

Supplemental Information

Defective fractalkine-CX3CR1 signaling aggravates neuroinflammation and affects recovery from cuprizone-induced demyelination

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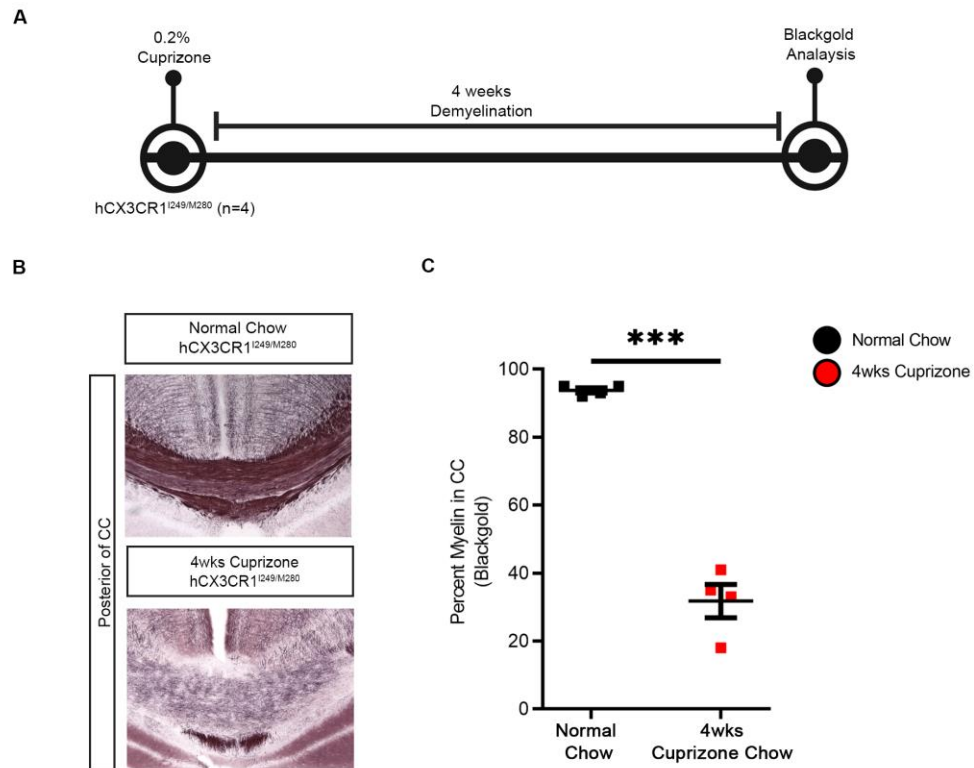
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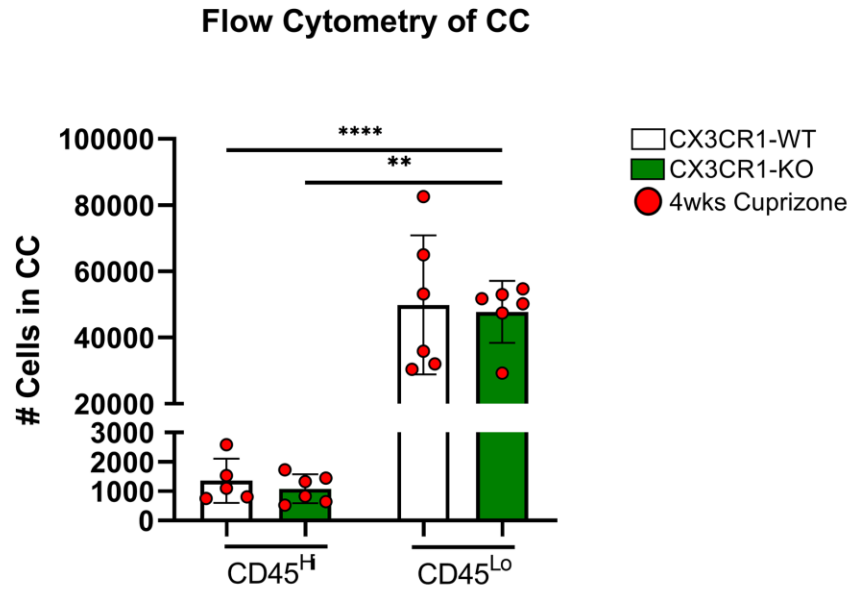
Contributed equally to this work.

