

Infectious Diseases Labs

ID LABS

## ID LABS SEMINAR SERIES

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## Dr Ashley Lauren St. John

Programme in Emerging Infectious Diseases, Duke-NUS Medical School, Singapore

> Join zoom meeting <u>here</u> Meeting ID: 941 8662 2906 Passcode: 739814

Friday, 18th February 2022 11am to 12pm (SGT)



Webinar is open to all No registration required

## Maternal and fetal immune factors influencing Zika virus congenital syndrome

Zika virus (ZIKV) causes a mild febrile illness in adults, yet is of greatest concern because it can vertically transmit from mother to fetus. The fetuses of infected pregnant women are at risk for Zika congenital syndrome (ZCS), which can involve microcephaly, brain calcifications, absence of brain gyri, and other neurological defects. Emergence of ZIKV in French Polynesia in 2013 and Brazil in 2016 coincided with a sharp increase in cases of microcephaly and the viruses that were isolated were determined to be of a new "Asian" lineage, contrasting the milder "African" lineage strains. Using a mouse model of vertical transmission of ZIKV, we determined that both Asian and African lineage ZIKV strains induce ZCS involving disproportionate microcephaly, but that Asian lineage strains, including an isolate from Singapore, induce more severe neurological defects owing to their preferential targeting of neural progenitor cells and suppressed activation of microglia in the fetal brain. Microglia were determined to be neuroprotective in this model and their depletion led to enhanced infection and neuroinflammation in the fetal brain. Furthermore, we found that in mice and primates, maternal antibodies that are specific for the closely related virus, dengue, and cross-reactive to ZIKV can enhance fetal infection and ZCS severity in a mechanism dependent on ZIKV/immune complex translocation across the placenta and into the fetus. These findings show that both maternal and fetal immune factors can influence ZIKV infection outcomes and disease severity in fetuses.

**Dr. Ashley St. John** is an Associate Professor in the Programme in Emerging Infectious Diseases at Duke-NUS Medical School. She also holds appointments in the Department of Microbiology and Immunology, National University of Singapore, the Department of Pathology, Duke University and the SingHealth Duke-NUS Global Health Institute. Her research program focuses on viral immunology with emphasis on developing novel vaccination strategies, diagnostics, and therapeutics for infectious diseases.