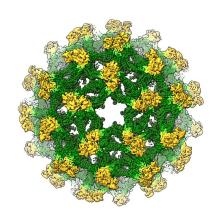


Scientists rewrite the genesis of mosquitoborne viruses

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Cryo-electron microscopy reconstruction of Binjari virus. The projecting spikes are a typical feature of immature flaviviruses such as dengue virus but reveal an unexpected organization. Credit: Associate Professor Fasseli Coulibaly

Better designed vaccines for insect-spread viruses like dengue and Zika are likely after researchers discovered models of immature flavivirus particles were originally misinterpreted.

Researchers from The University of Queensland and Monash University have now determined the first complete 3-D molecular structure of the immature flavivirus, revealing an unexpected organisation.

UQ researcher Associate Professor Daniel Watterson said the team was studying the insect-specific Binjari <u>virus</u> when they made the discovery.

"We were using Australia's safe-to-handle Binjari virus, which we combine with more dangerous viral genes to make safer and more effective vaccines,"

Dr. Watterson said.

"But when analysing Binjari we could clearly see that the <u>molecular structure</u> we've all been working from since 2008 wasn't quite correct.

"Imagine trying to build a house when your blueprints are wrong—that's exactly what it's like when you're attempting to build effective vaccines and treatments and your molecular 'map' is not quite right."

The team used a technique known as <u>cryogenic</u> <u>electron microscopy</u> to image the virus, generating high resolution data from Monash's Ramaciotti Centre for Cryo-Electron Microscopy facility.

With thousands of collected two-dimensional images of the virus, the researchers then combined them using a high-performance computing platform called 'MASSIVE' to construct a high-resolution 3-D structure.

Monash's Associate Professor Fasséli Coulibaly, a co-leader of the study, said the revelation could lead to new and better vaccines for flaviviruses, which have a huge disease burden globally.

"Flaviviruses are globally distributed and <u>dengue</u> <u>virus</u> alone infects around 400 million people annually," Dr. Coulibaly said.

"They cause a spectrum of potentially severe diseases including hepatitis, vascular shock syndrome, encephalitis, acute flaccid paralysis, congenital abnormalities and foetal death.

"This structure defines the exact wiring of the immature virus before it becomes infectious, and we now have a better understanding of the levers and pulleys involved in viral assembly.

"This is a continuation of fundamental research by us and others and, without this hard-won basic



knowledge, we wouldn't have the solid foundation needed to design tomorrow's treatments."

More information: The structure of an infectious immature flavivirus redefines viral architecture and maturation, *Science Advances* (2021). DOI: 10.1126/sciadv.abe4507

Provided by Monash University

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