

# The Arboviruses Next Door: Orphaned and Emerging Arboviruses of the US

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APHL annual meeting

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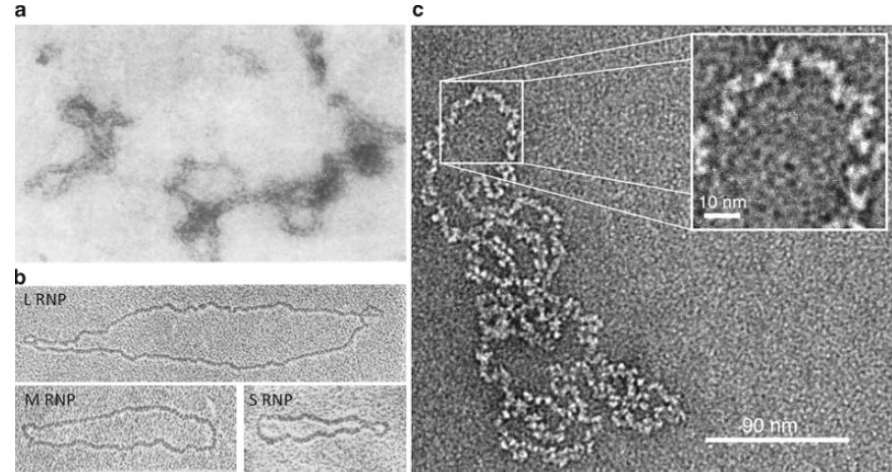


# Orphaned arboviruses

Orphan  $\neq$  neglected

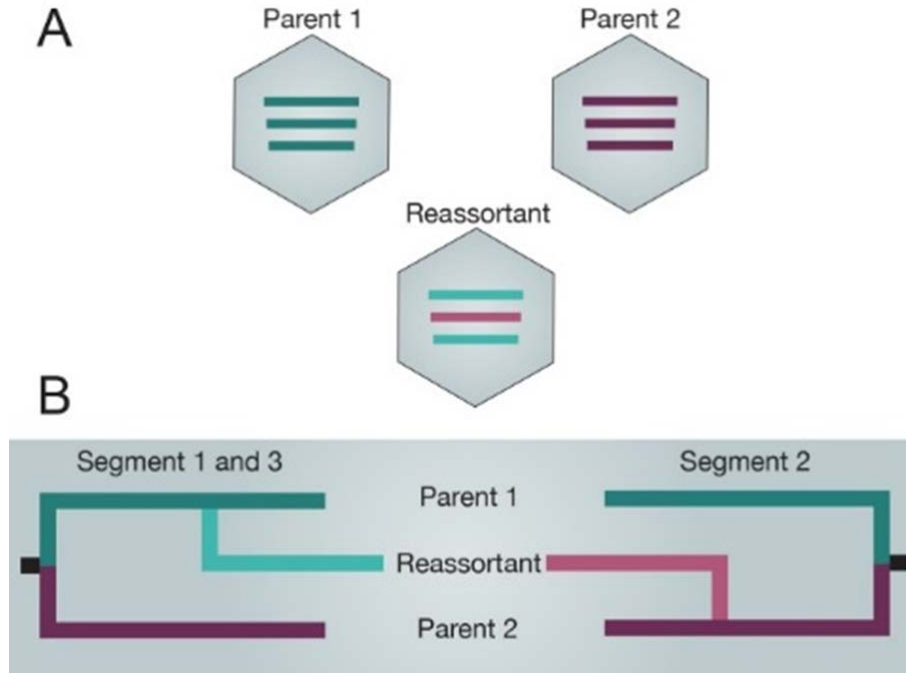
# *Orthobunyavirus* (Order *Bunyavirales*, Family *Peribunyaviridae*)

- Negative sense, single-stranded, segmented, RNA genome
  - Large (L) : RNA-dependent RNA polymerase
  - Medium (M) : structural polyprotein Gn-NSm-Gc
  - Small (S): nucleocapsid and NSs
- 20 serogroups
  - California (US/North America)
  - Bunyamwera (US/North America, South America, Africa)



Guu et al. Bunyavirus: structure and replication. In *Advances in Experimental Medicine and Biology*. 2012

# Segment reassortment (rapid evolution)



Vijaykrishna, D et al. [PLoSPathog.](https://doi.org/10.1371/journal.ppat.1004902) 2015 Jul; 11(7): e1004902

