CDC/IDSA Clinician Call

August 28, 2024

Welcome & Introductions



Dana Wollins, DrPH, MGC Senior Vice President, Strategy Infectious Diseases Society of America

- About the Clinician Call: Initiated in 2020 as a forum for information sharing among frontline clinicians caring for patients with COVID-19. Now expanded to address timely topics in infectious diseases—all from a clinical perspective.
- The views and opinions expressed here are those of the presenters and do not necessarily reflect the official policy or position of the CDC or IDSA. Involvement of CDC and IDSA should not be viewed as endorsement of any entity or individual involved.
- This webinar is being recorded and can be found online at <u>www.idsociety.org/cliniciancalls</u>.

CDC/IDSA Clinician Call: Update on Dengue & Other Vector Borne Diseases

1. Dengue Virus Update



Situational Update on Dengue Joshua M. Wong, MD Medical Officer Dengue Branch Division of Vector Borne Diseases U.S. Centers for Disease Control and Prevention



Dengue in Puerto Rico: A Federally Qualified Health Center Experience Hector Villanueva, MD Senior Clinical Advisor HealthproMed San Juan, Puerto Rico

2. West Nile Virus & Oropouche Update



J. Erin Staples, MD, PhD Lead for Surveillance, Epidemiology, and Clinical Care Task Force Oropouche in 2024 Dengue-Oropouche Response U.S. Centers for Disease Control and Prevention

3. Q&A/Discussion – All Presenter Plus



Bonnie Word, MD, FIDSA Member, IDSA Board of Directors Houston Travel Medicine Clinic

Question? Use the "Q&A" Button





Comment? Use the "Chat" Button



Dengue Virus Update

Joshua M. Wong, MD U.S. Centers for Disease Control & Prevention

Hector Villanueva, MD HealthproMed **Centers for Disease Control and Prevention** National Center for Emerging and Zoonotic Infectious Diseases



Situational Update on Dengue

Joshua M Wong, MD

Medical Officer, Dengue Branch, Division of Vector Borne Diseases, NCEZID, CDC

IDSA Conference Call

Aug 28, 2024

Why should we care about dengue?

📭 NPR

Dengue fever is – unfortunately – having a banner year. Can it be quelled?

In the Americas alone, almost 10.4 million suspected cases of dengue, a mosquitoborne viral disease, were reported to the World Health...



1 week ago

on Africa News

Burkina Faso: more than 350 deaths from dengue fever in a month

An epidemic of dengue fever, a mosquito-borne disease, claimed 356 lives in Burkina Faso between mid-October and mid-November, bringing the...

🔘 Reuters

Dengue outbreak in Argentina on track to break records

A major outbreak in Argentina of dengue, a mosquito-borne illness that can be fatal, is on track to smash previous records, reflecting wider...



Mar 21, 2024

Our Discussion Today

- Dengue virus review
- Dengue epidemiology globally and in the U.S.
- Steps to prepare for increased dengue cases:
 - Recognize dengue in your clinical practice
 - Know the **A** warning signs for progression to severe dengue
 - Test appropriately for dengue

Dengue Review

Dengue Viruses (DENVs)

- DENV-1, 2, 3, 4
 - Lifelong DENV type-specific immunity
 - **Short-term** cross-immunity (~1–2 years)
 - Individuals can be infected up to 4 times in their life.
- Clinical Course
 - ~3 in 4 DENV infections are asymptomatic.
 - If symptomatic, onset occurs abruptly after an incubation period of 5–7 days (Range: 3–10).
 - Early clinical findings are nonspecific
 - Can be difficult to distinguish from other pathogens.
 - Can be life-threatening
 - Specific "warning signs" predict progression to severe disease









Dengue 2



DENV Transmission

Vector-borne

• Saliva of infected Aedes spp. mosquito

• Other modes

- Perinatal
- Blood transfusion or organ transplantation
- Needle stick, mucocutaneous, or hospital/laboratory accident
- Breast milk



Aedes aegypti



Aedes albopictus

Dengue Globally

2024 Global Dengue by the Numbers





- 103 countries/territories reporting cases
 - All 6 WHO regions



• >20 countries reporting outbreaks

https://www.who.int/emergencies/disease-outbreak-news/item/2024-DON518

Geographical distribution of dengue cases as reported to WHO from January to April 2024



https://www.who.int/emergencies/disease-outbreak-news/item/2024-DON518

Dengue cases in the Americas, 1980–2023*

More than 4.6 million cases reported in 2023



*Data from PAHO PLISA Health Information Platform for the Americas

Dengue cases in the Americas, 1980–2024*

More than 11 million cases reported as of Aug 13, 2024



*Data from PAHO PLISA Health Information Platform for the Americas

Dengue in the US

In the United States, dengue is endemic in 6 U.S. territories and freely associated states.



Puerto Rico locally acquired dengue cases reported by week, 2024 compared to the historical median (1986-202?)



*Data from: Data and Statistics on Dengue in the United States | Dengue | CDC. Accessed 8/4/2024. Cases for 2023 and 2024 are preliminary.

