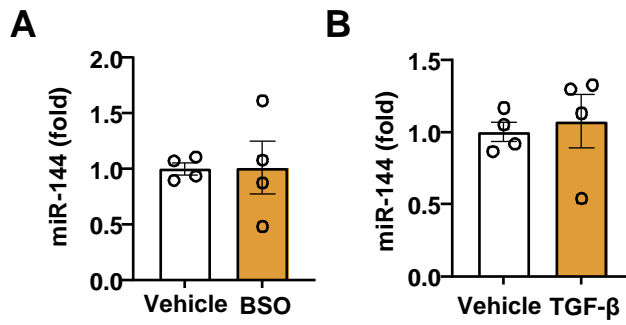
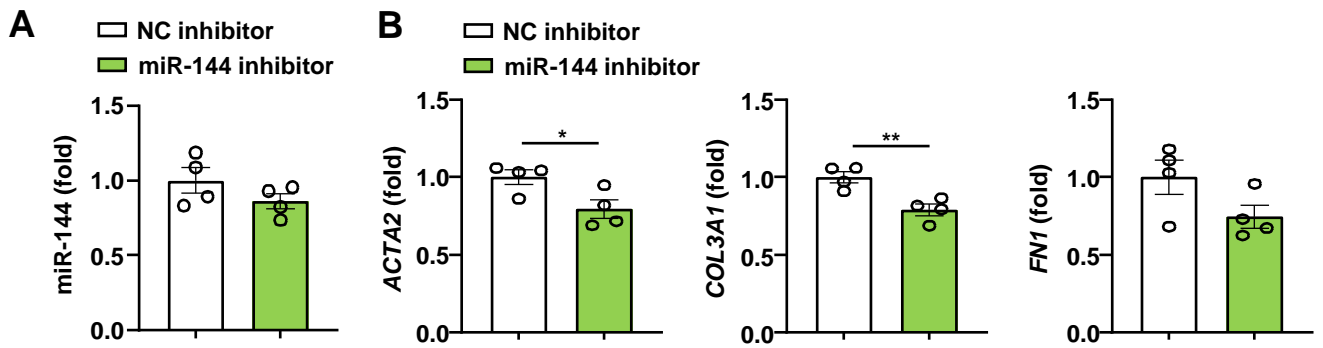