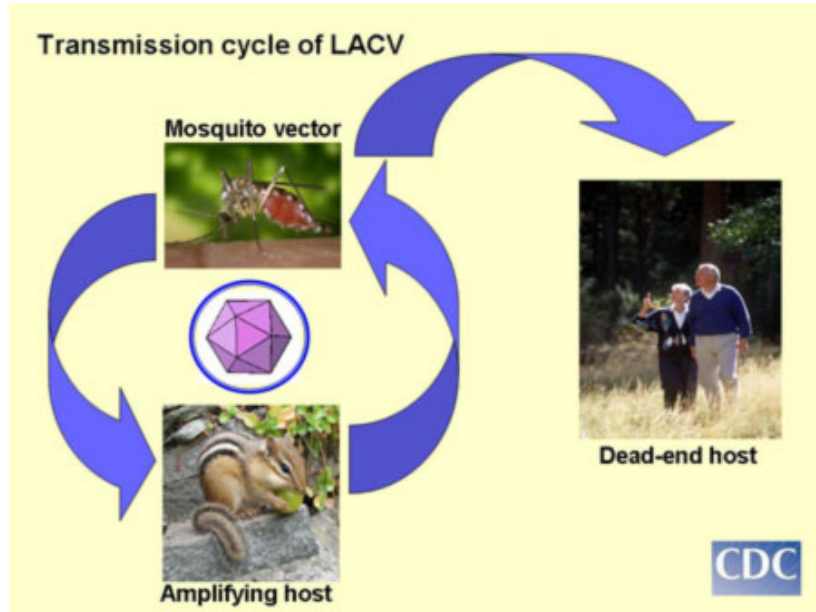
- **Superinfection of closely related viruses**
  - Mosquitoes
- **Increased pathogenesis**
  - Ngari (Garissa) virus
  - Schmallenberg virus
- **Vector/host expansion**
- **New ecological niches**
- **Prevalent in bunyaviruses**

# California serogroup

- Endemic viruses of public health importance
  - La Crosse virus – pediatric encephalitis
  - Jamestown canyon virus
  - Snowshoe hare virus
  - California encephalitis virus
- Human infection
  - Encephalitis
  - Febrile illness
  - Asymptomatic
- Mosquito borne
  - “strict” vector and small mammalian vertebrate host preferences
- Widely distributed



# La Crosse virus



- ***Aedes triseriatus* (treehole mosquito)**
  - Aggressive daytime-biting
- **Vertebrate hosts**
  - Chipmunks and squirrels
- **Human infection**
  - Fever (low viremia), headache, nausea
  - Severe neuroinvasive disease

# La Crosse virus

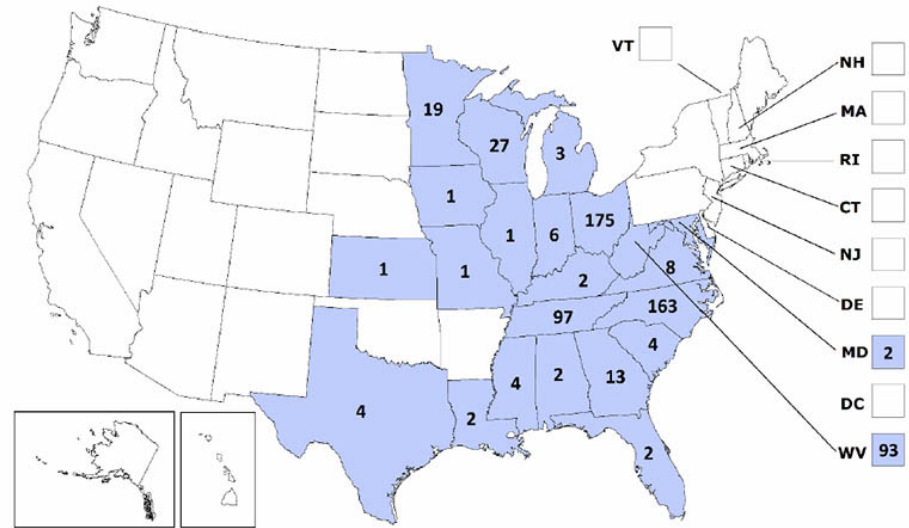
- **Changing distribution**

- Historically upper Midwest, now central Atlantic
- Widespread Midwest – Southeast

- **Clinical Lab Diagnosis**

- Difficult to culture and identify with molecular methods
- Serological diagnosis remains primary method

