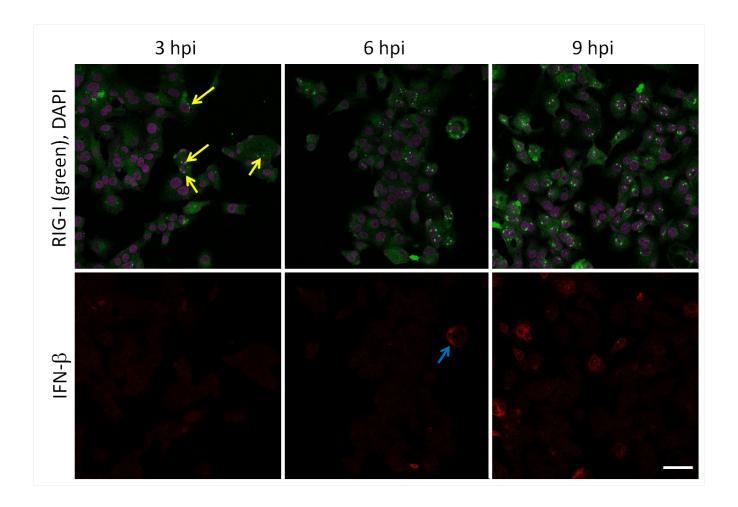


**Fig. S1.** Gaussian fitting of the intensity distribution of single RIG-I transcripts detected in (A) control cells, (B) IFN- $\beta$  treated (6 hours) cells.



**Fig. S2.** Images of *RIG-I* and *IFNB1* mRNA in SeV infected (3, 6, and 9 hours) HepG2 cells obtained with a low magnification objective (40x). Yellow arrows show few RIG-I overexpressing cells with bright RIG-I transcription sites. Blue arrow shows an IFN-b overexpressing cell.

Table S1 N gene **ADAR** APOBEC3A4 C6orf150 DDIT4 **HPSE** IFNA1 IFNA4 IRF1 IRF2 IRF3 IRF7 IRF9 MAP3K14 Mov10 NLRC5 NLRX1 **RNAseL** STAT1 TLR3 **TRAIL Table S1.** Number of transcripts detected by Nanostring nCounter gene expression system in control, IFN-β treated (2 hours), and SeV infected (2, 3, 6, and 9 hours) HepG2 cells. Number of transcripts is normalized to housekeeping gene GAPDH.

Gene

Control

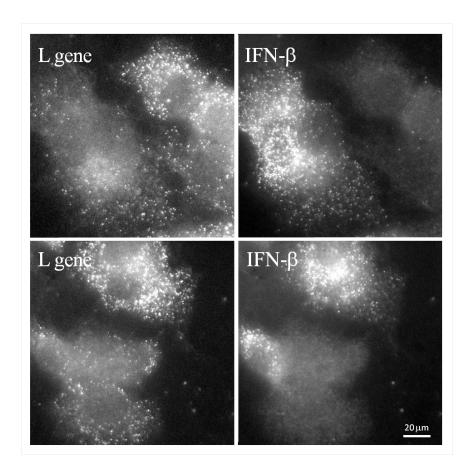
IFN-β st.

2 hpi

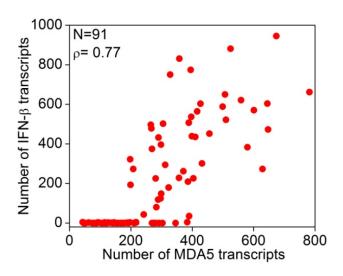
3 hpi

6 hpi

9 hpi



**Fig. S3**. Images of L gene and *IFNB1* mRNA in SeV infected (9 hours) HepG2 cells. Each row represents a different imaging area.



**Fig. S4** Scatter plots of number of MDA5 against *IFNB1* transcripts in single HepG2 cells at 9 hours post SeV infection. N: Number of cells analyzed, r: Pearson's correlation coefficient.