

**Supplemental information**

**Exosomes mediated fibrogenesis in dilated  
cardiomyopathy through a MicroRNA pathway**

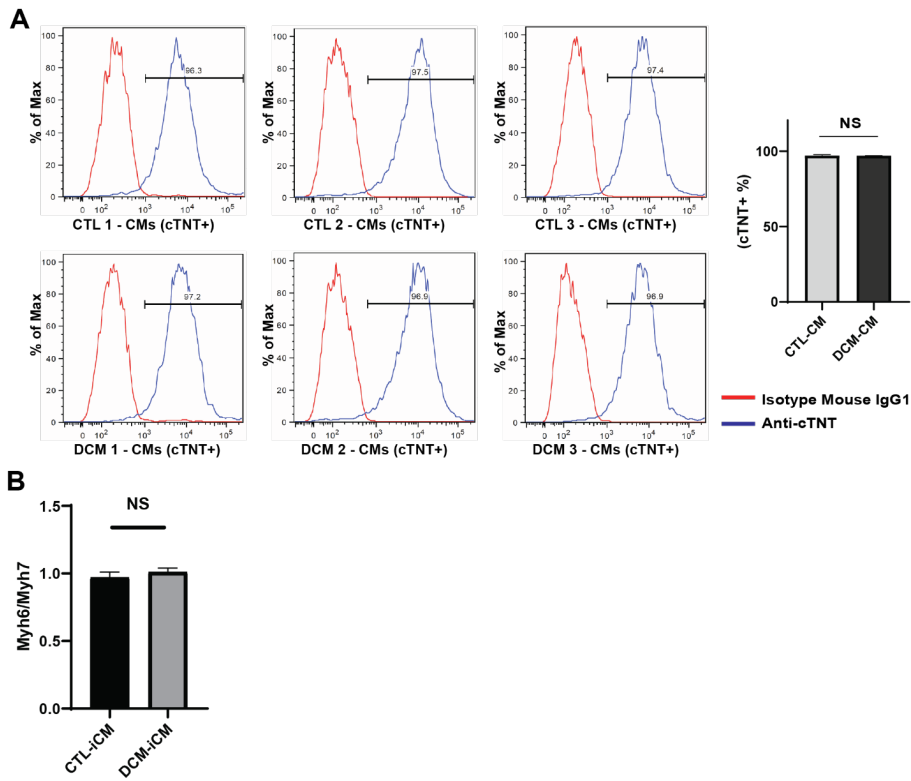
**Xuebin Fu, Rachana Mishra, Ling Chen, Mir Yasir Arfat, Sudhish Sharma, Tami Kingsbury, Muthukumar Gunasekaran, Progyaparamita Saha, Charles Hong, Peixin Yang, Deqiang Li, and Sunjay Kaushal**

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**Supplemental Information**

Exosomes Mediated Fibrogenesis in Dilated Cardiomyopathy Through a MicroRNA Pathway

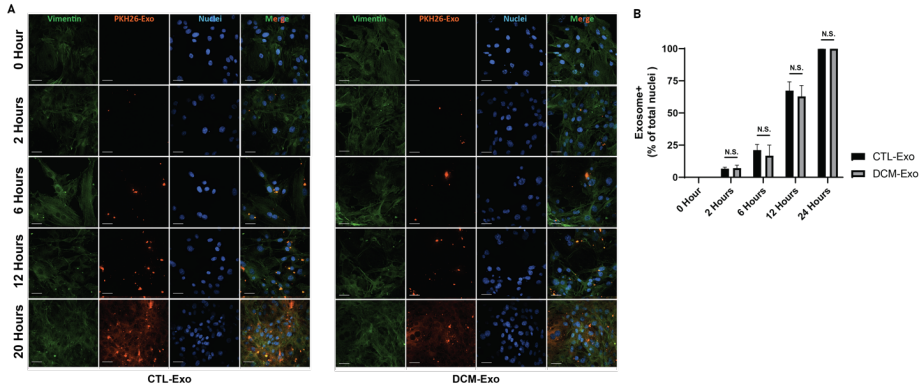
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21 **Figure S1.** The purity and maturation of CTL- and DCM-iPSC differentiated cardiomyocytes were  
22 determined by flow cytometry and RT-PCR. (A) Histograms for flow cytometric analysis of  
23 cTNT<sup>+</sup> in CTL-iCMs and DCM-iCMs. (B) Quantification of myh6 to myh7 ratio in CTL- and  
24 DCM-iCMs. Three CTL lines and 3 DCM lines were used in the experiments. Related to Figure  
25 1.