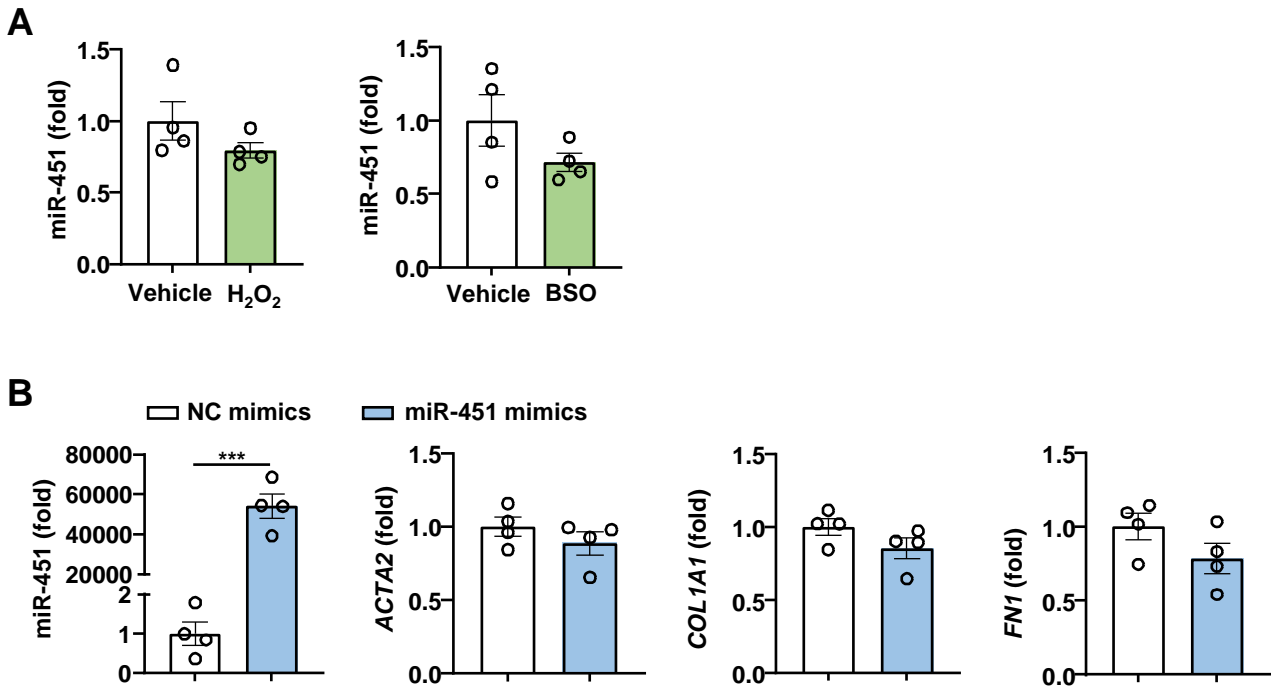
Supporting Figure S1. Hepatic oxidative stress was elevated in the liver of mice with CCl₄ and BDL-induced liver fibrosis. (A, C) C57BL/6J mice were injected intraperitoneally with 2 ml/kg weight of 10% CCl₄ in olive oil twice a week for 6 weeks and were sacrificed 48 hours after the last injection of CCl₄. (A) Hepatic H₂O₂ content was measured. (B) C57BL/6J mice were injected intraperitoneally with 2 ml/kg weight of 10% CCl₄ for 24 hours. Hepatic H₂O₂ content was measured. (C) RT-qPCR analyses of several oxidant related genes. (D-E) C57BL/6J mice were performed BDL or sham operation for 14 days. (D) Hepatic H₂O₂ content was measured. (E) The expression of *Ho-1* and *Nqo-1* were analyzed by RT-qPCR analyses. **P* < 0.05, ***P* < 0.01.



