

Lack of apoA-I in apoEKO mice causes skin xanthomas, worsening of inflammation and increased coronary atherosclerosis in the absence of hyperlipidemia

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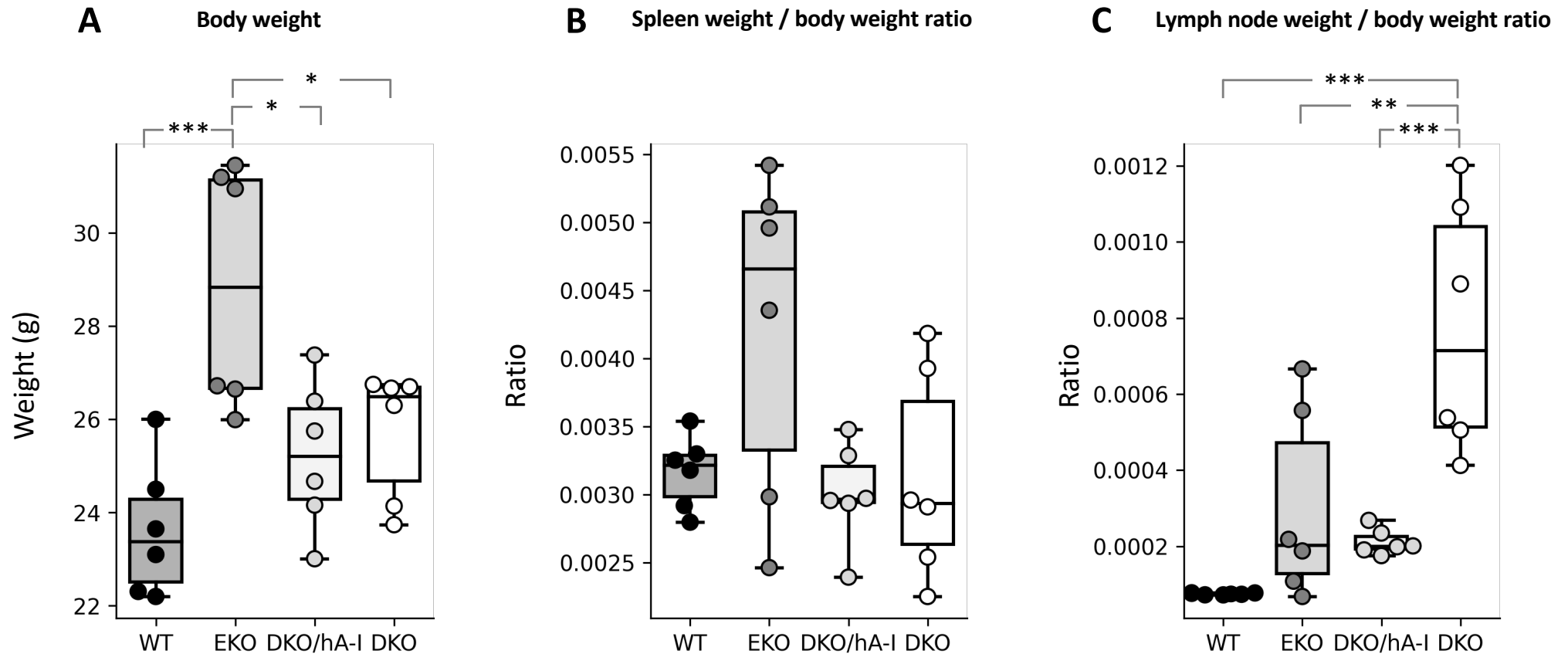
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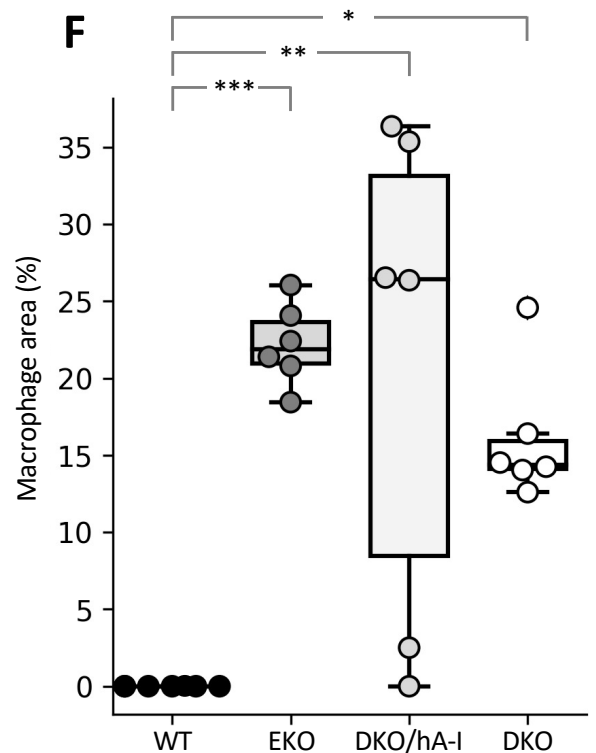
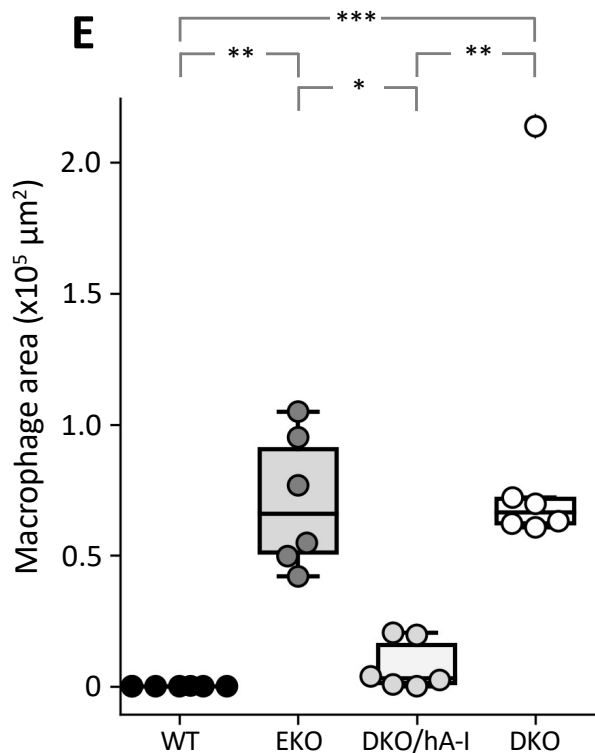
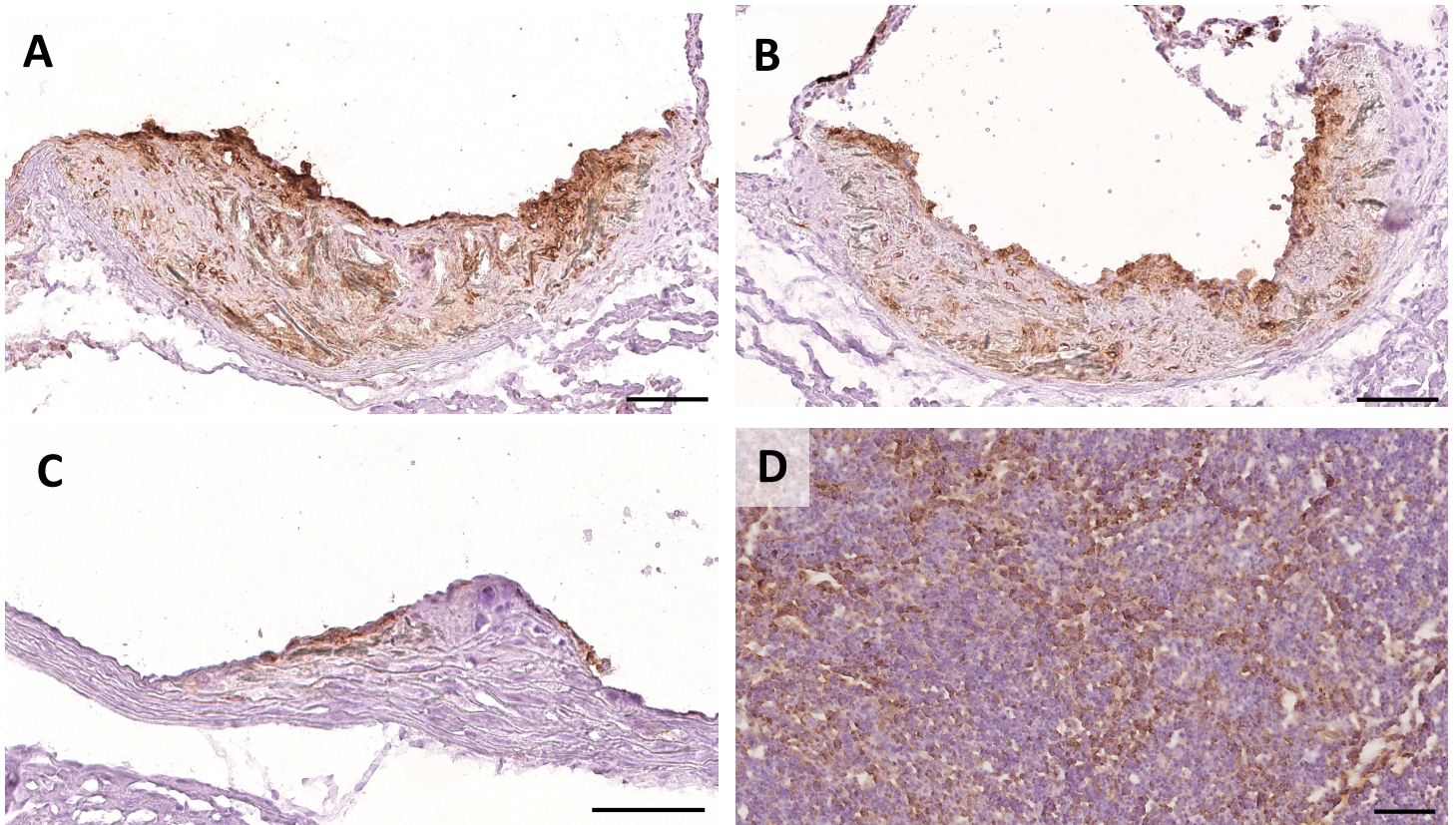
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Figure S1



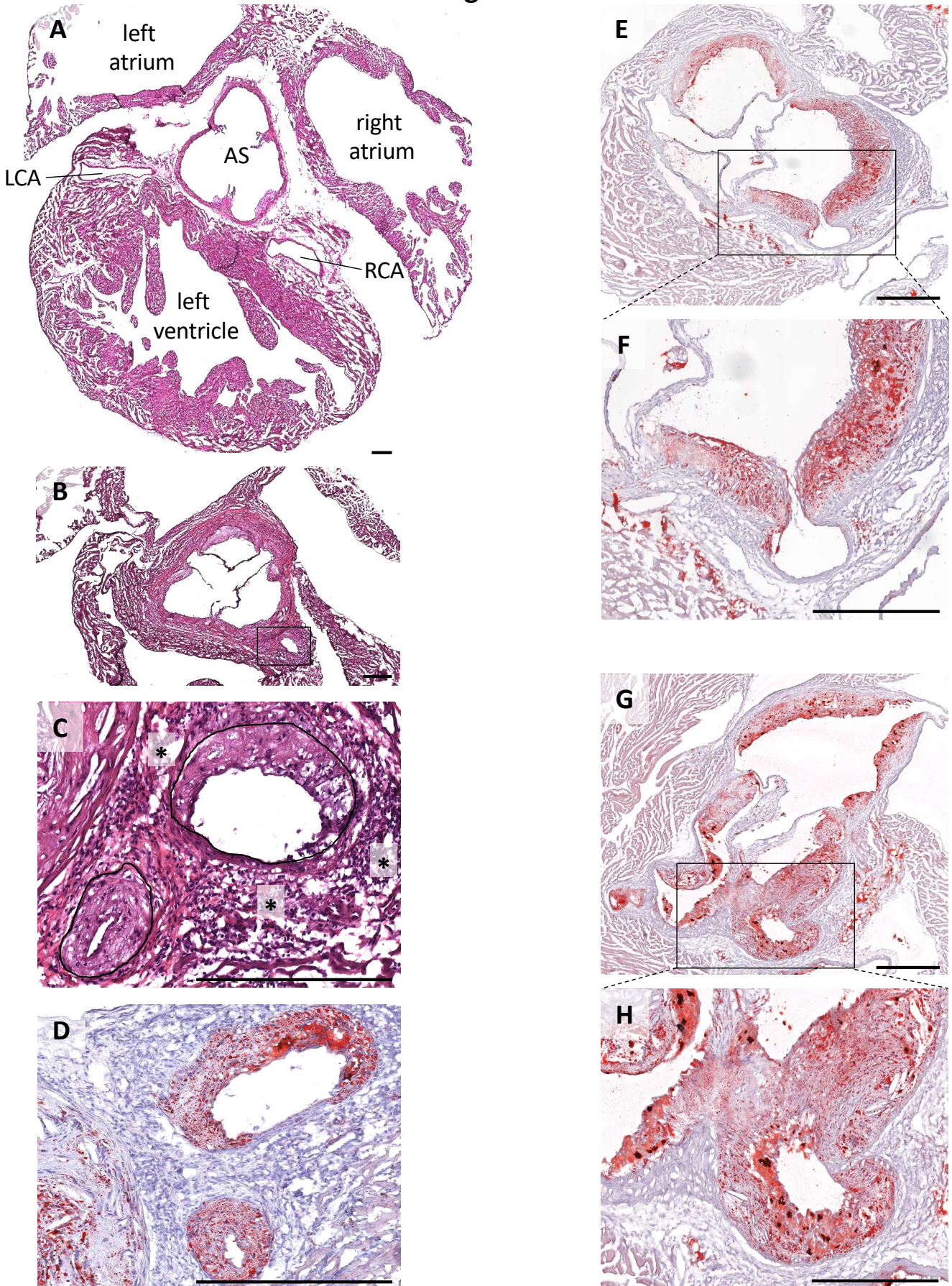
Body weight (**A**), spleen weight/body weight ratio (**B**), lymph node weight/body weight ratio (**C**) in the four genotypes, $n=6$. * $p<0.05$; ** $p<0.01$; *** $p<0.001$. Statistically significant differences were determined with ANOVA followed by Tukey's post-hoc test. The upper and lower ends of the boxes indicate the 25th and 75th percentiles, respectively. The length of the box shows the interquartile range within which 50% of the values are located. The solid grey lines denote the median. Exact adjusted p -values are shown in Table S1.

