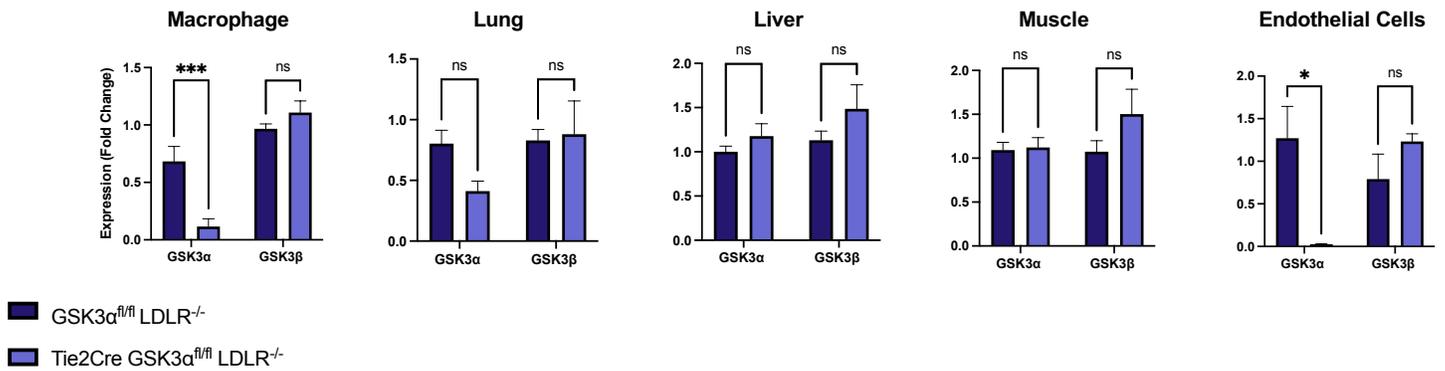
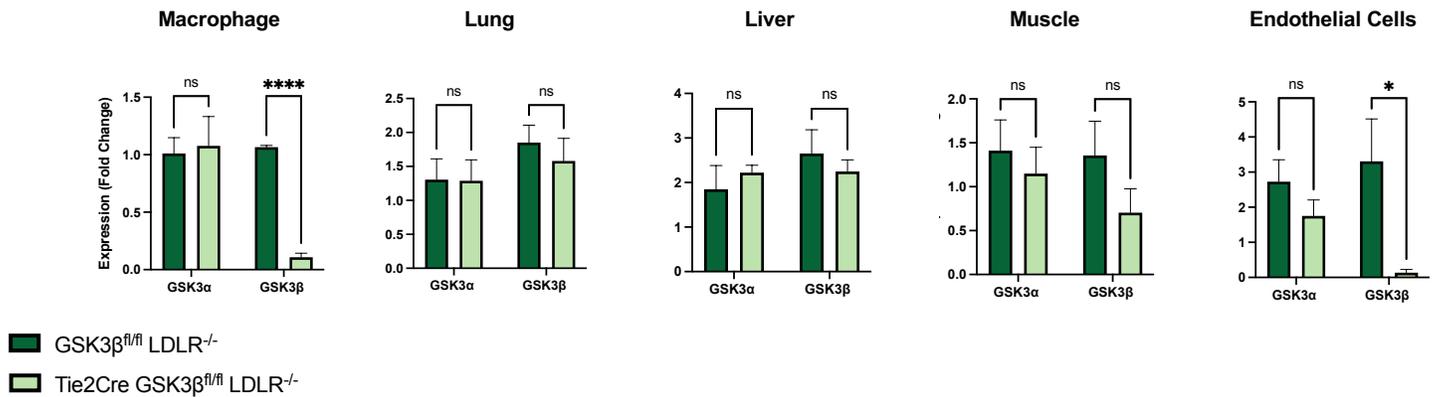


