Figure EV1. The IFN response is delayed during ZIKV infection.

A549 cells were infected with ZIKV (MOI = 2) for 18, 24 and 48 h. At each time point, relative levels of Ifnb and Ifit1 mRNA (compared to mock-infected cells) were determined by qRT–PCR (upper panel). As a positive control for IFN induction, cells were transfected with 2 μg of poly(I:C) for 12 h (lower panel). Levels of Ifnb and Ifit1 mRNA (compared to untreated cells) are shown. Values are expressed as mean ± standard error. N = 3.
ZIKV NS5 reduces NF-κB reporter activity.

HEK293T cells were transfected with plasmids encoding different ZIKV proteins. Sixteen hours later, they were transfected with an NF-κB promoter-driven firefly luciferase plasmid (pNF-κB-luciferase) and a constitutively active Renilla luciferase construct (pRL-TK), as well as 1 μg of poly(I:C). Twenty-four hours later, cell lysates were harvested and subjected to luciferase assay. C, capsid; E, envelope. Values are expressed as mean ± standard error. *P < 0.05, **P < 0.01 (Student’s t-test), N = 3.
**Figure EV3.** ZIKV infection does not affect STAT1 levels.

A  A549 cells or mouse embryonic fibroblasts (MEFs) were infected with ZIKV (MOI = 2) for 24 and 48 h after which IFN-α (100 U/ml) was added for 2 h. Cells were then fixed and processed for indirect immunofluorescence. Representative panels are shown. The experiment was repeated two times.

B  A549 cells were infected with ZIKV (MOI = 5) for 24 h and then treated with DMSO or bafilomycin-A1 (Baf-A1) (400 nM) for 12 h. Cell lysates were processed by SDS–PAGE and immunoblotting. Representative panels are shown. The experiment was repeated three times.
Figure EV4. Expression of ZIKV NS5 induces degradation of STAT2.
A549 cells were transfected with control plasmid pcDNA 3.1 or plasmids encoding FLAG-tagged ZIKV proteins. At 48 h, post-transfection cells were treated with IFN-α (100 U/ml) for 2 h and then processed for indirect immunofluorescence. Images were acquired on a spinning disk confocal microscope with a 40× objective. NS5-positive cells are indicated by dashed white circles. Representative panels are shown. The experiment was repeated three times.