

Figure. Temperature and heart rate relationship: scatter plot for patients with dengue fever and nondengue febrile illnesses.

the etiology of dengue-associated relative bradycardia.

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## West Nile Virus, Venezuela

**To the Editor:** West Nile virus (WNV; genus *Flavivirus*; family *Flaviviridae*) has been perpetuating in North America since 1999 (1). However, its status as a self-perpetuating pathogen in South America remains uncertain. Infected horses and birds have been reported in various Caribbean Islands, Mexico, and northern Central America (2,3). In South America, isolated reports of infected dead-end hosts (horses) have come from northern Colombia and Argentina, but they lack evidence for infection in avian amplifying hosts (4,5). We report serologic evidence of establishment of WNV in South America.

Serum samples from birds and horses from 33 locations in Venezuela (Online Appendix Table, available from <http://www.cdc.gov/EID/content/13/4/651-appT.htm>) were screened for immunoglobulin G (IgG) antibodies against WNV antigen by ELISA (6) and confirmed by plaque reduction neutralization test (PRNT) as previously described (7). The flavivirus generating the IgG response was identified by using the following criteria: 90% inhibition of virus in serum diluted at least 1:40 and 4-fold greater neutralizing antibody titer compared with closely related flaviviruses. IgG antibody against flavivirus was detected by ELISA in 14 of 576 resident birds, including 5 *Turdus leucomelas*, 3 *Gallus gallus* (captive), 2 *Campylorhamphus trochilirostris*, and 1 each of *Elaenia flavogaster*, *Coereba flaveola*, *Thraupis palmarum*, and *Anisognathus flavinucha*.

WNV was confirmed as the etiologic agent of infection in 5 adult birds (3 *T. leucomelas* [pale-breasted thrush], 1 *C. flaveola* [bananaquit], and 1 *G. gallus* [domestic chicken] with the earliest collection date in February 2006); virus neutralization

titers ranged from 80 to 320. One serum sample cross-reacted with other flaviviruses tested, with equivalent titers to WNV, Saint Louis encephalitis virus (SLEV), and Ilheus virus (ILHV) and was thus considered infected with an undetermined flavivirus. Seven serum samples were negative (antibody titers <20), and 1 sample was not tested because of insufficient sample volume.

Antibody against flavivirus was detected by ELISA in 141 of 791 horses, and 34 (4.3%) were confirmed positive for WNV infection by PRNT; viral titers  $\geq 640$  occurred in half of these horses. The earliest collection date for a WNV-positive horse was February 2004 and the most recent was May 2006. Specific WNV-reactive equine serum samples were distributed in valley regions (prevalence 1.3%), savannah grasslands (2.4%), the western region of Zulia (0.4%) and the Central Lake Basin (0.3%). A total of 46 (5.8%) equine serum samples were positive for neutralizing antibody to SLEV, and 8 (1.0%) samples were positive for neutralizing antibodies to ILHV. Forty-nine samples neutralized at least 2 of the 3 viruses and were classified as undetermined flaviviruses. Serum samples from 2 horses were negative in neutralization assays; 2 others were not tested because of insufficient sample volume.

Detection of WNV-infected resident birds provides strong evidence of the establishment, rather than importation, of WNV in South America. We hypothesize that ornithophilic mosquitoes (such as some *Culex* spp.), which are present in the area in consistently high numbers, acquired the virus through hematophagous feeding on recently infected, migrating birds. Once introduced to local mosquitoes, virus is amplified among susceptible resident birds fed upon by ornithophilic mosquitoes. This pattern allows perpetuation and subsequent establishment of virus in a continuous transmission cycle, as opposed to

infection of dead-end hosts, e.g., horses. This is the first report of WNV infection in South American birds and definitive establishment of the virus in South America.

We observed varying WNV seroprevalence rates in birds and horses across regions in Venezuela (Figure). These differences reflect the focal and stochastic nature of arbovirus transmission, which depends upon many ecologic factors. One possible explanation for the greater seroprevalence in the central and eastern llanos (savannahs) and valley regions, compared with the coastal western region of Zulia State ( $p < 0.0001$ , by Pearson's  $\chi^2$  test) would be virus introduction by migrating birds by an eastern migration route.

Existence of several closely related flaviviruses in the American tropics (8–10) may convey cross-protection in animals (e.g., ILHV and SLEV) or humans (dengue viruses, yellow fever virus), thereby potentially diminishing disease caused by a newly introduced flavivirus such as WNV. Although ILHV infection has

not been detected in Venezuela, this flavivirus is prevalent in Brazil, Peru, French Guyana, Trinidad, and Colombia. Our study demonstrated widespread distribution of ILHV in Venezuela. Other South American flaviviruses, such as Bussuquara, Cacipacore, and Iguape, and as yet undiscovered viruses may also circulate in Venezuela.

We encourage those involved in the public and animal health systems in Venezuela to consider zoonotic flaviviruses in the differential diagnoses of encephalitis and to consider ecologic surveillance for zoonotic flaviviruses in mosquito and vertebrate host populations. We recommend monitoring blood and organ donations for flavivirus infections. Our study sheds light on flavivirus distribution in Venezuela. However, nothing else is known about the ecology of zoonotic flaviviruses in this country. Such knowledge will be essential for designing effective surveillance and control should these viruses be shown to cause human illnesses.