In 2023, **DENV-1** was the predominant serotype in PR.



Vigilancia Enfermedades Arbovirales. Semana Epidemiológica Número 18, 2024. https://www.salud.pr.gov/CMS/DOWNLOAD/8937 accessed 5/28/2024.

As of August 2024, DENV-1 and DENV-3 now account for

most new infections in PR.



Vigilancia Enfermedades Arbovirales. Semana Epidemiológica Número 31, 2024. https://www.salud.pr.gov/CMS/DOWNLOAD/9209 accessed 8/26/2024.

Puerto Rico declared a public health emergency due to dengue on March 25, 2024.

Salud declara emergencia de salud pública por dengue en Puerto Rico

La última epidemia de dengue fue en 2012

Wilmarielys Agosto, El Vocero 25/03/2024



Agosto, W. Salud Declara emergencia de salud publica por dengue en Puerto Rico. El Vocero. 3/25/2024. Available form: https://www.elvocero.com/actualidad/salud/salud-declara-emergencia-de-salud-p-blica-por-dengue-en-puerto-rico/article_ca9be43c-eade-11ee-b034-bb87cd739f53.html

CDC activated a program level dengue emergency response in April and escalated to center level in July 2024.



A dengue health advisory was released in the health alert network on June 25.



HAN available at: <u>https://emergency.cdc.gov/han/2024/han00511.asp</u>

Among dengue cases reported to ArboNET from 2010–2022,

annun .

most dengue cases in U.S. states (>94%) were associated with travel to endemic areas.

Travel associated dengue cases (N = 12,057) reported in the US by year, 2010–2024*



*Data from: https://www.cdc.gov/dengue/statistics-maps/data-and-maps.html. Accessed 8/19/2024. Cases for 2023 and 2024 are preliminary.

Travel associated dengue cases, Jan 1–Aug 19, 2024 (n=1,145)

- Travel associated cases reported from 50 jurisdictions
- Highest case numbers:
 - Florida (n=261)
 - New York (n=169)
 - California (n=88)



Map from: Current Year Data (2024) | Dengue | CDC

New Global Dengue Webpage **Travel Health** Notices

https://wwwnc.cdc.gov/tra vel/notices/level1/dengueglobal



Dengue THN by WHO Region

AMERICAS	
1. Argentina	11. Honduras
2. Brazil	12. Martinique
3. Colombia	13. Mexico
4. Costa Rica	14. Nicaragua
5. Curaçao	15. Panama
6. Ecuador	16. Paraguay
7. French Guiana	17. Peru
8. Guadeloupe	18. Uruguay
9. Guatemala	
10. Guyana	

AFRICA
19. Burkina Faso
20. Ethiopia
21. Mali
22. Mauritius
23. Sudan

SOUTH-EAST	WESTERN PACIFIC
ASIA	26. Cambodia
24. Indonesia	27. Fiji
25. Sri Lanka	28. Laos
	29 Samoa

30. Singapore



Names and boundary representation are not necessarily authoritative

Locally Acquired Dengue in U.S. States

Dengue vectors are present across much of the US.



Locally Acquired Dengue in US States, 2010– 2023

- Sporadic cases historically limited to outbreaks in:
 - Florida, Hawaii, Texas
- Recently, more states are reporting local DENV transmission.
 - Arizona, n=2 (2022)
 - California, n=2 (2023)



Map from: <u>https://www.cdc.gov/dengue/statistics-maps/current-data.html</u>

Multiple DENV introductions in Florida from returning travelers resulted in increased local transmission.

Reported dengue cases in Florida, by month — Florida Department of Health, 2022–2023



How can we prepare for increasing dengue cases?

Emerging Tools to Prevent Dengue

- Novel vector control strategies under evaluation in multiple countries
- Dengue vaccines
 - **Dengvaxia**: recommended in US (children 9-16 years old with previous DENV infection and living in endemic areas)
 - TAK-003: licensed in several countries
 - Butantan-DV: high efficacy in early phase 3 results
- No dengue vaccines are currently recommended for use in US travelers.



How can we prepare for increasing dengue cases?

- **Recognize** dengue in your clinical practice
- Know the warning signs for progression to severe dengue
- Test appropriately for dengue

How can we prepare for increasing dengue cases?

• **Recognize** dengue in your clinical practice


Early recognition and appropriate treatment of dengue saves lives.

- No specific antiviral treatment available.
- Standard of care is protocolized IV fluid management, according to WHO guidelines.
- Up to 13% mortality if severe disease is untreated but can be reduced to <0.05% mortality with appropriate management.



For further dengue training resources, visit: <u>https://www.cdc.gov/dengue/hcp/clinical-care/index.html</u>

All clinicians should be asking:

Could this patient have dengue?

How can we prepare for increasing dengue cases?

• **Recognize** dengue in your emergency rooms

• Know the warning signs for progression to severe dengue



Dengue Clinical Course



Febrile Phase



Critical Phase



Convalescent Phase



How do I know if my patient will progress to severe dengue?