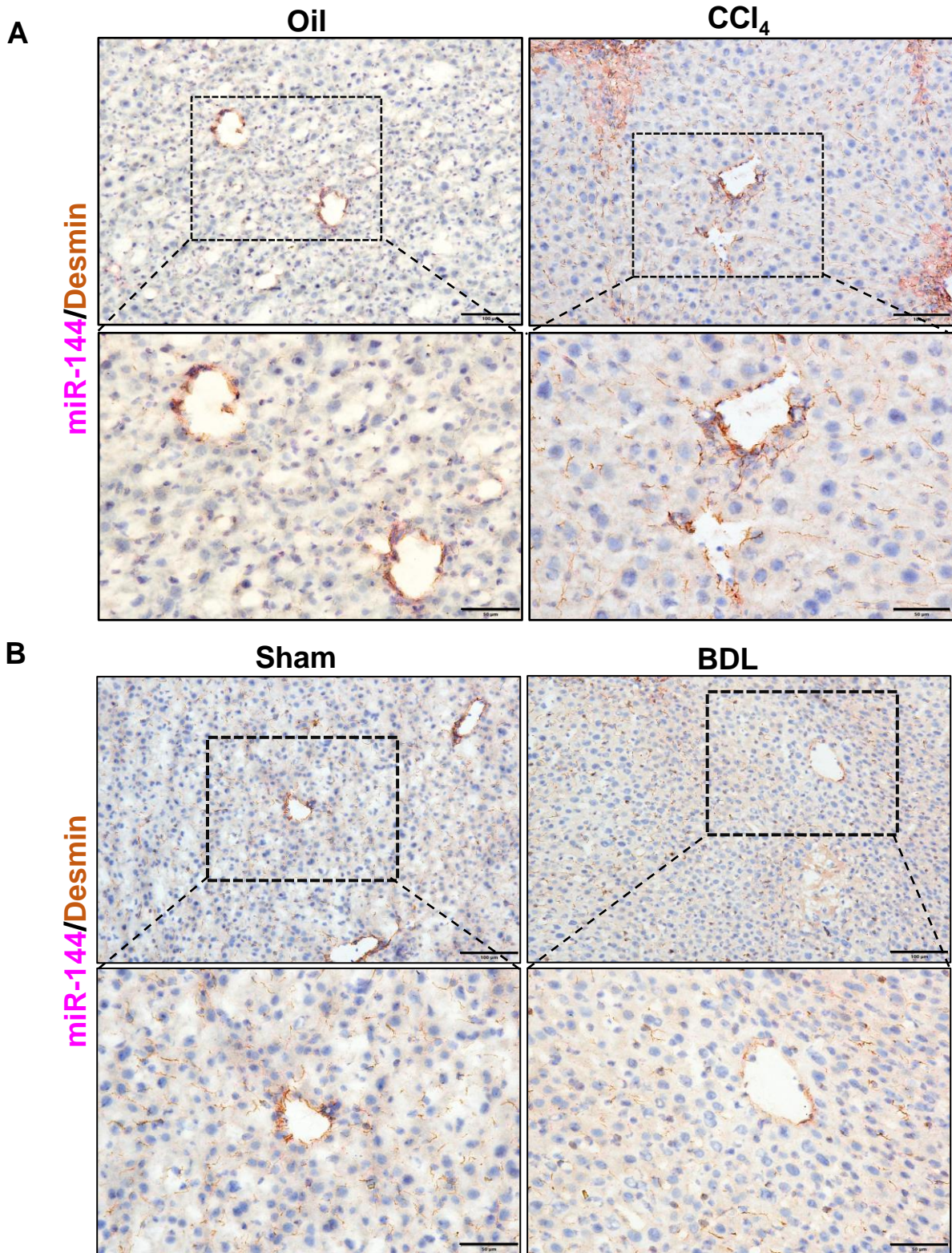
Supporting Figure S2. BSO and TGF- β did not elevated miR-144 in HSCs. (A) LX-2 cell lines were administered with BSO (500 μ M). The expression of miR-144 were analyzed by RT-qPCR analyses. (B) LX-2 cell lines were administered with TGF- β (10ng/ml). The expression of miR-144 were analyzed by RT-qPCR analyses.



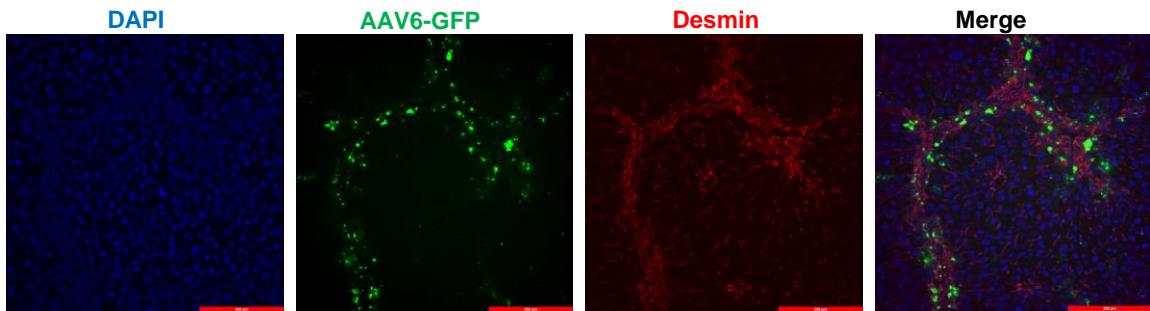
Supporting Figure S3. Blockage of miR-144 in LX-2 cells inhibits HSC activation. LX-2 cell lines were transfected with miR-144 inhibitor and NC inhibitor (100nM) for 48 hours. (A) The expression of miR-144 were analyzed by RT-qPCR analyses. (B) The expression of *ACTA2*, *COL3A1* and *FN1* were analyzed by RT-qPCR analyses. * $P < 0.05$, ** $P < 0.01$.



