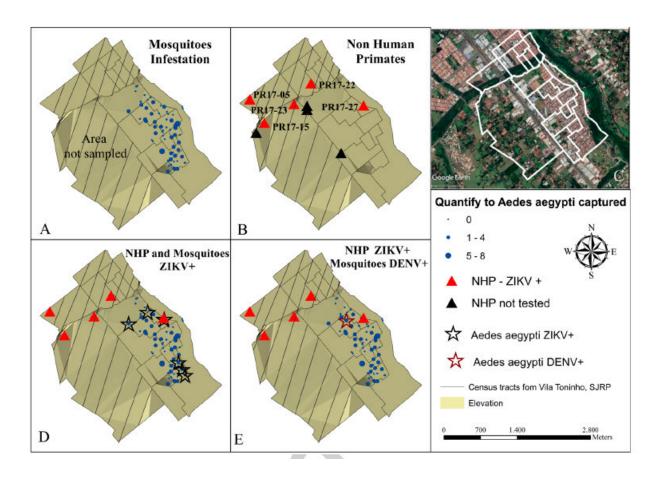


Discovery of Zika virus in monkeys suggests disease may also have wild cycle

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Sylvatic cycle would give virus a natural reservoir from which it could more frequently reinfect humans. Study on subject has been published by Brazilian researchers in *Scientific Reports*. Credit: *Scientific Reports*



Zika virus has been detected in dead monkeys found in Brazil near São José do Rio Preto, São Paulo State, and Belo Horizonte, Minas Gerais. The animals had been shot or beaten to death by locals who thought they had yellow fever. In fact, the monkeys were bearers of Zika virus, which had made them sick and more vulnerable to attack by humans.

"The discovery shows the potential exists for Zika to establish a sylvatic transmission cycle [involving wild animals] in Brazil, as already occurs in the case of <u>yellow fever</u>. If the wild cycle is confirmed, it completely changes the epidemiology of Zika because it means there's a natural reservoir from which the <u>virus</u> can reinfect the human population much more frequently," said Maurício Lacerda Nogueira, principal investigator for the study. Nogueira is a professor at São José do Rio Preto Medical School (FAMERP) and chairs the Brazilian Society for Virology (SBV).

Zika virus was detected in monkeys accustomed to the presence of humans in the state of Ceará some years ago, but this is the first detection in the wild since the ongoing epidemic began. An article on the study signed by Nogueira and researchers affiliated with several other institutions has just been published in *Scientific Reports*.

"During the yellow fever epidemic, we realized there were many dead monkeys, killed not by the disease but by people who were afraid of contagion. The monkeys had been shot, cudgeled or bitten to death by dogs. When healthy, these primates—mainly marmosets (Callithrix sp.) and capuchins (Sapajus sp.)—are very hard to capture," Nogueira explained. "So we concluded that if they were being killed relatively easily, it was probably because they were sick, not from yellow fever, which is lethal to monkeys, but from some other disease that didn't kill them but made them weak and vulnerable."

The researchers analyzed the dead monkeys and found them to have



been infected by Zika in São José do Rio Preto and Belo Horizonte. Whole genome sequencing showed the virus to be very similar to the type that was infecting humans. In the same week, moreover, the researchers collected mosquitoes infected by Zika from the same places as those in which the dead monkeys were found.

"We proceeded with the study by experimentally infecting live monkeys with Zika," Nogueira said. "Inoculation with the virus caused viremia [presence of the virus in the blood]. The monkeys displayed behavioral alterations, confirming our initial hypothesis that the infection made them more susceptible to being captured and killed."

Yellow fever cycle

Natural and experimental infection of monkeys with Zika shows these animals can be vertebrate hosts, participating in transmission and circulation of the virus in tropical urban environments. According to the article just published, however, "more studies are necessary to understand the role they may play in maintaining the urban cycle of Zika virus and how they can be a channel for the establishment of an enzootic [nonhuman] transmission cycle in tropical Latin America."

"As yellow fever has shown, epizootic diseases [occurring at the same time in many animals in the same geographic area] will always be a source of epidemics among humans, even after a possible control and eradication of the urban transmission cycle through the development of successful countermeasures such as vaccines and antivirals. It's a game changer, and policy makers, public health officials and vaccine developers should take notice," said Nikos Vasilakis, a professor at UTMB's Center for Tropical Diseases and one of the lead authors of the article just published.

Zika virus first appeared in monkeys in Africa. From time to time it left



the jungle and infected human populations. It eventually spread to Asia, where it circulated only among humans. It apparently retained this property when it came to the Americas, suggesting a cycle similar to that of dengue virus.

The new discovery, however, points to the possibility of a different epidemiology, more like that of yellow fever. If this epidemiology is confirmed, it means combating Zika will be much harder than has been assumed to date.

"Our observations will have important implications in our understanding of the ecology and transmission of Zika in the Americas," Vasilakis said. "Although this is among the first steps in the establishment an enzootic transmission cycle among New World nonhuman primates and arboreal mosquitoes, the implications are enormous, because it is impossible to eradicate this transmission cycle."

Vasilakis stressed the importance of the funding for the study received from FAPESP and the US National Institutes of Health (NIH), which "recognized the significance of the subject early on," and supported research that also serves as an example of the "seamless and long-established collaboration between groups at UTMB and FAMERP."

More information: Ana Carolina B. Terzian et al, Evidence of natural Zika virus infection in neotropical non-human primates in Brazil, *Scientific Reports* (2018). DOI: 10.1038/s41598-018-34423-6

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