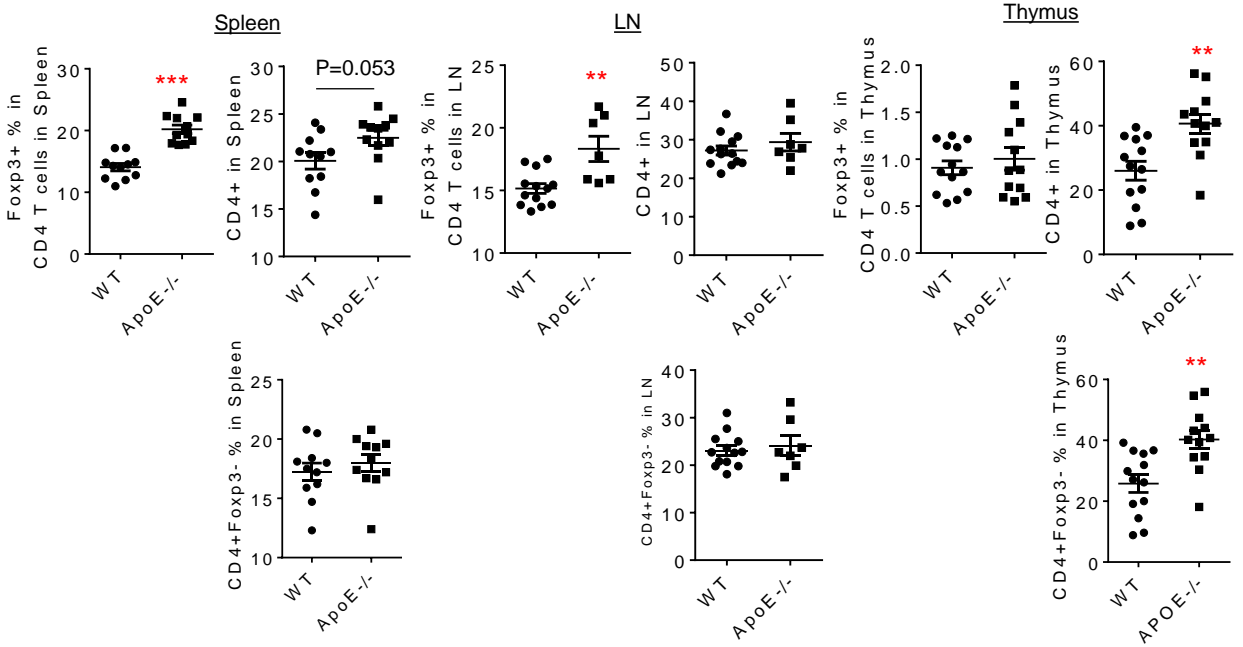
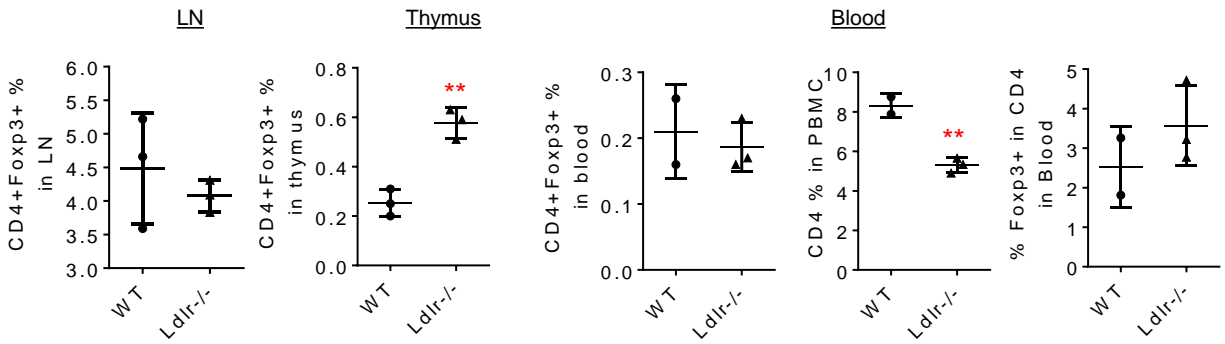


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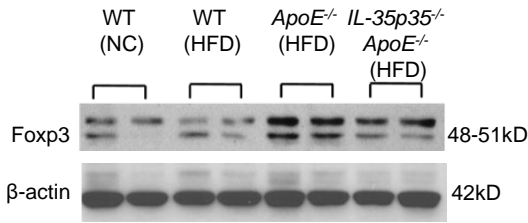