La Crosse disease 2007-2016 (ArboNet)  
(~70 per year)



# La Crosse virus

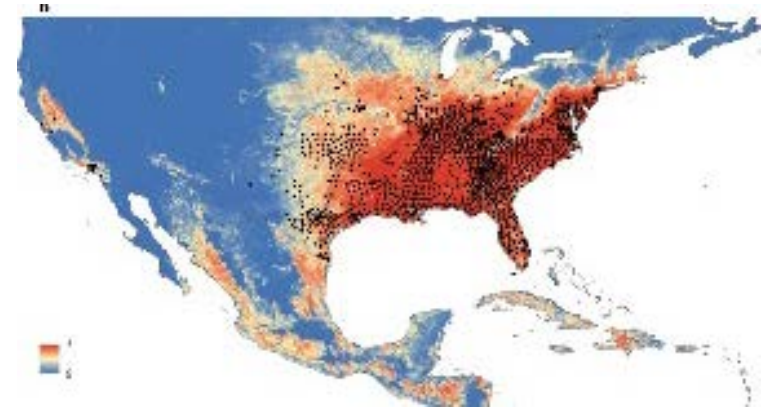
- Changing distribution

DISPATCHES

## **La Crosse Virus in *Aedes albopictus* Mosquitoes, Texas, USA, 2009**

Amy J. Lambert, Carol D. Blair, Mary D'Anton,  
Winnann Ewing, Michelle Harborth,  
Robyn Seiferth, Jeannie Xiang,  
and Robert S. Lanciotti

Predicted occurrence of *Ae. albopictus*



Kraemer et.al. eLIFE 2015; 4; e08347



# Jamestown Canyon virus

- **California serogroup**
- ***Aedes* and *Ochlerotatus***
  - 26 species of mosquito
  - 3 tabanid (horse) flies
- **Vertebrate host**
  - White-tailed deer (likely)
  - Wild ungulates and domestic livestock
- **Human infection**
  - Febrile, headache, nausea, photophobia
  - Respiratory symptoms
    - Not often reported
  - Meningitis and encephalitis
- **Predominantly in adults**

# Jamestown Canyon virus

## Jamestown Canyon 2000-2013 (31 cases)

Pastula et al. Am. J. Trop. Med. Hyg., 93(2), 2015, pp. 384–389



11 cases (2015), 15 (2016)

## ■ Widely distributed

- US and Canada
- Likely under-reported
  - 15 reported in 2016

## ■ Clinical Lab diagnosis

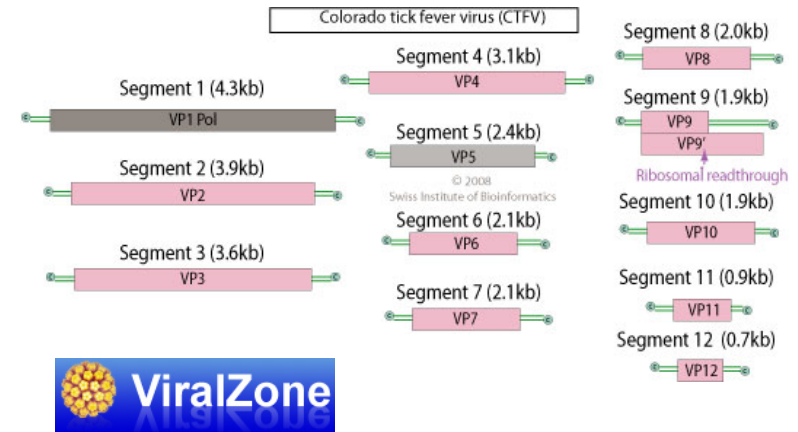
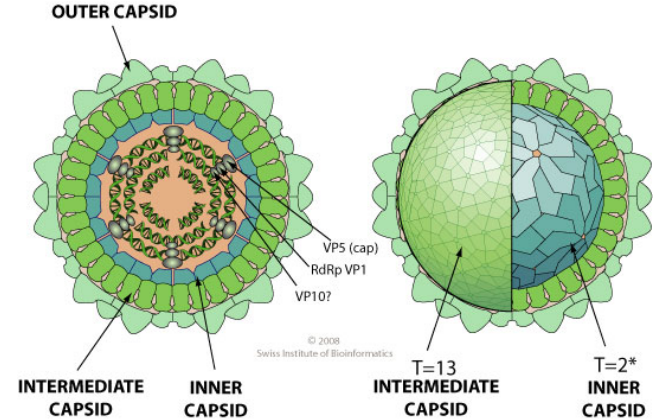
- No human isolations to date
  - RT-PCR of S segment
- Serological diagnosis primary method
  - Persistent IgM possible

