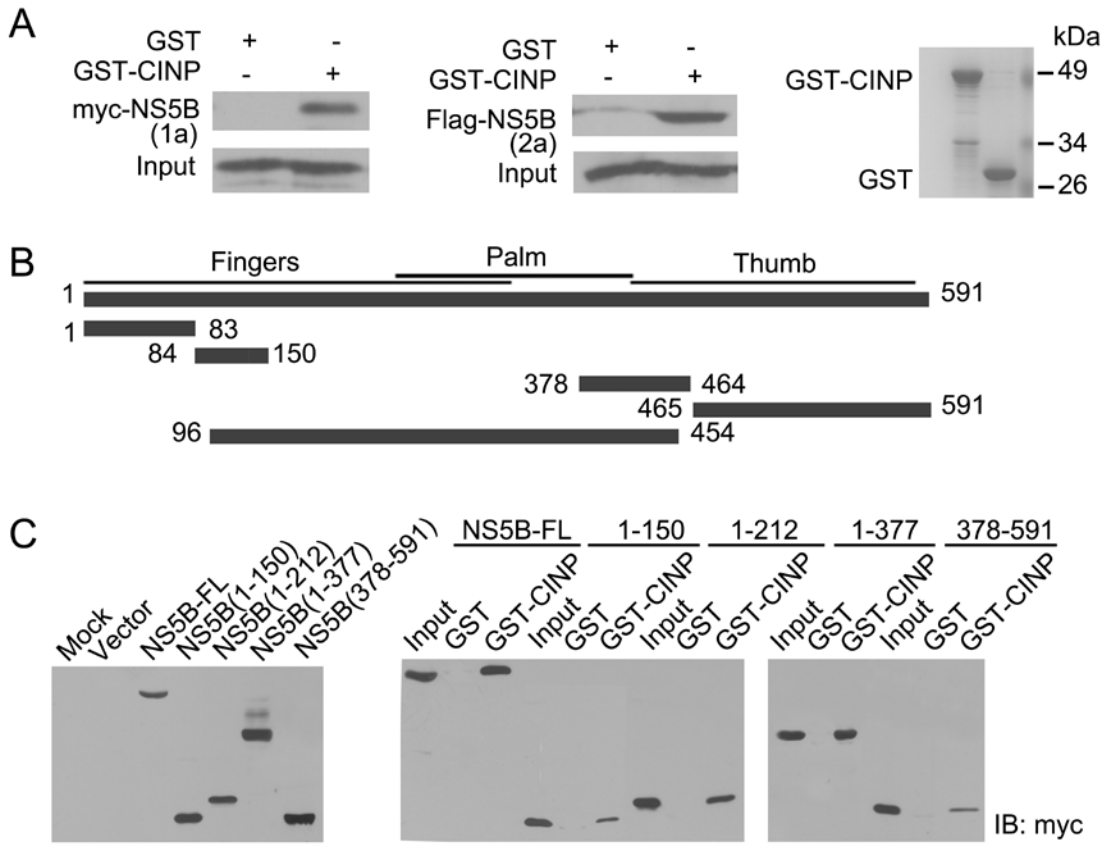
SUPPLEMENTAL DATA

Fig. S1. The physical association of CINP with NS5B. *A,* GST pull-down assay to identify the CINP/NS5B (1a and 2a subtype) interaction. 293T cells were grown in 6 cm dishes and transfected with the pCMV/myc-NS5B (genotype 1a from H77) or pCDNA 3.1/myc-His-3×Flag-NS5B (genotype 2a from JFH-1) plasmids. Forty-eight hours later, the cells were harvested as described in the Materials and methods section, and the supernatant was divided into two equal parts and incubated with GST or GST-CINP beads for 2 h. Protein-bound Glutathione-Sepharose 4B beads were washed and boiled for 5 min, and then detected by western blot using anti-myc or anti-Flag antibody. Coomassie-stained GST and GST-CINP are shown. *B,* A schematic diagram of the NS5B deletion mutants used in this study. *C,* Full-length or deleted myc-tagged NS5B was transiently transfected into 293T cells and detected by western blot (left panel). Proteins pulled down by GST-CINP were examined by western blot with an anti-myc antibody (right panel).

