Review of: "Classical swine fever virus NS5A inhibits NF-kB signaling by targeting NEMO"

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Potential competing interests: The author(s) declared that no potential competing interests exist.

The manuscript entitled "*Classical swine fever virus NS5A inhibits NF-κB signaling by targeting NEMO*" by Li and colleagues described an observation that the CSFV NS5A protein interacts NEMO. Mechanistically, NS5A mediated proteasomal degradation of NEMO by inducing K27-linked polyubiquitination of NEMO. Together, these finding proved that NS5A inhibits NF-κB signaling by facilitating NEMO degradation, which is a novel mechanism that CSFV inhibits host innate immunity. Overall, the idea is clear and concision and the manuscript is well written. The experiments were well-performed and appropriate control was included.

There are a number of concerns with this study which include the following:

- The title and the conclusion of this study is that CSFV NS5A inhibits NF-κB signaling. However, little experiment was performed to measure the inhibition of NF-κB pathway, for instance the translocation of NF-κB(p65). In addition, using gene expression of IFNa to indicate NF-κB signaling activation or inhibition seems not enough. More experiments were need to demonstrate NS5A inhibits NF-κB signaling.
- 2. Fig 5B, different MOI of CSFV should be used to infect PK-15 cell and detect the degradation of NEMO.
- 3. The effect of NF-κB against CSFV infection should be measured. How about the viral load in NS5A overexpressed cells?
- 4. In some figure legend, "Evidence that..." is not necessary and it is better to delete it.