## **Supplementary Material**

## *Synechococcus elongatus* PCC7942 secretes extracellular vesicles to accelerate cutaneous wound healing by promoting angiogenesis

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Figure S1. Effects of *S. elongatus* PCC 7942 on functional properties of keratinocytes and fibroblasts. (A) CCK-8 analysis of proliferation of human keratinocyte cell line HaCaT and human skin fibroblasts (HSFs). n = 4 per group. (B-C) Representative images of wound healing assay in HaCaT (B) and quantitative analysis of the migration rates (C). Scale bar: 200 µm. n = 3 per group. (D-E) Representative images of transwell migration assay in HSFs (D) and quantitative analysis of the migrated cells (E). Scale bar: 100 µm. n = 3 per group. Data are plotted as mean  $\pm$  SD. \**P* < 0.05, \*\**P* < 0.01, \*\*\**P* < 0.001.



Figure S2. Effects of *S. elongatus*-EVs on functional properties of human keratinocytes and fibroblasts. (A) CCK-8 analysis of proliferation of keratinocyte cell line HaCaT and HSFs. n = 4 per group. (B-C) Representative images (B) and quantification (C) of wound healing assay for HaCaT. Scale bar: 200 µm. n = 3 per group. (D-E) Representative images (D) and quantification (E) of wound healing assay for HSFs. Scale bar: 200 µm. n = 3 per group. Data are plotted as mean  $\pm$  SD. \**P* < 0.05, \*\**P* < 0.01, \*\*\**P* < 0.001.



Figure S3. Effects of *S. elongatus* PCC 7942 and *S. elongatus*-EVs on functional properties of mouse epidermal cells and fibroblasts. (A) CCK-8 analysis of proliferation of mouse epidermal JB6 cells and NIH3T3 fibroblasts. n = 4 per group. (B-C) Representative images (B) and quantification (C) of wound healing assay for JB6 cells. Scale bar: 200 µm. n = 3 per group. (D-E) Representative images (D) and quantification (E) of wound healing assay for NIH3T3 fibroblasts. Scale bar: 200 µm. n = 3 per group. Data are plotted as mean  $\pm$  SD. \*P < 0.05, \*\*P < 0.01, \*\*\*P < 0.001.



Figure S4. Effects of *S. elongatus* PCC 7942 and *S. elongatus*-EVs on morphology and growth of endothelial cells. (A) Morphology of HMECs under optical microscopy. Scale bar: 100  $\mu$ m. (B) CCK-8 analysis of HMECs' viability. n = 4 per group. Data are plotted as mean  $\pm$  SD. <sup>\*</sup>*P* < 0.05, <sup>\*\*</sup>*P* < 0.01, <sup>\*\*\*</sup>*P* < 0.001.