# Cache Valley virus

- **Bunyamwera serogroup**
- ***Anopheles***
  - >30 mosquito species
- **Vertebrate host**
  - White-tailed deer
- **Widely distributed**
  - North and Central America
- **Livestock disease**
  - Abortion and still birth
- **Human infection**
  - Neuroinvasive encephalitis
  - 5 human cases identified in US (2015 most recent)
  - 18% seropositive in Yucatan
    - Febrile illness
- **Clinical Lab diagnosis**
  - PRNT primary method
  - Virus isolated from CSF and serum
    - Real-time RT-PCR

# Colorado tick fever (CTF)

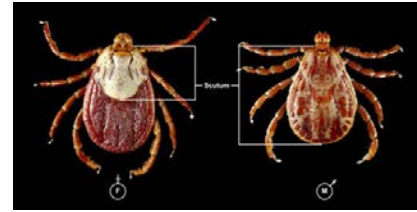
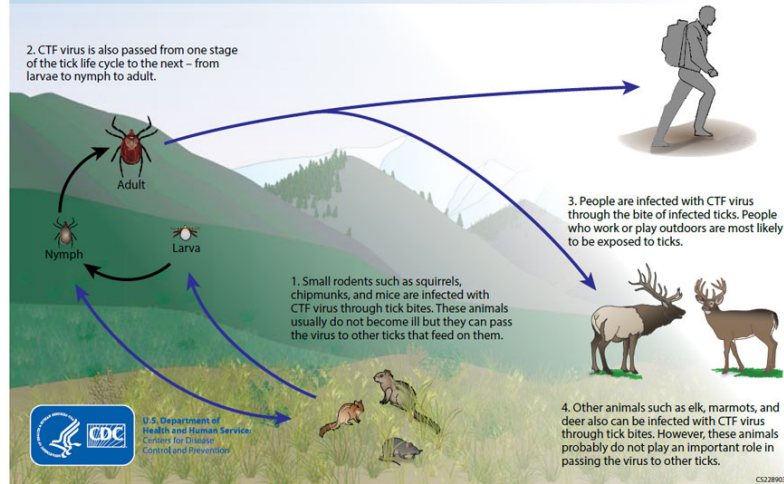
- *Reoviridae*, genus *Coltivirus*
- Non-enveloped virion
- 12 segments
- Double-stranded RNA
- Tropism for hematopoietic stem cells
  - Mature erythrocytes
- Virus found in RBCs ~ 6 weeks after symptom onset



# Colorado tick fever (CTF)

## Ecology of Colorado Tick Fever Virus

Colorado tick fever (CTF) virus is spread by Rocky Mountain wood ticks (*Dermacentor andersoni*). Rocky mountain wood ticks are found in the western United States and Canada at 4,000–10,000 feet above sea level. Here are the steps in how the virus is spread:

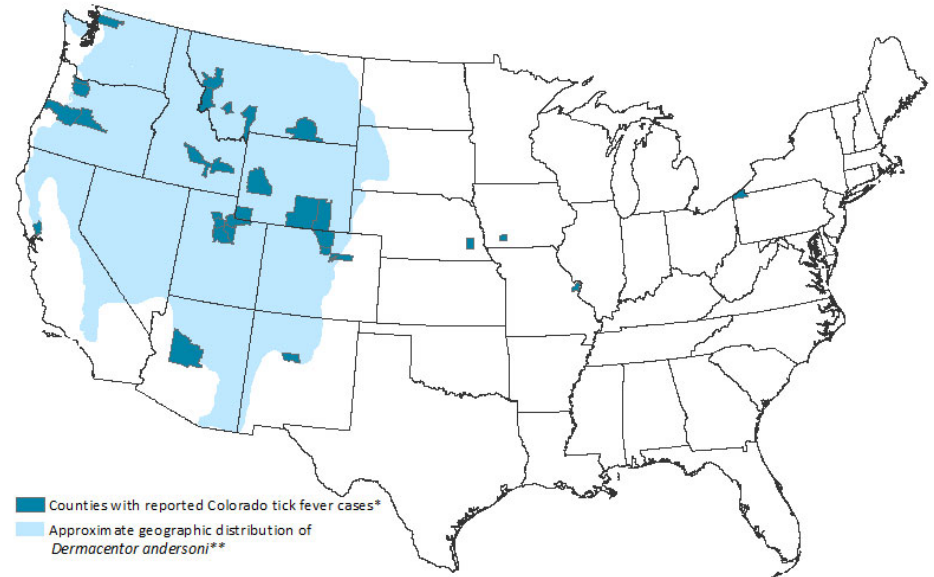


- ***Dermacentor andersoni***  
**(Rocky Mountain wood tick)**
- **Vertebrate hosts**
  - Chipmunks, squirrels, and mice
- **Human infection**
  - Febrile, head and body aches
  - Biphasic fever
  - Leukopenia
  - Rarely neuroinvasive

# Colorado tick fever (CTF)

- **Distributed widely in Western States (4,000-10,000 feet)**
- **Clinical lab diagnosis**
  - Real-time RT-PCR
    - Sensitive <21 dpo
    - RNA detected up to 42 dpo
  - Delayed antibody response
    - PRNT
    - >15 dpo

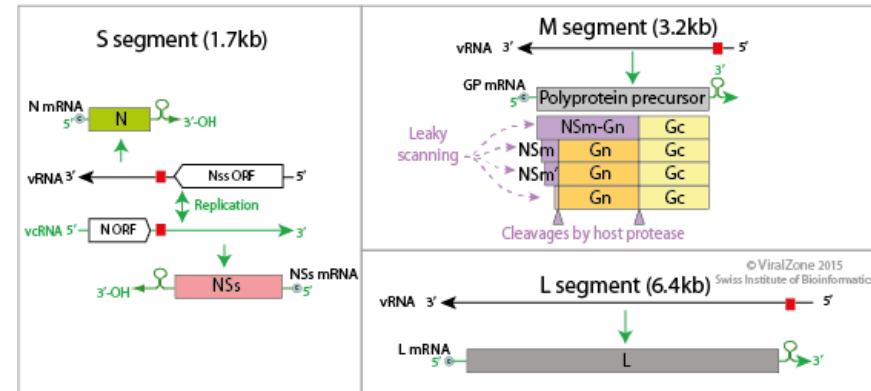
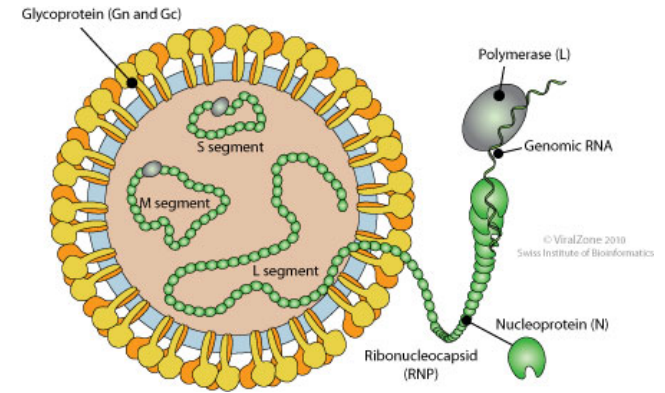
Distribution of *D. andersoni* and CTF cases 2002-2012 (83)



**Emerging arboviruses**

# Heartland virus

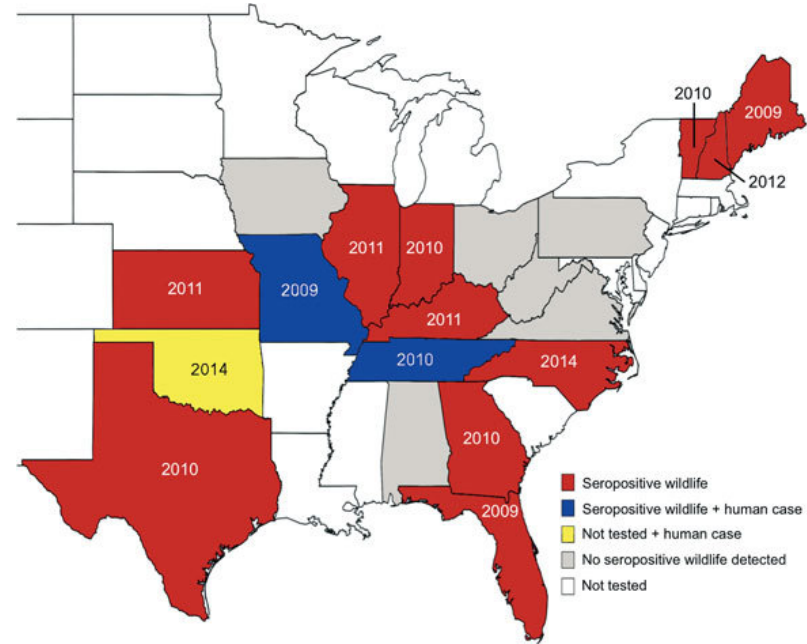
- Order *Bunyavirales*, Family *Phenuiviridae*, genus *Phlebovirus*
  - Ukuniemi serogroup
- 3 segment, negative sense RNA genome
  - Ambisense S segment
- 2009, Missouri
- Genetically similar to Severe Fever with Thrombocytopenia Syndrome virus (SFTS)
  - 2009 China





