

Supporting Information

(Replaces the original version published on 20 Dec 2021 – Fig. S2 and
Fig. S3 updated)

Clodronate-nintedanib-loaded exosome-liposome hybridization enhances liver fibrosis therapy by inhibiting Kupffer cell activity

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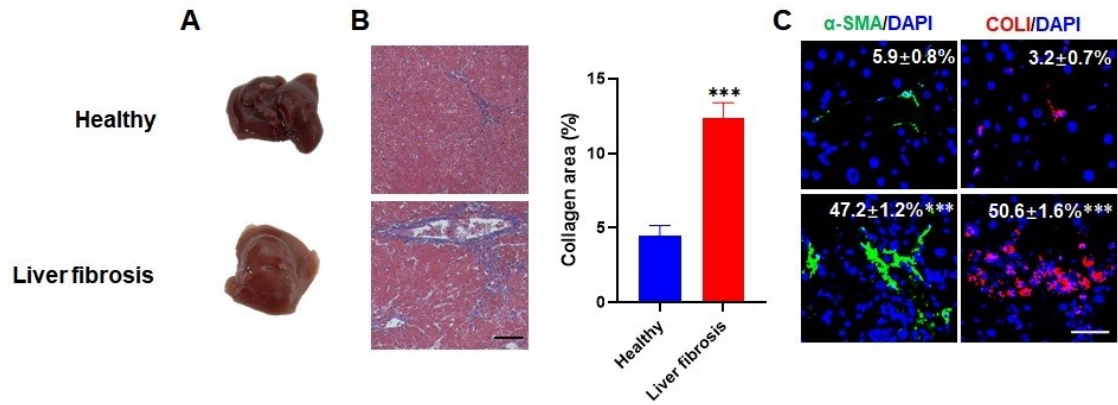


Fig. S1 Liver fibrosis model was successfully established in mice. (A) Representative photos of livers of healthy mice and in mice with liver fibrosis. (B) Masson-stained of liver sections and quantification of the presence of collagen. $*P < 0.05$, $**P < 0.01$, $***P < 0.001$. (C) Immunofluorescent staining of α -SMA and collagen I. Data are mean \pm SD, $n = 3$; $*P < 0.05$, $**P < 0.01$, $***P < 0.001$, compared to healthy mice. Scale bar = 50 μ m.

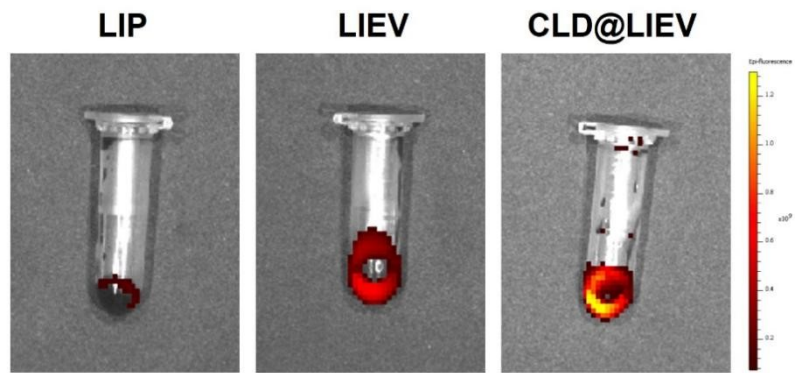


Fig. S2 Fluorescence images of blood samples 4 h after intravenous injection of DiD labeled liposomes in healthy mice.