Mosquito

bite





- Intense continuous abdominal pain or tenderness
- Persistent vomiting
 - \geq 3 episodes in 1 hr or \geq 4 in 6 hrs
- Fluid accumulation
 - pleural effusion, ascites, or pericardial effusion
- Mucosal bleeding
 - gums, nose, vagina (metrorrhagia or hypermenorrhea), kidney (macroscopic hematuria)
- Altered mental status
 - irritability, drowsiness, Glasgow Coma Scale score <15
- Hepatomegaly
 - ≥2cm below costal margin
- Progressive increase of hematocrit
 - in at least 2 consecutive measurements taken 6 hours apart



For further dengue training resources, visit: https://www.cdc.gov/dengue/hcp/clinical-care/index.html

How can we prepare for increasing dengue cases?

- **Recognize** dengue in your emergency rooms
- Know the warning signs for progression to severe dengue
- Test appropriately for dengue



Laboratory testing is most sensitive when performed within the first 7 days of illness.

• Within 7 days of symptom onset, test with:

RT-PCR + IgM ELISA

or

NS1 antigen ELISA + IgM ELISA

Using this test combination provides a laboratory diagnosis in >90% of dengue cases.*



For more information on testing, visit: <u>www.cdc.gov/dengue/healthcare-providers/testing/</u>

*Hunsperger, E. A., et al. (2016). "Performance of Dengue Diagnostic Tests in a Single-Specimen Diagnostic Algorithm." J Infect Dis 214(6): 836-844.

Laboratory testing is most sensitive when performed within the first 7 days of illness.

• Within 7 days of symptom onset, test with:

Most patients present for clinical care and testing during the first 7 days of illness.

NS1 antigen ELISA + IgM ELISA

Using these test combinations provides a laboratory diagnosis in >90% of dengue cases.*
Image: Second se

For more information on testing, visit: www.cdc.gov/dengue/healthcare-providers/testing/

*Hunsperger, E. A., et al. (2016). "Performance of Dengue Diagnostic Tests in a Single-Specimen Diagnostic Algorithm." J Infect Dis 214(6): 836-844.

Testing >7 days After Illness Onset

• Test with:

IgM serology

and consider*

NS1 antigen ELISA or RT-PCR

(lower sensitivity compared to days 0–7)



*Testing guidance may vary by jurisdiction, especially in endemic areas.

For more information on testing, visit: <u>www.cdc.gov/dengue/healthcare-providers/testing/</u>

In Summary

- Dengue cases globally are increasing.
- Dengue cases in the US are increasing.
 - Outbreak declared in PR
 - CDC has stood up an emergency response.
- Steps to prepare for increased dengue cases:
 - Recognize dengue in your clinical practice
 - Know the **A**warning signs for progression to severe dengue
 - Test appropriately for dengue

The new dengue clinical management pocket guide available.

- Early recognition and appropriate treatment of dengue saves lives.
- The pocket guide describes dengue case management, including recommendations by patient group (outpatient, inpatient, and inpatient with compensated or hypotensive shock).
- To download the pocket guide or request printed copies, visit:
 - <u>https://www.cdc.gov/dengue/hcp/pocketguide/index.html</u>



• Don't forget:

- **Recognize** dengue in your clinical practice

- Know the warning signs for progression to severe dengue

Test appropriately for dengue

For more information, contact CDC 1-800-CDC-INFO (232-4636) TTY: 1-888-232-6348 www.cdc.gov

Joshua M Wong, MD nof9@cdc.gov

The findings and conclusions in this report are those of the authors and do not necessarily represent the official position of the Centers for Disease Control and Prevention.





DENGUE IN PUERTO RICO: A FEDERALLY QUALIFIED HEALTH CENTER EXPERIENCE

HEALTHPROMED, SAN JUAN, P.R.

BACKGROUND

Dengue is endemic in PR – clinicians are used to managing Dengue cases throughout the year.

PR declared a public health emergency due to dengue on March 25/2024.

On June 21, it was extended up to December 31/2024.

Evidence based written protocols where reviewed and discussed with clinicians during faculty meetings.

BACKGROUND

Education was provided to patients and the community regarding dengue prevention and warning signs.

According to a recent administrative order, all clinicians are required to take a Dengue Management webinar, created by the Panamerican Health Organization (PAHO).

HealthproMed has an in-house laboratory to process diagnostic tests rapidly and efficiently.





FACTS ABOUT DENGUE

CLINICAL PRESENTATION 75% ASYMPTOMATIC 25% SYMPTOMATIC AMONG SYMPTOMATIC PERSONS

- ~5% SEVERE DENGUE
- ~95% MILD TO MODERATE



Four Serotypes – second infection is associated to highest probability of severe dengue.

2009 WHO Case definition

-Hemorrhagic dengue - discontinued.

-Severe dengue - accepted.

Shock, not hemorrhage, is the main cause of death in Dengue.

Instead, increased capillary permeability \rightarrow plasma leakage \rightarrow dehydration \rightarrow hemoconcentration \rightarrow hypovolemic shock \rightarrow metabolic acidosis \rightarrow multiple organ failure \rightarrow death.