Figure S2



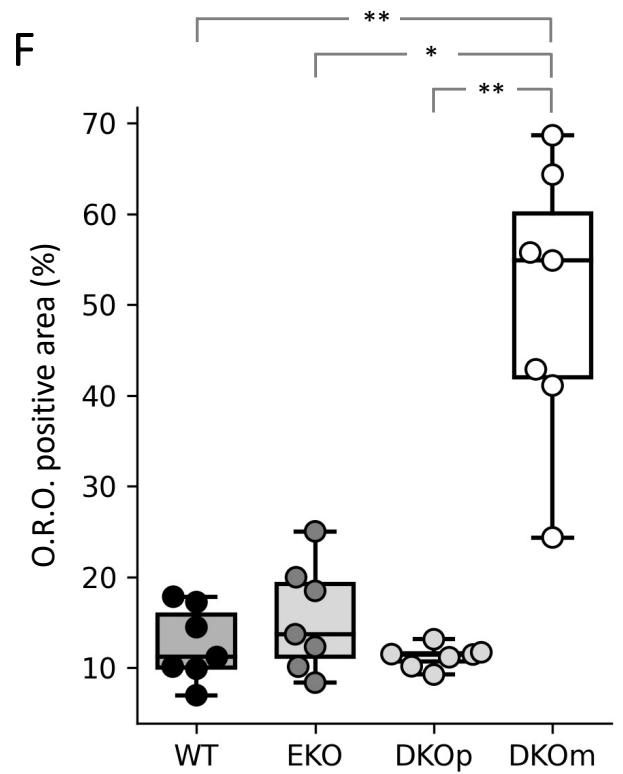
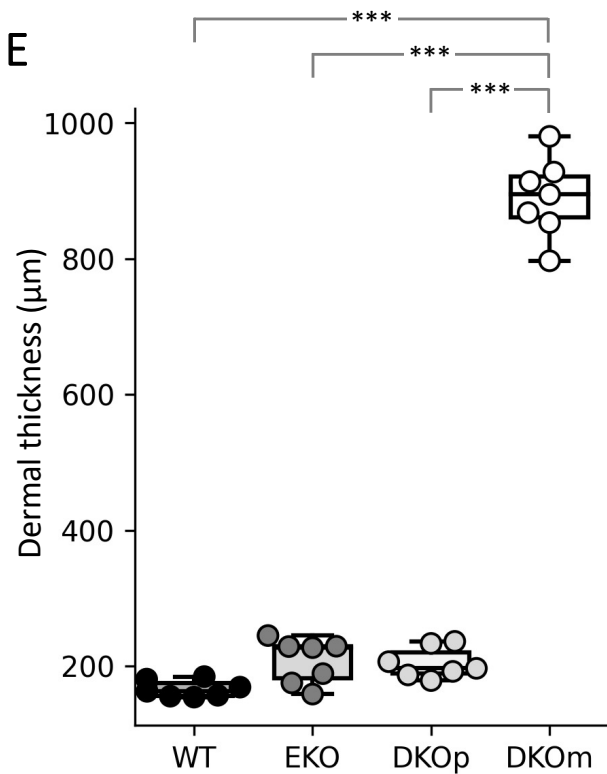
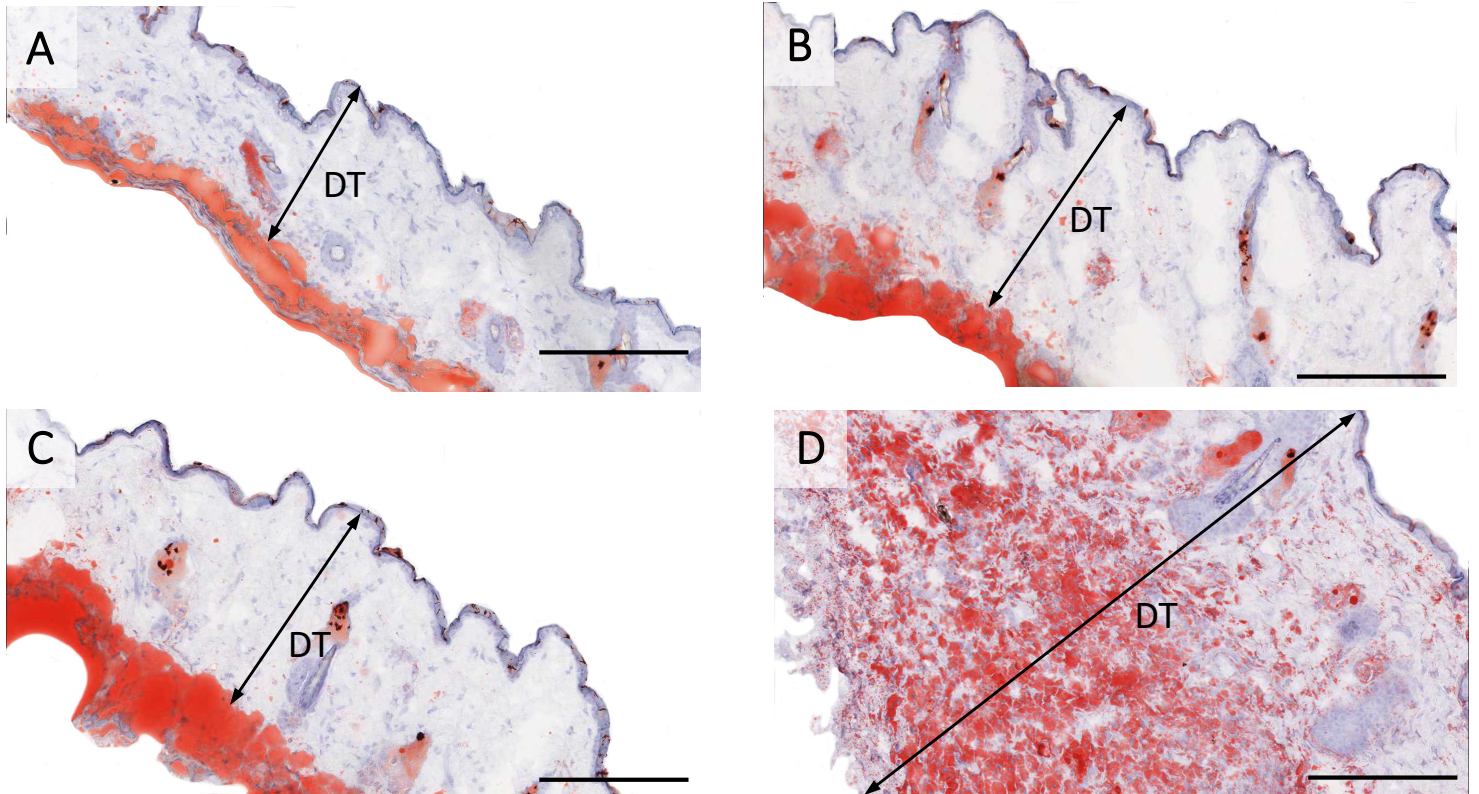
Immunohistochemical staining of macrophages in the aortic sinus of DKO (A), EKO (B) and DKO/hA-I (C). The area occupied by macrophages was significantly reduced in DKO/hA-I vs EKO and DKO mice (E), although plaque % composition was not different among EKO, DKO/hA-I and DKO (F). A positive control from a lymph node cryosection is shown (D). Statistically significant differences were determined by ANOVA followed by Tukey's post-hoc. * $p < 0.05$; ** $p < 0.01$; *** $p < 0.001$. Bar length: 100 μm . The upper and lower ends of the boxes indicate the 25th and 75th percentiles, respectively. The length of the box shows the interquartile range within which 50% of the values are located. The solid grey lines denote the median. Exact adjusted p-values are shown in Table S1.

Figure S4



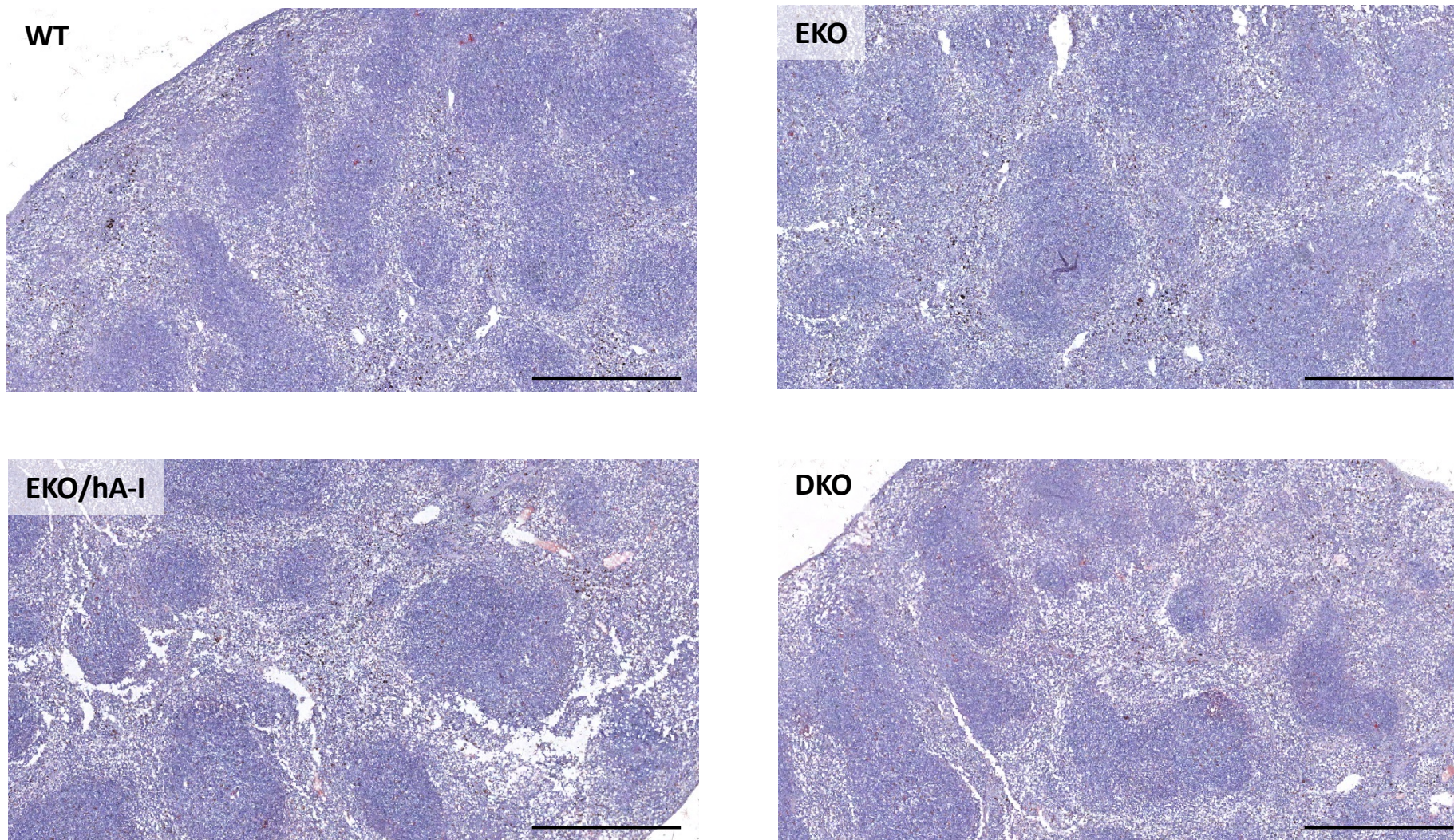
Representative H&E staining of the aortic sinus (AS) with the origin of left (LCA) and right coronary arteries (RCA) (A). Aortic sinus from a DKO mouse with coronary plaques (B). Magnifications showing lymphocytes infiltrating the myocardium around the coronary arteries (asterisks, C) and neutral lipid accumulation in coronary plaques by O.R.O. staining (D). Representative O.R.O. stained pictures of plaque development at the ostia of the right coronary artery at the aortic root of EKO (E) and DKO mice (G). Magnifications of the same plaques are shown (F, H). Bar: 250 μ m.

Figure S5



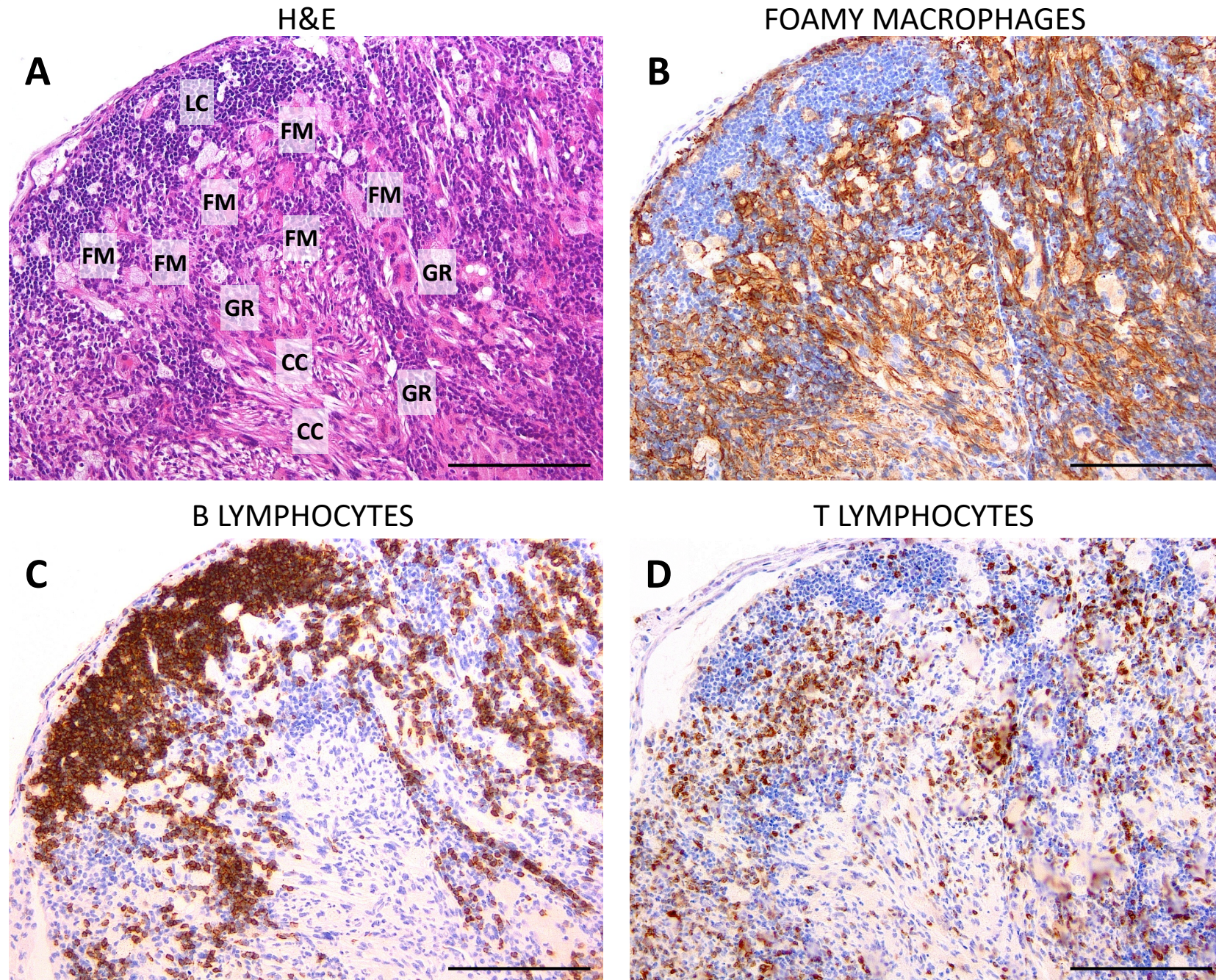
Representative photomicrographs of skin cryosections stained with O.R.O. from WT (A), EKO (B), DKO/hA-I (C) and DKO (D). The epidermis was unaltered regardless of the genotype, whereas, only in DKO, the dermis was thickened (E) and showed increased neutral lipid deposition (F). DT = dermal thickness (n=7). Statistically significant differences were determined by ANOVA followed by Tukey's post-hoc in (E) and by Kruskal-Wallis followed by Dunn's post-hoc test in (F). *p<0.05; **p<0.01; ***p<0.001. Bar length = 100 µm. The upper and lower ends of the boxes indicate the 25th and 75th percentiles, respectively. The length of the box shows the interquartile range within which 50% of the values are located. The solid grey lines denote the median. Exact adjusted p-values are shown in Table S1.