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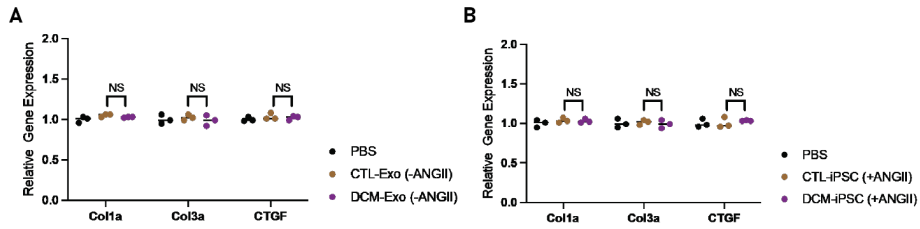


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28 **Figure S2.** Time course assay of PKH26-labeled Exos treatment in vitro. The pictures were taken  
 29 from different time-points after Exos treatment (A). Bar = 50um. The Exos uptake at different time  
 30 points was analyzed (B). Related to Figure 2.

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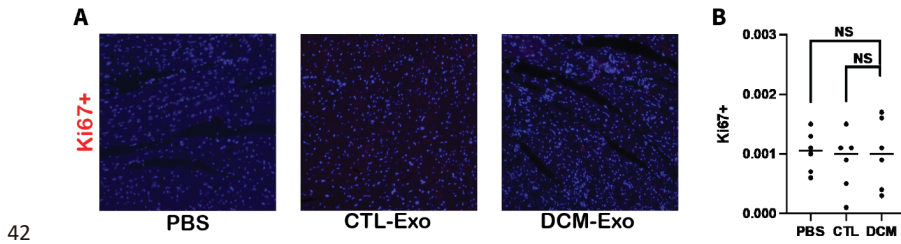
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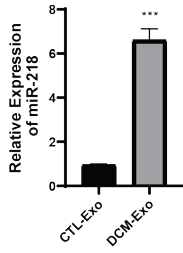
34 **Figure S3.** Neither Exos derived from basal media nor the Exos derived from iPSCs promoted  
 35 fibrogenesis. The expression of fibrotic markers was determined by qRT-PCR. (A), CFs were  
 36 treated with vehicle (PBS), CTL-Exo, or DCM-Exo derived from cardiomyocytes cultured in basal  
 37 media. N= 3; One-way ANOVA. (B), CFs were treated with vehicle (PBS) or Exos derived from  
 38 ANG II stimulated CTL-iPSC or DCM-iPSC. N= 3; One-way ANOVA; NS = no significance.  
 39 Related to Figure 2.  
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**Figure S4.** The injection of Exos did not induce proliferation in the hearts. (A) Immunofluorescent staining of detecting Ki67 (Red) in the vehicle, CTL-Exo, or DCM-Exo injection heart sections. The positive Ki67 indicated the proliferative cells in the hearts. (B) Statistical analysis showed positive Ki67 showed no significant differences. N= 6; One-way ANOVA; NS = nonsignificant. Related to Figure 3.

**A**



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50 **Figure S5.** RT-PCR determined the expression of miR-218 in Exos. (A) Compared with CTL-  
51 Exos, miR-218-5p expression in DCM-Exos was significantly increased. N=3; [Student's \*t\*-test](#);  
52 \*\*\*=P < 0.005. Related to Figure 4.

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56 **Table S1.** iPSCs used in this study. Related to Figure 1.

Patient No.	Cell Type	Disease	Age	Gender	Race	Passage
SCVI479	Sendai virus reprogrammed hiPSC	Healthy Donor	24	Male	African American	10
SCVI480	Sendai virus reprogrammed hiPSC	Healthy Donor	18	Female	African American	8
SCVII15	Sendai virus reprogrammed hiPSC	Healthy Donor	16	Male	Caucasian	8
SCVII17	Sendai virus reprogrammed hiPSC	DCM,TN NT2 <sup>R173W</sup>	75	Female	Caucasian	8
SCVII18	Sendai virus reprogrammed hiPSC	DCM,TN NT2 <sup>R173W</sup>	39	Male	Caucasian	11
SCVII19	Sendai virus reprogrammed hiPSC	DCM,TN NT2 <sup>R173W</sup>	46	Male	Asian	9

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59 **Table S2.** Sequence of wild-type 5'UTR and mutated 5'UTR. Related to Figure 7.

Name	Sequence
Wild-type	gcagtctgca gtcttcgtgg cgggccaagc gagcttggag ccgcgggggg cggagcggtg agagcggccg ccaagagaga tcacacccc agccgaccct gccagcgagc gagcccgacc ccaggcgctc atggagcgtc gcctccgcc ggtccctgcc ccgacccccg cctgcggcgc gctctgcct tgaccaggac ttgggacttt gcgaaaggat cgcggggccc ggagaggtaa ccgccgcgc tccgggagag gtgttggaga gcacaatggc tgaacaagtc ctctcagg
Mutation	gcagtctgca gtcttcgtgg cgggccaagc gagcttggag ccgcgggggg cggagcggtg agagcggccg ccaagagaga tcacacccc agccgaccct gccagcgagc gagcccgacc ccaggcgctc atggagcgtc gcctccgcc ggtccctgcc ccgacccccg cctgcggcgc gctctgcct tgaccaggac ttgggacttt gcgaaaggat cgcggggccc ggagaggtaa ccgccgcgc tccgggagag ctataccgc tcttggc tgaacaagtc ctctcagg

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