

Supplementary Data

Dapagliflozin does not modulate atherosclerosis in mice with insulin resistance

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Supplementary Tables

Table S1. Sequences of primers used for gene expression analysis by qPCR

Gene	Primer	Sequence 5'→3'
<i>Cyclophilin</i>	Forward	AGATGGAGAACCGACGTATCA
	Reverse	ACTGAGCGTGCTGACAAGTC
<i>Cd11c</i>	Forward	ATGGAGCCTCAAGACAGGAC
	Reverse	GGATCTGGGATGCTGAAATC
<i>Cd206</i>	Forward	GCATGGGTTTTACTGCTACTTGATT
	Reverse	CAGGAATGCTTGTTTCATATCTGTCTT
<i>Il6</i>	Forward	CCCAACAGACCTGTCT
	Reverse	CCAGTTTGGTAGCATCC
<i>Mcp1</i>	Forward	CTTCTCCACCACCATGCA
	Reverse	CCAGCCGGCAACTGTGA
<i>Tnfa</i>	Forward	CCCACACCGTCAGCCGATTT
	Reverse	GTCTAAGTACTTGGGCAGATTGACC

Supplemental Figures

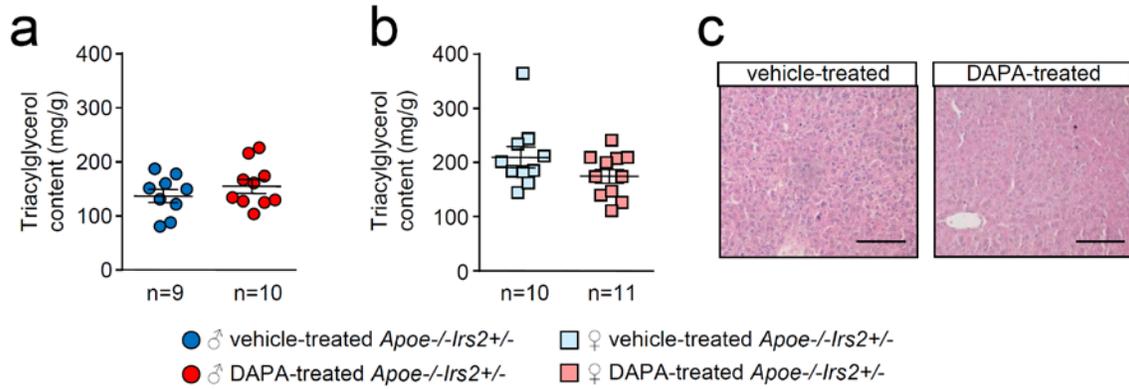


Figure S1. Hepatic triglyceride content in vehicle- and DAPA-treated *Apoe-/-Irs2+/-* mice. Hepatic triglyceride content (mg/g liver tissue) in (a) male and (b) female mice. (c) Representative images of hematoxylin-eosin stained cross-sections of vehicle- and DAPA-treated *Apoe-/-Irs2+/-* mice. Scale bar: 100 μ m. Data is represented as individual points with mean \pm sem. The statistical analysis for normality was D'Agostino-Pearson omnibus test and for differences were the student's t-test (a) and Mann-Whitney U test (b).

