

Supplementary data. Non-specific trans cleavage of the SV4 (NS4-NS5) peptide in the presence of the reticulocyte lysates. The peptide SV4 (NH<sub>2</sub>-TRGSRTEEAKGKTKH-COOH, 30  $\mu$ M) was incubated with purified NS6 or NS6-7 or their respective active site mutants  $\mu$ NS6 and  $\mu$ NS6-7 at 30 °C for 17 h in the presence or absence of reticulocyte lysates, as indicated. Cleavage products are monitored by reversed-phase HPLC as described in Materials and Methods. **A, B** and **C**, incubation of peptide SV4 with NS6, NS6 in the presence of reticulocyte lysates, and  $\mu$ NS6 in the presence of reticulocyte lysates, respectively. The peptide is completely cleaved in the presence of reticulocyte lysates independently of the presence of reticulocyte lysates, and  $\mu$ NS6-7 in the presence of reticulocyte lysates, and  $\mu$ NS6-7 in the presence of reticulocyte lysates, respectively. The peptide is completely cleaved in the presence of reticulocyte lysates independently of the presence of the active protease NS6-7 or its active site mutant  $\mu$ NS6-7. RL, reticulocyte lysates.