Fever Headache Joint pain Myalgia Retroorbital pain



FREQUENT SYMPTOMS: NOT ALL OF THEM NEED TO BE PRESENT

- Fever is of acute and sudden onset.
- Retroorbital pain is a red flag for diagnosis of dengue.
- Lack of URI Symptoms helps to differentiate from influenza.

LABORATORY WORK UP

- Diagnostic tests for dengue according to onset of symptoms.
- CBC, CMP, Serum Protein and Albumin, Liver Enzymes, Coagulation Panel.
- Dengue should be considered in any patient who presents a high fever of acute onset and has a history of recent travel (< 2 weeks) to a country where dengue is endemic.
- If dengue is suspected, there is no need to wait for diagnostic test results to treat the patient accordingly.



LABORATORY WORK UP CONT.

• **CBC** – should be repeated daily during the febrile and critical phases.

Patients usually present with leukopenia and thrombocytopenia, but its absence does not rule out dengue.

- WBC count decreases gradually during the febrile phase and then increases after defervescence.



LABORATORY WORK UP CONT.

- Platelet count decreases gradually and usually increases after the WBC increase.
- Hemorrhage, if it occurs, is not related to a determined platelet count level.
- Decreasing platelet count, which is secondary to transient platelet destruction is indicative of active viral disease.

AMBULATORY MANAGEMENT OF PATIENTS

Once diagnosed or suspected, patients should be reevaluated daily in clinic or ER.

Vital signs, hydration state and CBC results should be monitored daily.

Watch closely for warning signs and educate patients regarding the importance of early identification.

Referral to ER for further evaluation and management if warning signs are present or for any concerning symptom.

AMBULATORY MANAGEMENT OF PATIENTS

Watch for resolution of fever and provide close follow-up during next 48 hours.

Instructions to patients:

- -Acetaminophen PRN for fever or pain.
- -Aspirin, NSAID's and steroids are contraindicated.
- -Bed rest
- -Take plenty of fluids.
- -Use insect repellent and mosquito nets at home.
- -Watch closely for warning signs.



PATIENT OUTCOMES

- Most of our patients with dengue received ambulatory treatment in our clinic, showing gradual complete recovery and their lab tests returned to normal values.
- Most common warning signs of those who required ER evaluation and hospitalization:
- Abdominal pain
- Persistent vomiting
- Hemoconcentration



THANK YOU

Héctor Villanueva Rodríguez

Clinical Advisor HealthproMed



West Nile Virus and Oropouche Update

Erin Staples, MD. PhD U.S. Centers for Disease Control & Prevention









J. Erin Staples, MD, PhD Surveillance, Epidemiology, and Clinical Care Task Force 2024 Dengue Oropouche Response CDC/IDSA Clinician Call August 26, 2024

> Slides developed by Sarah Guagliardo, PhD, CDC https://www.freepik.com/premium-ai-image/river-jungle-seen-from-nearby-manaus-amazon-jungle_40943049.htm

Oropouche virus

Oropouche virus (oro-POU-che, oro-POOCH, oro-POOSH)



Elliott RM. Orthobunyaviruses: recent genetic and structural insights. Nat Rev Microbiol. 2014 Oct;12(10):673-85. doi: 10.1038/nrmicro3332. Epub 2014 Sep 8. PMID: 25198140.

- Simbu serogroup, Orthobunyavirus genus, Peribunyaviridae family
- Tripartite genome susceptible to reassortment
- Current strain some evidence of successive reassortment and mutations
- Influence on vector competence, disease severity, immune protective status, and virus transmissibility is unknown



Current Oropouche Outbreak



Clinical Overview

- Initial presentation similar to infections caused by dengue, Zika, and chikungunya viruses
- Incubation period: 3-10 days
- Symptomatic attack rate: ~60%



Initial Clinical Illness

- Acute onset of fever
- Severe headache
- Chills
- Myalgia
- Arthralgia
- Fatigue
- Photophobia
- Dizziness
- Conjunctival injection
- Maculopapular rash
Clinical Laboratory Findings

- Initial leukopenia, lymphopenia, and slightly elevated liver enzymes
- Can have leukocytosis later in course





Clinical Course

- Relapse of symptoms in up to 70% of cases
 - Days to weeks after first occurrence
- Disease is typically mild, deaths are rare



Clinical Presentation – Severe Disease

Neuroinvasive disease



Vernal et al, 2019

- Occipital pain
- Dizziness
- Confusion
- Lethargy
- Photophobia
- Nausea
- Vomiting
- Nuchal rigidity
- Nystagmus

Hemorrhagic manifestations

- Epistaxis
- Gingival bleeding
- Melena
- Menorrhagia
- Petechiae



Clinical Management

- No specific antiviral treatment
- Treat symptoms: rest, fluids, and use of analgesics and antipyretics
 - Acetaminophen preferred treatment for fever and pain
 - Aspirin and other non-steroidal antiinflammatory drugs (NSAIDS) should not be used to reduce risk of hemorrhage
- Patients with more severe symptoms might need hospitalization for observation and supportive treatment



