#covissummaries.