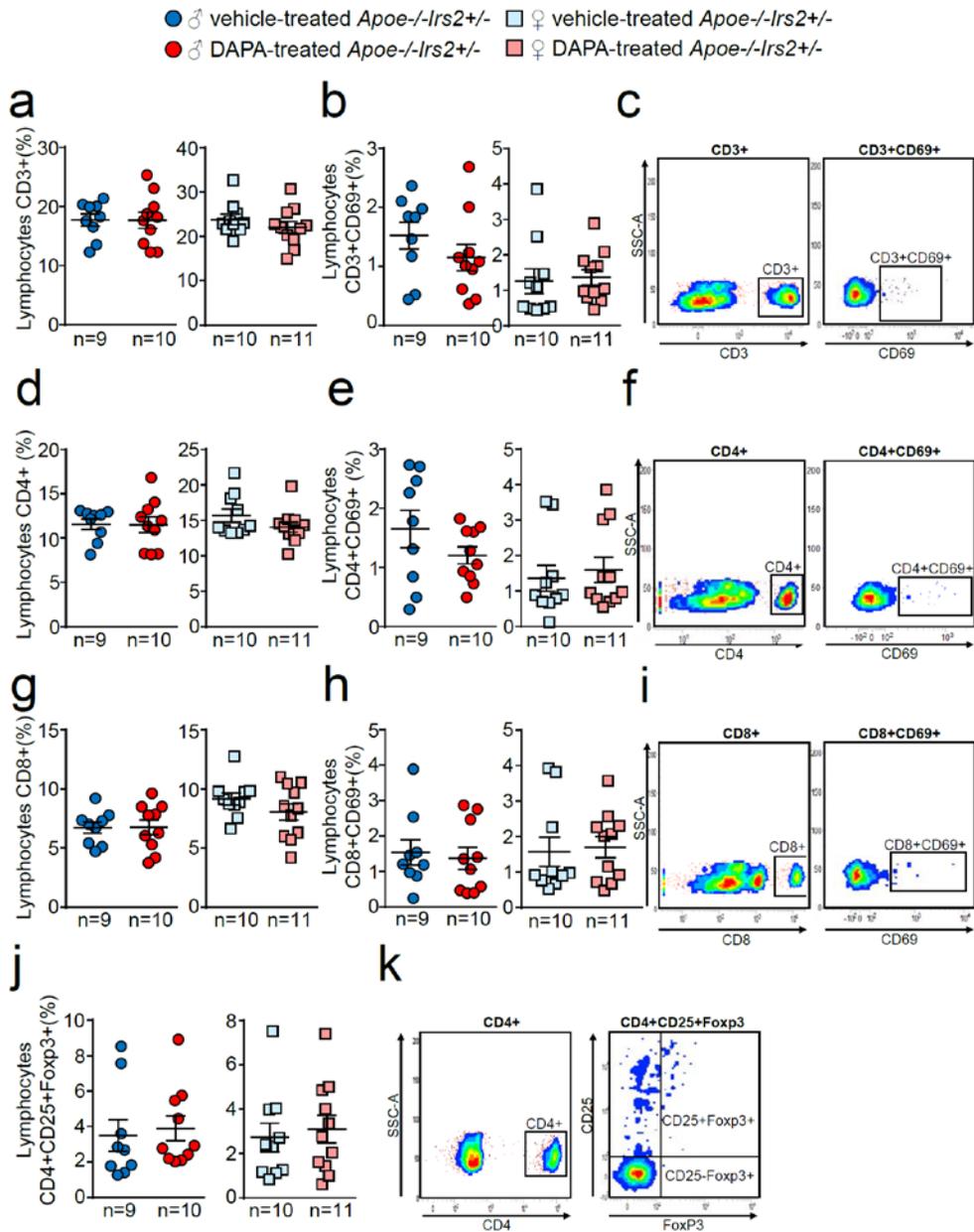


Figure S2. Analysis of circulating blood levels of total and subtypes of lymphocytes in male and female vehicle- and DAPA-treated *ApoE-/-Irs2+/-* mice. Circulating blood percentages of total (a) CD3+, (b) activated CD3+CD69+, (d) CD4+, (e) activated CD4+CD69+, (g) CD8+, and (h) activated CD8+CD69+ lymphocyte subsets in vehicle and DAPA-treated mice. (j) Levels of CD4+CD25+Foxp3+ Treg lymphocytes in circulating blood of vehicle and DAPA-treated *ApoE-/-Irs2+/-* mice. (c,f,i,k) Representative plots of the gating strategy used for flow cytometry analysis in the different lymphocyte subpopulations in blood samples are shown. The statistical analysis for normal distribution was D'Agostino-Pearson test and for differences were Student's t-test (right panels in a,b,d and h,e,g and left panel in j) and Mann-Whitney U test (left panels in a,b,d and h, and right panel in j).

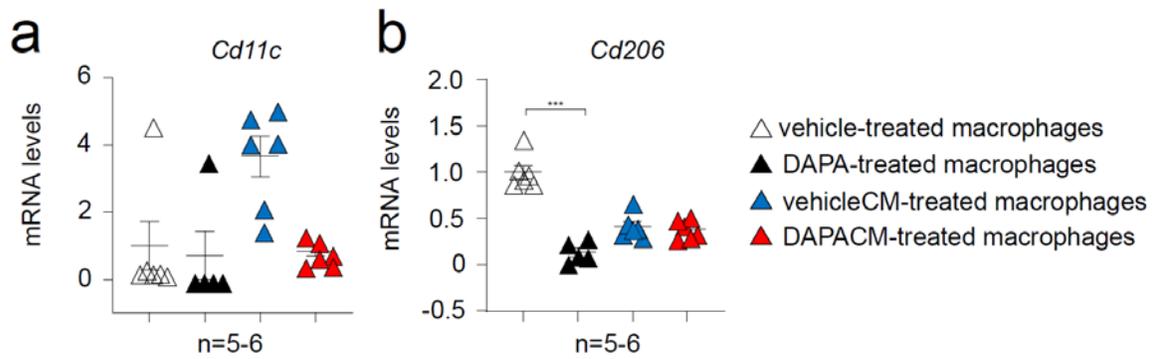


Figure S3. Effect of dapagliflozin in *ApoE*^{-/-}*Irs2*^{+/-} mouse macrophage phenotype. mRNA expression of (a) *Cd11c* integrin and the (b) *Cd206* mannose receptor in *ApoE*^{-/-}*Irs2*^{+/-} macrophages treated with vehicle (vehicle-treated), 1 μ M of dapagliflozin (DAPA-treated), conditioned media containing 10% of plasma from vehicle-treated mice (vehicleCM), or conditioned media containing 10% of plasma from DAPA-treated mice (DAPACM). The statistical analysis for normal distribution was Saphiro-Wilk test and for differences was Kruskal-Wallis followed by Dunn's multiple comparison test.***p<0.001.