# Heartland virus

- Tick associated disease
- *Amblyomma americanum* (Lone star)
  - Virus isolation
  - Laboratory transmission
- Vertebrate host unknown
  - Seropositive: white-tailed deer, raccoon, coyote, moose



Riemersma and Komar 2015 EID

# Heartland virus

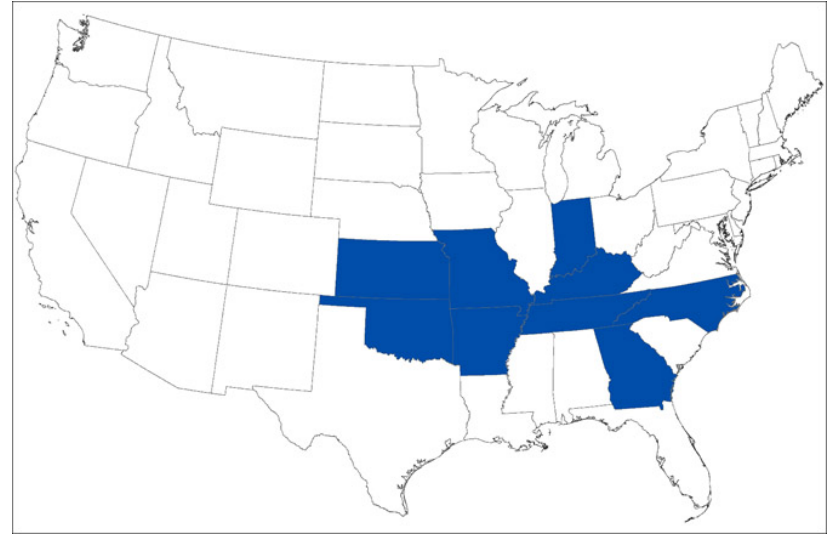
## ■ Human infection

- Febrile, headache, nausea, muscle or joint pain, leukopenia, and thrombocytopenia
- Often confused with ehrlichiosis

## ■ Clinical lab diagnosis

- Real-time RT-PCR
- Serology

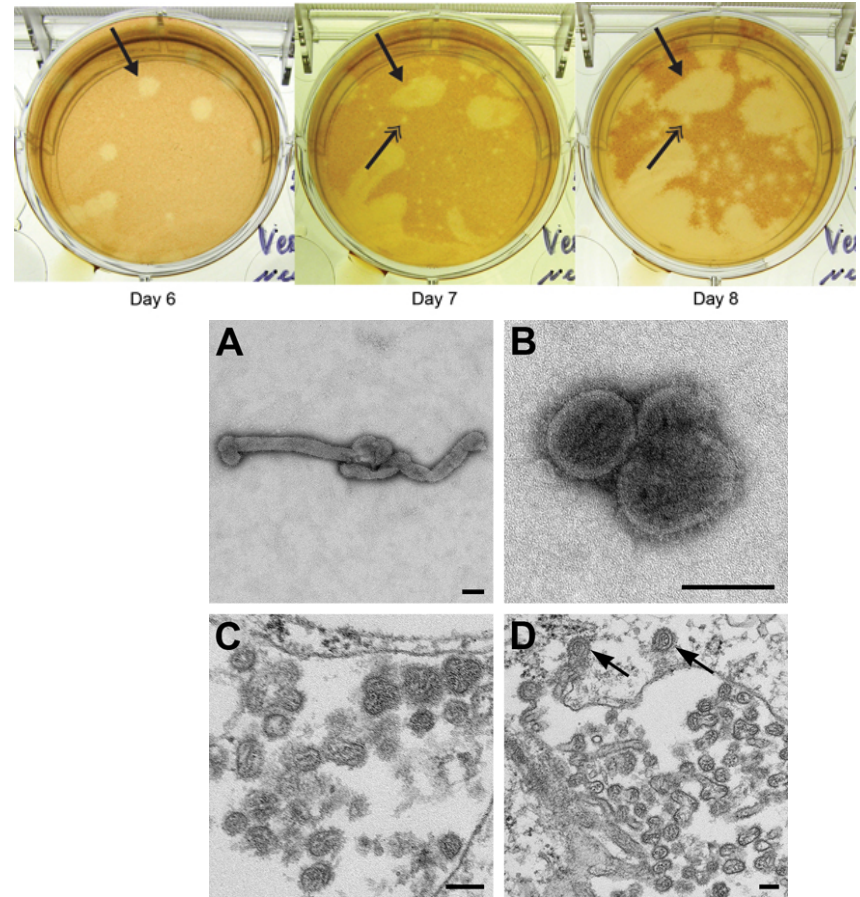
Heartland virus cases as of July 2017 (30)