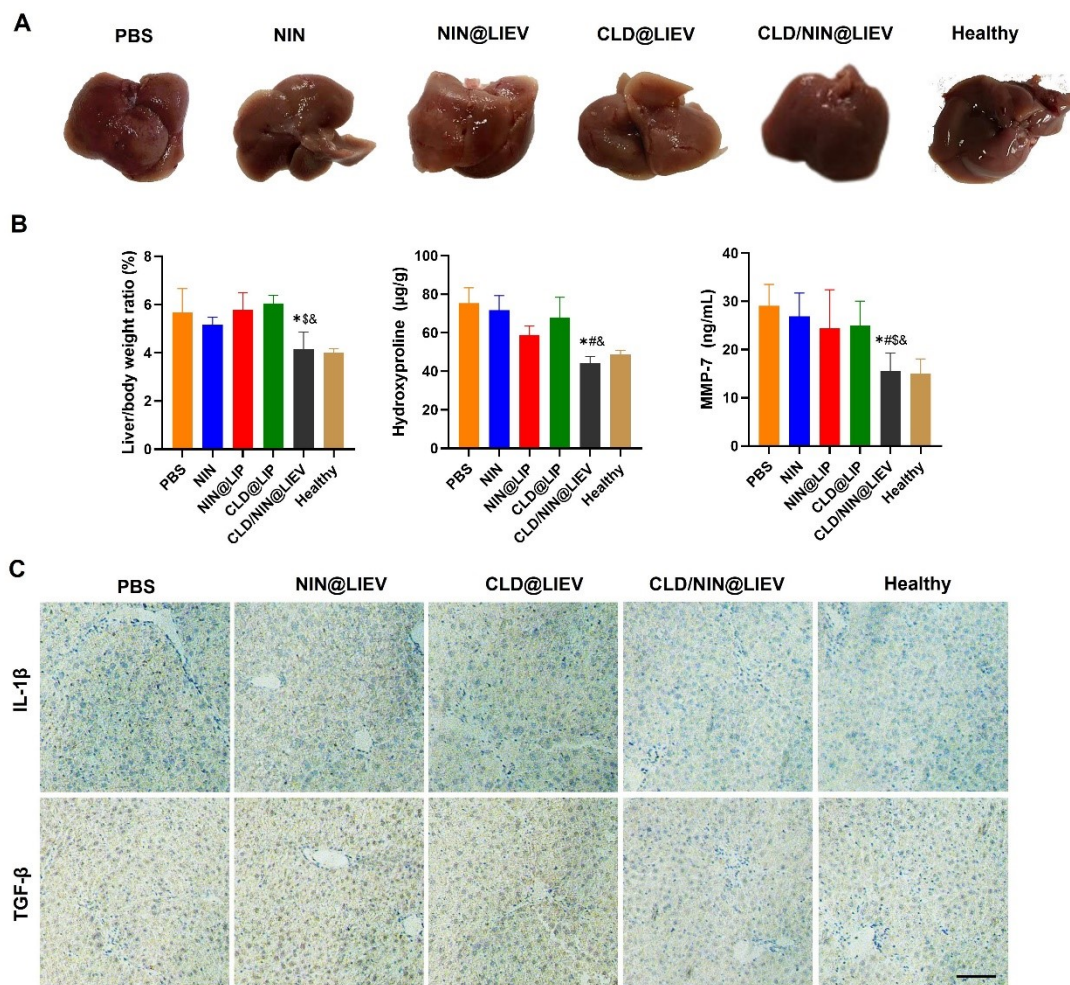


Fig. S3 Anti-fibrosis efficacy of CLD/NIN@LIEV. (A) Representative photos of mouse livers. (B) The liver weight-to-body weight ratio, hydroxyproline content, and MMP-7 expression after treatment. * $P < 0.05$, # $P < 0.05$, § $P < 0.05$, & $P < 0.05$, compared to PBS, NIN, NIN@LIEV and CLD@LIEV treatment, respectively. (C) IL-1 β and TGF- β immunohistochemistry. Data are mean \pm SD, $n = 3$. Scale bar = 50 μm .

Table S1. Antibodies used in the study.

Antibody	Company	Catalog	Application
Anti-F4/80	Servicebio	GB11027	IF
Anti- α -SMA	Servicebio	GB13044	IF
Anti-CD31	Servicebio	GB11063-1	IF
Anti-HNF-4 α	Abcam	ab201460	IF
Anti-Collagen I	Servicebio	GB11022-3	IF
Anti-CD86	Abcam	ab238468	IF
Anti-CD163	Servicebio	GB11340-1	IF
Anti-IL-1 β	Servicebio	GB11113	IHC
Anti-TGF- β	Servicebio	GB13028	IHC