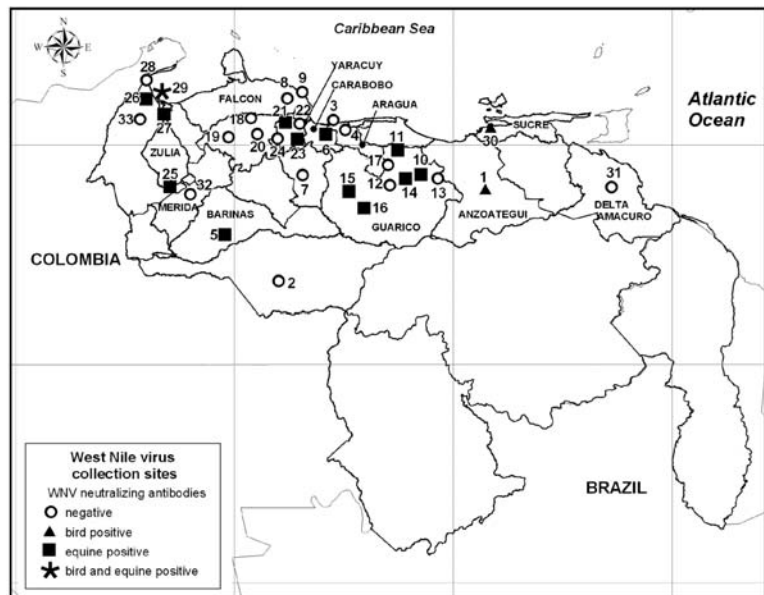


Figure. West Nile virus (WNV) collection sites in Venezuela, indicated by number (see online Appendix Table). Symbols represent results of tests for specific antibodies to WNV in serum samples of birds and horses (viral titers in a 90% plaque reduction neutralization test  $>40$  and a 4-fold differential inhibition in a neutralization assay to WNV compared with other related flaviviruses). Source: Instituto Geográfico de Venezuela Simón Bolívar, Caracas.

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## Novel Extended-spectrum $\beta$ -Lactamase in *Shigella sonnei*

**To the Editor:** A 38-year-old French man with a history of chronic juvenile arthritis was referred to the Necker-Enfants Malades University hospital (Paris, France) with a dysenteric syndrome. The patient had returned the day before from a 1-month stay in Port-au-Prince, Haiti, where he spent most of his time in close contact with young children from an orphanage, most of whom had diarrhea. Clinical examination at admission showed fever (39°C), chills, diffuse abdominal pain, bloody diarrhea, and vomiting. The patient received ceftriaxone, which was stopped on day 4 because initial blood and stool cultures were negative for pathogens and clinical signs had completely resolved.

Ten days later, he reported the recurrence of diarrhea without fever. A novel stool culture grew *Shigella sonnei*. An extended-spectrum  $\beta$ -lactamase (ESBL) was detected by double-disk synergy test; the isolate was also resistant to aminoglycosides (except amikacin), tetracycline, and cotrimoxazole. The strain was susceptible to fluoroquinolones and fosfomicin. It also appeared susceptible to azithromycin (MIC 4  $\mu$ g/mL), although azithromycin MIC for *Shigella* spp. should be interpreted with caution (1). The patient was successfully treated with azithromycin at a dose of 500 mg/day for 5 days. Azithromycin was preferred to fluoroquinolones to avoid the risk for tendinopathy because of the patient's history of chronic juvenile arthritis and because this antimicrobial agent was shown to be effective in the treatment of shigellosis caused by multidrug-resistant strains (2).

To identify the molecular basis of this ESBL, a series of PCR primers

<sup>1</sup>Deceased.

Online Appendix Table. Locations in Venezuela sampled for West Nile virus

ID	State	County, Location	Regional ecosystem*	No. serum samples tested (no. positive)†	
				Birds	Equines
4	Aragua	Guirardot, Portachuelos	Coast	96 (0)	7 (0)
8	Falcón	Manuere, Guayabal	Coast		
25	Zulia	Francisco J. Pulgar	Coast	12 (0)	6 (1)
26	Zulia	Mara	Coast		9 (1)
33	Zulia	La C. de Urdaneta	Coast	327 (1)	6 (0)
27	Zulia	Maracaibo	Coast		
28	Zulia	Paez, Goajira	Coast	25 (0)	43 (1)
2	Apure	Romulo Gallegos	Llanos		
10	Guarico	Rivas	Llanos	6 (3)	5 (0)
11	Guarico	Monagas, Altgr. Orituco	Llanos		
13	Guarico	Ortiz, Zaraza	Llanos	18 (3)	50 (3)
14	Guarico	Infante, V. de la Pascua	Llanos		
15	Guarico	Miranda, Guarico Dam	Llanos	13 (5)	27 (2)
16	Guarico	Miranda	Llanos		
21	Yaracuy	Bolivar	Valley	75 (0)	24 (8)
23	Yaracuy	S. Felipe, Guaquira farm	Valley		
22	Yaracuy	Veroes	Valley	9 (4)	18 (0)
5	Barinas	Pedraza	Llanos		
7	Cojedes	Pao	Llanos	3 (0)	4 (0)
12	Guarico	Las Mercedes	Llanos		
17	Guarico	Monagas	Llanos	1 (0)	3 (0)
18	Lara	Urdaneta	Valley		
19	Lara	Torres	Valley	3 (0)	1 (0)
20	Lara	Iribarren	Valley		
24	Yaracuy	Sucre	Valley	5 (0)	1 (0)
32	Merida	Libertador, Merida	Mountain		
3	Aragua	Girardot, Cata farm	Coast	52 (0)	1 (0)
6	Carabobo	Valencia, Lake	Valley		
29	Zulia	Paez, Sinamaica Lagoon	Coast	142 (1)	173 (2)
30	Sucre	Sucre, Cumana	Coast		
1	Anzoategui	Miranda, F. El Tigre	Llanos	7 (1)	56 (3)
9	Falcón	Iturriza, Cuare Reserve	Coast		
31	Delta Amacuro	Tucupita	Delta system	72 (0)	791 (34)
	Total			576 (5)	

\*Llanos, savannah.

†Positive by 90% plaque reduction neutralization test.