SUPPLEMENTARY FIGURES



Supplementary Figure 1. Xenograft tumor of GSCs in chimeric mice (Model II). (A) Under confocal microscopy, close contact could be observed between the bone marrow-derived EGFP+ cells and RFP+ GSCs (white arrow); (B) RFP expressing GSCs under confocal microscopy; (C) GFP expressing bone marrow-derived EGFP+ cells; (D) DAPI labeled cell nucleus. Scale bar: 20 µm.



Supplementary Figure 2. (A) Magnetic resonance image of a 34-year-old male patient diagnosed with glioblastoma; (B) Tumorigenicity of the three transformed cell lines: TMEC1, TMEC2 and TMEC3; (C) Fluorescence-activated EGFP⁺ cell sorting of the intracerebral xenograft of GSCs; (D) Fluorescence-activated EGFP⁺ cell sorting of the intracerebral xenograft of clinical tumor specimens; (E) Immunofluorescence of the three tMSC lines. Scale bar: (C) 2 μ m



Supplementary Figure 3. Direct co-culture of GSCs and MSCs *in vitro*. (A) Time-lapse photography of a live-cell imaging station revealed tight contacts between RFP^+ GSCs and $EGFP^+$ MSCs; B-F) Both GSCs and MSCs protruded cell processes toward each other (B), which released vesicle-like mini-particles (C–E) and turned yellow (F) after the two kinds of particles had fused (black arrow). Scale bar: 200 µm.



Supplementary Figure 4. GSCs and MSCs were co-cultured *in vitro***.** RFP⁺ GSCs communicated with EGFP⁺ MSCs via long, tiny microtube-like structures (A–C, black arrow). After the intercellular cytoplasm exchange, these microtubes turned yellow (D–F, white arrow). Scale bar: 200 μm.





GFP-expressing MSCs



Supplementary Figure 5. Transwell indirect co-culture of GSCs and MSCs revealed certain morphological changes in MSCs, but no activation of MSC proliferation. (A) Schematic overview of indirect co-culture in Transwell chambers; (B) Day 2, no obvious proliferation of MSCs when indirectly co-cultured with GSCs, just certain morphological changes compared with MSCs cultured alone (E, control); (C) Day 12, MSC proliferation could be observed in both the co-culture and the control plate (F); (D) Day 22, the proliferation of MSCs decreased gradually, either when the cells were indirectly co-cultured with GSCs, or when they were cultured alone (G). Scale bar: 50 µm.



Supplementary Figure 6. (A) Clone formation assay with HNRNPD-overexpressing cells; (B) Flow cytometry assay detecting the cell cycle in HNRNPD-overexpressing cells.



Supplementary Figure 7. Functional recovery assay. (A) qRT-PCR after overexpression of HNRNPD; (B) CCK-8 cell proliferation curve; (C) Colony formation assay of tMSC1 cells overexpressing HNRNPD and miRNA-146a-5p; (D) Statistical analysis of the colony formation assay.



Supplementary Figure 8. GO and KEGG analyses of several dysregulated miRNAs. (A) Top 30 GO terms; (B) Top 20 KEGG pathway classifications.