Supplementary Material



Figure S1. OVX mice show decreased size and weight of uterus. (A) Representative images of uteruses from Sham, OVX and OVX + hucMSC-EVs mice. (B) Quantitative analysis of uterus weight in (A). n = 8 per group for OVX + hucMSC-EVs; n = 10 for other groups. *P < 0.05, **P < 0.01, ***P < 0.001.



Figure S2. hucMSC-EVs slightly reverse the osteoporotic phenotypes of aged mice. (A-C) Quantitative μ CT analysis of Tb. BV/TV, Tb. Th and Tb. N in femora from hucMSC-EVs- or PBS-treated 19-month-old mice. n = 7 per group (PBS); n = 8 per group (hucMSC-EVs).



Figure S3. Identification of BMSCs. (A) BMSCs showed a fusiform morphology. Scale bar: 100 µm.

(**B**) BMSCs had the ability to differentiate into osteoblasts, adipocytes or chondrocytes under osteogenic, adipogenic or chondrogenic culture condition, confirmed by Alizarin Red S staining, Oil Red O staining and Alcian Blue staining. Scale bars: 50 μ m. (**C**) The characteristic surface markers on BMSCs analyzed by flow cytometry. Blank curves: the isotype controls; solid gray curves: the test samples.