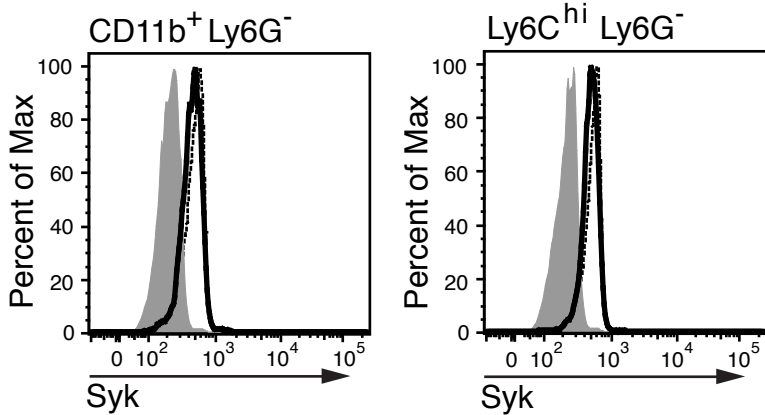
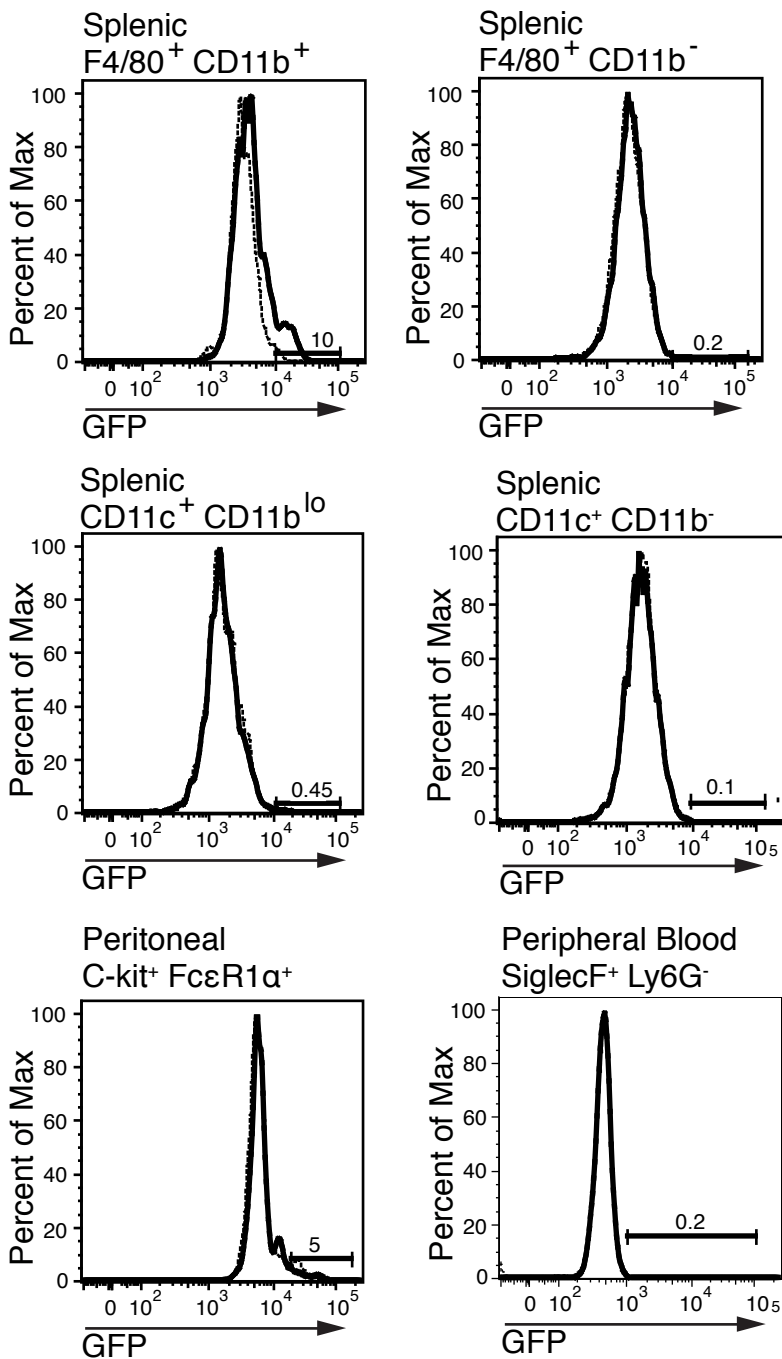