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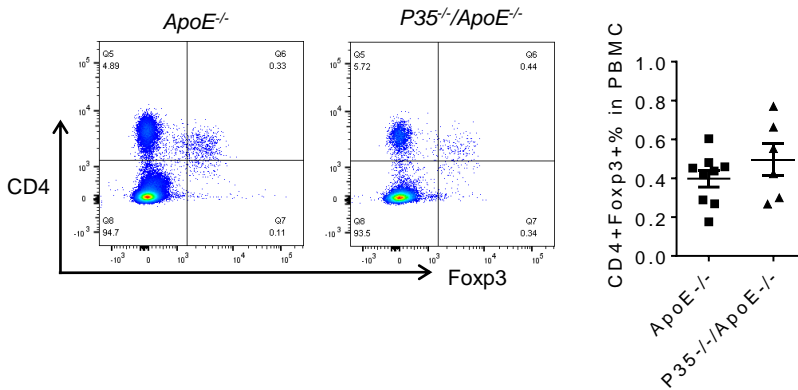


Supplementary Figure 1. The distribution of CD4⁺Foxp3⁺ and CD4⁺Foxp3⁻ T cells in spleen, lymph node and thymus from *ApoE*^{-/-} mice and *Ldlr*^{-/-} mice. (A) The *ApoE*^{-/-} mice of 20 weeks old fed with high fat diet (HFD) for 12 weeks were investigated. The flow cytometry detections were performed for analyzing the populations of CD4⁺ T cells and Foxp3⁺ Tregs, respectively (n=10). (B) The flow cytometry detections were performed for analyzing the populations of CD4⁺ T cells and Foxp3⁺ Tregs in The *Ldlr*^{-/-} mice of 20 weeks old fed with HFD for 12 weeks were investigated, respectively (n=3) (T-test, *p<0.05, ** p<0.01).

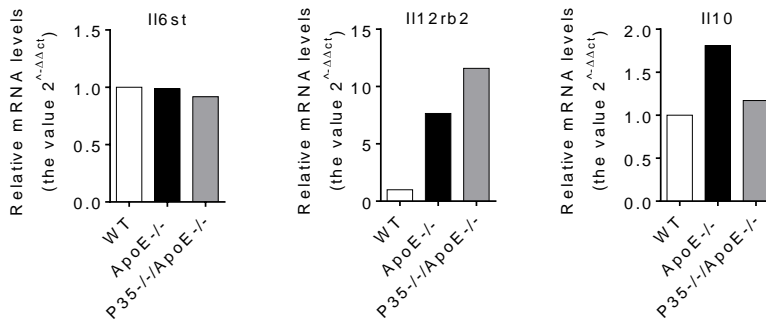
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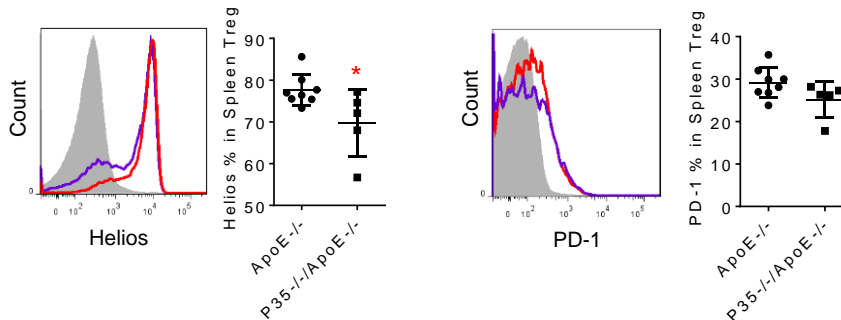
B



C



D



Supplementary Figure 2. Anti-inflammatory immune checkpoint receptors, such as programmed death-1 (PD-1) and Helios (*Ikzf2*) are found in a decrease trend in splenic Tregs in the deficiency of *IL-35p35*. (A) Western blots were performed in detecting Foxp3 protein levels in spleens from four groups of mice; (B) Blood Treg in *ApoE*^{-/-} mice were not significantly changed by *IL-35p35* deficiency. The *IL-35p35*^{-/-}/*ApoE*^{-/-} mice of 20 weeks old ($n = 8$) and *APOE*^{-/-} mice ($n=6$) fed with high fat diet for 12 weeks were investigated. (C) Real-time PCRs were performed for detecting *Il6st* (encodes CD130), *Il12rb2*, and *Il10* gene expression levels in pooled aorta samples ($n = 5$ /group) from *ApoE*^{-/-} and *IL-35p35*^{-/-}/*ApoE*^{-/-} mice, respectively. (D) The expressions of Helios and PD-1 were examined in Treg subpopulations in freshly isolated spleens from *IL-35p35*^{-/-} *ApoE*^{-/-} mice ($n=5$) and *ApoE*^{-/-} mice, respectively, ($n=8$) by flow cytometry (T-test, * $p<0.05$).