Fig. S2. **RNA polymerase activities of the deletion mutants**. The RdRp activities were monitored by *in vitro de novo* transcription assay. Briefly, 0.1 mM purified HCV RdRps were pre-incubated for 30 min at 29℃, and then incubated with 50 mM Tris/HCl (pH 8.0), 200 mM monopotassium glutamate, 3.5 mM MnCl2, 1mM DTT, 0.5 mM GTP, 50 mM ATP, 50 mM CTP, 5 mM [α -^{32}P]UTP, 0.02 mM RNA template (SL12-1S) and 100 U/ml human placental RNase inhibitor at 29℃ for 90 min. [^{32}P]-RNA products (HCV RdRp activity) were resolved by SDS-PAGE (6% gel, 8 M urea) and subjected to autoradiography.

Fig. S3. NS5B has no effect on CINP protein level. *A,* CINP expression was compared in Huh7 cells and the HCV subgenomic replicon. *B,* CINP protein level in Huh7.5 cells was examined after mock infection or infection with different MOI (0.1 and 0.5). *C* and *D,* The effect of NS5B on CINP was examined in HepG2 Tet-on NS5B stable cells. (C) Different time points after induction by 1 μg/ml Dox and (D) 48 hours after induction with different Dox concentrations are presented.

Figure. S1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GST
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Input
Flag-NS5B (2a)
kDa
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34
26
B
Fingers
Palm
Thumb
1
591
83
84
150
378
464
465
96
454
C
Mock
Vector
NS5B-FL
NS5B(1-150)
NS5B(1-212)
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1-377
378-591
Input
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Figure. S2

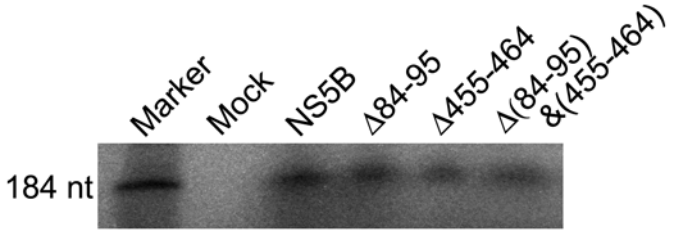
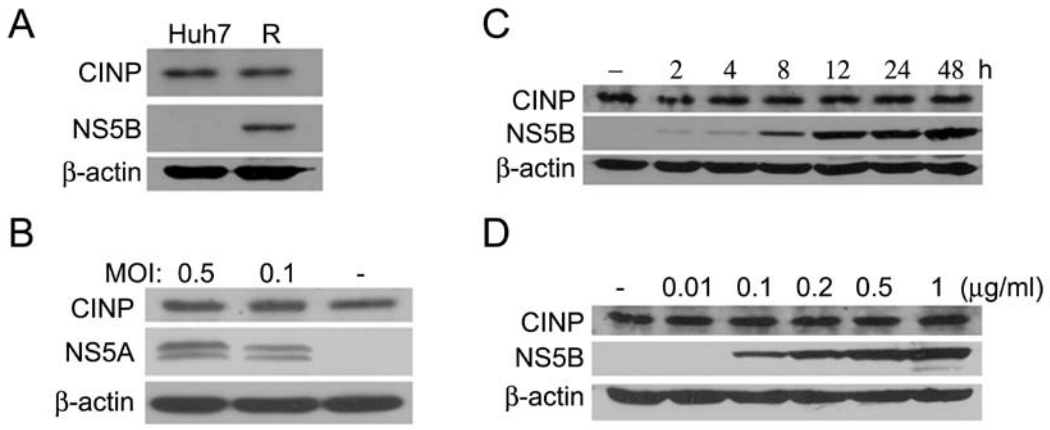


Figure. S3


A
Huh7
R
CINP
NS5B
β-actin
B
MOI: 0.5
0.1
-
CINP
NS5A
β-actin
C
–
2
4
8
12
24
48 h
CINP
NS5B
β-actin
D
-
0.01
0.1
0.2
0.5
1 (μg/ml)
CINP
NS5B
β-actin