## 1. Supplementary Figures



Supplementary Figure S1. Successful establishment of cardiac fibrosis model in vivo and in vitro.

(A) The transcriptional expression of PML was examined in RNA-sequencing data (GSE116250) including patients with normal myocardium (No Failing; n = 14), patients with dilated cardiomyopathy (DCM; n = 37), and patients with ischemic cardiomyopathy (ICM; n = 13). (B) Masson's trichrome staining of the LV sections of mouse hearts at 7 days after MI operation. Scale bar, 50 µm (n = 5). (C-E) qRT-PCR showing increased mRNA levels of Col-1 $\alpha$ 1, Col-3 $\alpha$ 1 and  $\alpha$ -SMA in the infarct border zone of mouse left ventricular tissues at 7 days after MI operation (n = 6). (F-G) Following treatment with TGF- $\beta$ 1, Col-1 $\alpha$ 1 and Col-3 $\alpha$ 1 mRNA levels in CFs were examined by qRT-PCR (n = 4-6). \*\**P* < 0.01, \*\*\**P* < 0.001.





(A-D) qRT-PCR and western blot assay were used to verify PML knockdown or overexpression efficiency in CFs (n = 4-6). \*\*\*P < 0.001.



Supplementary Figure S3. The transfection efficiency of p53 knockdown and overexpression in cardiac fibroblasts.

(A) The protein level of phospho-p53 in CFs treated with TGF- $\beta$ 1 at the indicated time points (n = 6). (B-E) qRT-PCR and western blot assay were used to verify p53 knockdown or overexpression efficiency in CFs (n = 3). \*P < 0.05, \*\*P < 0.01, \*\*\*P < 0.001.



Supplementary Figure S4. Overexpression of PML cancelled the anti-fibrotic effect of p53 depletion.

(A) After transfection with PML plasmid, CFs were treated with or without Pifithrin- $\alpha$  in the presence of TGF- $\beta$ 1. Cell viabilities were examined by the CCK8 assay (n = 4). (B-D) qRT-PCR showing the mRNA levels of Col-1 $\alpha$ 1, Col-3 $\alpha$ 1 and  $\alpha$ -SMA (n = 4). (E) Western blot quantification of  $\alpha$ -SMA expression (n = 3). \**P* < 0.05, \*\**P* < 0.01, \*\*\**P* < 0.001.

## 2. Supplementary Tables

Table S1 Sequences of the	specific siRNAs
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Gene	Sense (5'-3')	Antisense (5'-3')
si-PML	GCUGAUCUCCGCGACA UUTT	AAUUGUCGCGGAGAUC AGCTT
si-p53	CAUUUUCAGGCUUAUG GAATT	UUCCAUAAGCCUGAAA AUGTT

Gene	Forward primer (5'-3')	Reverse primer (5'-3')
PML	CCTTTTCTTTTGACGGA CCA	TGCAACACAGAGGCTTG GC
p53	TGGAGGAGTCACAGTC GGAT	CAGTGAGGTGATGGCAG GAT
Col-1a1	AAGAAGACATCCCTGAA GTCA	TTGTGGCAGATACAGAT CAAG
Col-3a1	TTGGGATGCAGCCACCT TG	CGCAAAGGACAGATCCT GAG
<del>TGF-β1</del>	CCTGAGTGGCTGTCTTT TGACG	AGTGAGCGCTGAATCGA AAGC
α-SMA	CCCAGACATCAGGGAGT AATGG	TCTATCGGATACTTCAGC GTCA
GAPDH	GACAGCAGTTGGTTGGA GCA	TTGGGAGGGTGAGGGA CTTC

Table S2 Quantitative real-time PCR primer sequences of genes

Table S3 ChIP-qPCR primer sequences

Gene	Forward primer (5'-3')	Reverse primer (5'-3')
PML	CTCACAGACAGGGAAAA	CAAGCAAGTAAACAAGC
	GCC	CCG