



Fig S1. m⁶A methylome of SARS-CoV-2 genomic RNA.

(a) IGV showing the RIP-seq reads of known m⁶A residues at 4,190 position of 28S rRNA, suggesting good enrichment of RIP-seq.

(b) Correlation analysis of two biological replicates of SARS-CoV-2 RIP-seq data.

(c) Validation of the m⁶A-marked viral RNA (13 m⁶A peaks identified by RIP-seq) by performing m⁶A-IP-qPCR using a different m⁶A antibody (as an orthogonal evidence to the originally used Millipore m⁶A antibody in RIP-seq) in Vero cells at 56 h post infection.

Data are represented as mean \pm SD; N = 3.

(d) m⁶A peak intensity of SARS-CoV-2-infected Vero cells between 24h and 56h post infection. “peak intensity” is calculated as $RPKM_{IP}/RPKM_{Input}$ in each peak.

(e) Immunofluorescence of SARS-CoV-2-infected Huh7 cells (S protein, green) and nuclei (DAPI, blue) at 120h after infection.

(f) Refined RIP-seq of SARS-CoV-2 RNA harvested from Huh7 cells at 120 hpi showing the distribution of m⁶A reads mapped to SARS-CoV-2 genome (red line). The baseline signal of input samples is represented by grey line and m⁶A peaks are represented by green rectangles along the x axis. A schematic diagram of the SARS-CoV-2 genome is shown below to indicate the location of the m⁶A-enriched sequences. Data are representative of N = 2 determinations.