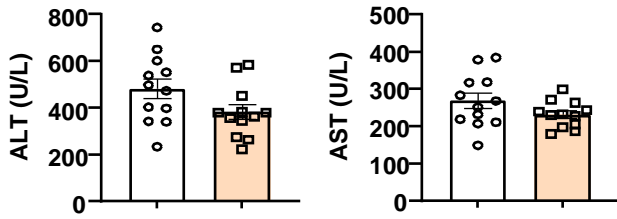
Supporting Figure S4. miR-451 is not involved in HSC activation. (A) LX-2 cell lines were administered with BSO (500 μ M) or H_2O_2 (200 μ M) for 3 hours. The expression of miR-451 were analyzed by RT-qPCR analyses. (B) LX-2 cell lines were transfected with miR-451 mimics and NC mimics (20nM) for 48 hours. The expression of miR-451, *ACTA2*, *COL1A1* and *FN1* were analyzed by RT-qPCR analyses. *** $P < 0.001$.



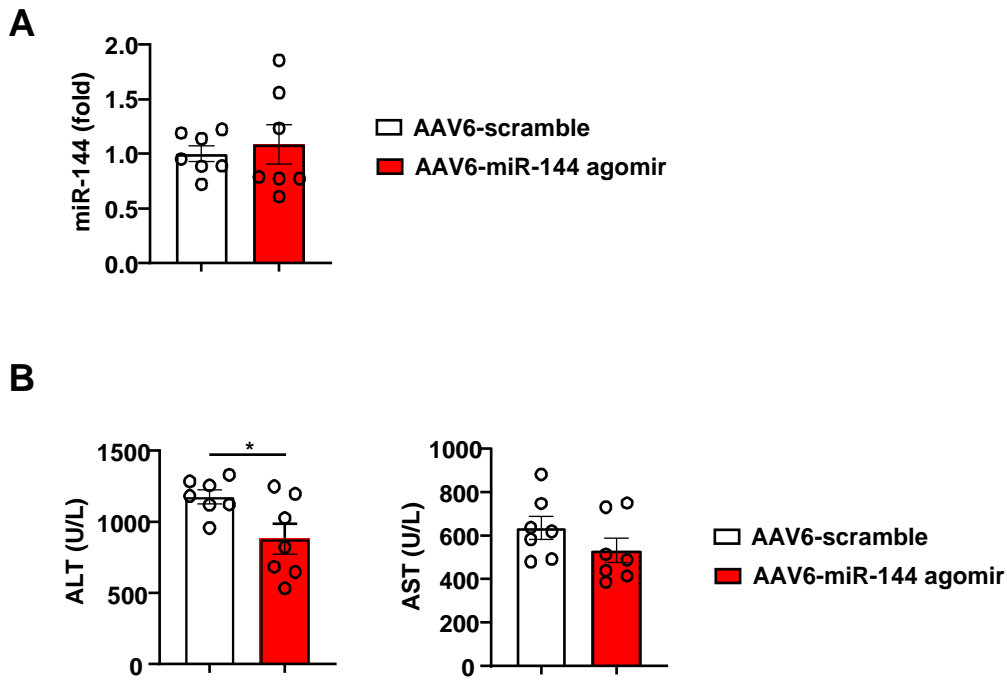
Supporting Figure S5. miR-144 is not elevated in non-fibrotic regions in liver fibrosis mouse models. C57BL/6J mice were injected intraperitoneally with 2 ml/kg weight of 10% CCl₄ in olive oil twice a week for 6 weeks or performed BDL operation for 2 weeks. Liver tissue samples were collected. Frozen liver tissue sections were subjected to miR-144 in situ hybridization with RNAscope probe. (A) Representative images of miR-144 expression in non-fibrotic regions in CCl₄ mouse model are shown. Scale bars: 100 μm or 50 μm. (B) Representative images of miR-144 expression in non-fibrotic regions in BDL model are shown. Scale bars: 100 μm or 50 μm.