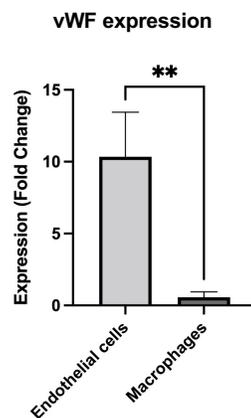
A

 $\alpha^{fl/fl}$ Mouse model

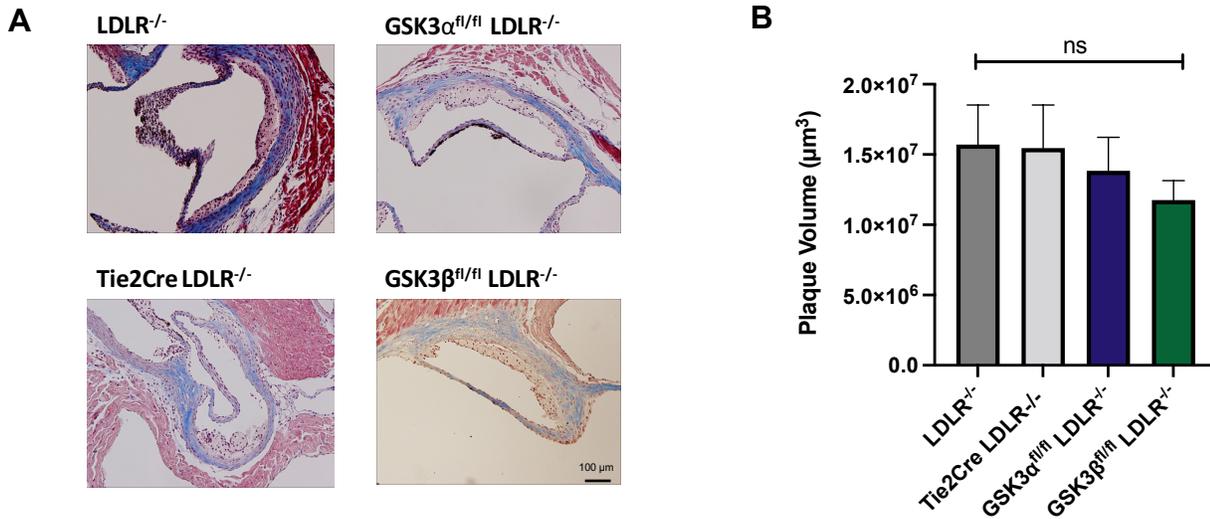
B

 $\beta^{fl/fl}$ Mouse model

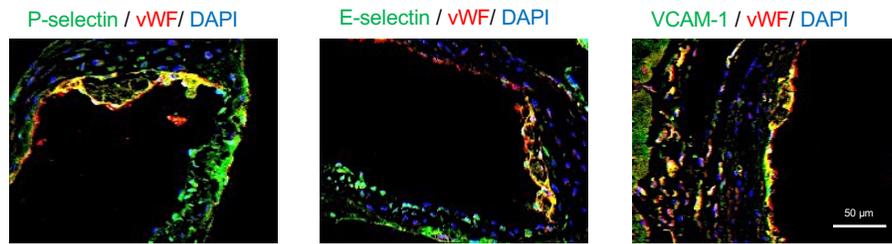
C



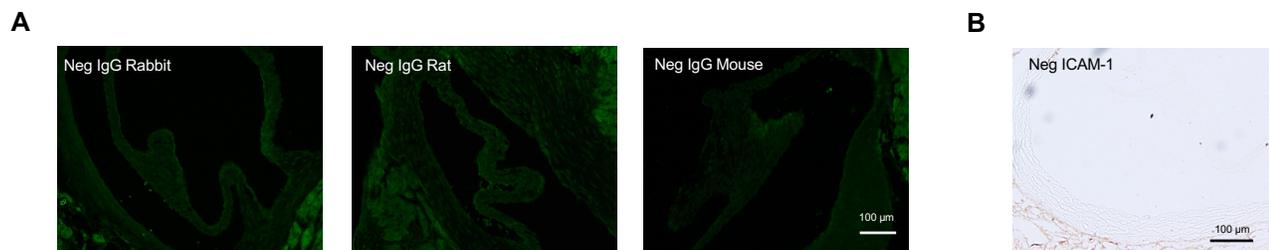
Supplementary Figure S1: Characterizing the endothelial/macrophage GSK3 α/β knockout mouse model. **A** Gene expression of GSK3 α/β was assessed in macrophages, lung, liver, skeletal muscle and endothelial cells from GSK3 $\alpha^{fl/fl}$ LDLR $^{-/-}$ and Tie2Cre GSK3 $\alpha^{fl/fl}$ LDLR $^{-/-}$ mice. **B** Gene expression of GSK3 α/β was assessed in macrophages, lung, liver, skeletal muscle and endothelial cells from GSK3 $\beta^{fl/fl}$ LDLR $^{-/-}$ and Tie2Cre GSK3 $\beta^{fl/fl}$ LDLR $^{-/-}$ mice. Gene expression was normalised to a β actin control. **C** To verify that the cells isolated from murine aortas were, in fact, endothelial cells, the expression of endothelial marker, vWF was assessed. n=3-7, * P<0.05, **P<0.005, ***P<0.0005, ****P<0.00005.



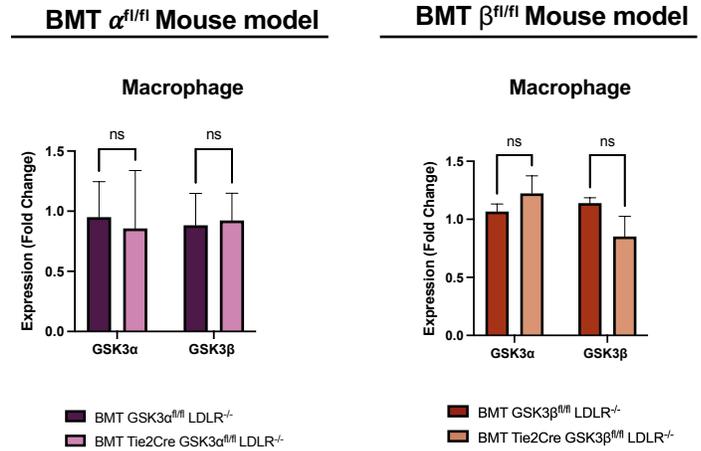
Supplementary Figure S2: Atherosclerosis in control mice from the endothelial/macrophage GSK3 α/β knockout mouse model. **A** Representative images of aortic sinus cross sections from 15 week old control