West Nile Virus Infection

1. DISEASE REPORTING

1.1 Purpose of Reporting and Surveillance

1. To identify areas in which West Nile virus (WNv) is being transmitted.
2. To provide education to the general public on means of reducing mosquito habitats and the risk of mosquito bites.
3. To recommend preventive measures based on risk.

1.2 Laboratory and Physician Reporting Requirements

Physicians and others providing health care must report confirmed or suspected cases to the Local Health Department (LHD) by telephone within 24 hours. Laboratories are asked to report within 1 working day, and to submit all samples for confirmation to the Oregon State Public Health Laboratory (OSPHL).

1.3 Local Health Department Reporting and Follow-Up Responsibilities

1. Report all confirmed cases to the Oregon Health Authority (OHA) (see definitions below) by the end of the calendar week of initial physician/lab report. Use the standard case report form.
3. Identify significant exposures within 24 hours of report.
4. If the case tests positive at a laboratory other than OSPHL, make sure that the sample (serum or CSF) is forwarded to the OSPHL for further testing and confirmation.

2. THE DISEASE AND ITS EPIDEMIOLOGY

2.1 Background

West Nile virus (WNv) was first isolated in 1937 from a febrile woman in the West Nile District of Uganda. WNv was recognized as a cause of severe human meningoencephalitis in elderly patients during an outbreak in Israel in 1957. It was first identified in the U.S. in New York in 1999; the equine disease was first recognized in Egypt and France in the early 1960s.

2.2 Etiologic Agent

West Nile virus is a single-stranded RNA virus of the family Flaviviridae, genus Flavivirus. It is a member of the Japanese encephalitis virus serocomplex, which contains several medically important viruses associated with human encephalitis: Japanese encephalitis, St. Louis encephalitis, Murray Valley encephalitis, and Kunjin virus (an Australian subtype of West Nile virus). The close antigenic relationship of the flaviviruses, particularly those belonging to the Japanese encephalitis complex, accounts for the serologic cross-reactions observed in the diagnostic laboratory.

2.3 Description of Illness

Serosurveys indicate that about 80% of those infected with WNv are asymptomatic. Only 10–20% of individuals present with febrile illness of sudden onset, often accompanied by malaise, anorexia, nausea, vomiting, eye pain, headache, myalgia, rash, and lymphadenopathy; these symptoms generally last 3 to 6 days, and most cases are never diagnosed. Approximately half of hospitalized U.S. patients have had severe muscle weakness and encephalopathy. Approximately 10% of patients with encephalitis develop a poliomyelitis-like syndrome with flaccid paralysis. Neurologic presentations other than encephalitis or
West Nile Virus

meningitis, which occur more rarely, include ataxia and extrapyramidal signs, cranial nerve abnormalities, myelitis, optic neuritis, polyradiculitis, and seizures. Myocarditis, pancreatitis, and fulminant hepatitis have been described in outbreaks occurring before 1990. Although recent outbreaks of West Nile virus seem to be associated with increased morbidity and mortality, severe neurologic disease remains uncommon. Two serosurveys conducted in New York City in 1999 and 2000 showed that approximately 1 in 150 infections resulted in meningitis or encephalitis, a result consistent with a 1996 Romanian serosurvey indicating that 1 in 140 to 320 infections led to these diseases.

2.4 Reservoirs

West Nile virus is maintained in an enzootic cycle involving culicine mosquitoes and different bird species; Corvids (crows, ravens, jays, magpies) seem to be most affected. After passing through three aquatic stages (egg, larva, pupa), adult mosquitoes begin to emerge in the spring in temperate regions. Viral amplification occurs in the bird–mosquito–bird cycle until early fall, when female mosquitoes bite infrequently. Many environmental factors affect this viral amplification cycle (for example, weather or climate, host and vector predators and parasites, and host immune status). When environmental conditions promote significant amplification, sufficient numbers of "bridge vector" mosquitoes—mosquitoes that bite both humans and birds—become infected in late summer and then pose an infection threat to humans. Year-round transmission is possible in more tropical climates. Through spring 2002, West Nile virus had been detected in 29 North American mosquito species; this number will undoubtedly increase as the virus spreads into new ecologic habitats. Although Culex pipiens, Culex restuans, Culex tarsalis and Culex quinquefasciatus are probably the most important maintenance vectors in the eastern United States, it is unknown which species are most responsible for transmission to humans. In Oregon, all the above-mentioned species are able and ready to bite!