Supporting Figure S6. AAV6 mainly infects HSCs. C57BL/6J mice were injected intraperitoneally with 2 ml/kg weight of 10% CCl₄ in olive oil twice a week for 2 weeks before administering with AAV6-GFP. After AAV6 injection, another 8 doses of CCl₄ were administered to the mice, which were sacrificed 48 hours after the last injection of CCl₄. (A) Representative images of GFP (green), Desmin (red), and nuclei (blue) are shown. Scale bars: 200 μ m.

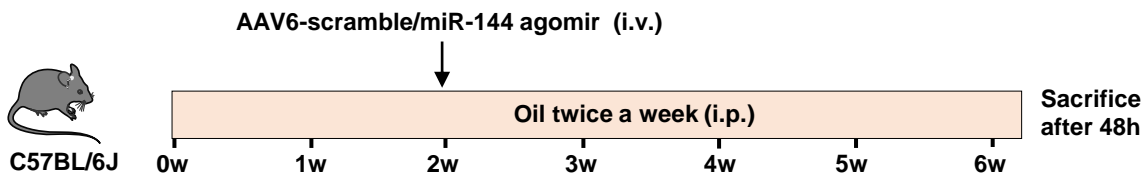


Supporting Figure S7. Blockade of miR-144 in hepatic stellate cells did not affect liver injury. C57BL/6J mice were administrated with AAV6-miR-144 antagomir. Serum ALT and AST levels were measured.

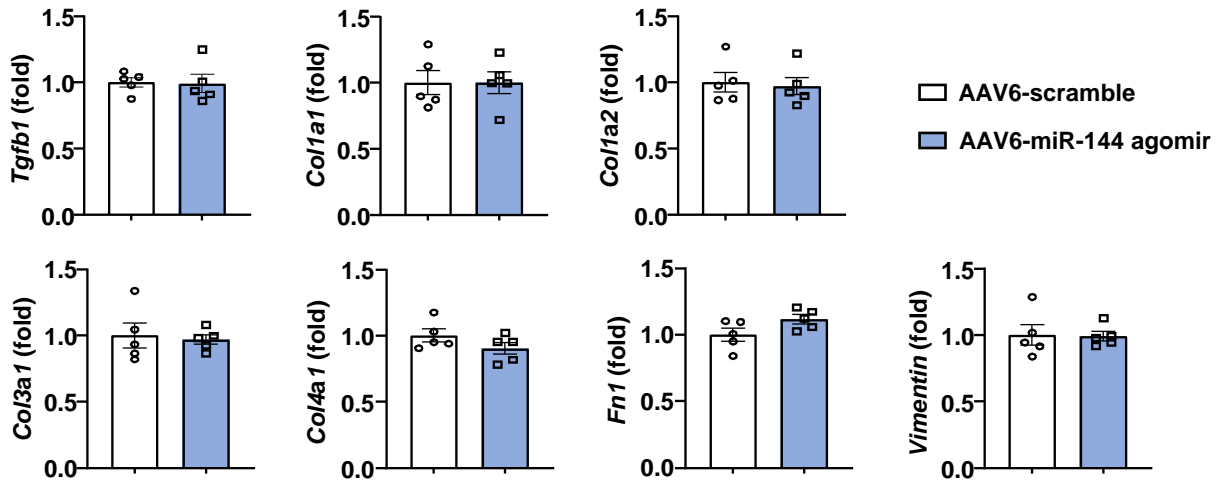


Supporting Figure S8. The liver injury after overexpression of miR-144 in HSCs. (A) The expression of miR-144 in liver were analyzed by RT-qPCR analyses. (B) Serum ALT and AST levels were measured. * $P < 0.05$.