LDLR^{-/-}, Tie2Cre LDLR^{-/-}, GSK3 $\alpha^{fl/fl}$ LDLR^{-/-} and GSK3 $\beta^{fl/fl}$ LDLR^{-/-} mice stained with Masson's Trichrome. Scale bar represents 100 μ m. **B** Plaque volume quantification in the aortic sinus of control mice, n=4-8.



Supplementary Figure S3: Co-staining of adhesion protein and vWF. Representative images of aortic cross sections stained with antibodies against vWF and P-selectin, E-selectin or VCAM-1. Green represents the adhesion proteins (P-selectin, E-selectin and VCAM-1), red represents vWF staining and blue represents DAPI. Yellow represents where both proteins are present. Scale bar represents 50 μ m.

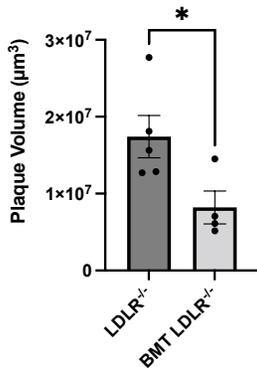


Supplementary Figure S4: Negative controls for immunofluorescence and immunohistochemistry staining. **A** Pre-immune IgG rabbit was used in place of primary antibody as a negative control for both E-selectin and VCAM-1 staining. Pre-immune IgG rat was used in place of primary antibody as a negative control for Mac-3 staining. Pre-immune IgG mouse was used in place of primary antibody as a negative control for P-selectin staining. **B** Pre-immune goat IgG was used in place of the primary antibody for a negative control for ICAM-1 staining. Scale bar represents 100 μ m.

