

MEDIA RELEASE FOR IMMEDIATE RELEASE

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UNVEILING THE MAP TO DESIGNING TREATMENTS FOR DENGUE AND ZIKA VIRUSES

Singapore – Scientists from the Agency for Science, Technology and Research's (A*STAR) Genome Institute of Singapore (GIS), together with scientists from Duke-NUS Medical School and A*STAR's Bioinformatics Institute (BII), have mapped out the structures of four dengue and four Zika viruses. The research focuses on understanding the genetic materials and shapes of these viruses, as well as their corresponding functions. Consequently, understanding their biology enables scientists to design better drugs and treatments.

Mosquito-borne infections are common and deadly in tropical and subtropical regions around the world. Estimates suggest up to 700 million infections and over one million deaths are caused by a variety of mosquito-borne viruses yearly¹. One major class of mosquito-borne viruses are the Flaviviruses², which include the dengue and Zika viruses.

Dengue is a well-known virus which affects more than 390 million people around the world. Singapore is one of the major hotspots of dengue infection³. Every year, 2,000 to 3,000 people are affected by dengue in Singapore⁴. Whilst most of these cases are fairly mild, dengue-associate deaths still occur each year. While the Zika virus has generally limited severity in healthy adults, there has been greater public interest due to the Zika epidemic from 2015 to 2016, and reports of adverse health effects observed in South America⁵.

Using structure probing, the scientists mapped the long genetic materials of four dengue and four Zika virus strains. The research studied the viruses' activities and vital parts to identify portions of the viruses' genetic materials that could be targeted for treatments. Through the identification of the common shapes and structures of the viruses, scientist will be able to design new drug treatments to target these areas.

² Flaviviruses: A family of positive, single-stranded, enveloped RNA viruses found in arthropods, (primarily ticks and mosquitoes), and can occasionally infect humans. (Source: CDC

https://www.cdc.gov/vhf/virus-families/flaviviridae.html)

⁵ Source: CDC <u>https://www.cdc.gov/zika/about/index.html</u>

¹ Source: "Emergency department management of mosquito-borne illness: malaria, dengue, and West Nile virus" <u>https://europepmc.org/abstract/med/25207355</u>

³ Source: "The global distribution and burden of dengue"

https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3651993/ ⁴ Source: The Straits Times <u>https://www.straitstimes.com/singapore/environment/20-more-dengue-</u> cases-in-2018-than-previous-year-207-cases-reported-in-first

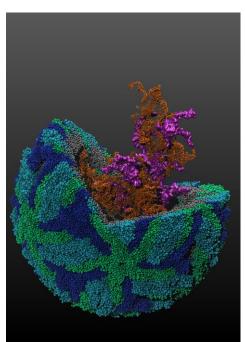
Dr Wan Yue, Principal Investigator at A*STAR's GIS and lead author of the study, said, "One needs to understand the shape and mechanism of a lock, in order to design a key that can open it. Similarly, mapping the shapes and structures of the dengue and Zika viruses will boost understanding of their biology, and contribute significantly towards designing drug treatments targeted at these viruses."

"Moving ahead, additional research needs to be performed to better understand how the viruses' structural mutations reduce their effect. It will also identify the roles that different structures play during their lifecycles," she added.

Dr Roland G. Huber, Assistant Principal Investigator at A*STAR's BII, said, "Understanding the structures of these viruses required us to develop innovative techniques to find synergies in the vast amount of data generated for this study. We developed new tools that will allow us to better understand the architecture of these and other viruses in the future. A strong team of researchers with complementary skills enabled us to succeed."

Professor Ng Huck Hui, Executive Director at A*STAR's GIS, said, "The collaborative study with BII and Duke-NUS unravels significant information on the genetics of the dengue and Zika viruses. This is a crucial step towards designing targeted therapeutics to tackle the diseases."

The study was published in *Nature Communications* on 29 March 2019.



IMAGES

Schematic representation of mature and infectious dengue virus (*Copyright: A*STAR's Genome Institute of Singapore*)

Notes to Editor:

The research findings described in this media release can be found in the scientific journal *Nature Communications*, under the title, "Structure mapping of dengue and Zika viruses reveals functional long-range interactions" by Roland G. Huber^{1*}, Xin Ni Lim^{2*}, Wy Ching Ng^{3*}, Adelene Y. L. Sim^{1*}, Hui Xian Poh², Yang Shen⁴, Su Ying Lim², Karin B. Sundstrom³, Xuyang Sun^{5,6}, Jong Ghut Aw², Horng Khit Too^{7,8}, Peng Hee Boey^{7,8}, Andreas Wilm⁴, Tanu Chawla³, Ming M. Choy⁹, Lu Jiang^{10,11}, Paola Florez de Sessions¹², Xian Jun Loh^{10,11}, Sylvie Alonso^{7,8}, Martin Hibberd⁹, Niranjan Nagarajan⁴, Eng Eong Ooi³, Peter J. Bond^{1†}, October M. Sessions^{3,13,14†}, Yue Wan^{2,15,16†}.

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About A*STAR's Genome Institute of Singapore (GIS)

The Genome Institute of Singapore (GIS) is an institute of the Agency for Science, Technology and Research (A*STAR). It has a global vision that seeks to use genomic sciences to achieve extraordinary improvements in human health and public prosperity. Established in 2000 as a centre for genomic discovery, the GIS will pursue the integration of technology, genetics and biology towards academic, economic and societal impact.

The key research areas at the GIS include Human Genetics, Infectious Diseases, Cancer Therapeutics and Stratified Oncology, Stem Cell and Regenerative Biology, Cancer Stem Cell Biology, Computational and Systems Biology, and Translational Research.

The genomics infrastructure at the GIS is utilised to train new scientific talent, to function as a bridge for academic and industrial research, and to explore scientific questions of high impact.

For more information about GIS, please visit www.a-star.edu.sg/gis.

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As a Science and Technology Organisation, A*STAR bridges the gap between academia and industry. Our research creates economic growth and jobs for Singapore, and enhances lives by contributing to societal benefits such as improving outcomes in healthcare, urban living, and sustainability.

We play a key role in nurturing and developing a diversity of talent and leaders in our Agency and research entities, the wider research community and industry. A*STAR's R&D activities span biomedical sciences and physical sciences and engineering, with research entities primarily located in Biopolis and Fusionopolis. For ongoing news, visit www.a-star.edu.sg.