Figure S6

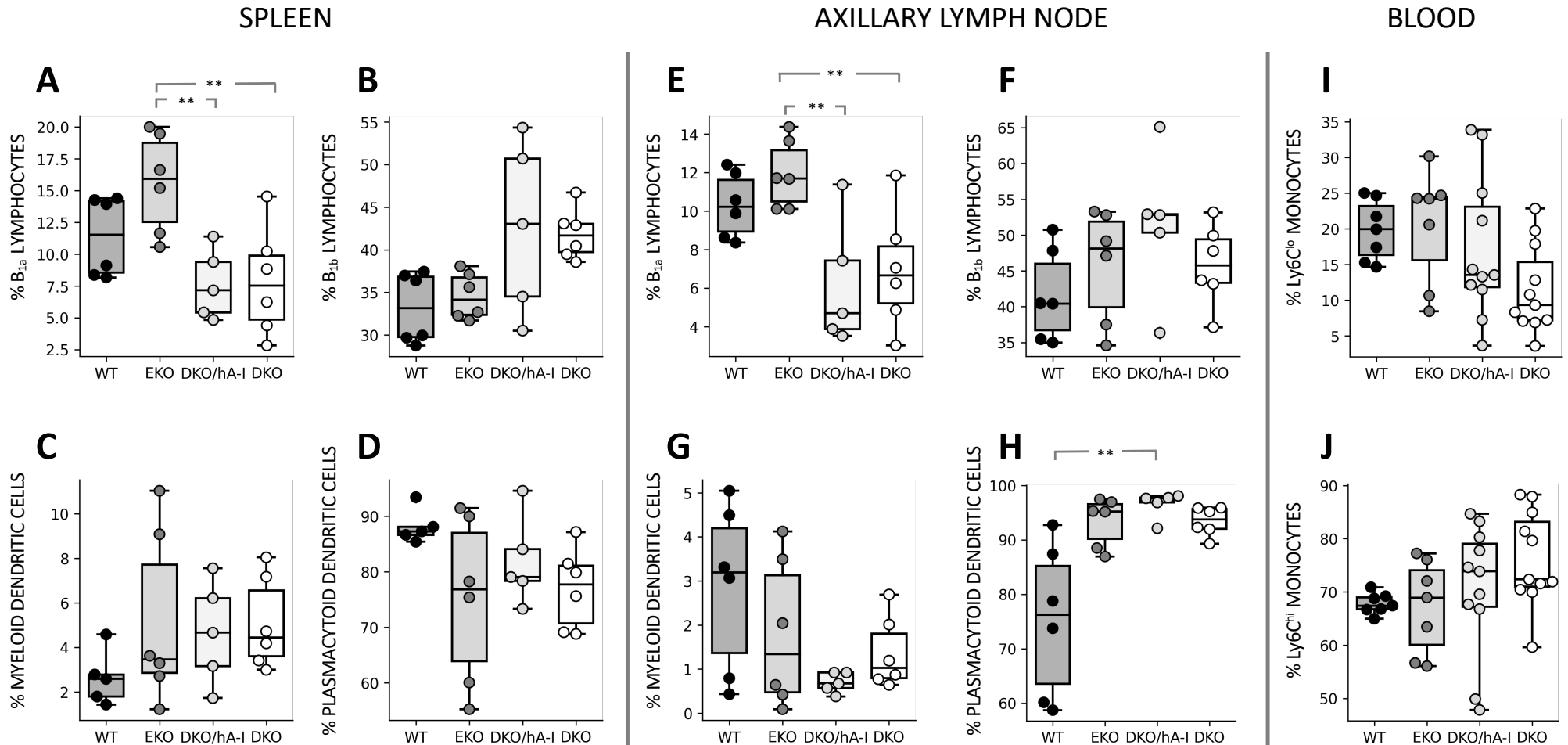


Representative photomicrographs of spleen cryosections stained with Oil Red O. Spleen histology and neutral lipid deposition were unaffected by genotype. Bar length = 200 μm).

Figure S7



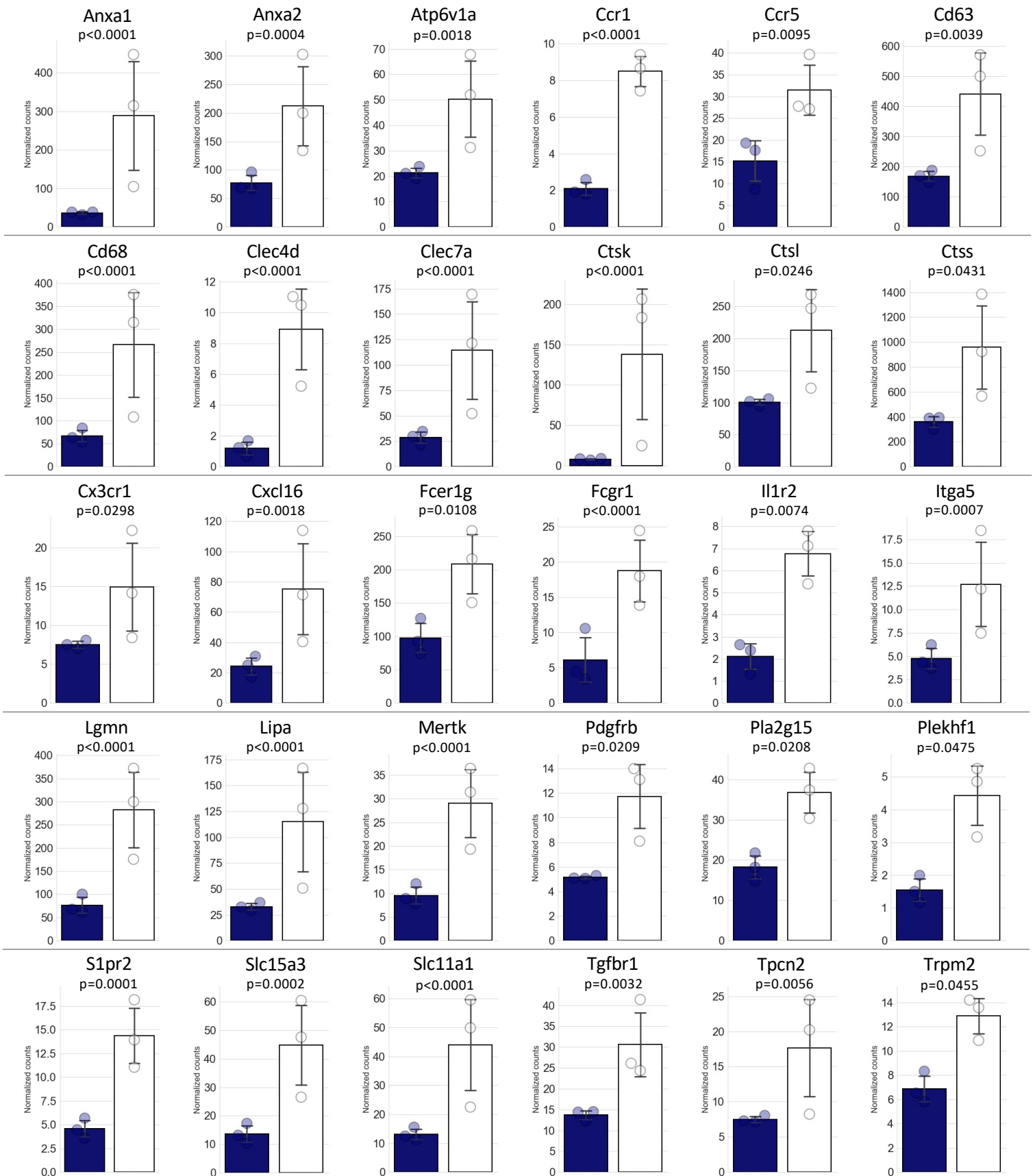
Histological and immunohistochemical characterization of a DKO lymph node. H&E staining (**A**) FM: foamy macrophages; LC: lymphoid cells; GR: granulomatous reactions; CC: cholesterol crystals. Foamy macrophages (IBA1⁺ cells; **B**), B lymphocytes (B220⁺ cells; **C**), T lymphocytes (CD3⁺ cells; **D**). Bar length=100 μ m.

Figure S9

FACS analyses results for immune cells subsets in spleen (left), axillary lymph node (middle) and blood (right) in the four genotypes. *p<0.05; **p<0.01. Statistically significant differences were determined with ANOVA followed by Tukey's post-hoc test except for **H**, where Kruskal-Wallis followed by Dunn's post-hoc was used. The upper and lower ends of the boxes indicate the 25th and 75th percentiles, respectively. The length of the box shows the interquartile range within which 50% of the values are located. The solid grey lines denote the median. Exact adjusted p-values are shown in Table S1.

Figure S10

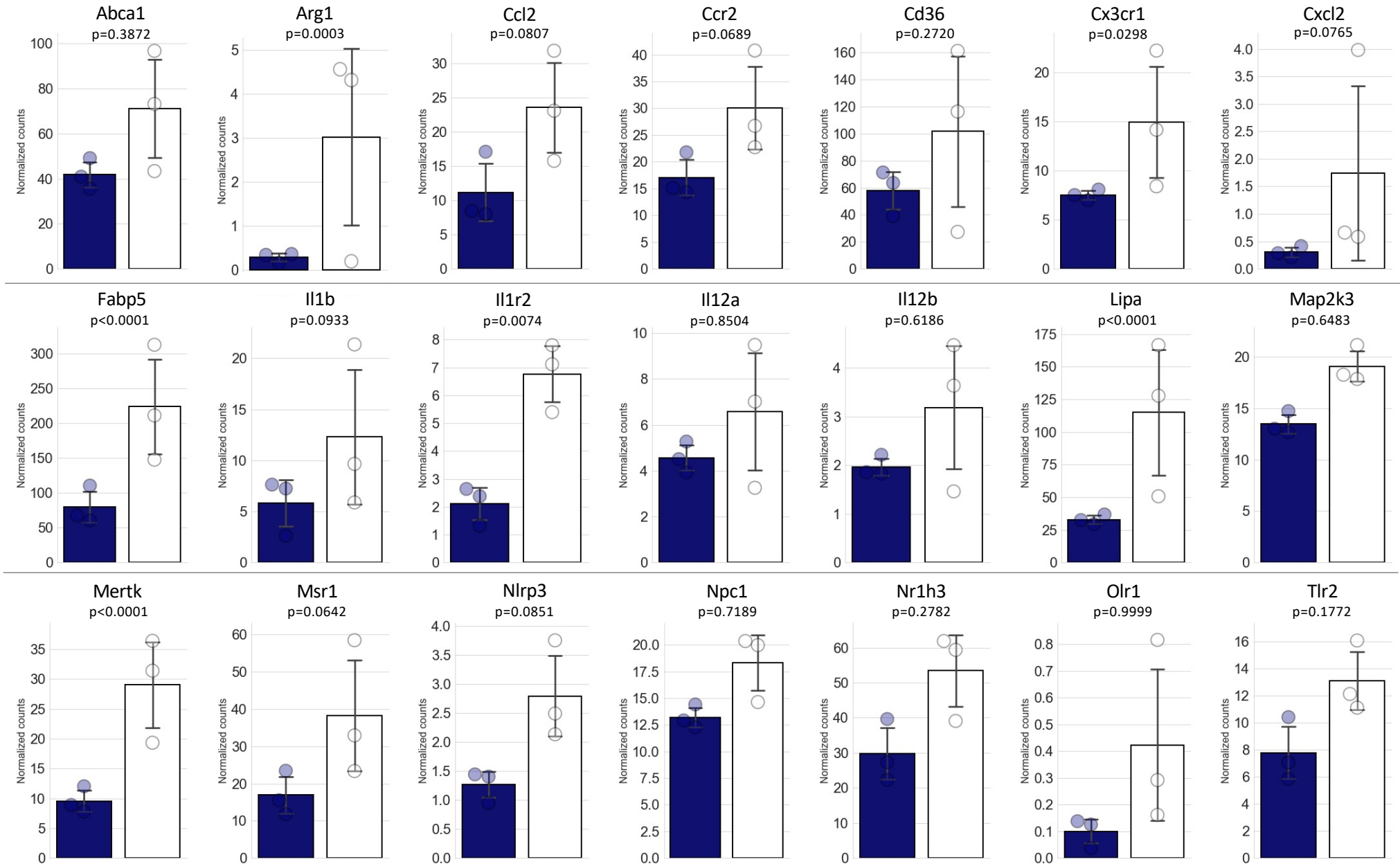
■ EKO/hA-I □ DKO



RNAseq expression levels (normalized counts) is charted for selected macrophage markers that show increased expression in DKO mice. Transcript abundance estimation and differential expression analyses were performed using the standard Bowtie-Tophat-Cuffdiff pipeline.

Figure S11

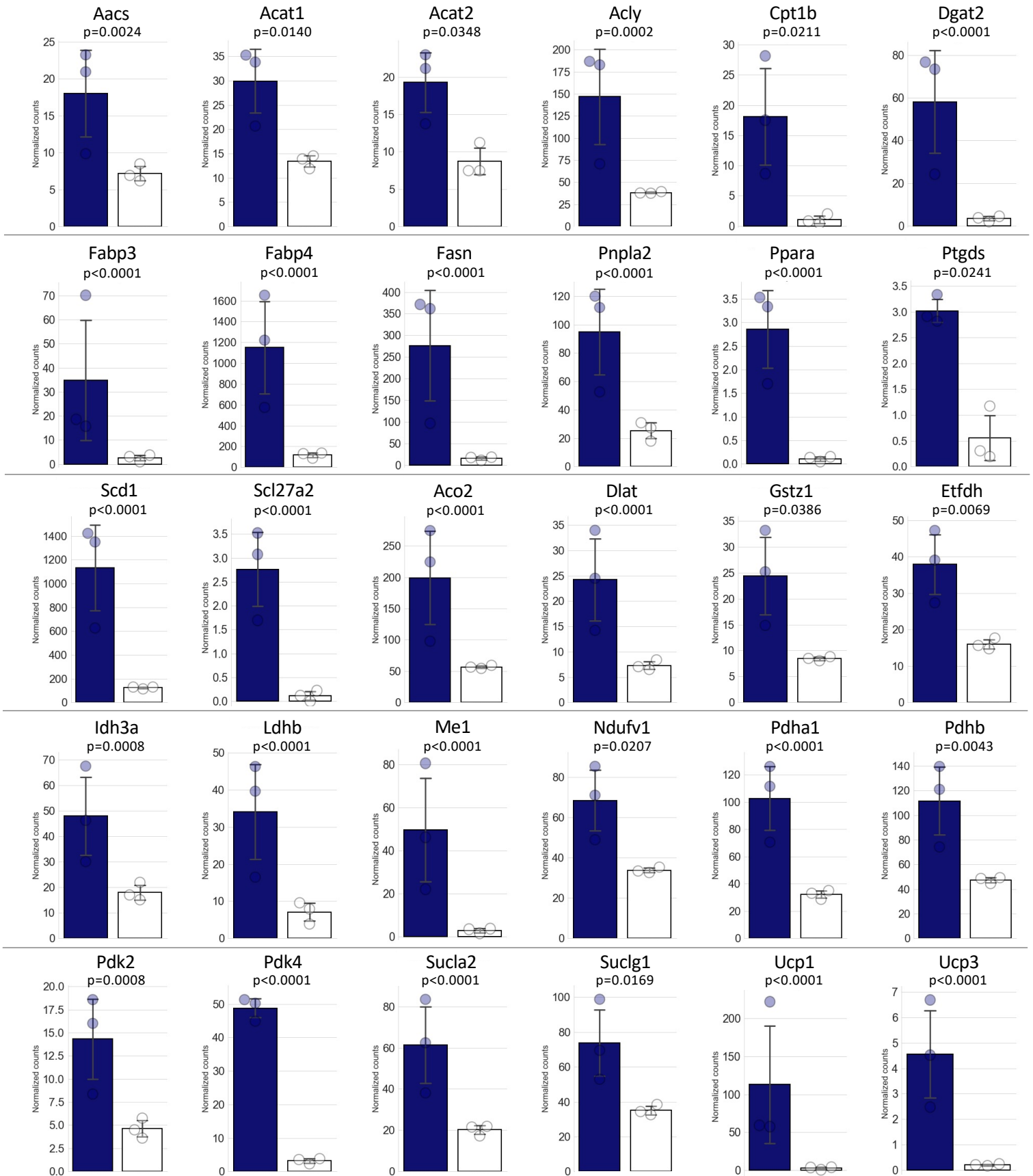
■ EKO/hA-I □ DKO



RNAseq expression levels (normalized counts) is charted for selected foam cell marker that show increased expression in DKO mice. Transcript abundance estimation and differential expression analyses were performed using the standard Bowtie-Tophat-Cuffdiff pipeline.

Figure S12

■ EKO/hA-I □ DKO



RNAseq expression levels (normalized counts) is charted for selected metabolic genes that show increased expression in DKO mice. Transcript abundance estimation and differential expression analyses were performed using the standard Bowtie-Tophat-Cuffdiff pipeline.