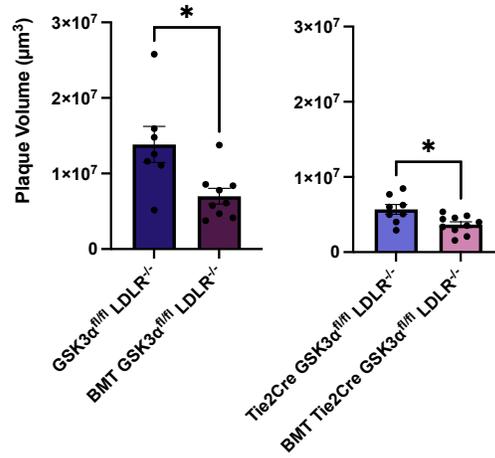
A**B**

Supplementary Figure S5: Confirmation that BMT restored GSK3 α and GSK3 β in macrophages. **A** PCR analysis of genomic DNA from blood cells from BMT GSK3 $\alpha^{fl/fl}$ LDLR $^{-/-}$, LDLR $^{-/-}$, GSK3 $\alpha^{fl/fl}$ LDLR $^{-/-}$, and Tie2Cre GSK3 $\alpha^{fl/fl}$ LDLR $^{-/-}$ mice. Results confirm that BMT mice have the GSK3 α wildtype (WT) gene restored. **B** RT-PCR analysis of GSK3 α/β expression in macrophages isolated from BMT GSK3 $\alpha^{fl/fl}$ LDLR $^{-/-}$, BMT Tie2Cre GSK3 $\alpha^{fl/fl}$ LDLR $^{-/-}$, BMT GSK3 $\beta^{fl/fl}$ LDLR $^{-/-}$, and BMT Tie2Cre GSK3 $\beta^{fl/fl}$ LDLR $^{-/-}$ mice after a BMT, n=3.

