Supplemental Figure 1. Representative images of immunostaining of FIBRONECTIN, COL1A1 and α -SMA in HPFs. The nuclei were stained blue by DAPI, and the images were taken under original magnification ×400. The data are represented as the mean ± SEM of three independent experiments. *, p < 0.05; **, p < 0.01; ***, p < 0.001.

Supplemental Figure 2. A: Representative results for Transwell assay in *SRPX2* siRNA or Scrambled siRNA treated HPFs. **B**: Representative results for EdU staining in *SRPX2* siRNA or Scrambled siRNA treated HPFs. The data are represented as the mean \pm SEM of three independent experiments. *, p < 0.05; ***, p < 0.001.

Supplemental Figure 3. Characteristics of the liposomes. A: The hydrodynamic diameter, PDI and zeta potential of the liposomes (blank or siRNA-loaded) were measured by DLS. SiRNA entrapment efficiency was measured by RiboGreen assay. **B**: Representative TEM image of siRNA-loaded liposomes. **C**: Hydrodynamic diameter distribution of siRNA-loaded liposomes. **D**: Colloid stability of siRNA-loaded liposomes in PBS.

Supplemental Figure 4. A: CCK-8 assay to check the effect of liposomes on the viability of fibroblasts. **B:** Representative results of Tunel assay in the mice lung sections. **C:** Histological analysis of lung, heart, liver, spleen, kidney, and intestine in mice after induction with liposomes. Representative images for H&E. Images were captured at ×400 magnification. **D:** liver, renal, and cardiac function of mice after induction with liposomes. Seven mice were included in each group. The data are represented as the mean ± SEM. ALT: Alanine aminotransferase; AST: Aspartate aminotransferase; BUN: Blood Urea Nitrogen; CR: Creatinine; CK: Creatine Kinase.

Supplemental Figure 5. RT-PCR analysis of *Fibronectin, Col1a1, α-SMA* and *Srpx2* expression in mice after BLM induction with Scrambled or *Srpx2* siRNA-loaded liposomes. Six mice were included in each study group. The data are represented as the mean \pm SEM. *, p < 0.05; **, p < 0.01; ***, p < 0.001.

Supplemental Figure 6. A: Western blot analysis of Arginase 1 (alternative activated macrophages marker) expression in mice after BLM induction with Scrambled or *Srpx2* siRNA-loaded liposomes. **B**: Western blot analysis of Srpx2 expression in macrophages following IL-4 induction. Six mice were included in each study group. The data are represented as the mean ± SEM.





FIB COL1A1 α-SMA SRPX2







Α			
_		Blank liposome	siRNA loaded liposome
_	Hydrodynamic diameter (nm)	115	102
	PDI	0.18	0.07
	Zeta-potential (mv)	22.1	2.8
_	Entrapment efficiency (%)	1	>95
B	С		D
	20- (%) 15- 0		
	200 nm	100 200 Diameter (nm)	0 2 4 6 8 10 12 24 Time (h)











0.0

PBS

IL-4