Figure S13

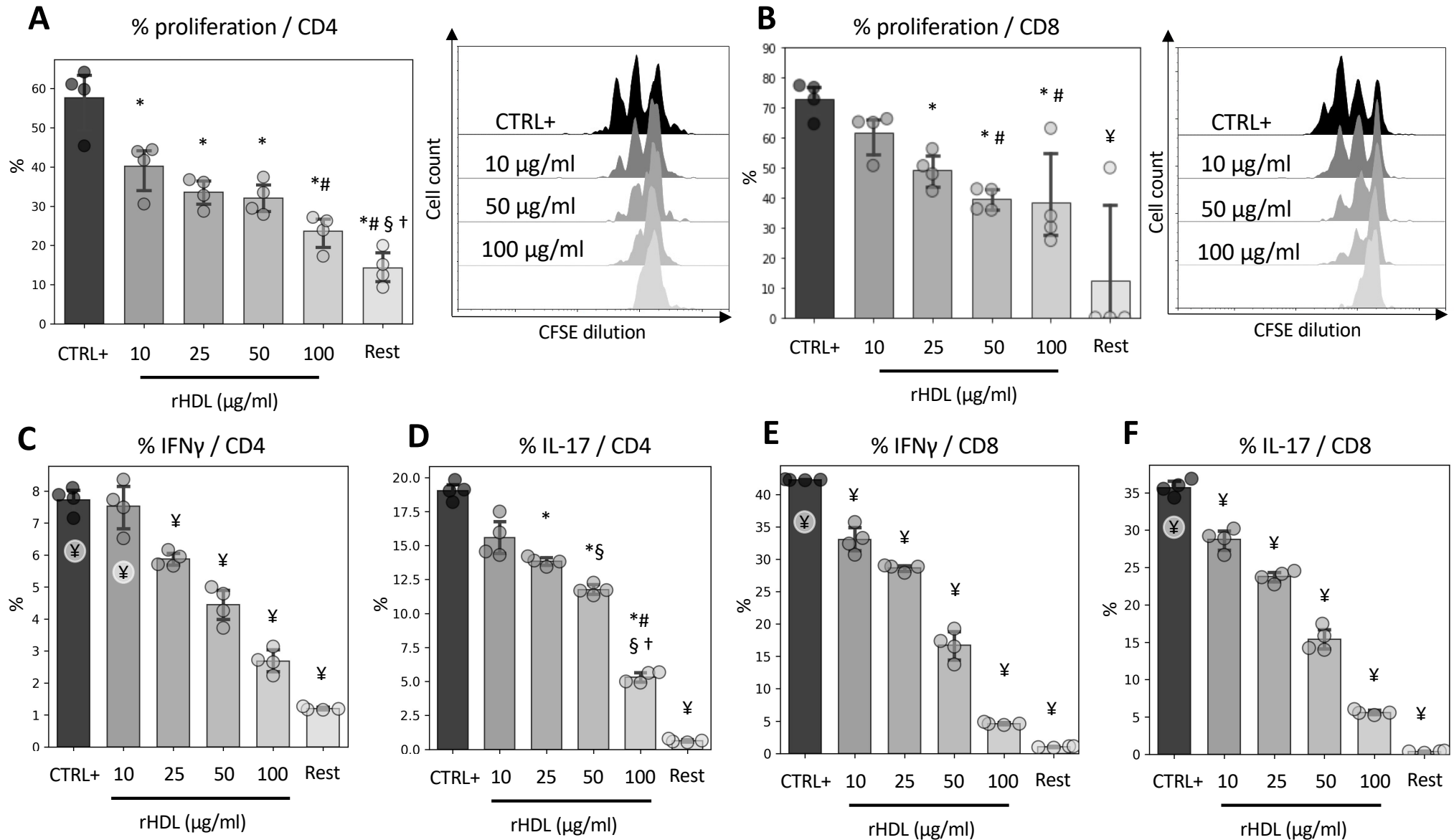
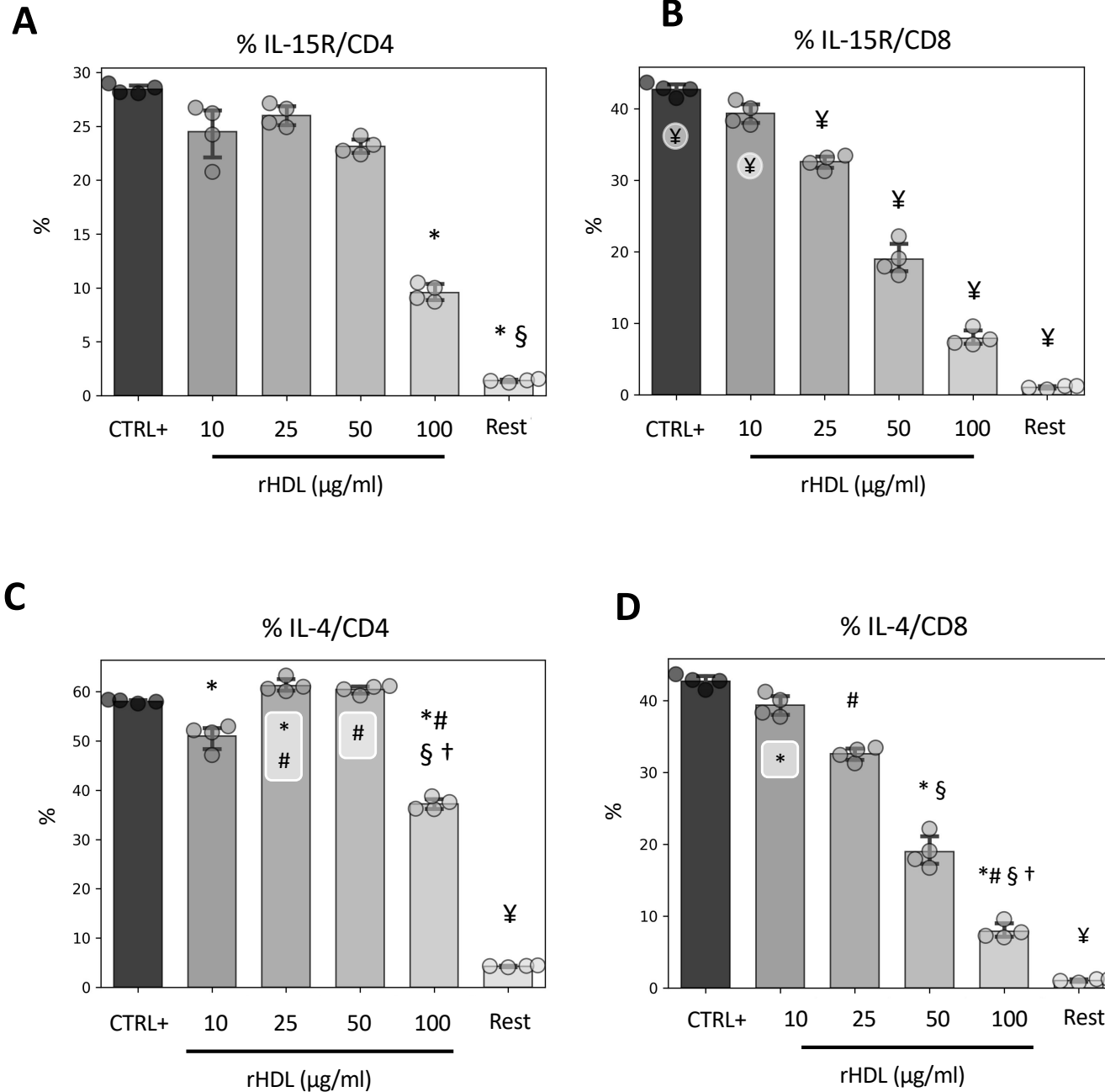


Figure 1. rHDL dampens T cell proliferation and cytokine production.

A-B: Impact of increasing concentration of rHDL (0, 10, 25, 50 and 100 $\mu\text{g/ml}$) on CD4+ and CD8+ T cell proliferation after 4 days of activation compared to resting (not stimulated sample). Representative histograms of CFSE dilution from flow cytometry analysis are shown.

C-F: Percentage of activated CD4+ or CD8+ T cells from apoE KO mice expressing interferon γ (IFN γ) (CD4 Panel C, and CD8 Panel E) or IL-17 (CD4 Panel D, and CD8 Panel F) T following incubation with rHDL (0, 10, 25, 50 and 100 $\mu\text{g/ml}$). Data for resting (rest) T cells are also presented (last column of each graph). $P < 0.05$: * vs CTRL+; # vs 10; § vs 25; † vs 50; ‡ vs 100; ¥: vs everything else. Exact adjusted p-values are shown in Table S1.

Figure S14

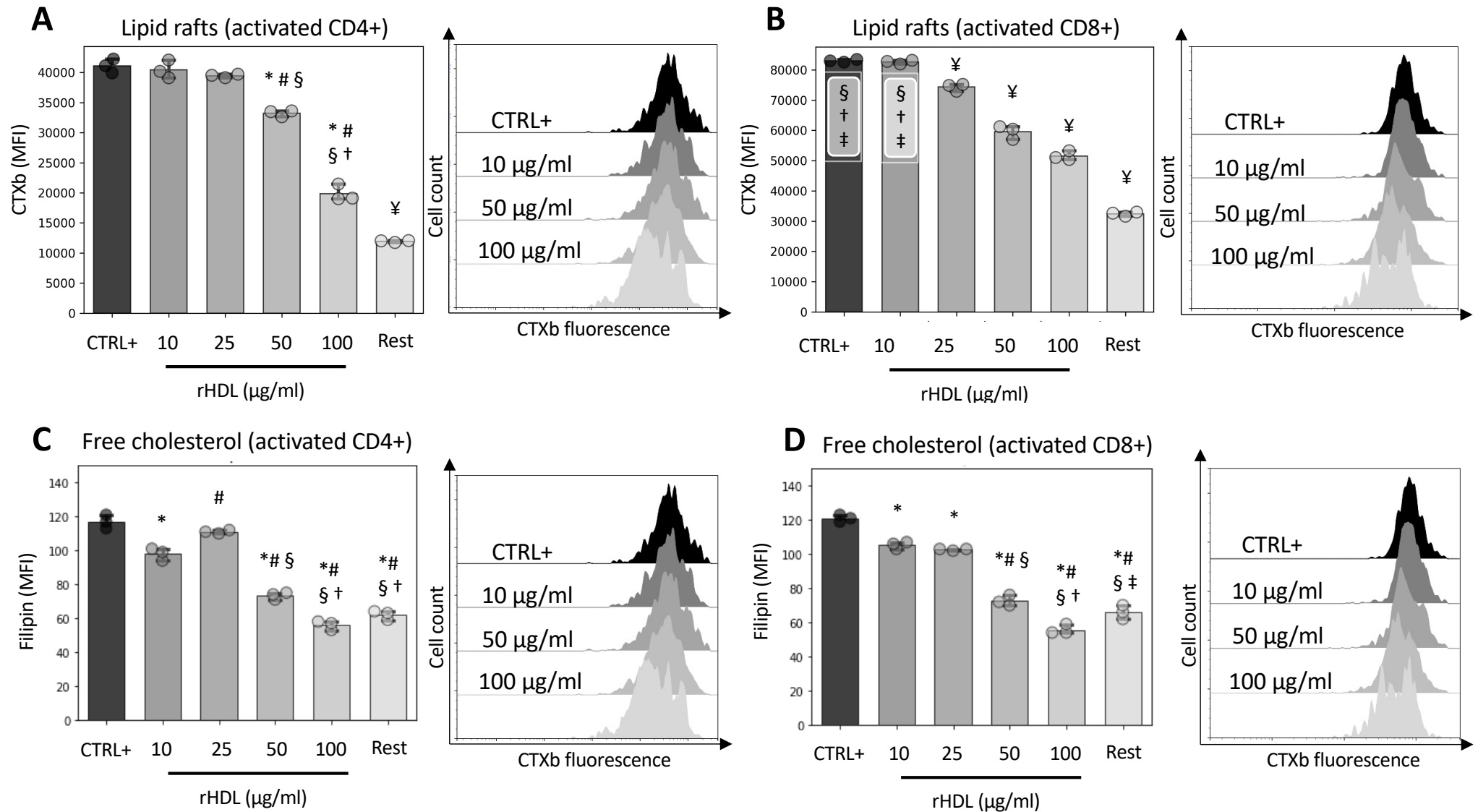


Impact of rHDL on IL15R and IL-4 production in T cells.

Percentage of activated CD4+ or CD8+ T cells from apoE KO mice expressing IL-15Ra, an indicator of IL-15 production as this interleukin is retained inside the cell and only secreted in complex with IL-15Ra, (CD4 Panel A, and CD8 Panel B) or IL-4 (CD4 Panel C, and CD8 Panel D) T following incubation with rHDL (0, 10, 25, 50 and 100 µg/ml). Data for resting (rest) T cells are also presented (last column of each graph).

P < 0.05: * vs CTRL+; # vs 10; § vs 25; † vs 50; ‡ vs 100; ¥: vs everything else. Exact adjusted p-values are shown in Table S1.

Figure S15

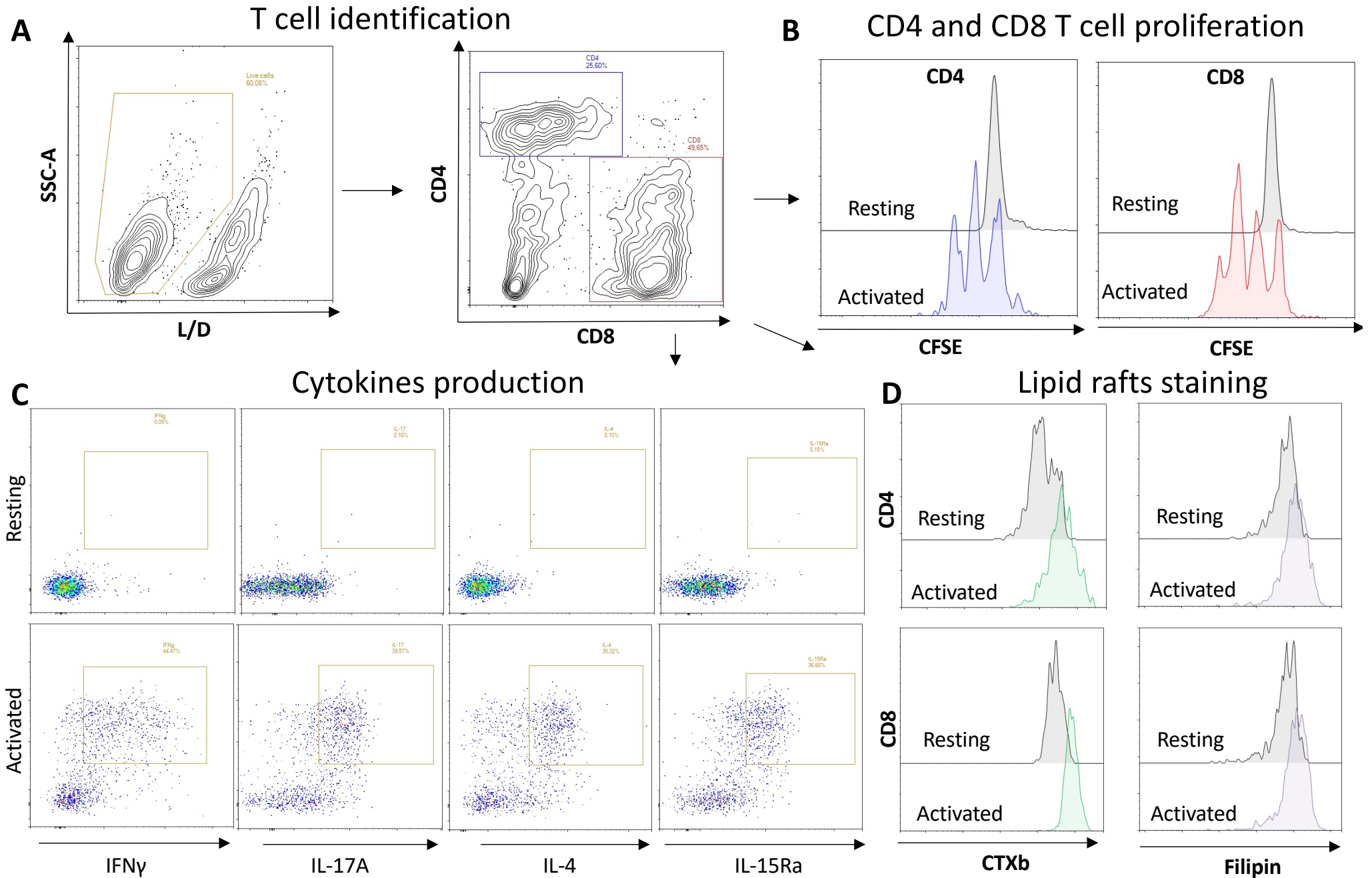


ApoA-I reduces lipid rafts and cholesterol abundance in T cells.

Cholera toxin-subunit B (CTXb, ligand of gangliosides enriched in lipid rafts) staining in activated CD4+ (A) and CD8+ (B) T cells from apoE KO mice following incubation with rHDL (0, 10, 25, 50 and 100 µg/ml). Data for resting (rest) T cells are also presented (last column of each graph). Representative histograms from flow cytometry analysis are shown.

