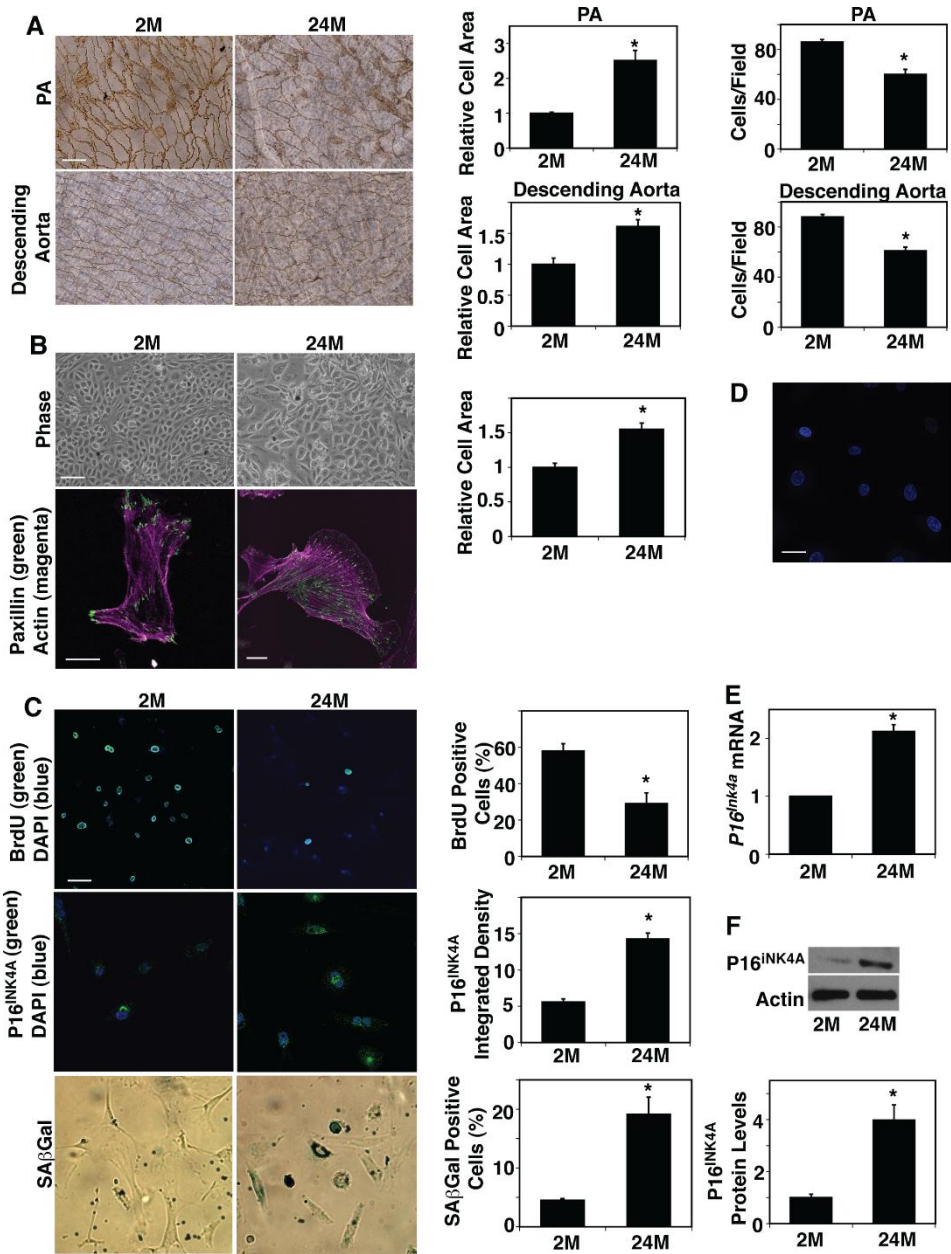
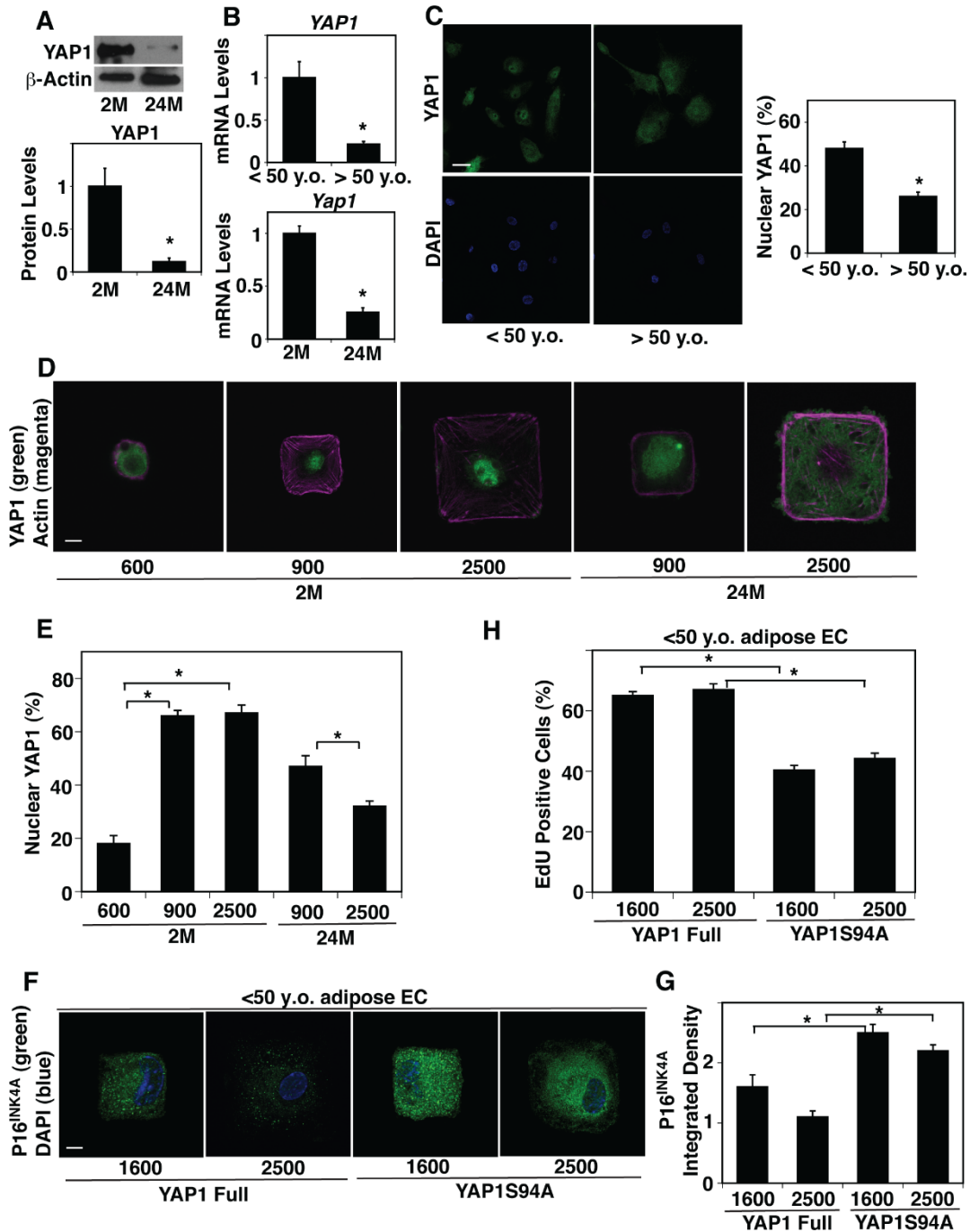


SUPPLEMENTARY FIGURES



Supplementary Figure 1. Age-dependent changes in mouse EC size, proliferation and senescence. (A) Silver nitrate-stained 2M and 24M old mouse PA (*top*) and descending aorta (*bottom*). Scale bar, 20 μ m. Graphs showing quantification of cell area and density in PA (*top*) and descending aorta (*bottom*) dissecting from 2M and 24M old mice (n=7, mean \pm s.e.m., *, p<0.05). (B) Phase contrast images of cultured ECs isolated from 2M vs. 24M old mouse lungs (*top*, scale bar, 100 μ m). IF micrographs showing paxillin-positive focal adhesions (green) and actin stress fiber formation (magenta, *bottom*). Scale bar, 20 μ m. Graph showing quantification of cell area of ECs isolated from 2M and 24M old mouse lungs (n=7, mean \pm s.e.m., *, p<0.05). (C) IF micrographs showing BrdU⁺ (*top*), P16^{INK4A} (*middle*), and SA β Gal-stained (*bottom*) ECs isolated from 2M vs. 24M old mouse lungs. Graphs showing quantification of BrdU⁺ (*top*), P16^{INK4A} (*middle*), and SA β Gal-stained (*bottom*) ECs isolated from 2M vs. 24M old mouse lungs (n=7, mean \pm s.e.m., *, p<0.05). Scale bar, 20 μ m. (D) IF micrograph showing the 2nd Ab alone (green) and DAPI (blue) staining. Scale bar, 20 μ m. (E) Graph showing P16^{INK4A} mRNA levels in ECs isolated from 2M vs. 24M old mouse lungs (n=5, mean \pm s.e.m., *, p<0.05). (F) Representative immunoblots showing P16^{INK4A} and β -actin protein levels in ECs isolated from 2M vs. 24M old mouse lungs (*top*). Graph showing the quantification of immunoblots (*bottom*, n=3, mean \pm s.e.m., *, p<0.05).



Supplementary Figure 2. Cell size-dependent changes in YAP1 activity in mouse lung ECs. (A) Representative immunoblots showing YAP1 and β -actin protein levels in ECs isolated from 2M vs. 24M old mouse lungs (*top*). Graph showing the quantification of immunoblots (*bottom*, $n=3$, $p<0.05$). (B) Graphs showing YAP1 mRNA levels in ECs isolated from <50 y.o. vs. >50 y.o. human adipose tissues (*top*) and 2M vs. 24M old mouse lungs (*bottom*, $n=5$, mean \pm s.e.m., $*$, $p<0.05$). (C) IF micrographs showing YAP1 localization (*top*) and DAPI (*bottom*) in ECs isolated from <50 y.o. and >50 y.o. human adipose tissues. Scale bar, 20 μ m. Graph