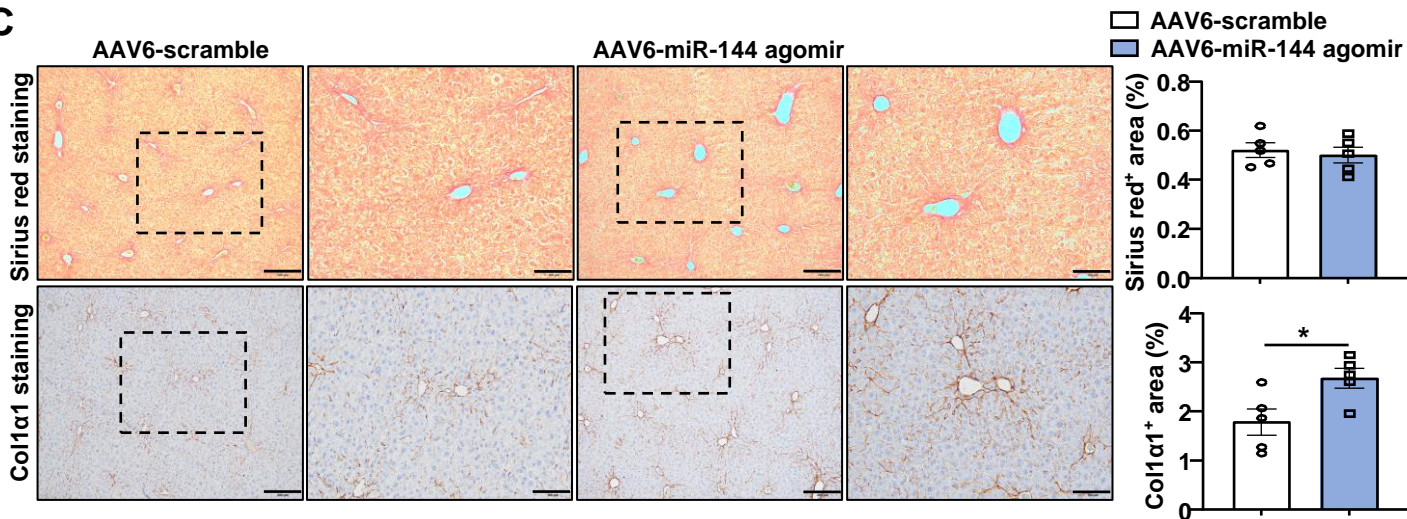
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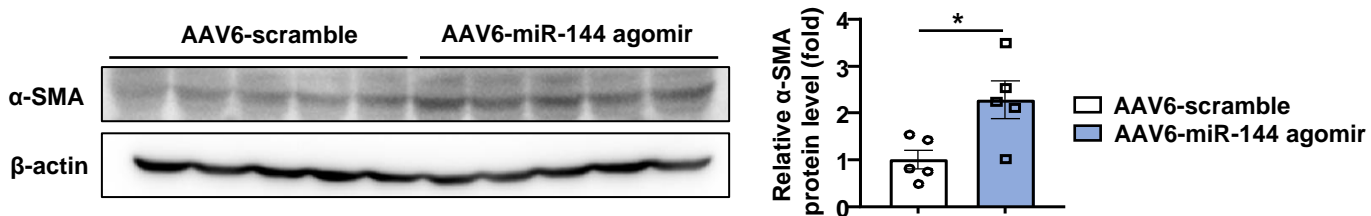
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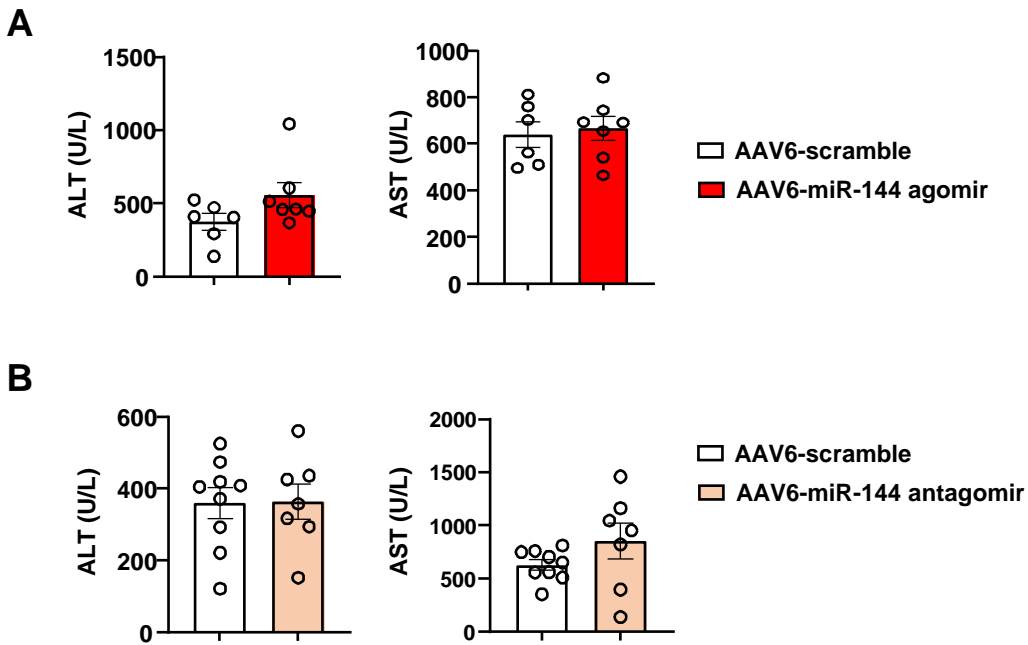
C



