

SIgN Immunology Seminar



Dr Velasco Cimica

Department of Molecular Genetics and Microbiology, State University of New York at Stony Brook, USA

The Andes Virus Nucleocapsid Protein is a Novel Virulence Factor in Innate Immunity **Evasion**

Host Dr Lisa Ng Singapore **Immunology** Network, A*Star

Date Monday 2 June 2014

Time 11am - 12pm

Venue SIgN Seminar Room **Immunos Building** Level 4 **Biopolis**

The Andes virus (ANDV) is the only known hantavirus strain able of person to person transmission, causing hantavirus pulmonary syndrome (HPS) with very high lethality (about 35%). Our studies revealed that ANDV nucleocapsid protein (N) uniquely inhibits the induction of IFN β -, IFN-stimulated response element-, or κ B- promoters mediated by pattern recognition receptors such as RIG-I and MDA5. In addition, ANDV N repressed RIG-I mediated induction of IFNB mRNA. Further analysis demonstrated that ANDV N protein specifically inhibited IFN downstream pathway directed by MAVS, Tank-Binding Kinase 1 (TBK1), and IκB Kinase ε(IKKε). Conversely, ANDV N was unable to signaling by IFN α or TNF α Biochemical studies showed that expression of ANDV N blocks TBK1 mediated IRF3 phosphorylation of serine 396 and TBK1 auto-phosphorylation of serine 172.

In conclusion, we have characterized ANDV N as a novel virulence determinant regulating innate immune signaling. Such findings suggest that ANDV N may contribute for the enhanced ANDV replication and spread between human.