Filipin staining (a marker of free cholesterol content) in activated CD4+ (C) and CD8+ (D) T cells from apoE KO mice following incubation with rHDL (0, 10, 25, 50 and 100 µg/ml). Data for resting (rest) T cells are also presented. Representative histograms from flow cytometry analysis are shown. MFI: mean fluorescence intensity. P < 0.05: * vs CTRL+; # vs 10; § vs 25; † vs 50; ‡ vs 100; ¥: vs everything else. Exact adjusted p-values are shown in Table S1.

Figure S16



Gating strategy for flow cytometry analysis. **A** - Live cells were discriminated as negative for live/dead staining and identified as CD4+ or CD8+ T cells; **B** - CD4+ and CD8+ T proliferation was evaluated by tracking CFSE dilution in resting and activated T cells; **C** - Cytokines production was detected by intracellular staining for IFN γ , IL-17A, IL-4 and IL-15Ra in resting and activated T cells; **D** - Stainings for Cholera toxin subunit B (CTXb) and filipin were performed in resting and activated CD4+ and CD8+ T cells.

Table S1

Figure 1A	Total Cholesterol					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-248.5	-281,3 to -215,7	Yes	****	<0,00000001
	WT vs. DKO/hA-I	-217	-249,8 to -184,2	Yes	****	<0,00000001
	WT vs. DKO	-18.52	-51,31 to 14,28	No	ns	0.42728105
	EKO vs. DKO/hA-I	31.5	-1,300 to 64,29	No	ns	0.06324018
EKO vs. DKO	230	197,2 to 262,8	Yes	****	<0,00000001	
DKO/hA-I vs. DKO	198.5	165,7 to 231,3	Yes	****	<0,00000001	
Figure 1B	Plasma apoA-I					
	Welch's ANOVA +Dunnnett's					
	Dunnnett's T3 multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	46.87	36.37 to 57.37	Yes	****	<0.0001
WT vs. DKOp	-56.87	-91.38 to -22.35	Yes	**	0.0014	
EKO vs. DKOp	-103.7	-138.3 to -69.20	Yes	****	<0.0001	
Figure 1C	Plasma HDL-C					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	55.65	45,54 to 65,75	Yes	****	<0,00000001
	WT vs. DKO/hA-I	18.12	8,022 to 28,22	Yes	***	0.00020413
	WT vs. DKO	75.47	65,37 to 85,57	Yes	****	<0,00000001
	EKO vs. DKO/hA-I	-37.52	-47,62 to -27,42	Yes	****	<0,00000001
EKO vs. DKO	19.82	9,723 to 29,92	Yes	****	0.00005889	
DKO/hA-I vs. DKO	57.35	47,25 to 67,45	Yes	****	<0,00000001	
Text	Plasma NON HDL-C					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-304.1	-333,3 to -275,0	Yes	****	<0,00000001
	WT vs. DKO/hA-I	-235.1	-264,2 to -206,0	Yes	****	<0,00000001
	WT vs. DKO	-93.99	-123,1 to -64,88	Yes	****	<0,00000001
	EKO vs. DKO/hA-I	69.02	39,91 to 98,12	Yes	****	0.00000297
EKO vs. DKO	210.2	181,1 to 239,3	Yes	****	<0,00000001	
DKO/hA-I vs. DKO	141.1	112,0 to 170,2	Yes	****	<0,00000001	
Figure 2K	Aortic sinus plaque area					
	Welch's ANOVA +Dunnnett's					
	Dunnnett's T3 multiple comparisons test	Mean Diff.	95,00% CI of diff.	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-291815	-418782 to -164848	Yes	***	0.0006
	WT vs. DKOp	-32902	-76228 to 10424	No	ns	0.0722
	WT vs. DKOm	-545253	-675156 to -415351	Yes	****	<0.0001
	EKO vs. DKOp	258913	135918 to 381908	Yes	***	0.0005
EKO vs. DKOm	-253439	-394156 to -112721	Yes	***	0.0004	
DKOp vs. DKOm	-512351	-637896 to -386806	Yes	****	<0.0001	
Figure 2L	Number of coronary plaques					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-4.588	-8,249 to -0,9263	Yes	*	0.01191354
	WT vs. DKO/hA-I	0	-3,661 to 3,661	No	ns	>0,99999999
	WT vs. DKO	-8.282	-11,94 to -4,620	Yes	****	0.00004161
	EKO vs. DKO/hA-I	4.588	0,9263 to 8,249	Yes	*	0.01191354
EKO vs. DKO	-3.694	-7,355 to -0,03279	Yes	*	0.04759952	
DKO/hA-I vs. DKO	-8.282	-11,94 to -4,620	Yes	****	0.00004161	
Figure 2M	Coronary plaque length					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-200.7	-300,7 to -100,7	Yes	***	0.00016071
	WT vs. DKO/hA-I	0	-99,98 to 99,98	No	ns	>0,99999999
	WT vs. DKO	-229.7	-329,7 to -129,7	Yes	****	0.00003465
	EKO vs. DKO/hA-I	200.7	100,7 to 300,7	Yes	***	0.00016071
EKO vs. DKO	-28.99	-129,0 to 70,99	No	ns	0.83968018	
DKO/hA-I vs. DKO	-229.7	-329,7 to -129,7	Yes	****	0.00003465	
Figure 4A	Spleen weight					
	Kruskal-Wallis + Dunn's					
	Dunn's multiple comparisons test	Mean rank diff,	Significant?	Summary	Adjusted P Value	
	WT vs. EKO	-14.09	No	ns	0.11922781	
	WT vs. DKO/hA-I	4.958	No	ns	>0,99999999	
	WT vs. DKO	0.3173	No	ns	>0,99999999	
	EKO vs. DKO/hA-I	19.05	Yes	**	0.00367468	
EKO vs. DKO	14.41	Yes	*	0.0494134		
DKO/hA-I vs. DKO	-4.641	No	ns	>0,99999999		
Figure 4B	Axillary lymph node weight					
	Kruskal-Wallis + Dunn's					
	Dunn's multiple comparisons test	Mean rank diff,	Significant?	Summary	Adjusted P Value	
	WT vs. EKO	-14.75	No	ns	0.0885624	
	WT vs. DKO/hA-I	-13.29	No	ns	0.07636121	
	WT vs. DKO	-28.25	Yes	****	0.00000045	
	EKO vs. DKO/hA-I	1.458	No	ns	>0,99999999	
EKO vs. DKO	-13.5	No	ns	0.0825309		
DKO/hA-I vs. DKO	-14.96	Yes	**	0.00833983		

Figure 4F	ORO positive area axillary lymph node					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-2.106	-8,289 to 4,076	No	ns	0.77870738
	WT vs. DKO/hA-I	-0.389	-6,346 to 5,568	No	ns	0.99778903
	WT vs. DKO	-13.06	-19,24 to -6,873	Yes	****	0.00004254
	EKO vs. DKO/hA-I	1.717	-4,240 to 7,675	No	ns	0.85201309
	EKO vs. DKO	-10.95	-17,13 to -4,766	Yes	***	0.0003774
	DKO/hA-I vs. DKO	-12.67	-18,62 to -6,709	Yes	****	0.00003885
Figure 5A	% CD4 N Blood					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	14.71	-3,498 to 32,91	No	ns	0.14881276
	WT vs. DKO/hA-I	18.92	2,450 to 35,38	Yes	*	0.019178
	WT vs. DKO	43.07	27,10 to 59,03	Yes	****	0.00000011
	EKO vs. DKO/hA-I	4.21	-12,26 to 20,68	No	ns	0.89997618
	EKO vs. DKO	28.36	12,39 to 44,33	Yes	***	0.00017725
	DKO/hA-I vs. DKO	24.15	10,20 to 38,10	Yes	***	0.00025379
Figure 5B	% CD4 EM Blood					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-10.89	-26,17 to 4,396	No	ns	0.23725878
	WT vs. DKO/hA-I	-13.5	-27,33 to 0,3231	No	ns	0.05764329
	WT vs. DKO	-33.12	-46,52 to -19,71	Yes	****	0.00000068
	EKO vs. DKO/hA-I	-2.614	-16,44 to 11,21	No	ns	0.95599853
	EKO vs. DKO	-22.23	-35,64 to -8,824	Yes	***	0.00044952
	DKO/hA-I vs. DKO	-19.62	-31,33 to -7,901	Yes	***	0.00039589
Figure 5C	% CD4 CM Blood					
	Kruskal-Wallis + Dunn's					
	Dunn's multiple comparisons test	Mean rank diff,	Significant?	Summary	Adjusted P Value	
	WT vs. EKO	-0.2857	No	ns	>0,99999999	
	WT vs. DKO/hA-I	-11.52	No	ns	0.1922359	
	WT vs. DKO	-10.35	No	ns	0.28158347	
	EKO vs. DKO/hA-I	-11.23	No	ns	0.21930571	
	EKO vs. DKO	-10.07	No	ns	0.32009386	
	DKO/hA-I vs. DKO	1.168	No	ns	>0,99999999	
Figure 5D	% CD4 N Spleen					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	12.84	1,886 to 23,80	Yes	*	0.01670787
	WT vs. DKO/hA-I	21.22	11,18 to 31,27	Yes	****	0.00001627
	WT vs. DKO	32.16	21,87 to 42,45	Yes	****	0.00000001
	EKO vs. DKO/hA-I	8.38	-2,053 to 18,81	No	ns	0.15084634
	EKO vs. DKO	19.32	8,648 to 29,99	Yes	***	0.00016262
	DKO/hA-I vs. DKO	10.94	1,209 to 20,66	Yes	*	0.02287739
Figure 5E	% CD4 EM Spleen					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-9.806	-24,09 to 4,475	No	ns	0.26334662
	WT vs. DKO/hA-I	-21.43	-34,52 to -8,343	Yes	***	0.00060017
	WT vs. DKO	-34.03	-47,44 to -20,62	Yes	****	0.00000068
	EKO vs. DKO/hA-I	-11.63	-25,22 to 1,973	No	ns	0.11495074
	EKO vs. DKO	-24.22	-38,13 to -10,32	Yes	***	0.00027346
	DKO/hA-I vs. DKO	-12.6	-25,28 to 0,08111	Yes	*	0.04997055
Figure 5F	% CD4 CM Spleen					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-1.066	-4,810 to 2,679	No	ns	0.86565008
	WT vs. DKO/hA-I	-1.855	-5,287 to 1,577	No	ns	0.46754166
	WT vs. DKO	-0.7368	-4,252 to 2,779	No	ns	0.94026456
	EKO vs. DKO/hA-I	-0.7897	-4,355 to 2,776	No	ns	0.93054485
	EKO vs. DKO	0.3287	-3,317 to 3,975	No	ns	0.99470894
	DKO/hA-I vs. DKO	1.118	-2,206 to 4,443	No	ns	0.79710676
Figure 5G	% CD4 N Lymph node					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	9.716	-11,32 to 30,75	No	ns	0.59716251
	WT vs. DKO/hA-I	14.96	-4,225 to 34,15	No	ns	0.16976379
	WT vs. DKO	22.69	3,168 to 42,22	Yes	*	0.01783477
	EKO vs. DKO/hA-I	5.248	-13,03 to 23,53	No	ns	0.86263774
	EKO vs. DKO	12.98	-5,656 to 31,61	No	ns	0.25213376
	DKO/hA-I vs. DKO	7.728	-8,791 to 24,25	No	ns	0.58725915
Figure 5H	% CD4 EM Lymph node					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-4.744	-21,47 to 11,99	No	ns	0.86755454
	WT vs. DKO/hA-I	-8.721	-23,98 to 6,541	No	ns	0.42062802
	WT vs. DKO	-21.83	-37,09 to -6,565	Yes	**	0.00271432
	EKO vs. DKO/hA-I	-3.978	-18,52 to 10,56	No	ns	0.8791265
	EKO vs. DKO	-17.08	-31,62 to -2,544	Yes	*	0.01630279

	DKO/hA-I vs. DKO	-13.11	-25,93 to -0,2832	Yes	*	0.04367608
Figure 5I	% CD4 CM Lymph node					
	Kruskal-Wallis + Dunn's					
	Dunn's multiple comparisons test	Mean rank diff,	Significant?	Summary	Adjusted P Value	
	WT vs. EKO	-11.05	No	ns	0.31572741	
	WT vs. DKO/hA-I	-10.61	No	ns	0.24836251	
	WT vs. DKO	-10.47	No	ns	0.26447247	
	EKO vs. DKO/hA-I	0.4416	No	ns	>0,99999999	
	EKO vs. DKO	0.5779	No	ns	>0,99999999	
	DKO/hA-I vs. DKO	0.1364	No	ns	>0,99999999	
Supplementary Figure IA	Body weight					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-5.2	-8,156 to -2,244	Yes	***	0.00044116
	WT vs. DKO/hA-I	-1.6	-4,557 to 1,356	No	ns	0.4474289
	WT vs. DKO	-2.09	-5,047 to 0,8659	No	ns	0.22881859
	EKO vs. DKO/hA-I	3.6	0,6432 to 6,556	Yes	*	0.01364612
	EKO vs. DKO	3.11	0,1532 to 6,066	Yes	*	0.03703952
	DKO/hA-I vs. DKO	-0.49	-3,446 to 2,466	No	ns	0.96606468
Supplementary Figure IB	Spleen weight / Body weight ratio					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-0.001051	-0,002271 to 0,0001702	No	ns	0.10763235
	WT vs. DKO/hA-I	0.0001609	-0,001060 to 0,001382	No	ns	0.982352
	WT vs. DKO	0.00003604	-0,001185 to 0,001257	No	ns	0.99979039
	EKO vs. DKO/hA-I	0.001211	-9,292e-006 to 0,002432	No	ns	0.05222043
	EKO vs. DKO	0.001087	-0,0001342 to 0,002307	No	ns	0.0920161
	DKO/hA-I vs. DKO	-0.0001249	-0,001346 to 0,001096	No	ns	0.99156132
Supplementary Figure IC	Lymph node weight / Body weight ratio					
	Welch's ANOVA +Dunnett's					
	Dunnett's T3 multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-0.000226	-0,0007021 to 0,0002501	No	ns	0.2989
	WT vs. DKOp	-0.0001365	-0,0002016 to -7,147e-005	Yes	***	0.0009
	WT vs. DKOm	-0.0006982	-0,001335 to -6,128e-005	Yes	*	0.0171
	EKO vs. DKOp	0.00008951	-0,0003910 to 0,0005700	No	ns	0.9245
	EKO vs. DKOm	-0.0004722	-0,001105 to 0,0001610	No	ns	0.1061
	DKOp vs. DKOm	-0.0005617	-0,001202 to 7,851e-005	Yes	*	0.0421
Supplementary Figure IIE	Macrophage area (%)					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-22.2	-35,79 to -8,609	Yes	***	0.00098127
	WT vs. DKO/hA-I	-21.19	-34,78 to -7,601	Yes	**	0.00157638
	WT vs. DKO	-16.07	-29,66 to -2,481	Yes	*	0.0169197
	EKO vs. DKO/hA-I	1.008	-12,58 to 14,60	No	ns	0.99672583
	EKO vs. DKO	6.128	-7,463 to 19,72	No	ns	0.59645546
	DKO/hA-I vs. DKO	5.12	-8,472 to 18,71	No	ns	0.72024786
Supplementary Figure IIF	Macrophage area					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-70510	-124378 to -16643	Yes	**	0.00772691
	WT vs. DKO/hA-I	-7804	-61672 to 46063	No	ns	0.97684746
	WT vs. DKO	-90234	-144101 to -36366	Yes	***	0.00075218
	EKO vs. DKO/hA-I	62706	8839 to 116574	Yes	*	0.01893913
	EKO vs. DKO	-19723	-73591 to 34144	No	ns	0.73714041
	DKO/hA-I vs. DKO	-82429	-136297 to -28562	Yes	**	0.00189757
Supplementary Figure IIIE	CD3+ lymphocytes plaque					
	Kruskal-Wallis + Dunn's					
	Dunn's multiple comparisons test	Mean rank diff,	Significant?	Summary	Adjusted P Value	
	WT vs. EKO	-11.67	Yes	*	0.02064806	
	WT vs. DKO/hA-I	-5	No	ns	>0,99999999	
	WT vs. DKO	-15.33	Yes	***	0.00072429	
	EKO vs. DKO/hA-I	6.667	No	ns	0.56762546	
	EKO vs. DKO	-3.667	No	ns	>0,99999999	
	DKO/hA-I vs. DKO	-10.33	Yes	*	0.04742317	
Supplementary Figure IIIF	CD3+ lymphocytes plaque normalized					
	ANOVA + Tukey's					
	Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
	WT vs. EKO	-1.657	-4,205 to 0,8912	No	ns	0.29347685
	WT vs. DKO/hA-I	-2.45	-4,999 to 0,09787	No	ns	0.06214867
	WT vs. DKO	-2.029	-4,577 to 0,5196	No	ns	0.14972924
	EKO vs. DKO/hA-I	-0.7933	-3,342 to 1,755	No	ns	0.81944933
	EKO vs. DKO	-0.3716	-2,920 to 2,177	No	ns	0.9764176
	DKO/hA-I vs. DKO	0.4218	-2,126 to 2,970	No	ns	0.96620105

