



Supplementary Figure 2: CAIs diminish vascular A^β burden and endothelial caspase-3 activation in CA1 area in TgSwDI animals. A) Representative immunofluorescence images of the CA1 hippocampal region of 16-month-old mice. Compared to WT, untreated TgSwDI animals exhibited massive amount of A β (red) and evident caspase-3 activation (green) in CA1, both significantly decreased by 8-month CAI treatment. Original magnification, 60x. Scale bar, 25µm. On the right, the relative quantification is plotted as the percentage of A β and active caspase-3 area per acquisition field. For %Aβ, WT, TgSwDI and MTZ: N=5, ATZ: N=4, n≥12 measurements acquired /group. For %active caspase-3, WT, TgSwDI and MTZ: N=5, ATZ: N=3, n≥9 measurements acquired /group. **p<0.01, **** and ++++p<0.0001, One-way ANOVA and Tukey's post-hoc test. The magnified images show the spatial overlap between the signals: Aß (red) in the microvasculature; CD31 (blue) (overlap, magenta), endothelial active caspase-3 (green) (overlap, cyan), AB overlapping active caspase-3 (yellow). Below, percentage of AB and active caspase-3 signals overlapping with CD31 are graphed. ATZ and MTZ significantly attenuate Aβ/CD31 overlap, and show a trend in reducing active caspase-3/CD31 overlap in TgSwDI mice. For A β /CD31 overlap, TgSwDI and MTZ: N=5, ATZ: N=4, n \geq 12 measurements acquired /group. *p<0.05 and **p<0.01, One-way ANOVA and Tukey's post-hoc test. For active caspase-3/CD31 overlap, =3-5 mice/group, $n \ge 5$ measurements/group. **B**) Representative 10x and 60x images of DAB-enhanced Perls Prussian Blue staining in meningeal arteries of 16-month-old mice, showing higher number of microhemorrhages (MH) (indicated with arrows) in TgSwDI mice, vs WT animals, which are reduced in CAI-treated groups. Colocalization of iron aggregates with CD31+ BVs (magenta) is shown in the 60x and magnified images. Scale bars are 250µm and 50µm, respectively for 10x and 60x magnification. The plot on the right represents the average number of MH counted. WT, TgSwDI and MTZ: N=4, ATZ: N=3, n≥10 counts/group. **p<0.01 and ***p<0.001, One-way ANOVA and Tukey's post-hoc test. Data are expressed as mean ± SEM.