Deaths Associated with Oropouche



- Two recent deaths in otherwise healthy young women
 - 21 and 24 years of age
 - Non-pregnant
- Similar presentation to severe dengue
- Clinical features
 - Signs of hemorrhage, rapid evolution
 - Died within ~4 days of symptom onset
 - High level viremia (based on low RT-PCR CT values)

Bandeira et al 2024, Clinical profile of Oropouche Fever in Bahia, Brazil: unexpected fatal cases (Preprint) https://preprints.scielo.org/index.php/scielo/preprint/view/9342

Oropouche and Pregnancy

- Based on limited data from Brazil, vertical transmission of Oropouche virus is possible
 - Several pregnant people with evidence of vertical transmission to their fetus associated with fetal death or congenital abnormalities, including microcephaly
 - Pregnant persons had symptoms during pregnancy; most had positive test results
 - Tissues from still births and one infant born with microcephaly tested positive by RT-PCR
- Frequency of vertical transmission is unknown



https://www.news-medical.net/health/What-is-Vertical-Transmission.aspx

Effect of timing of disease during pregnancy on risk of adverse outcome is unknown

What is the risk to the US?

60.000 routes

Possible Vectors in the United States

Culicoides paraensis

Competent vector in lab studies Implicated in field studies

Culex quinquefasciatus

Less competent vector in lab studies Implicated in field studies

"NACER355-12 Lateral" - BOLD:ABX5601 (cf. Culicoides paraensis) (licensed under http://creativecommons.org/licenses/by/4.0/)

CDC Public Health Image Library



Dominic

Known Distribution of Vectors in the United States



Guagliardo et al, 2024 (Under review)

Risk of Local Transmission in the US

 Risk for sustained local transmission in continental U.S. is likely low...

...but non-zero

- Risk is not uniformly distributed over space and time
 - Travel patterns
 - Vector distribution

CDC Response Activities

Updated Travel Health Notices

Pregnant people should reconsider non-essential travel to Cuba and take precautions to prevent bites in other areas where OROV may be circulating

Oropouche in Cuba

Level 4 - Avoid All Travel	L
Level 3 - Reconsider Nonessential Travel	L
Level 2 - Practice Enhanced Precautions	L
Level 1 - Practice Usual Precautions	1

Key points

- There is an outbreak of Oropouche in Cuba (see map).
 - A Level 1 Travel Health Notice has been issued for <u>Oropouche</u> in <u>South America</u>.
- Multiple cases of Oropouche have recently been reported in <u>U.S.</u> and <u>European</u> travelers returning from travel to Cuba indicating there is ongoing risk.
- Oropouche is spread by the bite of infected midges (small flies) and mosquitoes.
- All travelers to Cuba should take steps to prevent bug bites during travel to protect themselves from infection.
- Pregnant people should reconsider non-essential travel to Cuba. If travel is unavoidable, these travelers should strictly follow Oropouche <u>prevention</u> recommendations.
- Illness can occur in people of any age and is often mistaken for dengue.

Current Situation

 On August 1, 2024, the Pan American Health Organization (PAHO) issued an <u>epidemiological alert</u> to about Oropouche cases, including deaths, in the Americas. There were also concerns about an



Map: Areas in Cuba with reported cases of Oropouche (see larger map)

What is Oropouche?

<u>Oropouche</u> is a disease caused by Oropouche virus. It is spread through the bites of infected midges (small flies) and mosquitoes.

<u>Symptoms</u> of Oropouche include headache, fever, muscle aches, stiff joints, nausea, vomiting, chills, or sensitivity to light. Severe

Oropouche in South America

evel 4 - Avoid All Travel	
evel 3 - Reconsider Nonessential Travel	

evel 2 - Practice Enhanced Precautions

Level 1 - Practice Usual Precautions

Key points

- Parts of Bolivia, Brazil, Colombia, and Peru (see map) are reporting cases of Oropouche, although the case numbers are declining.
 A Level 2 Travel Health Notice has been issued for Oropouche
 - in Cuba.
- Oropouche is spread by the bite of infected midges (small flies) and mosquitoes.
- Travelers to affected areas should take steps to prevent bug bites.
- Illness can occur in people of any age and is often mistaken for dengue.

Current Situation

What Travelers Can Do to Protect Themselves and Others

- Travelers to affected areas should take steps to prevent bug bites during travel to protect themselves from infection. They should also prevent bug bites for 3 weeks after travel to avoid possibly spreading the virus to others in the U.S.
 - If travelers are pregnant, they should discuss travel plans, reasons for travel, steps to prevent bug bites, and potential risks with their healthcare provider.