Supplementary Figure 1



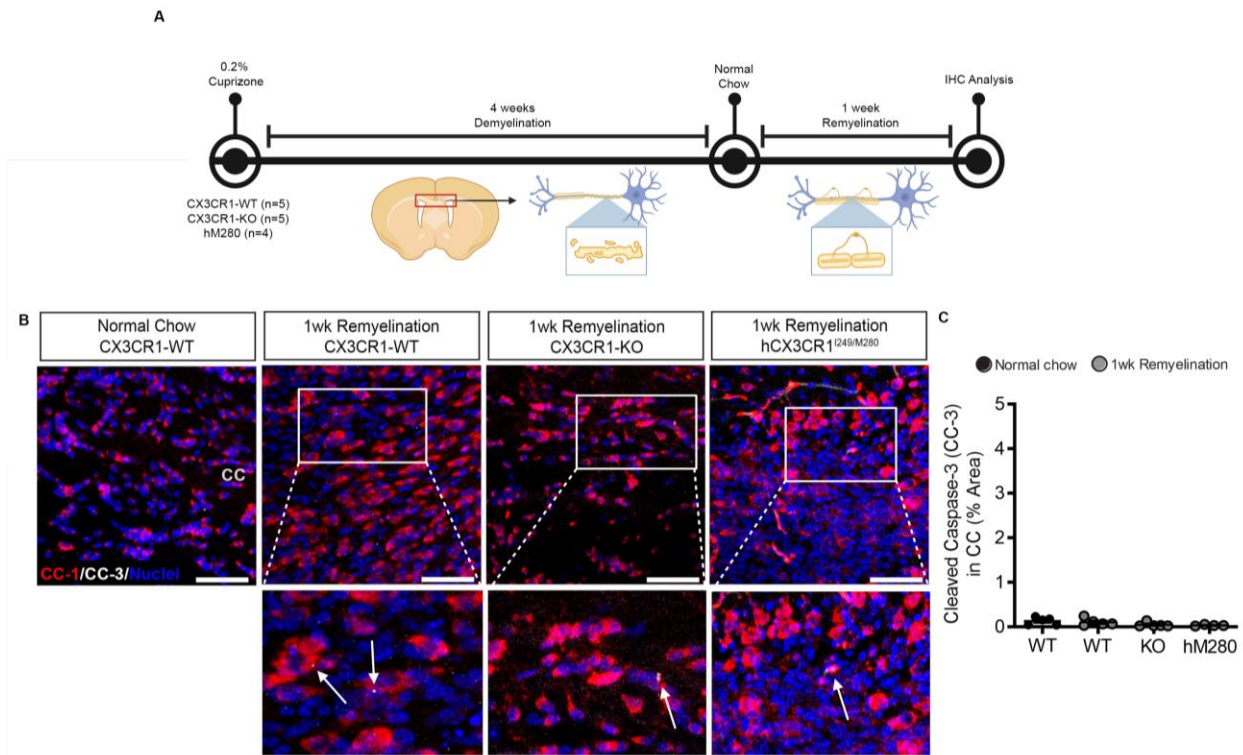
Supplementary Figure 1. Histological analysis of myelin in $hCX3CR1^{I249/M280}$ mice. A, Experimental design for 4 wks cuprizone-induced demyelination and downstream analyses in $hCX3CR1^{I249/M280}$ mice. **B,** Representative images of brain sections from $hCX3CR1^{I249/M280}$ mice fed normal chow or cuprizone, stained for Blackgold. **C,** Image quantification of myelin staining in the posterior of the corpus callosum as shown in **B**. Data are mean \pm SEM for $n = 4$ per group, each dot represents an individual mouse. *** <0.001 using Mann-Whitney two-tailed t-test.

Supplementary Figure 2



Supplementary Figure 2. Flow cytometric analysis of resident and infiltrating myeloid cells following acute cuprizone. Flow cytometric analysis showing cell numbers of CD45^{hi}CD11b⁺ (infiltrating) and CD45^{lo}CD11b⁺ (resident) cells from corpus callosum of CX3CR1-WT and CX3CR1-KO mice fed cuprizone. Data are mean ± SEM for $n = 5-6$ mice per group, each dot represents an individual mouse. **** <0.001 by one-way ANOVA followed by Tukey's posttest.

Supplementary Figure 3



Supplementary Figure 3. Histological analysis of cell death in mature oligodendrocytes. A, Experimental design for remyelination studies following cuprizone-induced demyelination. **B,** Representative images of brain sections from CX3CR1-WT, CX3CR1-KO and hCX3CR1^{I249/M280} mice following 1wk remyelination (4wks cuprizone + 1 wk normal chow) or fed normal chow immunostained for CC-1 (red) and CC-3 (white). **C,** Image quantification of CC-3 staining in the posterior of the corpus callosum as shown in **B**. Data are mean \pm SEM for $n = 4-5$ per group, each dot represents an individual mouse.