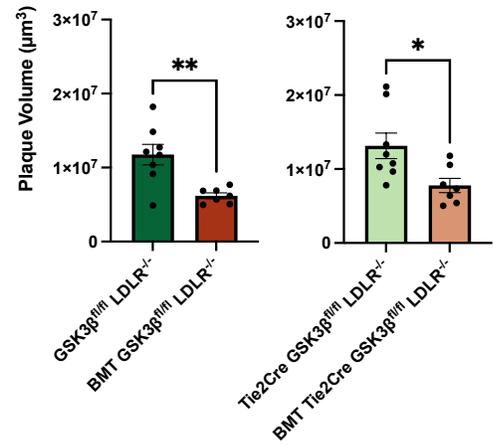
LDLR^{-/-} model



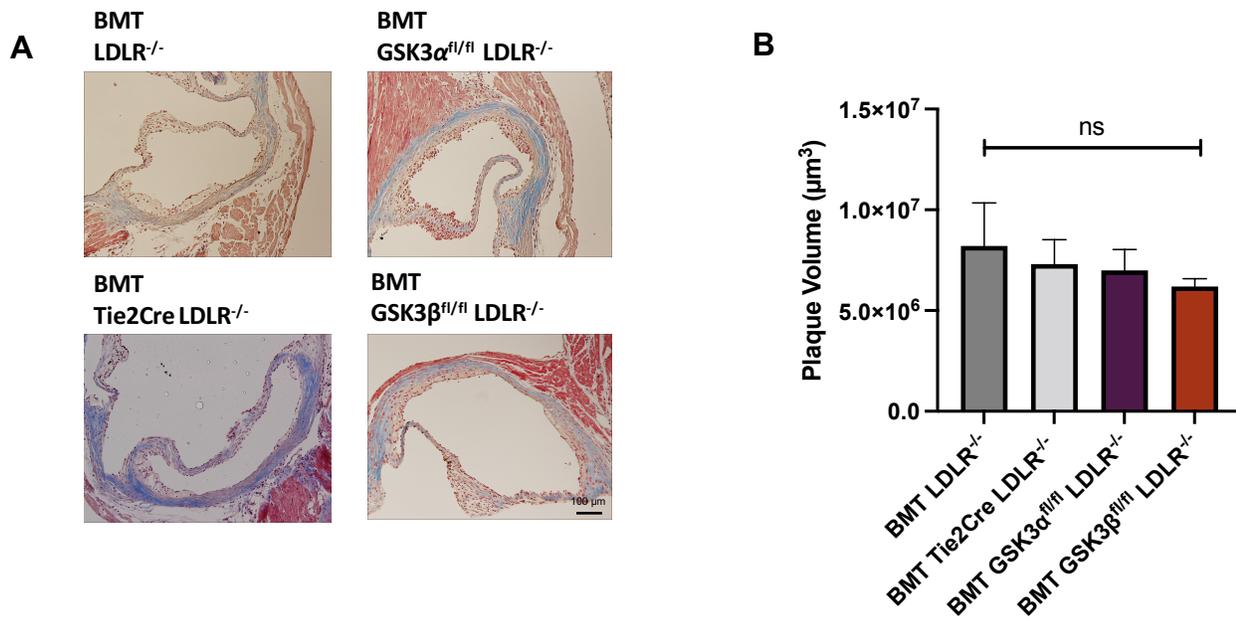
$\alpha^{\text{fl/fl}}$ Mouse model



$\beta^{\text{fl/fl}}$ Mouse model



Supplementary Figure S6: Atherosclerosis assessment comparison between non-BMT mice and BMT mice. Plaque volume quantification in the aortic sinus of non-BMT mice and BMT mice in LDLR^{-/-} mice, $\alpha^{\text{fl/fl}}$ mouse model and the $\beta^{\text{fl/fl}}$ mouse model. n=4-10, * $P < 0.05$, ** $P < 0.05$.

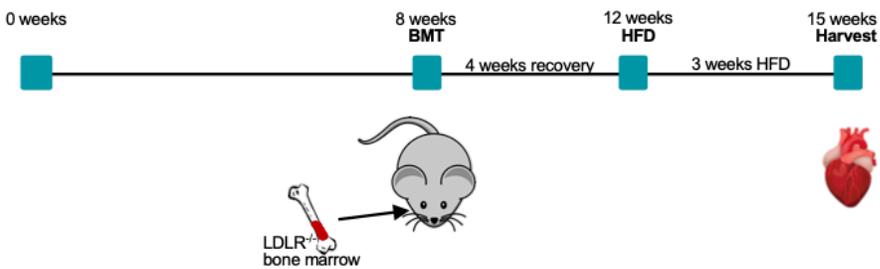


Supplementary Figure S7: Atherosclerosis assessment in control mice from the BMT endothelial GSK3 α/β knockout mouse model. **A** Representative images of Masson's Trichrome staining of aortic sinus cross sections from 15 week old control BMT LDLR^{-/-}, BMT Tie2Cre LDLR^{-/-}, BMT GSK3 α ^{fl/fl} LDLR^{-/-} and BMT GSK3 β ^{fl/fl} LDLR^{-/-} mice. Scale bar represents 100 μ m. **B** Plaque volume quantification in the aortic sinus of control mice, n=4-9.

A Endothelial/macrophage GSK3 α or GSK3 β knockout mouse model



B BMT- Macrophage GSK3 α or GSK3 β knockout mouse model



Supplementary Figure S8: Experimental design. **A** Experimental plan for analysis of endothelial/macrophage GSK3 knockout mice. Genotyped mice were placed on a HFD for 3 weeks and harvested at 15 weeks of age. **B** Experimental plan for BMT endothelial GSK3 α / β knockout mice. Genotyped mice received a BMT at 8 weeks of age. After 4 weeks of recovery the mice were placed on a HFD for 3 weeks and harvested at 15 weeks of age. *Figure created using Biorender.com 2022.*