

Dengue: Asymptomatic people transmit the virus to mosquitoes

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Scientists at the Institut Pasteur in Cambodia, the Institut Pasteur in Paris and the CNRS provided proof that people infected by dengue virus but showing no clinical symptoms can actually infect mosquitoes that bite them. It appears that these asymptomatic people - who, together with mildly symptomatic patients, represent three-quarters of all dengue infections - could be involved in the transmission chain of the virus. These findings, published in the journal *PNAS*, on the 9 of November, question established theories concerning the epidemiology of dengue.

Dengue virus infects 390 million people worldwide each year through the bite of mosquitoes of the Aedes genus. But estimates suggest that



300 million of these people do not present clinical symptoms that are severe enough to be detected by health care systems. It was previously thought that these asymptomatic or mildly symptomatic infections did not reach a high enough level of viremia (the concentration of the virus in the blood) to infect mosquitoes.

In this study published in *PNAS*, scientists from the Institut Pasteur in Cambodia, the Institut Pasteur in Paris, and the CNRS focused their attention on the hidden mass of the iceberg, in other words the people with few or no symptoms that represent 75% of all dengue infections, and experimentally tested whether these people could actually contaminate mosquitoes. The scientists carried out their study in a human population at risk of dengue infection in Cambodia, in the town of Kampong Cham, roughly 100km northeast of Phnom Penh. Their research strategy relied on their ability to detect dengue infections that had not been identified by traditional health care systems because the participants showed virtually no signs of the disease.

The scientists performed blood tests on people living in close proximity in the same household or in the immediate neighborhood - to patients with confirmed symptomatic dengue. People who tested positive for dengue virus in their blood tests but without clinical symptoms were then put into contact with healthy laboratory-bred mosquitoes. Subsequent analysis of the mosquitoes confirmed they had been infected and would be capable of transmitting the virus the next time they bit a human. The research data also confirmed that the viremia level is a critical factor in transmission of dengue virus from a human to a mosquito.

"This finding raises the possibility that people with few or no symptoms - in other words the majority of those infected by dengue - may actually be contributing to the spread of the virus without realizing it," explained Louis Lambrechts, a CNRS scientist in charge of the Insect-Virus Interactions Group at the Institut Pasteur in Paris. Moreover, people who



are virtually or completely unaffected by the virus are potentially exposed to more <u>mosquitoes</u> during their daily routines than those who are severely ill, bed-ridden or hospitalized.

"These data should lead us to revisit our approach to the early management of dengue epidemics. Transmission rate estimates will also have to be adjusted to ensure sufficient vaccination coverage for the vaccines currently under development," commented Veasna Duong, a scientist in the Virology Unit directed by Philippe Buchy at the Institut Pasteur in Cambodia, when this work had been done. Meanwhile, at the Institut Pasteur in Paris, the DENFREE European project, coordinated by Anavaj Sakuntabhai, is investigating the specific biological characteristics of asymptomatic infections.

More information: V. Duong et al. Asymptomatic humans transmit dengue virus to mosquitoes, *Proceedings of the National Academy of Sciences* (2015). DOI: 10.1073/pnas.1508114112

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