ID No.	K3004
Project Title	Recognition of influenza virus by DNA sensors
Principal Investigator	Akiko Iwasaki (Prof./Yale School of Medicine)
Project Members IMSUT Host Researcher	Takeshi Ichinohe (Associate Prof./IMSUT)
Members	Miyu Moriyama (Postdoctoral Fellow/ Yale School of Medicine) Shota Chino (Graduate Student/IMSUT)

Report

We showed that viroporin activity of influenza virus M2 or encephalomyocarditis virus (EMCV) 2B protein triggers translocation of mtDNA into the cytosol in a MAVS-dependent manner. Although influenza virus-induced cytosolic mtDNA stimulated cGAS- and DDX41-dependent innate immune responses, the nonstructural protein 1 (NS1) of influenza virus associated with mtDNA to evade the STING-dependent antiviral immunity. The STING-dependent antiviral signaling was amplified in neighboring cells through gap junctions. In addition, we found that STING-dependent recognition of influenza virus is essential for limiting virus replication in vivo. These results show a mechanism by which influenza virus stimulates mtDNA release and highlight the importance of DNA sensing pathway in limiting influenza virus replication.