Map: Areas in South America with reported cases of Oropouche (see larger map)

Health Alert Network (HAN) Health Advisory

Notifying clinicians and public health authorities of increase in Oropouche virus disease in the Americas

Increased Oropouche Virus Activity and Associated Risk to Travelers

Print



Distributed via the CDC Health Alert Network August 16, 2024, 4:00 PM ET CDCHAN-00515

- Advises on evaluating and testing travelers returning from impacted areas with signs and symptoms consistent with Oropouche virus infection
- Raises awareness of possible risk of vertical transmission and associated adverse effects on pregnancy
- Highlights prevention measures to mitigate additional spread of virus and potential importation into unaffected areas

CDC's Oropouche pages related to pregnant persons and their infants *Considerations for Clinicians*



Oropouche EXPLORE TOPICS

Oropouche EXPLORE TOPICS

Q SEARCH

AUGUST 27, 2024

Interim Clinical Considerations for Pregnant People with Confirmed or Probable Oropouche Virus Disease

AT A GLANCE

CDC is working to learn more about the potential risks of Oropouche virus disease during pregnancy, in close collaboration with the American College of Obstetricians and Gynecologists, Society for Maternal-Fetal Medicine, and the American Academy of Pediatrics. Interim clinical considerations are based on recent reports from Brazil describing Oropouche virus vertical transmission associated with fetal deaths or congenital abnormalities and based on other congenital viral infections with similar clinical manifestations. This page describes considerations for clinical management of confirmed or probable Oropouche virus disease in pregnancy.



Manifestations and clinical management of pregnant people

Manifestations and clinical manage.

ON THIS PAGE

https://www.cdc.gov/oropouche/hcp/clinical-care-pregnancy https://www.cdc.gov/oropouche/hcp/clinical-care/infants Interim Guidance for Evaluating and Managing Infants Born to Pregnant People with Confirmed or Probable Oropouche Virus Disease

AT A GLANCE

- Data on congenital Oropouche virus disease (Oropouche) is currently limited.
- Findings among people in Brazil with Oropouche during pregnancy have included stillbirth and severe microcephaly.
- Infants born to people with confirmed or probable Oropouche during pregnancy should receive a comprehensive evaluation by a healthcare provider at birth and at each well-child visit.



 Infants and children with congenital anomalies might benefit from multidisciplinary care.



Oropouche Virus Disease Among U.S. Travelers – United States, 2024

Early Release / August 27, 2024 / 73

- As of August 16, 2024, 21 Oropouche virus disease cases were identified among U.S. travelers from Cuba.
- Most patients presented with fever, myalgia, and headache, often with other symptoms such as arthralgia, diarrhea, retroorbital pain, abdominal pain, and nausea/vomiting.
- At least three patients had recurrent symptoms after the initial illness

Q



For more information, contact CDC 1-800-CDC-INFO (232-4636) TTY: 1-888-232-6348 www.cdc.gov

Acknowledgements

Surveillance, Epidemiology, and Clinical Care Task Force

Carolyn Gould Sarah Guagliardo Shelby Lyons Stacey Martin Hannah Padda Laura Platt Kathryne Walker Jessica Winberg Van Tong Romeo Galang Rachel Alade Haben Debessai Jennita Reefhuis Nicki Roth Christina Sancken

The findings and conclusions in this report are those of the authors and do not necessarily represent the official position of the Centers for Disease Control and Prevention.





What's new for West Nile virus and eastern equine encephalitis virus

J. Erin Staples, MD, PhD Surveillance and Epidemiology Team Arboviral Diseases Branch

CDC/IDSA Clinician Call August 26, 2024



Overview

- West Nile virus (WNV) epidemiology
- Risk factor for severe disease
- Knowledge of diagnostic testing
- Eastern equine encephalitis (EEEV) epidemiology









WNV leading cause of mosquito-borne disease in continental U.S. – 1999-2023



Human Disease Cases	Hospitalizations	Deaths
59,141 Cases from year(s) and type of case selected above	27,617 Hospitalizations from year(s) and type of case selected above	2,958 Deaths from year(s) and type of case selected above

West Nile virus human disease cases reported by month of illness onset, 1999-2023, All disease cases



Current case counts for 2024* similar to prior years with several focal areas of concern

- Case counts tracking with 10-year median
 - Deaths increased
 - Human infections (blood donors screening positive) increased
- Increased activity
 - Texas (Houston and Dallas-Fort Worth areas),
 - Nevada (Las Vegas),
 - NYC area
- Remind patients to use insect repellent, alert high-risk patients, and order correct testing to diagnose disease

West Nile virus human disease cases reported by state of residence, 2024



*Preliminary data shared by state health department

Risk factor for severe disease: Rituximab and other Bcell depleting therapies



- Most clinical experience for Rituximab
 - First approved for treating non-Hodgkin's lymphoma in 1997
 - Approvals expanded to other conditions
 - Widely used off-label for autoimmune and inflammatory conditions
- 90% circulating B cells killed within 3 days of first infusion
- Near complete B cell depletion persists for 6–12 months
 - Prolonged depletion for >3 years reported
 - Quantification of CD19+ cells using flow cytometry preferred method of monitoring depletion



Fatal Cache Valley virus meningoencephalitis associated with rituximab maintenance therapy

Yuanguan Yang¹ | Jingxin Qiu² | Abigail Snyder-Keller³ | Yongping Wu³ Shufeng Sun³ | Haixin Sui³ | Amy B. Dean³ | Laura Kramer³ Francisco Hernandez-Ilizaliturri¹