[www.cdc.gov](http://www.cdc.gov)

# Bourbon virus

- *Orthomyxoviridae*, genus *Thogotovirus*
- 6 segments, negative stranded RNA genome
- Thogoto and Dhori viruses known human pathogens
  - Europe, Asia, Africa
  - Tick transmission
- **Kansas 2014**



# Bourbon virus

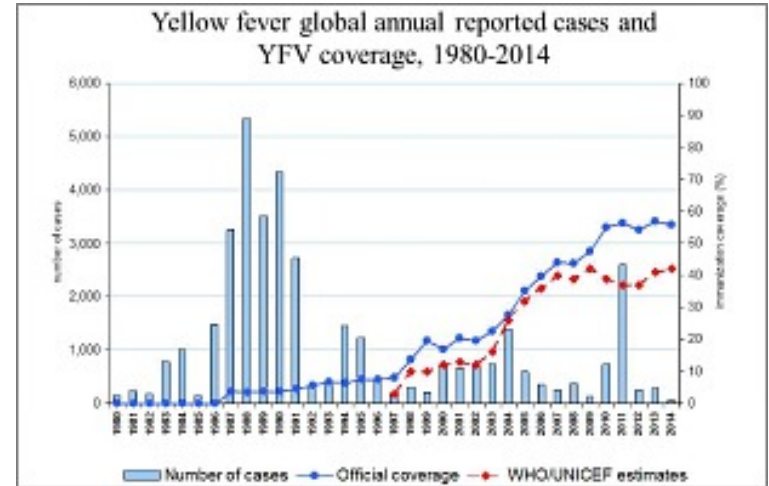
- Tick associated
- ***Amblyomma americanum* (Lone star)**
  - Isolated in ticks from MO 2013 and KS 2015
  - Nymph and adult
- **Human infection**
  - Febrile, thrombocytopenia, leukopenia
- **Clinical lab diagnosis**
  - Real-time RT-PCR
  - PRNT