Supplementary

Figure III G

D3+ lymphocytes around the aortic sinus

ANOVA + Tukey's

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-24.45	-32,67 to -16,22	Yes	****	0.00000036
WT vs. DKO/hA-I	-8.5	-16,72 to -0,2776	Yes	*	0.04115602
WT vs. DKO	-25	-33,22 to -16,78	Yes	****	0.00000025
EKO vs. DKO/hA-I	15.95	7,724 to 24,17	Yes	***	0.00014141
EKO vs. DKO	-0.5534	-8,776 to 7,669	No	ns	0.99754764
DKO/hA-I vs. DKO	-16.5	-24,72 to -8,278	Yes	****	0.000093

Supplementary

Figure VE

Dermal thickness

ANOVA + Tukey's

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-41.28	-94,33 to 11,78	No	ns	0.16741871
WT vs. DKO/hA-I	-38.09	-91,14 to 14,96	No	ns	0.2230497
WT vs. DKO	-724.4	-777,4 to -671,3	Yes	****	<0,00000001
EKO vs. DKO/hA-I	3.186	-49,87 to 56,24	No	ns	0.99833584
EKO vs. DKO	-683.1	-736,1 to -630,0	Yes	****	<0,00000001
DKO/hA-I vs. DKO	-686.3	-739,3 to -633,2	Yes	****	<0,00000001

Supplementary

Figure VF

Dermal ORO

Kruskal-Wallis + Dunn's

Dunn's multiple comparisons test	Mean rank diff,	Significant?	Summary	Adjusted P Value
WT vs. EKO	-3.429	No	ns	>0,99999999
WT vs. DKO/hA-I	0.8571	No	ns	>0,99999999
WT vs. DKO	-14.86	Yes	**	0.00436577
EKO vs. DKO/hA-I	4.286	No	ns	>0,99999999
EKO vs. DKO	-11.43	Yes	*	0.04606588
DKO/hA-I vs. DKO	-15.71	Yes	**	0.00211029

Supplementary

Figure VIIIA

CD45+ cells Blood

Kruskal-Wallis + Dunn's

Dunn's multiple comparisons test	Mean rank diff,	Significant?	Summary	Adjusted P Value
WT vs. EKO	-1.2	No	ns	>0,9999
WT vs. DKO/hA-I	-3.057	No	ns	>0,9999
WT vs. DKO	-11.53	Yes	*	0.0299
EKO vs. DKO/hA-I	-1.857	No	ns	>0,9999
EKO vs. DKO	-10.33	No	ns	0.0712
DKO/hA-I vs. DKO	-8.476	No	ns	0.1481

Supplementary

Figure VIIIB

CD45+ cells Spleen

ANOVA + Tukey's

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	11250	-29146 to 51646	No	ns	0.86116388
WT vs. DKO/hA-I	3890	-33510 to 41289	No	ns	0.99100545
WT vs. DKO	4952	-33724 to 43628	No	ns	0.98352924
EKO vs. DKO/hA-I	-7361	-44760 to 30039	No	ns	0.94447857
EKO vs. DKO	-6298	-44974 to 32378	No	ns	0.96725354
DKO/hA-I vs. DKO	1063	-34472 to 36597	No	ns	0.99977888

Supplementary

Figure VIIIC

CD45+ cells Lymph node

ANOVA + Tukey's

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-538.7	-19805 to 18728	No	ns	0.9998191
WT vs. DKO/hA-I	-2059	-19896 to 15778	No	ns	0.98780306
WT vs. DKO	-18678	-37124 to -231,9	Yes	*	0.04655067
EKO vs. DKO/hA-I	-1520	-19358 to 16317	No	ns	0.99498798
EKO vs. DKO	-18139	-36585 to 306,8	Yes	*	0.04992312
DKO/hA-I vs. DKO	-16619	-33567 to 329,1	Yes	*	0.04978933

Supplementary

Figure VIIID

CD45+ cells Blood

ANOVA + Tukey's

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	92.81	-152,7 to 338,4	No	ns	0.71554463
WT vs. DKO/hA-I	38.83	-188,5 to 266,2	No	ns	0.96255613
WT vs. DKO	11.65	-223,4 to 246,8	No	ns	0.99899784
EKO vs. DKO/hA-I	-53.98	-281,3 to 173,4	No	ns	0.90794432
EKO vs. DKO	-81.15	-316,2 to 153,9	No	ns	0.76746846
DKO/hA-I vs. DKO	-27.17	-243,2 to 188,8	No	ns	0.98435537

Supplementary

Figure VIIIE

CD45+ cells Spleen

ANOVA + Tukey's

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	508.5	-1284 to 2301	No	ns	0.85460828
WT vs. DKO/hA-I	-346	-2006 to 1314	No	ns	0.93503416
WT vs. DKO	-221.3	-1938 to 1495	No	ns	0.98319429
EKO vs. DKO/hA-I	-854.5	-2514 to 805,1	No	ns	0.48670166
EKO vs. DKO	-729.8	-2446 to 986,5	No	ns	0.63683648
DKO/hA-I vs. DKO	124.7	-1452 to 1702	No	ns	0.99598693

**Supplementary
Figure VIII D**

**CD45+ cells Lymph node
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	495.1	-4085 to 5075	No	ns	0.9899302
WT vs. DKO/hA-I	-36.82	-4277 to 4204	No	ns	0.99999457
WT vs. DKO	-3232	-7617 to 1153	No	ns	0.1977574
EKO vs. DKO/hA-I	-531.9	-4772 to 3708	No	ns	0.98447751
EKO vs. DKO	-3727	-8112 to 657,9	No	ns	0.11312667
DKO/hA-I vs. DKO	-3195	-7224 to 833,7	No	ns	0.15094957

**Supplementary
Figure IX A**

**% B1a lymphocytes spleen
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-4.208	-10,05 to 1,631	No	ns	0.21343879
WT vs. DKO/hA-I	3.735	-2,388 to 9,859	No	ns	0.34355326
WT vs. DKO	3.527	-2,311 to 9,366	No	ns	0.35160026
EKO vs. DKO/hA-I	7.943	1,819 to 14,07	Yes	**	0.00851368
EKO vs. DKO	7.735	1,896 to 13,57	Yes	**	0.0071761
DKO/hA-I vs. DKO	-0.208	-6,332 to 5,916	No	ns	0.99967623

**Supplementary
Figure IX B**

**% B1b lymphocytes spleen
Welch's ANOVA +Dunnett's**

Dunnett's T3 multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-1.372	-8.897 to 6.152	No	ns	0.9776
WT vs. DKOp	-9.414	-32.15 to 13.32	No	ns	0.4042
WT vs. DKOm	-8.656	-16.41 to -0.8996	Yes	*	0.0132
EKO vs. DKOp	-8.041	-33.12 to 17.04	No	ns	0.5207
EKO vs. DKOm	-7.283	-13.27 to -1.293	Yes	**	0.0073
DKOp vs. DKOm	0.758	-21.32 to 22.83	No	ns	>0.9999

**Supplementary
Figure IX C**

**% myeloid dendritic cells spleen
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-2.526	-7,055 to 2,003	No	ns	0.415926
WT vs. DKO/hA-I	-2.028	-6,758 to 2,702	No	ns	0.627649
WT vs. DKO	-2.453	-6,982 to 2,076	No	ns	0.440785
EKO vs. DKO/hA-I	0.4982	-4,031 to 5,027	No	ns	0.989229
EKO vs. DKO	0.07315	-4,245 to 4,391	No	ns	0.999959
DKO/hA-I vs. DKO	-0.425	-4,954 to 4,104	No	ns	0.993235

**Supplementary
Figure IX D**

**% plasmacytoid dendritic cells spleen
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	13.1	-3,422 to 29,63	No	ns	0.14993384
WT vs. DKO/hA-I	6.294	-10,97 to 23,56	No	ns	0.73416205
WT vs. DKO	11.16	-5,372 to 27,68	No	ns	0.25976275
EKO vs. DKO/hA-I	-6.811	-23,34 to 9,716	No	ns	0.65575427
EKO vs. DKO	-1.95	-17,71 to 13,81	No	ns	0.9848342
DKO/hA-I vs. DKO	4.861	-11,67 to 21,39	No	ns	0.8389462

**Supplementary
Figure IX E**

**% B1a lymphocytes lymph node
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-1.633	-5,397 to 2,131	No	ns	0.62239988
WT vs. DKO/hA-I	4.529	0,5814 to 8,477	Yes	*	0.02117234
WT vs. DKO	3.373	-0,3912 to 7,137	No	ns	0.08872849
EKO vs. DKO/hA-I	6.162	2,214 to 10,11	Yes	**	0.00164713
EKO vs. DKO	5.006	1,242 to 8,770	Yes	**	0.00695507
DKO/hA-I vs. DKO	-1.156	-5,104 to 2,792	No	ns	0.84262528

**Supplementary
Figure IX F**

**% B1b lymphocytes lymph node
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-4.089	-16,45 to 8,268	No	ns	0.78902293
WT vs. DKO/hA-I	-9.85	-22,81 to 3,110	No	ns	0.17735653
WT vs. DKO	-4.169	-16,53 to 8,188	No	ns	0.77936795
EKO vs. DKO/hA-I	-5.761	-18,72 to 7,199	No	ns	0.60423408
EKO vs. DKO	-0.07958	-12,44 to 12,28	No	ns	0.99999778
DKO/hA-I vs. DKO	5.681	-7,279 to 18,64	No	ns	0.61464209

**Supplementary
Figure IX G**

**% myeloid dendritic cells lymph node
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	1.055	-1,176 to 3,286	No	ns	0.55629545
WT vs. DKO/hA-I	2.163	-0,1771 to 4,503	No	ns	0.07616841
WT vs. DKO	1.493	-0,7379 to 3,724	No	ns	0.26851061
EKO vs. DKO/hA-I	1.108	-1,232 to 3,448	No	ns	0.55523982
EKO vs. DKO	0.4383	-1,793 to 2,669	No	ns	0.94476499
DKO/hA-I vs. DKO	-0.6697	-3,010 to 1,670	No	ns	0.85140099

**Supplementary
Figure IXH**

**Plasmacytoid dendritic cells lymph node
Kruskal-Wallis + Dunn's**

Dunn's multiple comparisons test	Mean rank diff,	Significant?	Summary	Adjusted P Value
WT vs. EKO	-8.5	No	ns	0.17972126
WT vs. DKO/hA-I	-14.13	Yes	**	0.0034728
WT vs. DKO	-7.833	No	ns	0.27270832
EKO vs. DKO/hA-I	-5.633	No	ns	>0,99999999
EKO vs. DKO	0.6667	No	ns	>0,99999999
DKO/hA-I vs. DKO	6.3	No	ns	0.750175

**Supplementary
Figure IXI**

**% Ly6C low blood
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-0.623	-11,63 to 10,38	No	ns	0.99868739
WT vs. DKO/hA-I	2.637	-7,319 to 12,59	No	ns	0.88933359
WT vs. DKO	8.308	-1,648 to 18,26	No	ns	0.12882225
EKO vs. DKO/hA-I	3.26	-6,696 to 13,22	No	ns	0.81146942
EKO vs. DKO	8.931	-1,025 to 18,89	No	ns	0.09148822
DKO/hA-I vs. DKO	5.671	-3,109 to 14,45	No	ns	0.31563317

**Supplementary
Figure IXJ**

**% Ly6C high blood
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	0.6076	-12,90 to 14,11	No	ns	0.99933806
WT vs. DKO/hA-I	-2.747	-14,96 to 9,467	No	ns	0.92836804
WT vs. DKO	-8.366	-20,58 to 3,848	No	ns	0.26697312
EKO vs. DKO/hA-I	-3.354	-15,57 to 8,859	No	ns	0.8784717
EKO vs. DKO	-8.974	-21,19 to 3,240	No	ns	0.2126567
DKO/hA-I vs. DKO	-5.619	-16,39 to 5,152	No	ns	0.50049149

Table 1

**Eotaxin
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-292.8	-894,2 to 308,6	No	ns	0.52938895
WT vs. DKO/hA-I	-245.5	-821,3 to 330,3	No	ns	0.63154074
WT vs. DKO	-565	-1141 to 10,72	No	ns	0.05548456
EKO vs. DKO/hA-I	47.27	-528,5 to 623,0	No	ns	0.99543676
EKO vs. DKO	-272.3	-848,0 to 303,5	No	ns	0.55285708
DKO/hA-I vs. DKO	-319.5	-868,5 to 229,4	No	ns	0.37992357

Table 1

**G-CSF
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-1066	-2759 to 626,8	No	ns	0.31458526
WT vs. DKO/hA-I	-838.6	-2459 to 782,2	No	ns	0.47923145
WT vs. DKO	-73.38	-1694 to 1547	No	ns	0.99922241
EKO vs. DKO/hA-I	227.4	-1393 to 1848	No	ns	0.97821231
EKO vs. DKO	992.7	-628,1 to 2613	No	ns	0.33738563
DKO/hA-I vs. DKO	765.2	-780,1 to 2311	No	ns	0.51558465

Table 1

**IL-4
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	65.29	-40,16 to 170,7	No	ns	0.32839115
WT vs. DKO/hA-I	52.69	-48,28 to 153,7	No	ns	0.47211605
WT vs. DKO	76.81	-24,15 to 177,8	No	ns	0.17530475
EKO vs. DKO/hA-I	-12.61	-113,6 to 88,36	No	ns	0.98442358
EKO vs. DKO	11.52	-89,45 to 112,5	No	ns	0.98802662
DKO/hA-I vs. DKO	24.12	-72,14 to 120,4	No	ns	0.89249852

Table 1

**IL-5
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-0.8927	-81,86 to 80,07	No	ns	0.99998869
WT vs. DKO/hA-I	40.08	-37,44 to 117,6	No	ns	0.47990017
WT vs. DKO	31.34	-46,18 to 108,9	No	ns	0.66888057
EKO vs. DKO/hA-I	40.97	-36,55 to 118,5	No	ns	0.46143393
EKO vs. DKO	32.23	-45,28 to 109,7	No	ns	0.64949322
DKO/hA-I vs. DKO	-8.736	-82,65 to 65,17	No	ns	0.98671865

Table 1

**IL-6
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-11.88	-193,4 to 169,6	No	ns	0.9976657
WT vs. DKO/hA-I	-20.33	-194,1 to 153,4	No	ns	0.98710939
WT vs. DKO	-87.75	-261,5 to 86,00	No	ns	0.49931635
EKO vs. DKO/hA-I	-8.445	-182,2 to 165,3	No	ns	0.99903864
EKO vs. DKO	-75.87	-249,6 to 97,88	No	ns	0.61413261
DKO/hA-I vs. DKO	-67.43	-233,1 to 98,24	No	ns	0.66431798

Table 1

**IL-10
ANOVA + Tukey's**

Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	189.9	-59,54 to 439,4	No	ns	0.17484772
WT vs. DKO/hA-I	114.3	-124,5 to 353,1	No	ns	0.54330443
WT vs. DKO	176.9	-61,93 to 415,7	No	ns	0.19281211

EKO vs. DKO/hA-I	-75.62	-314,4 to 163,2	No	ns	0.80757918
EKO vs. DKO	-13.01	-251,8 to 225,8	No	ns	0.9986503
DKO/hA-I vs. DKO	62.61	-165,1 to 290,3	No	ns	0.86377004