JAMA Neurology | Brief Report

Fatal Powassan Encephalitis (Deer Tick Virus, Lineage II) in a Patient With Fever and Orchitis Receiving Rituximab

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EMERGING INFECTIOUS DISEASES®

DISPATCHES

Fatal Case of Chronic Jamestown Canyon Virus Encephalitis Diagnosed by Metagenomic Sequencing in Patient Receiving Rituximab

Isaac H. Solomon,¹ Vijay S. Ganesh,¹ Guixia Yu, Xian Ding Deng, Michael R. Wilson, Steve Miller, Tracey A, Milligan, Shibani S, Mukerii, Abigail Mathewson, Justin Linxweiler, Darlene Morse, Jana M, Ritter, J. Erin Staples, Holly Hughes, Carolyn V. Gould, Pardis C. Sabeti,² Charles Y. Chiu,² Anne Piantadosi^{2,3}

Open Forum Infectious Diseases

ID CASE



Fatal Eastern Equine Encephalitis in a Patient on Maintenance Rituximab: A Case Report

Isaac H. Solomon,¹ Pedro D. S. C. Ciarlini,¹ Sandro Santagata,¹ Asim A. Ahmed,³ Umberto De Girolami,¹ Sashank Prasad,⁴ and Shibani S. Mukerji^{2,4}

Two Cases of Severe Tick-Borne **Encephalitis in Rituximab-Treated** Patients in Germany: Implications for **Diagnosis and Prevention**

Philipp A. Steininger,¹ Tobias Bobinger,² Wenke Dietrich,³ De-Hyung Lee,² Michael Knott,⁴ Christian Bogdan,⁵ Klaus Korn,¹ and Roland Lang⁵

THE AMERICAN JOURNAL of **MEDICINE**®

> Seronegative West Nile Virus **Infection in a Patient Treated** with Rituximab for Rheumatoid Arthritis



Clinical Infectious Diseases







Severe Arboviral Neuroinvasive Disease in Patients on Rituximab Therapy: A Review

Ronak K. Kapadia,^{1,2} J. Erin Staples,³ Christine M. Gill,⁴ Marc Fischer,³ Ezza Khan,⁵ Janeen J. Laven,³ Amanda Panella,³ Jason O. Velez,³ Holly R. Hughes,³ Aaron Brault,³ Daniel M. Pastula,^{13,6} and Carolyn V. Gould³

- Identified 21 cases of arboviral disease in patients taking rituximab
 - WNV, tick-borne encephalitis virus, eastern equine encephalitis virus, Powassan virus, Cache Valley virus, and Jamestown Canyon virus
 - Often indolent clinical course; presenting off-season
- 18/19 (95%) patients had negative serologic results
 - Collected 3 days–10 months after illness onset
 - 2 patients seroconverted at 6-month follow-up
- 15 (71%) died and at least 4 (19%) with long-term sequelae

Arboviral disease in patients on B cell-depleting monoclonal antibodies can be severe → counsel on mosquito and tick prevention

- Severe clinical presentations with high mortality
- Seasonal onset may not apply as disease can be protracted
- Lack of serologic response
- Patients who are being prescribed rituximab should be counseled about mosquito and tick prevention!
- Molecular testing is preferred
- CDC has resources on website for patients and providers









Assessment of Healthcare Provider (HCP) Knowledge of WNV Diagnostic Testing*

- Nation web-based survey distribute August-September 2022 to 2,167 HCP been in practice ≥3 years selected by specialty
- Two multi-select questions on diagnostic testing for patients with suspected WNV neuroinvasive disease:
 - Patient A: 70 yo male immunocompetent patient
 - Patient B: 49 yo female immunocompromised patient on anti-B-cell monoclonal antibody therapy

• Answer selections included:

RT-PCR on serum or CSF
IgM antibody test on CSF
IgG antibody test on CSF

 $\hfill\square$ IgM antibody test on serum

□ IgG antibody test on serum

□ Not sure



*Information presented by Shelby Lyons, CDC, at Council of State and Territorial Epidemiologist Meeting, June 2024

Fewer than half of surveyed HCPs were able to correctly diagnosis WNV disease (N=1,502)

Respondents exclusively selecting most appropriate test

Respondents able to diagnose WNV



Ensure correct testing ordered for WNV to make diagnosis

- Response influenced by patient characteristics, specialty type, location of practice, and work setting
 - More challenges with ordering correct tests for immunocompromised patient
 - Internists often able to make diagnosis compared to FPs, NPs, PAs, and Peds
 - HCPs practicing in inpatients settings able to make diagnosis compared to outpatient
 - HCPs in areas with more WNV able to make diagnosis compared to lower incidence areas
- 1,025 excess (incorrect) tests ordered by HCPs



Access to provider education on appropriate WNV diagnostic testing and clinical care

- Decrease delays in diagnosis and unnecessary testing or treatment
- Improve reporting and target prevention measures

Medscape training: "Diagnosis and Management of West Nile Virus Infection: A Case-Based Approach"

CDC's WNV diagnostic testing algorithm webpage



CDC's WNV Disease Therapeutics: Review of the Literature



EEE most severe arbovirus with ~30% case fatality rate and 50% of survivors with long-term disabilities

• To date, four cases reported to CDC in 2024 but in peak transmission season



Clinical and testing inquiries

State and local health department vector-borne disease coordinator

CDC

Oropouche virus clinical inquires: <u>eocevent495@cdc.gov</u> Other arboviral disease clinical inquires: <u>ADBClinicalTeam@cdc.gov</u>

For more information, contact CDC 1-800-CDC-INFO (232-4636) TTY: 1-888-232-6348 www.cdc.gov

The findings and conclusions in this report are those of the authors and do not necessarily represent the official position of the Centers for Disease Control and Prevention.



