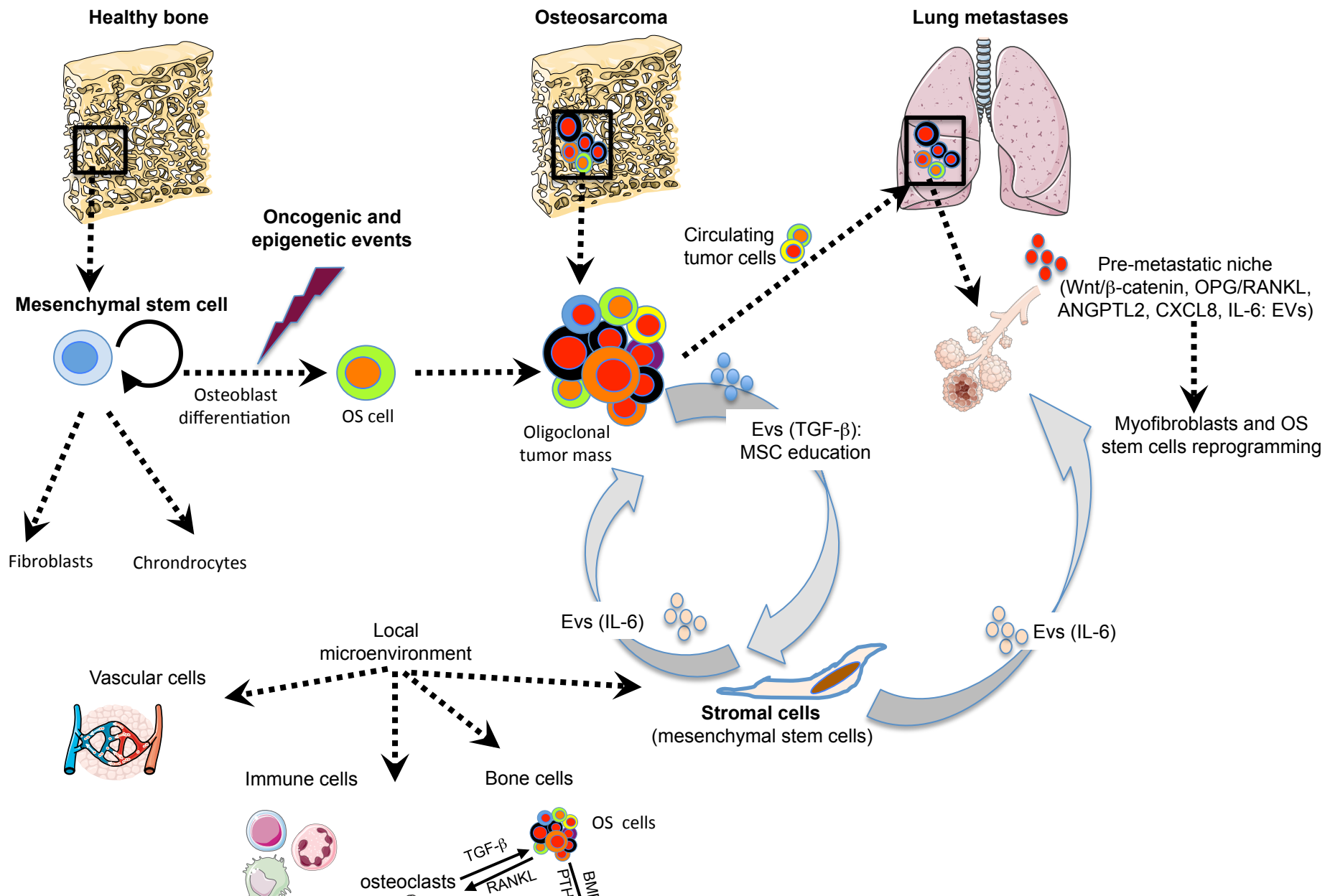
Healthy bone
Osteosarcoma
Lung metastases
Oncogenic and epigenetic events
Mesenchymal stem cell
Osteoblast differentiation
OS cell
Oligoclonal tumor mass
Circulating tumor cells
Pre-metastatic niche (Wnt/β-catenin, OPG/RANKL, ANGPTL2, CXCL8, IL-6: EVs)
Myofibroblasts and OS stem cells reprogramming
Evs (TGF-β): MSC education
Fibroblasts
Chrondrocytes
Evs (IL-6)
Evs (IL-6)
Local microenvironment
Vascular cells
Stromal cells (mesenchymal stem cells)
Immune cells
Bone cells
OS cells
TGF-β
RANKL
osteoclasts
PTH
BM