

#### **Supporting Information**

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Exosome-Guided Phenotypic Switch of M1 to M2 Macrophages for Cutaneous Wound Healing

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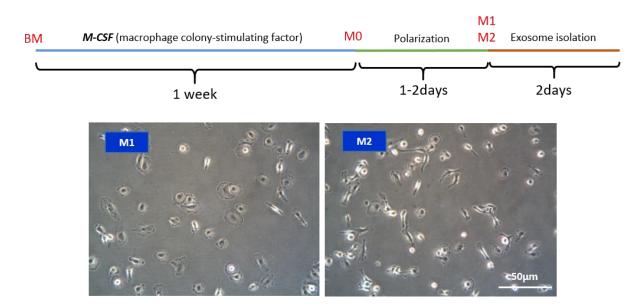


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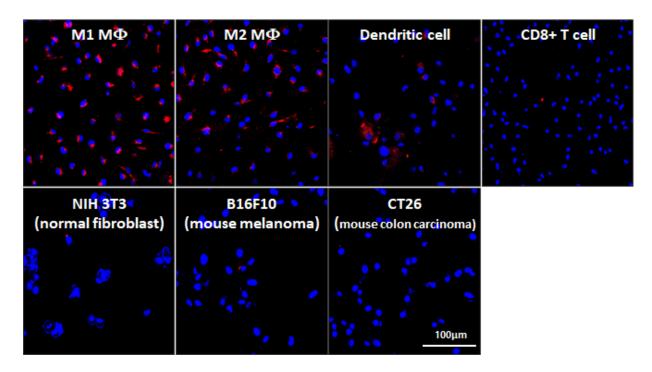
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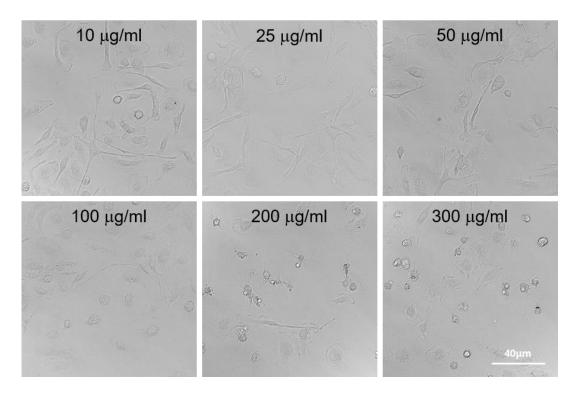
Hyosuk Kim, Sun Young Wang, Gijung Kwak, Yoosoo Yang, Ick Chan Kwon\*, and Sun Hwa Kim\*



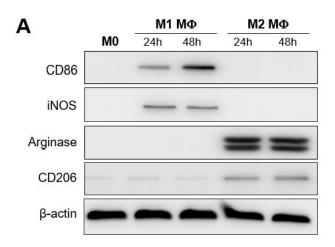
**Figure S1.** Establishment of M1 and M2 M $\phi$ s. Experimental schedules of M $\phi$  polarization and the differences in morphology of M1 and M2 M $\phi$ s.

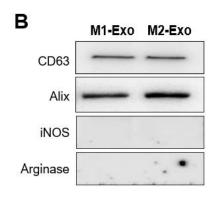


**Figure S2.** Confocal images of various types of cells after 4 h incubation with 50  $\mu$ g/ml of DiD-labeled M2-Exo. Images of DiD-labeled M2-Exo (red) with DAPI (blue) were visualized by merging the confocal images.

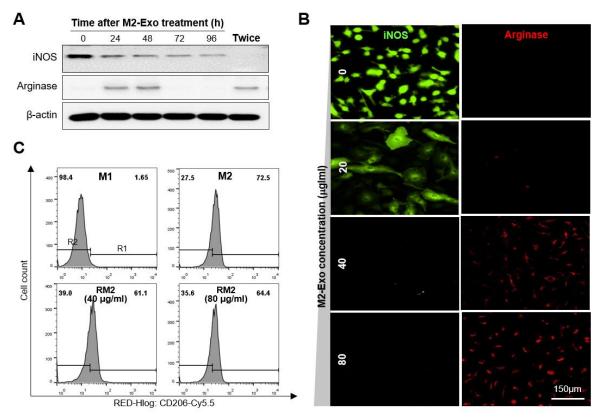


**Figure S3.** Morphological change of M $\phi$ s with exosome concentration. Representative bright-field micrographs of M $\phi$ s after 24 h incubation with 10, 25, 50, 100, 200 and 300  $\mu$ g/ml of M2-Exo.

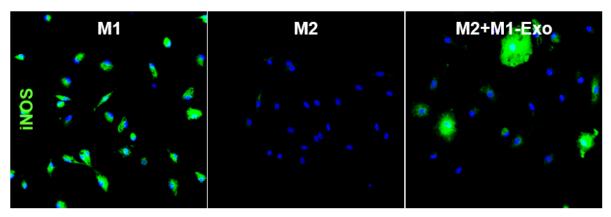




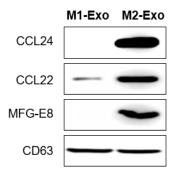
**Figure S4.** Establishment of M1 and M2 M $\phi$ s of C57BL/6 mice, and identification of exosomes released from M $\phi$ s. A) Western blot analysis demonstrating differences in expression of M $\phi$  markers by polarization time. B) Western blot analysis of exosomes. Equal amounts of total proteins extracted from exosomes were immunoblotted for CD63, Alix, iNOS, and Arginase.



**Figure S5.** M2-Exo-guided direct reprogramming using M $\phi$ s from C57BL/6 mice. A) Western blot analysis of M1 M $\phi$ s treated with 40  $\mu$ g/ml of M2-Exo over time. B) Immunostaining of iNOS and Arginase in M1 M $\phi$ s after 24 h incubation with 20, 40 and 80  $\mu$ g/ml of M2-Exo, respectively. C) FACS histogram showing reprogramming efficiency of M1 M $\phi$ s treated with 40 and 80  $\mu$ g/ml of M2-Exo.



**Figure S6.** M1-Exo-guided direct reprogramming of M2 M $\phi$ s to M1 M $\phi$ s. Immunostaining of iNOS in M2 M $\phi$ s after 24 h incubation with 40  $\mu$ g/ml of M1-Exo.



**Figure S7.** Western blot analysis of M1 and M2-derived exosomes. Equal amounts of total proteins extracted from exosomes were immunoblotted for CCL24, CCL22, MFG-E8 and CD63.

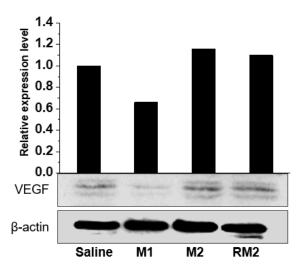
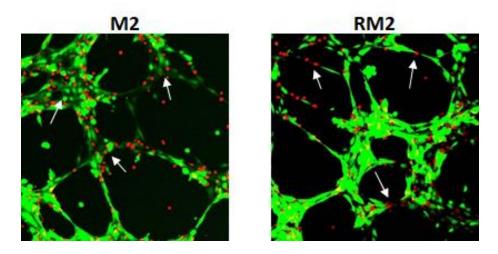


Figure S8. Western blot analysis of VEGF among the three different types of  $M\phi$  groups.



**Figure S9.** Co-localization of fluorescently labeled endothelial cells (green) and M $\phi$ s (red). Arrows indicate M $\phi$ s merging into the tubular structures.