2.5 Modes of Transmission

The main route of human infection with West Nile virus is through the bite of an infected mosquito. Mosquitoes become infected when they feed on infected birds, which may circulate the virus in their blood for a few days. The virus eventually gets into the mosquito’s salivary glands. During later blood meals (when mosquitoes bite), the virus may be injected into humans and animals, where it can multiply and possibly cause illness. In a very small number of cases, WNv also has spread through blood transfusions, organ transplants, breastfeeding and even during pregnancy from mother to baby. There is no evidence that WNv virus can be transmitted to humans through consuming infected birds or animals. In keeping with overall public health practice, and due to the risk of known food-borne pathogens, people should always follow procedures for fully cooking meat from either birds or mammals. WNv is not spread through casual contact such as touching or kissing a person with the virus.

2.6 Incubation Period

Usually 3 to 14 days.

2.7 Period of Communicability

Affected persons develop a short-lived viremia (2-3 days) and are considered to be dead-end hosts, except in the rare situation where transmission occurs through blood transfusions, organ transplants, breastfeeding, and perinatal spread. Human–mosquito transmission does not occur due to the low concentration of viruses during the short-lived viremic stage.

2.8 Treatment

Treatment is supportive, often involving hospitalization, intravenous fluids, respiratory support, and prevention of secondary infections for patients with severe disease. Ribavirin in high doses and interferon alpha-2b were found to have some activity against WNv in vitro, but no controlled studies have been completed on the use of these or other medications, including steroids, antiseizure drugs, or osmotic agents, in the management of WNv encephalitis.
West Nile Virus

3. CASE DEFINITIONS, DIAGNOSIS, AND LABORATORY SERVICES

3.1 Indication for Testing

1. Febrile, aseptic meningitis with prior testing to rule out enteroviruses by PCR or culture (recommended)
2. Febrile encephalitis of unknown etiology
3. Flaccid paralysis, myelitis or neurological symptoms following a febrile illness
4. Onset of fever, rash, encephalitis or meningitis symptoms within two weeks of receiving blood products
5. Compatible illness and exposure history in a pregnant or breast-feeding woman
6. Clinically compatible illness during transmission season

3.2 Case Definition

**Confirmed case:** A clinical illness characterized by acutely altered mental status (e.g., disorientation, obtundation, stupor, or coma); or other acute signs of neurologic dysfunction (e.g., paresis or paralysis, nerve palsies, sensory deficits, abnormal reflexes, convulsions, or abnormal movements); or clinical signs or symptoms of meningitis with CSF pleocytosis

and

WNv-specific antibodies in serum or CSF confirmed by OSPHL or other state or federal public-health laboratory according to CDC guidelines;

or

WNv-specific genomic sequence identified in a clinical specimen; or isolation of WNv from a clinical specimen.

**Suspect case:** A neurologic illness as characterized above;

and

WNv-specific antibodies in serum or CSF identified by a commercial laboratory but not confirmed by a public-health laboratory.

3.3 Laboratory Confirmation

The most efficient diagnostic method is detection of IgM antibody to WNv in serum collected within 4–14 days of illness onset or cerebral spinal fluid (CSF) collected within 8 days of illness onset using the IgM antibody capture enzyme-linked immunosorbent assay (MAC-ELISA).

Any sample testing positive by MAC-ELISA will be forwarded to CDC or the California State Laboratory Division of Viral Disease for further confirmation by Plaque Reduction Neutralization Test (PRNT).

Since IgM antibody does not cross the blood-brain barrier, the presence of IgM antibody in CSF strongly suggests central nervous system infection. IgG positive and IgM negative tests are indicative of previous infection and these cases do not need to be investigated.

