Supplementary information

Resveratrol enhances the functionality and improves the regeneration of

mesenchymal stem cell aggregates

Yi-Jing Wang <sup>1, 2, †</sup>, Pan Zhao <sup>3, 4, †</sup>, Bing-Dong Sui <sup>3, †</sup>, Nu Liu <sup>3, 5</sup>, Cheng-Hu Hu <sup>3, 4</sup>, Ji Chen <sup>3</sup>,

Chen-Xi Zheng<sup>3</sup>, An-Qi Liu<sup>3</sup>, Kun Xuan<sup>3</sup>, Ya-Ping Pan<sup>1,\*</sup>, and Yan Jin<sup>3,\*</sup>

<sup>1</sup> Department of Periodontics and Oral Biology, School of Stomatology, China Medical University,

Shenyang, Liaoning 110002, China.

<sup>2</sup> General Hospital of Shenyang Military Region, Shenyang, Liaoning 110016, China.

<sup>3</sup> State Key Laboratory of Military Stomatology & National Clinical Research Center for Oral

Diseases & Shaanxi International Joint Research Center for Oral Diseases, Center for Tissue

Engineering, School of Stomatology, Fourth Military Medical University, Xi'an, Shaanxi 710032,

China.

<sup>4</sup> Xi'an Institute of Tissue Engineering and Regenerative Medicine, Xi'an, Shaanxi 710032, China.

<sup>5</sup> Department of Periodontology, Stomatological Hospital, Zunyi Medical College, Zunyi, Guizhou

563003, China.

<sup>†</sup> Equal contribution; <sup>\*</sup> Corresponding authors.

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Supplementary information: 2 figures with their legends.

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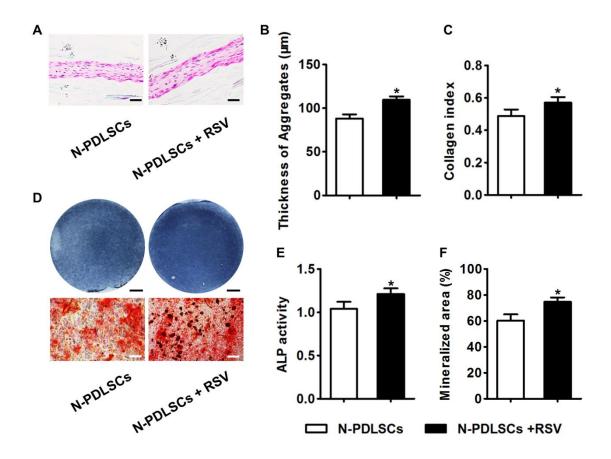


Figure S1. Resveratrol (RSV) treatment improved N-PDLSC-aggregate formation and osteogenic potential. (A-C) Histological analysis of PDLSC-aggregates (A) showed that N-PDLSCs formed thicker aggregates (B) with more collagen deposition (C) under RSV treatment. Bars:  $50 \, \mu m$ . (D) Osteogenesis of N-PDLSC-aggregates demonstrated by alkaline phosphatase (ALP) and alizarin red staining. Bars:  $5 \, mm$  (up) and  $50 \, \mu m$  (down). (E, F) N-PDLSC-aggregates showed higher ALP activity (E) and stronger mineralized capability (F) in osteogenic differentiation under RSV treatment. RSV was applied at  $10 \, nM$  throughout the *in vitro* treatments. n = 3 per group. Data represents mean  $\pm$  standard deviation (SD). \*P < 0.05.

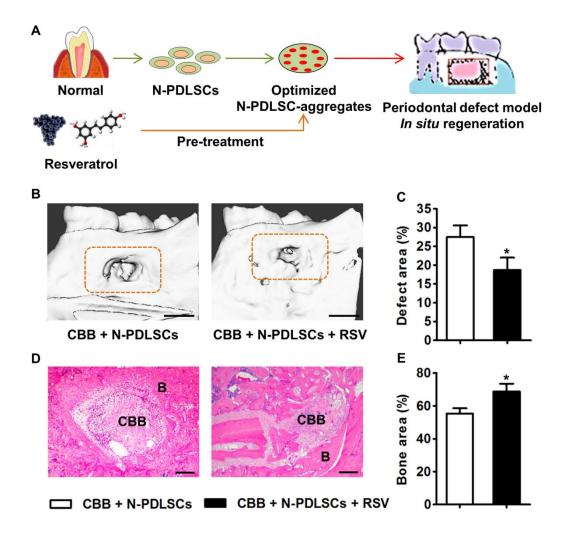


Figure S2. RSV treatment facilitated alveolar bone regeneration of N-PDLSC-aggregates in a rat periodontal defect model. (A) A schema illustrating the experimental design. (B, C) Representative micro-CT figures of periodontal defects after N-PDLSC-aggregate-based regeneration (B) and statistical analysis of bone volume per tissue volume (BV/TV) (C). The orange brackets indicate the original defects that were selected as region of interests in statistical analysis. (D, E) Histological analysis of periodontal bone regeneration (denoted by B) by N-PDLSC-aggregates by HE staining and the statistical analysis of regenerated bone area (E). RSV was applied at 10 nM throughout the *in vitro* treatments. Calcined bovine bone (CBB) was used as the scaffold. n = 4 per group. Data represents mean  $\pm$  SD. \*P < 0.05.