Risk Associated with Possible Vectors

Culicoides paraensis

Low abundance in the US Different ecology in North America Not detected in Puerto Rico

Culex quinquefasciatus

Dominic

Unknown contribution to transmission Target of vector control programs Insecticide resistance is an issue

Culicoides sonorensis

Low abundance in the US Found in rural areas Not detected in Puerto Rico Unknown contribution to transmission

"NACER355-12 Lateral" - BOLD:ABX5601 (cf. Culicoides paraensis) (licensed under http://creativecommons.org/licenses/by/4.0/,

CDC Public Health Image Library

Q&A/ Discussion

Selected Resources

Program Links:

- This webinar is being recorded and can be found with the slides online at https://www.idsociety.org/cliniciancalls
- COVID-19 Real-Time Learning Network: <u>https://www.idsociety.org/covid-19-real-time-learning-network/</u>

Joshua Wong, MD

- <u>https://www.who.int/emergencies/disease-outbreak-news/item/2024-DON518</u>
- <u>https://www.who.int/emergencies/disease-outbreak-news/item/2023-DON498</u>
- <u>https://www.cdc.gov/dengue/data-research/facts-stats/?CDC_AAref_Val=https://www.cdc.gov/dengue/statistics-maps/data-and-maps.html</u>
- <u>https://www.salud.pr.gov/CMS/DOWNLOAD/8937</u>
- <u>https://www.salud.pr.gov/CMS/DOWNLOAD/9209</u>
- <u>https://wwwnc.cdc.gov/travel/notices/level1/dengue-global</u>
- <u>https://www.cdc.gov/dengue/statistics-maps/data-and-maps.html</u>
- <u>https://www.cdc.gov/dengue/data-research/facts-stats/current-</u>

data.html?CDC_AAref_Val=https://www.cdc.gov/dengue/statistics-maps/current-data.html

- <u>https://www.cdc.gov/mosquitoes/pdfs/Aedes-mosquito-maps.pdf</u>
- <u>https://www.cdc.gov/dengue/statistics-maps/current-data.html</u>
- <u>https://www.cdc.gov/dengue/hcp/clinical-care/index.html</u>
- <u>https://www.cdc.gov/dengue/healthcare-providers/testing/</u>
- <u>https://www.cdc.gov/dengue/healthcare-providers/testing/testing-guidance.html</u>
- <u>https://www.cdc.gov/dengue/hcp/pocketguide/index.html</u>

Selected Resources

Erin Staples, MD, PhD

- <u>https://preprints.scielo.org/index.php/scielo/preprint/view/9342</u>
- <u>https://www.cdc.gov/oropouche/hcp/clinical-</u> <u>care/pregnancy.html?CDC_AA_refVal=https%3A%2F%2Fwww.cdc.gov%2Foropouche%2Fhcp%2Fclinical-care-</u> pregnancy%2Findex.html
- <u>https://www.cdc.gov/mmwr/volumes/73/wr/mm7335e1.htm?s_cid=mm7335e1_e&ACSTrackingID=USCDC_921-</u> DM135007&ACSTrackingLabel=Early%20Release%20%E2%80%93%20Vol.%2073%2C%20August%2027%2C%202024&deliveryNam e=USCDC_921-DM135007
- Oropouche virus clinical inquires: <u>eocevent495@cdc.gov</u> Other arboviral disease clinical inquires: <u>ADBClinicalTeam@cdc.gov</u>

COVID-19 Real-Time Learning Network

Brought to you by CDC and \bigcirc

An online community bringing together information and opportunities for discussion on latest research, guidelines, tools and resources from a variety of medical subspecialties around the world.



Specialty Society Collaborators

American Academy of Family Physicians American Academy of Pediatrics American College of Emergency Physicians American College of Obstetricians and Gynecologists American College of Physicians American Geriatrics Society American Thoracic Society Pediatric Infectious Diseases Society Society for Critical Care Medicine Society for Healthcare Epidemiology of America Society of Hospital Medicine Society of Infectious Diseases Pharmacists

www.COVID19LearningNetwork.org @RealTimeCOVID19 #RealTimeCOVID19

THANK YOU

We want to hear from you! Please complete the post-call survey.

A recording of this call, slides and the answered Q&A will be posted at <u>www.idsociety.org/cliniciancalls</u>

-- library of all past calls available --

Contact Us:

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