showing quantification of nuclear YAP1 in ECs isolated from <50 y.o. and >50 y.o. human adipose tissues ($n=7$, mean \pm s.e.m., $*$, $p<0.05$). (D) IF micrographs showing YAP1 localization (green) and actin structure (magenta) in ECs isolated from 2M vs. 24M old mouse lungs and cultured on FN-coated island of different sizes. Scale bar, 10 μ m. (E) Graph showing quantification of nuclear YAP1 ($n=7$, mean \pm s.e.m., $*$, $p<0.05$). (F) IF micrographs showing P16^{INK4A} expression (green) and DAPI (blue) in ECs isolated from <50 y.o. human adipose tissues overexpressing full-length YAP1 or YAP1S94A, cultured on FN-coated island of different sizes. Scale bar, 10 μ m. (G) Graph showing quantification of P16^{INK4A} integrated density in ECs isolated from <50 y.o. human adipose tissues overexpressing full-length YAP1 or YAP1S94A, cultured on FN-coated island of different sizes ($n=7$, mean \pm s.e.m., $*$, $p<0.05$). (H) Graph showing quantification of EdU-positive ECs isolated from <50 y.o. human adipose tissues overexpressing full-length YAP1 or YAP1S94A, cultured on FN-coated island of different sizes ($n=7$, mean \pm s.e.m., $*$, $p<0.05$).