

Supplementary Table 1. sgRNAs used for CRISPR/Cas9 depletion in human hepatocytes.

Target-sg#	Target sequence
Control (non-targeting “Scramble” sgRNA)	GCACTCACATCGCTACATCA
<i>NLRX1</i> -sg#2 (T2)	GCGAAGTAGTCATCAG
<i>NLRX1</i> -sg#3 (T3)	GATACGGCCAATGGCAG
<i>MAVS</i> -sg#5	TATAAGTCCGAGGGCACCTT
<i>IRF1</i> -sg#2	GAGCCAGATCCCAAGACG
<i>IRF3</i> -sg#3	AGTATTCTCCAGGGAGG
<i>RELA</i> -sg#1	CAATGATCTCCACATAG
<i>RELA</i> -sg#3	TCCAGTGTGTGAAGAAG
<i>PKR</i> -sg#1	GATGGAAGAGAATTTCCAGA
<i>PKR</i> -sg#3	TGGTACAGGTTCTACTAAAC

Supplementary Table 2. Oligonucleotide primers

Primer/Purpose	5'-Sequence-3'
pLOC-ClaXbaI/ Removal of TurboGFP-2A	F:ATCGATTCTAGAATGAAGACCTTCAACATCTCTCAGCAG GATCTGGAG R:ATCGATTCTAGAGGTATTATCGTGTTTTTCAAAGGAAAA CCACGTCCC
pLOC-NLRX1/ Cloning human NLRX1	F:TTCTGCAGATATCACAAAGTTTGTACAAAAAAGTTGGCAT GAGGTGGGGCCACCATT R:TTAGCTAGCAACCACTTTGTACAAGAAAGTTGGGTATCA GCTTCCAGAGCTTCC
X1T3-TarMut/ Mutating and sequencing NLRX1-T3 Underline labeling represents same sense mutation (ATT→AT <u>C</u> , GGC→GG <u>T</u>) to prevent Cas9 editing of reconstituted NLRX1	F:GACCACTCGGCCCTCTGCCAT <u>CGG</u> TCGTATCCCCAGCAA GTACGTGG R:CCACGTACTTGCTGGGGATACG <u>ACC</u> GATGGCAGAGGGCC GAGTGGTC SEQ:CCTGATGGCTGCTGCTG
Bsd-NLRX1/ Switching selection marker in sgRNA vector to blasticidin	F:GCTGGAGATGTTGAGAGCAACCCAGGTCCCATGAAAACC TTCAACATCTCTCAG R:GTAATCCAGAGGTTGATTGTCGACGTTTAAACTCAGTTC CTGGTGTACTTGAG
hsIL6-3UTR/ Inserting IL6-3'UTR into psiCHECK2	F:CGTGGAGCGCGTGCTGAAGAACGAGCAGTAACATGGGC ACCT R:CGAATTC ³ CGGGCTCGAGCGATCGCTAGAAGGTATAAA AACCATT SEQ:GTACATCAAGAGCTTCGTGG
<i>IFNB1</i> / q-PCR of human <i>IFNB1</i> cDNA	F:CATTACCTGAAGGCCAAGGA R:CAATTGTCCAGTCCCAGAGG
<i>IFNL1</i> / q-PCR of human <i>IFNL1</i> cDNA	F:CGCCTTGAAGAGTCACTCA R:GAAGCCTCAGGTCCCAATTC
<i>IL1B</i> / q-PCR of human <i>IL1B</i> cDNA	F:GAAGCTGATGGCCCTAAACAG R:AGCATCTTCTCAGCTTGTC
<i>IL6</i> / q-PCR of human <i>IL6</i> cDNA	F:CGGGAACGAAAGAGAAGCTCTA R:GGCGCTTGTGGAGAAGGAG
<i>IRF1</i> / q-PCR of human <i>IRF1</i> cDNA	F:AGCTCAGCTGTGCGAGTGTA R:CATGACTTCTCTTGGCCTT
<i>IRF3</i> / q-PCR of human <i>IRF3</i> cDNA	F:ACCAGCCGTGGACCAAGAG R:TACCAAGGCCCTGAGGCAC
<i>ACTB</i> / q-PCR of human <i>ACTB</i> cDNA	F:GACCCAGATCATGTTTGAGACC R:GTCACCGGAGTCCATCACGA
HAV/ q-PCR of HAV genomic cDNA	F:GGTAGGCTACGGGTGAAAC R:AACAAC ³ TACCAATATCCGC
HCV/ Reverse transcription of HCV genomic RNA and q-PCR of HCV cDNA	RT:GGCCAGTATCAGCACTCTCTGCAGTC F:CATGGCGTTAGTATGAGTGTCGT R:CCCTATCAGGCAGTACCACAA
<i>Tnf</i> / q-PCR of mouse <i>Tnf</i> cDNA	F:CATCTTCTCAA ³ AATTCGAGTGACAA R:TGGGAGTAGACAAGGTACAACCC
<i>Ifnb</i> , <i>Il6</i> , <i>Actb</i> / q-PCR of mouse <i>Ifnb</i> , <i>Il6</i> , <i>Actb</i> cDNAs	Primers were described previously ⁸ .