Patients who have been recently vaccinated against or recently infected with related flaviviruses (e.g., yellow fever, Japanese encephalitis, dengue) may have positive WNv MAC-ELISA results.

3.4 Services Available at the Oregon State Public Health Laboratory

OSPHL will test serum samples and CSF samples for the WNv by ELISA and forward positive samples for confirmatory testing.

4. ROUTINE INVESTIGATION

4.1 Case Interview

Interview the case (or parents) and others who may be able to provide pertinent information.

1. Identify Source of Infection

Obtain a travel history and about mosquito exposures during the likely exposure period to assist vector control. It is also important to ask about receiving blood products or about organ or tissue transplants.
2. Identify Potentially Exposed Persons
Determine if patient donated blood or organs, breastfed, or gave birth during the communicable period. If patient donated blood or organs, inform the blood or tissue bank of the potential exposure. In cases of potential mother-to-infant transmission, monitor the infant for compatible signs and symptoms for 14 days after last possible exposure

4.2 Environmental Evaluation
Generally none, although in outbreak settings, an investigation may be warranted to identify factors favoring transmission

5. CONTROLLING FURTHER SPREAD

5.1 Education
Potentially exposed persons should be instructed to watch for fever, rash, lethargy, headache, loss of appetite, or vomiting. Should signs or symptoms develop within the next two weeks, they should seek medical care as needed and advise the physician of the potential for exposure. Persons exposed should be advised not to donate blood. Pregnant women should discuss breast-feeding with their medical care provider.

5.2 Isolation
No isolation needed.

5.3 Case Follow up
No case follow-up needed.

5.4 Protection of Contacts
None.

5.5 Precautions
Precautions against the disease are based on protection against mosquito bites.
Reduce the number of mosquitoes in breeding sites outdoors by draining sources of standing water.

• At least once or twice a week, empty water from flowerpots, pet food and water dishes, birdbaths, swimming pool covers, buckets, barrels, and cans.
• Check for clogged rain gutters and clean them out.
• Remove discarded tires, and other items that could collect water.
• Be sure to check for containers or trash in places that may be hard to see, such as under bushes or under the home.
• Place mosquito netting over infant carriers when outdoors.
• Consider staying indoors at dawn, dusk, and in the early evening, which are peak mosquito biting times.
• Install or repair window and door screens so that mosquitoes cannot get indoors

Reduce exposure to mosquitoes

• Repellants are an important tool to assist people in protecting themselves from mosquito-borne diseases. CDC recommends the use of products containing active ingredients which have been registered by the U.S. Environmental Protection Agency (EPA) for use as repellents applied to skin and clothing. EPA registration of repellent active ingredients indicates the materials have been reviewed and approved for efficacy and human safety when applied according to the instructions on the label.

Repellents for use on skin and clothing

• CDC evaluation of information contained in peer-reviewed scientific literature and data available from the EPA has identified several EPA registered products that provide repellent activity sufficient to help people avoid the bites of disease carrying mosquitoes. Products containing these active ingredients typically provide reasonably long-lasting protection:
  o DEET (Chemical Name: N,N-diethyl-m-toluamide or N,N-diethly-3-methyl-benzamide)
  o Picaridin (KBR 3023, Chemical Name: 2-(2-hydroxyethyl)-1-piperidinecarboxylic acid 1-methylpropyl ester)
West Nile Virus

- Oil of Lemon Eucalyptus* or PMD (Chemical Name: para-Menthane-3,8-diol) the synthesized version of oil of lemon eucalyptus
- IR3535 (Chemical Name: 3-[N-Butyl-N-acetyl]-aminopropionic acid, ethyl ester)

EPA characterizes the active ingredients DEET and Picaridin as "conventional repellents" and Oil of Lemon Eucalyptus, PMD, and IR3535 as "biopesticide repellents", which are derived from natural materials. For more information on repellent active ingredients see (http://www.epa.gov/pesticides/health/mosquitoes/ai_insectrp.htm).

UPDATE LOG

March 2009. Changes included expansion of repellant information under 5.5 Precautions. (Emilio DeBess)
West Nile Virus