Table 1

IL-12 (p70)					
ANOVA + Tukey's					
Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-225.8	-711,2 to 259,6	No	ns	0.56580086
WT vs. DKO/hA-I	88.05	-376,7 to 552,8	No	ns	0.94921884
WT vs. DKO	46.95	-417,8 to 511,7	No	ns	0.99159713
EKO vs. DKO/hA-I	313.8	-150,9 to 778,5	No	ns	0.25931832
EKO vs. DKO	272.7	-192,0 to 737,4	No	ns	0.37293745
DKO/hA-I vs. DKO	-41.1	-484,2 to 402,0	No	ns	0.99346142

Table 1

IL-15					
ANOVA + Tukey's					
Tukey's multiple comparisons test	Mean Diff.	95,00% CI of diff.	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-5.047	-14,04 to 3,946	No	ns	0.41388152
WT vs. DKO/hA-I	2.849	-5,726 to 11,42	No	ns	0.78701435
WT vs. DKO	-1.316	-9,891 to 7,260	No	ns	0.97234305
EKO vs. DKO/hA-I	7.897	-1,097 to 16,89	No	ns	0.09766843
EKO vs. DKO	3.732	-5,262 to 12,73	No	ns	0.65420871
DKO/hA-I vs. DKO	-4.165	-12,74 to 4,410	No	ns	0.53477839

Table 1

IL-17					
Kruskal-Wallis + Dunn's					
Dunn's multiple comparisons test	Mean rank diff,	Significant?	Summary	Adjusted P Value	
WT vs. EKO	2.3	No	ns	>0,99999999	
WT vs. DKO/hA-I	-2.767	No	ns	>0,99999999	
WT vs. DKO	0.4833	No	ns	>0,99999999	
EKO vs. DKO/hA-I	-5.067	No	ns	>0,99999999	
EKO vs. DKO	-1.817	No	ns	>0,99999999	
DKO/hA-I vs. DKO	3.25	No	ns	>0,99999999	

Table 1

IP-10					
ANOVA + Tukey's					
Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-57.18	-173,6 to 59,29	No	ns	0.52260039
WT vs. DKO/hA-I	-96.1	-207,6 to 15,40	No	ns	0.10570928
WT vs. DKO	-50.75	-162,3 to 60,75	No	ns	0.58272884
EKO vs. DKO/hA-I	-38.93	-150,4 to 72,58	No	ns	0.75874142
EKO vs. DKO	6.423	-105,1 to 117,9	No	ns	0.99840502
DKO/hA-I vs. DKO	45.35	-60,97 to 151,7	No	ns	0.63133468

Table 1

KC					
ANOVA + Tukey's					
Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	-18.18	-237,0 to 200,6	No	ns	0.99527597
WT vs. DKO/hA-I	-33.97	-243,5 to 175,5	No	ns	0.96713239
WT vs. DKO	-68.59	-278,1 to 140,9	No	ns	0.79182687
EKO vs. DKO/hA-I	-15.79	-225,3 to 193,7	No	ns	0.99645599
EKO vs. DKO	-50.41	-259,9 to 159,1	No	ns	0.90331531
DKO/hA-I vs. DKO	-34.62	-234,4 to 165,1	No	ns	0.96036873

Table 1

LIX					
ANOVA + Tukey's					
Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	960.8	-7841 to 9762	No	ns	0.98946439
WT vs. DKO/hA-I	-2800	-11227 to 5626	No	ns	0.78442792
WT vs. DKO	-3035	-11462 to 5392	No	ns	0.74128912
EKO vs. DKO/hA-I	-3761	-12188 to 4665	No	ns	0.59771516
EKO vs. DKO	-3996	-12422 to 4431	No	ns	0.55065009
DKO/hA-I vs. DKO	-234.5	-8269 to 7800	No	ns	0.99979077

Table 1

MIG					
ANOVA + Tukey's					
Tukey's multiple comparisons test	Mean Diff,	95,00% CI of diff,	Below threshold?	Summary	Adjusted P Value
WT vs. EKO	18.63	-51,70 to 88,96	No	ns	0.87606188
WT vs. DKO/hA-I	-1.657	-69,00 to 65,68	No	ns	0.99987444
WT vs. DKO	-15.47	-82,81 to 51,87	No	ns	0.91439098
EKO vs. DKO/hA-I	-20.29	-87,63 to 47,05	No	ns	0.82924088
EKO vs. DKO	-34.1	-101,4 to 33,24	No	ns	0.49709968
DKO/hA-I vs. DKO	-13.82	-78,02 to 50,39	No	ns	0.92816812

**Supplementary
Figure XIII A**

CD4/prolif					
ANOVA + Tukey's					
Tukey's multiple comparisons test	Mean Diff.	95,00% CI of diff.	Below threshold?	Summary	Adjusted P Value
Control vs. ApoA-I 10 µg/ml	17.46	4,976 to 29,93	Yes	**	0.00359481
Control vs. ApoA-I 25 µg/ml	24.03	11,55 to 36,51	Yes	***	0.00011127
Control vs. ApoA-I 50 µg/ml	25.56	13,08 to 38,04	Yes	****	0.00005165
Control vs. ApoA-I 100 µg/ml	33.99	21,51 to 46,46	Yes	****	0.00000104
Control vs. Resting	43.4	30,92 to 55,87	Yes	****	0.00000003
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	6.578	-5,901 to 19,06	No	ns	0.56363273
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	8.103	-4,376 to 20,58	No	ns	0.34785893
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	16.53	4,051 to 29,01	Yes	**	0.00592282
ApoA-I 10 µg/ml vs. Resting	25.94	13,46 to 38,42	Yes	****	0.00004273
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	1.525	-10,95 to 14,00	No	ns	0.99864816

ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	9.953	-2.526 to 22.43	No	ns	0.16568977
ApoA-I 25 µg/ml vs. Resting	19.36	6.884 to 31.84	Yes	**	0.00128655
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	8.428	-4.051 to 20.91	No	ns	0.30875223
ApoA-I 50 µg/ml vs. Resting	17.84	5.359 to 30.32	Yes	**	0.00292403
ApoA-I 100 µg/ml vs. Resting	9.41	-3.069 to 21.89	No	ns	0.20901075

Supplementary
Figure XIII B

CD8/prolif						
ANOVA + Tukey's						
Tukey's multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value	
Control vs. ApoA-I 10 µg/ml	11.07	-8.559 to 30.69	No	ns	0.48891273	
Control vs. ApoA-I 25 µg/ml	23.63	4.004 to 43.26	Yes	*	0.01358581	
Control vs. ApoA-I 50 µg/ml	33.47	13.85 to 53.10	Yes	***	0.00051712	
Control vs. ApoA-I 100 µg/ml	34.68	15.05 to 54.30	Yes	***	0.00035036	
Control vs. Resting	72.92	51.72 to 94.12	Yes	****	0.00000005	
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	12.56	-7.064 to 32.19	No	ns	0.35785153	
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	22.41	2.779 to 42.03	Yes	*	0.0203711	
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	23.61	3.984 to 43.24	Yes	*	0.01367636	
ApoA-I 10 µg/ml vs. Resting	61.86	40.66 to 83.05	Yes	****	0.00000055	
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	9.843	-9.784 to 29.47	No	ns	0.60687097	
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	11.05	-8.579 to 30.67	No	ns	0.49078807	
ApoA-I 25 µg/ml vs. Resting	49.29	28.09 to 70.49	Yes	****	0.00001246	
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	1.205	-18.42 to 20.83	No	ns	0.99995115	
ApoA-I 50 µg/ml vs. Resting	39.45	18.25 to 60.65	Yes	***	0.00019448	
ApoA-I 100 µg/ml vs. Resting	38.25	17.05 to 59.44	Yes	***	0.00027723	

Supplementary
Figure XIII C

% INFγ/CD4						
ANOVA + Tukey's						
Tukey's multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value	
Control vs. ApoA-I 10 µg/ml	0.2025	-0.8327 to 1.238	No	ns	0.98790112	
Control vs. ApoA-I 25 µg/ml	1.858	0.8223 to 2.893	Yes	***	0.0002591	
Control vs. ApoA-I 50 µg/ml	3.285	2.250 to 4.320	Yes	****	0.00000011	
Control vs. ApoA-I 100 µg/ml	5.043	4.007 to 6.078	Yes	****	<0.00000001	
Control vs. Resting	6.53	5.495 to 7.565	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	1.655	0.6198 to 2.690	Yes	***	0.0009395	
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	3.083	2.047 to 4.118	Yes	****	0.00000028	
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	4.84	3.805 to 5.875	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. Resting	6.328	5.292 to 7.363	Yes	****	<0.00000001	
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	1.428	0.3923 to 2.463	Yes	**	0.00410872	
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	3.185	2.150 to 4.220	Yes	****	0.00000017	
ApoA-I 25 µg/ml vs. Resting	4.673	3.637 to 5.708	Yes	****	<0.00000001	
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	1.758	0.7223 to 2.793	Yes	***	0.00048746	
ApoA-I 50 µg/ml vs. Resting	3.245	2.210 to 4.280	Yes	****	0.00000013	
ApoA-I 100 µg/ml vs. Resting	1.488	0.4523 to 2.523	Yes	**	0.00278037	

Supplementary
Figure XIII D

% IL17/CD4						
Welch's ANOVA +Dunnett's						
Dunnett's T3 multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value	
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	1.743	-4.305 to 7.790	No	ns	0.48889158	
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	3.83	-2.282 to 9.942	No	ns	0.08978227	
Control vs. ApoA-I 10 µg/ml	3.46	-1.639 to 8.559	No	ns	0.0931755	
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	10.28	4.144 to 16.41	Yes	**	0.00560205	
ApoA-I 10 µg/ml vs. Resting	14.95	9.017 to 20.87	Yes	**	0.00167402	
ApoA-I 100 µg/ml vs. Resting	4.67	2.974 to 6.366	Yes	**	0.00128639	
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	8.533	7.246 to 9.819	Yes	****	0.00000057	
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	2.088	0.8360 to 3.339	Yes	**	0.00176556	
ApoA-I 25 µg/ml vs. Resting	13.2	12.13 to 14.28	Yes	****	0.00000136	
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	6.445	5.048 to 7.842	Yes	****	0.00000493	
ApoA-I 50 µg/ml vs. Resting	11.12	9.838 to 12.39	Yes	****	0.00000054	
Control vs. ApoA-I 100 µg/ml	13.74	11.59 to 15.88	Yes	****	0.00000347	
Control vs. ApoA-I 25 µg/ml	5.203	2.860 to 7.545	Yes	**	0.00122097	
Control vs. ApoA-I 50 µg/ml	7.29	5.172 to 9.408	Yes	****	0.00007563	
Control vs. Resting	18.41	15.67 to 21.14	Yes	****	0.00008883	

Supplementary
Figure XIII E

% INFγ/CD8						
ANOVA + Tukey's						
Tukey's multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value	
Control vs. ApoA-I 10 µg/ml	9.233	6.330 to 12.13	Yes	****	0.00000001	
Control vs. ApoA-I 25 µg/ml	13.6	10.70 to 16.50	Yes	****	<0.00000001	
Control vs. ApoA-I 50 µg/ml	25.53	22.63 to 28.43	Yes	****	<0.00000001	
Control vs. ApoA-I 100 µg/ml	37.64	34.74 to 40.54	Yes	****	<0.00000001	
Control vs. Resting	41.24	38.34 to 44.14	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	4.368	1.465 to 7.270	Yes	**	0.00175952	
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	16.3	13.40 to 19.20	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	28.41	25.51 to 31.31	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. Resting	32.01	29.11 to 34.91	Yes	****	<0.00000001	
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	11.93	9.030 to 14.83	Yes	****	<0.00000001	
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	24.04	21.14 to 26.94	Yes	****	<0.00000001	
ApoA-I 25 µg/ml vs. Resting	27.64	24.74 to 30.54	Yes	****	<0.00000001	
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	12.11	9.208 to 15.01	Yes	****	<0.00000001	
ApoA-I 50 µg/ml vs. Resting	15.71	12.81 to 18.61	Yes	****	<0.00000001	
ApoA-I 100 µg/ml vs. Resting	3.6	0.6979 to 6.502	Yes	*	0.01041919	

Supplementary
Figure XIII F

% IL17/CD8		
ANOVA + Tukey's		

Tukey's multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value
Control vs. ApoA-I 10 µg/ml	6.938	4.567 to 9.308	Yes	****	0.00000036
Control vs. ApoA-I 25 µg/ml	11.91	9.534 to 14.28	Yes	****	<0.00000001
Control vs. ApoA-I 50 µg/ml	20.31	17.94 to 22.68	Yes	****	<0.00000001
Control vs. ApoA-I 100 µg/ml	30.09	27.72 to 32.46	Yes	****	<0.00000001
Control vs. Resting	35.34	32.96 to 37.71	Yes	****	<0.00000001
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	4.968	2.597 to 7.338	Yes	****	0.0000386
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	13.37	11.00 to 15.74	Yes	****	<0.00000001
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	23.16	20.78 to 25.53	Yes	****	<0.00000001
ApoA-I 10 µg/ml vs. Resting	28.4	26.03 to 30.77	Yes	****	<0.00000001
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	8.403	6.032 to 10.77	Yes	****	0.00000002
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	18.19	15.82 to 20.56	Yes	****	<0.00000001
ApoA-I 25 µg/ml vs. Resting	23.43	21.06 to 25.80	Yes	****	<0.00000001
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	9.785	7.414 to 12.16	Yes	****	<0.00000001
ApoA-I 50 µg/ml vs. Resting	15.03	12.66 to 17.40	Yes	****	<0.00000001
ApoA-I 100 µg/ml vs. Resting	5.243	2.872 to 7.613	Yes	****	0.00001908

**Supplementary
Figure XIVA**

% IL15/CD4					
Kruskal-Wallis + Dunn's					
Dunn's multiple comparisons test	Mean rank diff.	Significant?	Summary	Adjusted P Value	
Control vs. ApoA-I 10 µg/ml	7.75	No	ns	>0.99999999	
Control vs. ApoA-I 25 µg/ml	5.25	No	ns	>0.99999999	
Control vs. ApoA-I 50 µg/ml	11	No	ns	0.41710343	
Control vs. ApoA-I 100 µg/ml	16	Yes	*	0.02061414	
Control vs. Resting	20	Yes	***	0.00095014	
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	-2.5	No	ns	>0.99999999	
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	3.25	No	ns	>0.99999999	
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	8.25	No	ns	>0.99999999	
ApoA-I 10 µg/ml vs. Resting	12.25	No	ns	0.21428432	
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	5.75	No	ns	>0.99999999	
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	10.75	No	ns	0.47332822	
ApoA-I 25 µg/ml vs. Resting	14.75	Yes	*	0.04766609	
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	5	No	ns	>0.99999999	
ApoA-I 50 µg/ml vs. Resting	9	No	ns	>0.99999999	
ApoA-I 100 µg/ml vs. Resting	4	No	ns	>0.99999999	

**Supplementary
Figure XIVB**

% IL15/CD8					
ANOVA + Tukey's					
Tukey's multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value
Control vs. ApoA-I 10 µg/ml	3.345	0.2830 to 6.407	Yes	*	0.02769205
Control vs. ApoA-I 25 µg/ml	10.11	7.043 to 13.17	Yes	****	0.00000006
Control vs. ApoA-I 50 µg/ml	23.71	20.65 to 26.77	Yes	****	<0.00000001
Control vs. ApoA-I 100 µg/ml	34.78	31.71 to 37.84	Yes	****	<0.00000001
Control vs. Resting	41.64	38.58 to 44.70	Yes	****	<0.00000001
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	6.76	3.698 to 9.822	Yes	****	0.00001948
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	20.37	17.31 to 23.43	Yes	****	<0.00000001
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	31.43	28.37 to 34.49	Yes	****	<0.00000001
ApoA-I 10 µg/ml vs. Resting	38.3	35.23 to 41.36	Yes	****	<0.00000001
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	13.61	10.55 to 16.67	Yes	****	<0.00000001
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	24.67	21.61 to 27.73	Yes	****	<0.00000001
ApoA-I 25 µg/ml vs. Resting	31.54	28.47 to 34.60	Yes	****	<0.00000001
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	11.06	8.000 to 14.12	Yes	****	0.00000001
ApoA-I 50 µg/ml vs. Resting	17.93	14.87 to 20.99	Yes	****	<0.00000001
ApoA-I 100 µg/ml vs. Resting	6.865	3.803 to 9.927	Yes	****	0.00001587