D



Supporting Figure S9. Overexpression of miR-144 in HSCs barely influences the development of liver fibrosis. (A) Schematic overview of the experimental design. Liver tissue samples were collected. (B) RT-qPCR analyses of several hepatic fibrogenesis genes. (C) Liver tissues were subjected to Sirius red staining (Scale bars: 200 μ m or 100 μ m) and Col1 α 1 staining (Scale bars: 200 μ m or 100 μ m). (D) α -SMA expression was analyzed by Western blot. Values represent means \pm SEM (n=5 per group).



Supporting Figure S10. Overexpression or inhibition of miR-144 in hepatic stellate cells did not affect liver injury. (A-B) C57BL/6J mice were administrated with AAV6-miR-144 agomir/antagomir after bile duct ligation. Then the mice were sacrificed 17 days later and their serum was collected. Serum ALT and AST levels were measured.

Supporting Table S1. The list general characteristics of the liver fibrosis human samples.

Patient #	Gender	Age	Primary Diagnosis	ALT (IU/L)	AST (IU/L)
1	F	63	Liver fibrosis, autoimmune hepatitis	60	73
2	F	58	Cirrhosis, CHB, HCC	109	371
3	M	49	Cirrhosis, CHB, HCC	28	61
4	F	30	Liver fibrosis, autoimmune hepatitis	11	19
5	M	61	Cirrhosis, HCC	33	119