Distribution of *A. americanum*



# Yellow fever virus

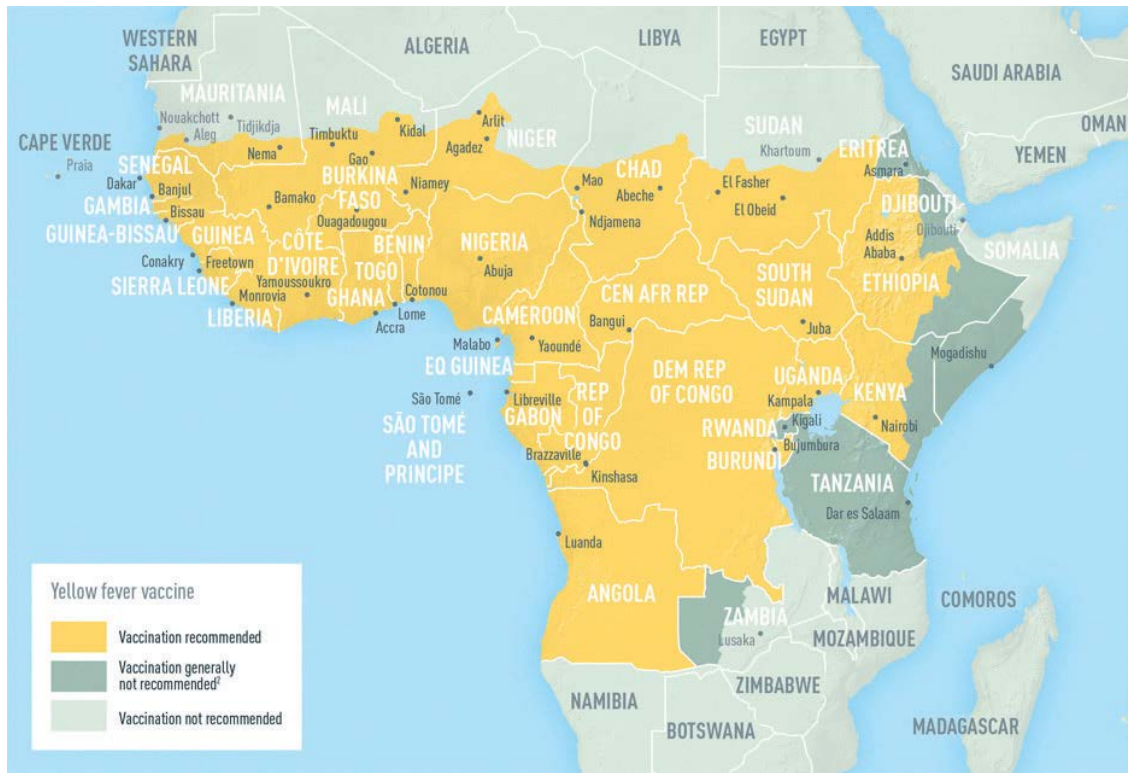
- **Flavivirus**
  - Positive RNA genome
- **Aedes aegypti**
  - urban transmission
- **Haemagogus and Sabethes**
  - sylvatic transmission (most common)
- **Human infection**
  - Febrile ILI with jaundice
  - 12-15% severe disease
  - High fever, bleeding, shock, organ failure



[www.who.int/emergencies/yellow-fever](http://www.who.int/emergencies/yellow-fever)

- **Clinical lab diagnosis**
  - Serology
  - Real-time RT-PCR

# Risk areas (vaccine recommended)

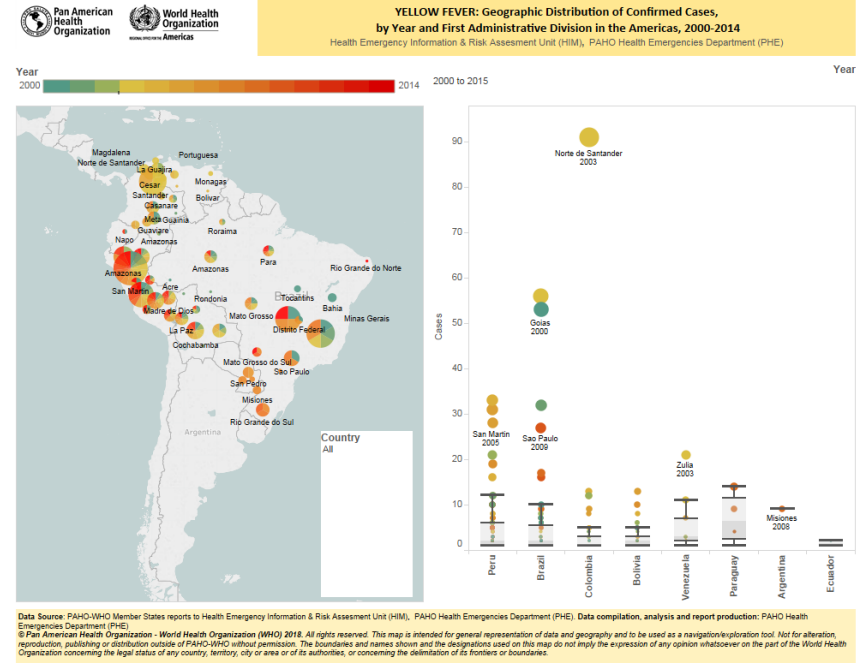




# Yellow fever virus

## Recent notable outbreaks

- 2016 Angola, Democratic Republic of the Congo (3,137 suspect)
- 2016/2017 Brazil (777 confirmed)
- 2017 Nigeria (341 suspect)
- 2018 Brazil (920 confirmed)
  - 11 international travelers



# Thank you

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