**Supplementary
Figure XIVC**

% IL4/CD4					
ANOVA + Tukey's					
Tukey's multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value
Control vs. ApoA-I 10 µg/ml	7.043	3.953 to 10.13	Yes	****	0.00001269
Control vs. ApoA-I 25 µg/ml	-3.208	-6.297 to -0.1183	Yes	*	0.03923849
Control vs. ApoA-I 50 µg/ml	-2.423	-5.512 to 0.6667	No	ns	0.17817092
Control vs. ApoA-I 100 µg/ml	20.85	17.76 to 23.94	Yes	****	<0.00000001
Control vs. Resting	53.78	50.69 to 56.86	Yes	****	<0.00000001
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	-10.25	-13.34 to -7.161	Yes	****	0.00000005
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	-9.465	-12.55 to -6.376	Yes	****	0.00000018
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	13.81	10.72 to 16.90	Yes	****	<0.00000001
ApoA-I 10 µg/ml vs. Resting	46.73	43.64 to 49.82	Yes	****	<0.00000001
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	0.785	-2.304 to 3.874	No	ns	0.96239711
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	24.06	20.97 to 27.15	Yes	****	<0.00000001
ApoA-I 25 µg/ml vs. Resting	56.98	53.89 to 60.07	Yes	****	<0.00000001
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	23.28	20.19 to 26.36	Yes	****	<0.00000001
ApoA-I 50 µg/ml vs. Resting	56.2	53.11 to 59.29	Yes	****	<0.00000001
ApoA-I 100 µg/ml vs. Resting	32.92	29.83 to 36.01	Yes	****	<0.00000001

**Supplementary
Figure XIVD**

% IL4/CD8					
ANOVA + Tukey's					
Tukey's multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value
Control vs. ApoA-I 10 µg/ml	5.865	2.600 to 9.130	Yes	***	0.00025576
Control vs. ApoA-I 25 µg/ml	-2.673	-5.938 to 0.5925	No	ns	0.1475696
Control vs. ApoA-I 50 µg/ml	6.45	3.185 to 9.715	Yes	****	0.0000813
Control vs. ApoA-I 100 µg/ml	25.51	22.24 to 28.77	Yes	****	<0.00000001
Control vs. Resting	66.97	63.71 to 70.24	Yes	****	<0.00000001
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	-8.538	-11.80 to -5.272	Yes	****	0.00000188
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	0.585	-2.680 to 3.850	No	ns	0.99186646
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	19.64	16.38 to 22.91	Yes	****	<0.00000001
ApoA-I 10 µg/ml vs. Resting	61.11	57.84 to 64.37	Yes	****	<0.00000001

ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	9.123	5.857 to 12.39	Yes	****	0.00000072
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	28.18	24.91 to 31.45	Yes	****	<0.00000001
ApoA-I 25 µg/ml vs. Resting	69.65	66.38 to 72.91	Yes	****	<0.00000001
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	19.06	15.79 to 22.32	Yes	****	<0.00000001
ApoA-I 50 µg/ml vs. Resting	60.52	57.26 to 63.79	Yes	****	<0.00000001
ApoA-I 100 µg/ml vs. Resting	41.47	38.20 to 44.73	Yes	****	<0.00000001

**Supplementary
Figure XVA**

CTXb CD4						
ANOVA + Tukey's						
Tukey's multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value	
Control vs. ApoA-I 10 µg/ml	636.7	-2071 to 3344	No	ns	0.96403246	
Control vs. ApoA-I 25 µg/ml	1633	-1074 to 4340	No	ns	0.38282828	
Control vs. ApoA-I 50 µg/ml	7846	5138 to 10553	Yes	****	0.00000562	
Control vs. ApoA-I 100 µg/ml	21229	18522 to 23936	Yes	****	<0.00000001	
Control vs. Resting	29153	26446 to 31860	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	996.3	-1711 to 3704	No	ns	0.81173799	
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	7209	4502 to 9916	Yes	****	0.00001373	
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	20592	17885 to 23300	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. Resting	28516	25809 to 31224	Yes	****	<0.00000001	
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	6213	3505 to 8920	Yes	****	0.00006284	
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	19596	16889 to 22303	Yes	****	<0.00000001	
ApoA-I 25 µg/ml vs. Resting	27520	24813 to 30227	Yes	****	<0.00000001	
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	13383	10676 to 16091	Yes	****	0.00000001	
ApoA-I 50 µg/ml vs. Resting	21307	18600 to 24015	Yes	****	<0.00000001	
ApoA-I 100 µg/ml vs. Resting	7924	5217 to 10631	Yes	****	0.00000506	

**Supplementary
Figure XVB**

CTXb CD8						
ANOVA + Tukey's						
Tukey's multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value	
Control vs. ApoA-I 10 µg/ml	350	-3175 to 3875	No	ns	0.99929752	
Control vs. ApoA-I 25 µg/ml	8660	5134 to 12185	Yes	****	0.00003158	
Control vs. ApoA-I 50 µg/ml	23448	19922 to 26973	Yes	****	<0.00000001	
Control vs. ApoA-I 100 µg/ml	31436	27911 to 34961	Yes	****	<0.00000001	
Control vs. Resting	50577	47051 to 54102	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	8310	4784 to 11835	Yes	****	0.000048	
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	23098	19572 to 26623	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	31086	27561 to 34611	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. Resting	50227	46701 to 53752	Yes	****	<0.00000001	
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	14788	11263 to 18313	Yes	****	0.00000009	
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	22776	19251 to 26302	Yes	****	<0.00000001	
ApoA-I 25 µg/ml vs. Resting	41917	38392 to 45442	Yes	****	<0.00000001	
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	7988	4463 to 11514	Yes	****	0.00007125	
ApoA-I 50 µg/ml vs. Resting	27129	23604 to 30654	Yes	****	<0.00000001	
ApoA-I 100 µg/ml vs. Resting	19141	15615 to 22666	Yes	****	<0.00000001	

**Supplementary
Figure XVC**

Filipin CD4						
ANOVA + Tukey's						
Tukey's multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value	
Control vs. ApoA-I 10 µg/ml	19	11.22 to 26.78	Yes	****	0.0000337	
Control vs. ApoA-I 25 µg/ml	6	-1.784 to 13.78	No	ns	0.17349341	
Control vs. ApoA-I 50 µg/ml	43.67	35.88 to 51.45	Yes	****	<0.00000001	
Control vs. ApoA-I 100 µg/ml	61	53.22 to 68.78	Yes	****	<0.00000001	
Control vs. Resting	55	47.22 to 62.78	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	-13	-20.78 to -5.216	Yes	**	0.00123439	
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	24.67	16.88 to 32.45	Yes	****	0.00000215	
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	42	34.22 to 49.78	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. Resting	36	28.22 to 43.78	Yes	****	0.00000003	
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	37.67	29.88 to 45.45	Yes	****	0.00000002	
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	55	47.22 to 62.78	Yes	****	<0.00000001	
ApoA-I 25 µg/ml vs. Resting	49	41.22 to 56.78	Yes	****	<0.00000001	
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	17.33	9.549 to 25.12	Yes	****	0.00008473	
ApoA-I 50 µg/ml vs. Resting	11.33	3.549 to 19.12	Yes	**	0.00386256	
ApoA-I 100 µg/ml vs. Resting	-6	-13.78 to 1.784	No	ns	0.17349341	

**Supplementary
Figure XVD**

Filipin CD8						
ANOVA + Tukey's						
Tukey's multiple comparisons test	Mean Diff.	95.00% CI of diff.	Below threshold?	Summary	Adjusted P Value	
Control vs. ApoA-I 10 µg/ml	15.67	8.382 to 22.95	Yes	***	0.00011929	
Control vs. ApoA-I 25 µg/ml	18.33	11.05 to 25.62	Yes	****	0.00002465	
Control vs. ApoA-I 50 µg/ml	48.33	41.05 to 55.62	Yes	****	<0.00000001	
Control vs. ApoA-I 100 µg/ml	65.33	58.05 to 72.62	Yes	****	<0.00000001	
Control vs. Resting	55	47.72 to 62.28	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. ApoA-I 25 µg/ml	2.667	-4.618 to 9.952	No	ns	0.81490193	
ApoA-I 10 µg/ml vs. ApoA-I 50 µg/ml	32.67	25.38 to 39.95	Yes	****	0.00000004	
ApoA-I 10 µg/ml vs. ApoA-I 100 µg/ml	49.67	42.38 to 56.95	Yes	****	<0.00000001	
ApoA-I 10 µg/ml vs. Resting	39.33	32.05 to 46.62	Yes	****	<0.00000001	
ApoA-I 25 µg/ml vs. ApoA-I 50 µg/ml	30	22.72 to 37.28	Yes	****	0.00000012	
ApoA-I 25 µg/ml vs. ApoA-I 100 µg/ml	47	39.72 to 54.28	Yes	****	<0.00000001	
ApoA-I 25 µg/ml vs. Resting	36.67	29.38 to 43.95	Yes	****	0.00000001	
ApoA-I 50 µg/ml vs. ApoA-I 100 µg/ml	17	9.715 to 24.28	Yes	****	0.00005311	
ApoA-I 50 µg/ml vs. Resting	6.667	-0.6182 to 13.95	No	ns	0.08024647	
ApoA-I 100 µg/ml vs. Resting	-10.33	-17.62 to -3.048	Yes	**	0.00474365	

Table S2 - details of DEGS listed in Figure 6

Fig. 6E	Fig. 6F	Fig. 6G
Marco	Nr1h3	Slc25a20
Timd4	Cx3cr1	Ptgds
Cd209e	Lipa	Slc44a3
Cd209b	Npc1	Ldhd
Cd55	Il1r2	Dbi
Cd209f	Ccr2	Fabp4
Cd209d	Ccl2	Acox1
Adora1	Abca1	Cpt1b
S1pr2	Msr1	Pcca
Lipa	Il12a	Eci1
Itgax	Fabp5	Acly
Galc	Il12b	Adhfe1
Cd68	Arg1	Pdha1
Il1r2	Cxcl2	Gstz1
Itga5	Cd36	Decr1
Plekhf1	Tlr2	Ehhadh
Colec12	Olr1	Slc25a1
Tgfb1	Map2k3	Pdk2
Anxa2	Mertk	Fabp3
Adam8	Il1b	Sucla2
Trim29	Nlrp3	Scd1
Atp6v1a		Elovl6
C3ar1		Hadh
Ctss		Agpat2
Clec5a		Pnpla2
Clec7a		Chpt1
Anxa1		Etfhd
Gpr137b		Pdhd
Litaf		Suclg1
Gusb		Dlat
Hexa		Idh3a
Ccr5		Aco2
Ctsl		Acacb
Igf2r		Acadm
Clec4e		Gpd2
Cd63		Gpd1
Cxcl16		Pdk4
Gla		Ppara
Clec4a2		Acaa2
Fcer1g		Dgat2
Fcgr4		Acadvl
Ada		Slc27a2
Ccr1		Etfh
Slc15a3		Acaca
C5ar1		Cpt2
Tlr8		Bche
Tpcn2		Hadhb
Clec4d		Ecp1
Cx3cr1		Elovl3
Lgmn		Thrsp
Fcgr1		Cidea
Galns		Cidec
Clec4a3		Cyp2e1
Asah1		Fasn
Clec12a		Acss3
Fcgr3		Me1
Mpo		Crat
Trpm2		Fitm2
Clec4a1		Acsl1
Slc11a1		Cyp1a1
Pla2g15		Ucp3
Cd300a		Aacs
Mertk		Ephx2
Pdgfrb		Ndufv1
		Pccb
		Acat1
		Acot11
		Acat2
		Cyp4b1
		Etfb

Major Resources Table

In order to allow validation and replication of experiments, all essential research materials listed in the Methods should be included in the Major Resources Table below. Authors are encouraged to use public repositories for protocols, data, code, and other materials and provide persistent identifiers and/or links to repositories when available. Authors may add or delete rows as needed.

Animals (in vivo studies)

Species	Vendor or Source	Background Strain	Sex	Persistent ID / URL
mouse	The Jackson Laboratory	C57BL6/J	M	https://www.jax.org/strain/000664

Genetically Modified Animals

	Species	Vendor or Source	Background Strain	Other Information	Persistent ID / URL
	mouse	The Jackson Laboratory	C57BL6/J	Apoe ^{-/-}	https://www.jax.org/strain/002052
Parent - Male	mouse	Professor Chiesa's Lab	C57BL6/J	Apoe ^{-/-} Apoa1 ^{-/-}	
Parent - Female	mouse	Professor Chiesa's Lab	C57BL6/J	APOA1 ^{tg} Apoe ^{-/-} Apoa1 ^{-/-}	

Antibodies

Target antigen	Vendor or Source	Catalog #	Working conc.	Persistent ID / URL
Mac2	Cedarlane	CL8942	1:1000	https://www.cedarlanelabs.com/Contents/Files?filePath=pdf/16CLM%20Galectin-3%20Antibodies%20Flyer-unpriced_web.pdf
CD3	R&D Systems	MAB4841	1:200	https://www.rndsystems.com/products/mouse-cd3-antibody-17a2_mab4841
IBA-1	WAKO	019-19741		https://labchem-wako.fujifilm.com/us/product/detail/W01W0101-1974.html
CD45R/B220	BD Pharmingen	557390		https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/purified-rat-anti-mouse-cd45r.557390
CD3 epsilon	Santa Cruz Biotechnology	SC-1127		https://www.scbt.com/p/cd3-epsilon-antibody-m-20
CD4 (PE YTS 191.1.2)	Immunotools	22150044		http://www.immunotools.de/html/datas-pe-m/22150044.pdf
CD44 (FITC, IM7)	BD Pharmingen	553133		https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/fitc-rat-anti-mouse-cd44.553133

CD45 (FITC, 30-F11)	BD Pharmingen	553080		https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/fitc-rat-anti-mouse-cd45.553080
CD62L (APC, mMEL-14)	BD Pharmingen	561919		https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/apc-rat-anti-mouse-cd62l.561919
Ly-6C (FITC AL-21)	BD Pharmingen	553104		https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/fitc-rat-anti-mouse-ly-6c.553104
CD115 (APC AF598)	eBioscience / Thermo Fisher	17-1152-82		https://www.thermofisher.com/antibody/product/CD115-c-fms-Antibody-clone-AFS98-Monoclonal/17-1152-82
CD11b (PE M1/70.15)	Immunotools	22159114		http://www.immunotools.de/html/datas-pe-m/22159114.pdf
CD11c (APC HL3)	BD Pharmingen	561119		https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/apc-hamster-anti-mouse-cd11c.561119
B220 (PerCP Cy5,5 RA3-6B2)	BD Pharmingen	561101		https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/percp-cy-5-5-rat-anti-mouse-cd45r-b220.561101
CD19 (FITC 1D3)	BD Pharmingen	553785		https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/fitc-rat-anti-mouse-cd19.553785
CD5 (APC 53-7.3)	BD Pharmingen	553023		https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/pe-rat-anti-mouse-cd5.553023
Anti-mouse CD4 BUV737	BD Bioscience	612843	1:100	https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/buv737-rat-anti-mouse-cd4.612843
Anti-mouse CD8 BUV805	BD Bioscience	612898	1:100	https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/buv805-rat-anti-mouse-cd8a.612898
Anti-mouse IFN γ PE	BD Bioscience	554412	1:100	https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/pe-rat-anti-mouse-ifn.554412
Anti-mouse IL-17A APC-Cy7	BD Bioscience	560821	1:100	https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/apc-cy-7-rat-anti-mouse-il-17a.560821

Anti-mouse IL-4 PECy7	BD Bioscience	560699	1:100	https://wwwbdbiosciences.com/en-us/products/reagents/flow-cytometry-reagents/research-reagents/single-color-antibodies-ruo/pe-cy-7-rat-anti-mouse-il-4.560699
Anti-mouse IL-15Ra	Invitrogen	17-7149-82	1:100	https://www.thermofisher.com/antibody/product/CD215-IL-15Ra-Antibody-clone-DNT15Ra-Monoclonal/17-7149-82
Anti-mouse CD3	Biolegend	100205	N/A	https://www.biolegend.com/en-us/products/pe-anti-mouse-cd3-antibody-47?GroupID=BLG242
Anti-mouse CD28	Biolegend	102102	N/A	https://www.biolegend.com/en-us/products/purified-anti-mouse-cd28-antibody-117

Data & Code Availability

Not applicable

Other

Not applicable