CHB: chronic hepatitis B; HCC: hepatocellular carcinoma

Supporting table S2: RT-qPCR primer sequences

Genes (mouse)	Forward primer (5'-3')	Reverse primer (5'-3')
<i>18s</i>	AACTTTCGATGGTAGTCGCCGT	TCCTTGGATGTGGTAGCCGTTT
<i>Acta2</i>	TCCTGACGCTGAAGTATCCGATA	GGTGCCAGATCTTTTCCATGTC
<i>Col1a1</i>	TAGGCCATTGTGTATGCAGC	ACATGTTCCAGCTTTGTGGACC
<i>Col1a2</i>	GGTGAGCCTGGTCAAACGG	ACTGTGTCCTTTCCAGCCTTT
<i>Col3a1</i>	TAGGACTGACCAAGGTGGCT	GGAACCTGGTTTCTTCTCACC
<i>Col4a1</i>	CACATTTTCCACAGCCAGAG	GTCTGGCTTCTGCTGCTCTT
<i>Fn1</i>	TTCAAGTGTGATCCCCATGAAG	CAGGTCTACGGCAGTTGTCA
<i>Vimentin</i>	TCCACACGCACCTACAGTCT	CCGAGGACCGGGTCACATA
<i>Tgfb1</i>	CAACCCAGGTCCTTCTCTAAA	GGAGAGCCCTGGATACCAAC
<i>Sin3a</i>	AGCCGAGTGTCCAGCTATT	TCTGCACCTCAATTTTGTAGCC
<i>Dusp1</i>	GTTGTTGGATTGTCGCTCCTT	TTGGGCACGATATGCTCCAG
<i>Hgf</i>	ACTTCTGCCGGTCCTGTTG	CCCCTGTTCTGATACACCT
<i>Gsk3b</i>	TGGCAGCAAGGTAACCACAG	CGGTTCTTAAATCGCTTGTCTG
<i>Foxo1</i>	GGGTCCCACAGCAACGATG	CACCAGGGAATGCACGTCC
<i>Cav2</i>	TCACCAGCTCAACTCTCATCT	GCCAGAAATACGGTCAGGAACT
<i>Pten</i>	TGGATTGACTTAGACTTGACCT	GCGGTGTCATAATGTCTCTCAG
<i>Tgif2</i>	ATGTCGGACAGCGATCTAGG	TCCCGGAGGATCTTTACTGAC
<i>Ho-1</i>	AGGTACACATCCAAGCCGAGA	CATCACCAGCTTAAAGCCTTCT
<i>Nqo-1</i>	AGGATGGGAGGTAICTCGAATC	AGGCGTCCTTCTTATATGCTA

Genes (human)	Forward primer (5'-3')	Reverse primer (5'-3')
<i>18s</i>	GGCCCTGTAATTGGAATGAGTC	CCAAGATCCAACCTACGAGCTT
<i>ACTA2</i>	GTGACGAAGCACAGAGCAAA	CTTTTCCATGTGCTCCAGT
<i>COL1A1</i>	CAGATCACGTCATCGCACAA	TGTGAGGCCACGCATGAG
<i>COL3A1</i>	AGGACTGACCAAGATGGGAA	AGGGGAGCTGGCTACTTCTC
<i>COL4A1</i>	CCTTTTGTCCCTTCACTCCA	CTCCACGAGGAGCACAGC
<i>FN1</i>	CGGTGGCTGTCAGTCAAAG	AAACCTCGGCTTCTCCATAA
<i>TGFB1</i>	CAATTCCTGGCGATACCTCAG	GCACAACCTCCGGTGACATCAA
<i>SIN3A</i>	GGTGGAGGATGCGCTATCTTA	GGGTGTCGATGCTCTGAGATTT

Supporting table S3: miRNA primer sequences

miRNA	RT primer	qPCR primer
Sno202	GTCGTATCCAGTGCAGGGTCCGAGGTAT TCGCACTGGATACGACCATCAG	AAGTACTTTTGAACCCTT TTCCAT
RUN44	GTCGTATCCAGTGCAGGGTCCGAGGTAT TCGCACTGGATACGACAGTCAG	GAACATGAAGGTCTTAAT TAGCTCTAA
miR-144	GTCGTATCCAGTGCAGGGTCCGAGGTAT TCGCACTGGATACGACAGTACA	GCGCGCGTACAGTATAGA TGA
miR-451a	GTCGTATCCAGTGCAGGGTCCGAGGTAT TCGCACTGGATACGACAACCTCA	CGCGAAACCGTTACCATT AC