A. Syk Expression in Peripheral Blood Leukocytes

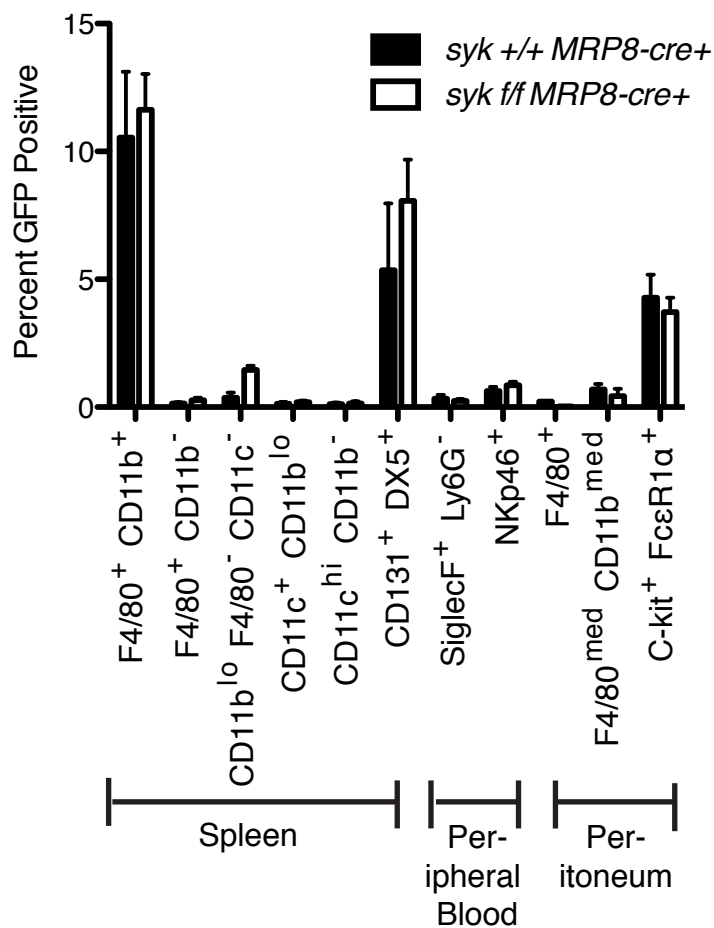


B. MRP8-cre-GFP Expression in Leukocytes



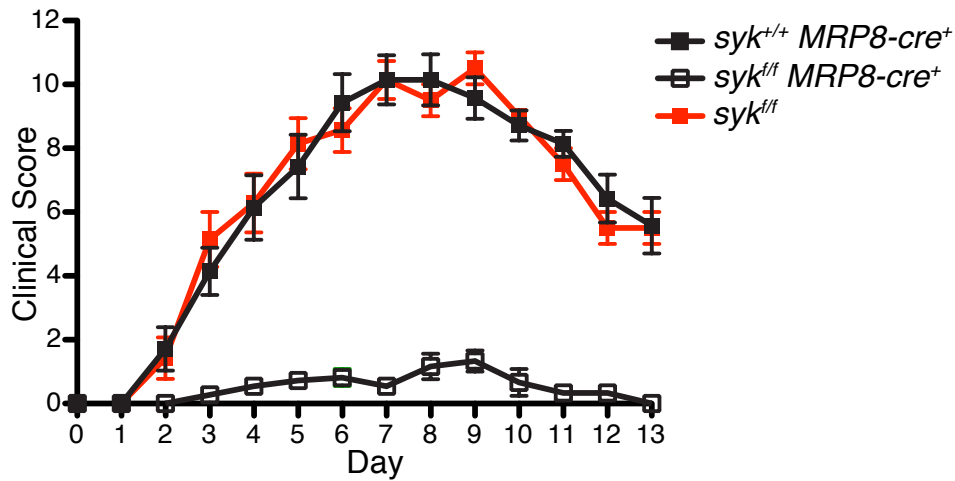
----- C57BL/6
 ■ *syk*^{-/-}
 — *syk*^{ff} *MRP8-cre*⁺

C. MRP8-cre-GFP Expression in Leukocytes

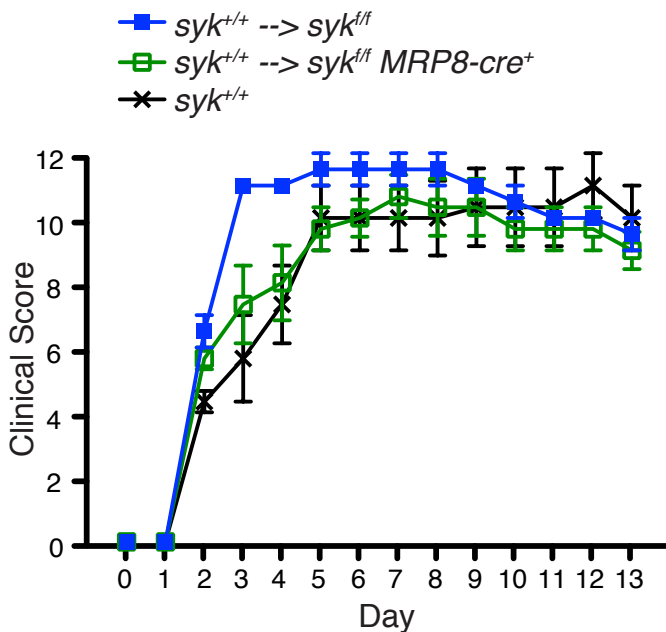


Supplemental Figure 1. MRP8-cre-GFP expression is limited mainly to neutrophils. **A**, Representative histograms of Syk expression of peripheral blood monocyte populations ($CD11b^+ Ly6G^-$ and $Ly6C^{hi} Ly6G^-$) from C57BL/6 (black dashed line), *syk*^{-/-} (solid gray), and *syk*^{ff} *MRP8-cre*⁺ (solid black line) mice. **B**, Representative histograms of GFP expression in splenic macrophages ($F4/80^+$) and dendritic cells ($CD11c^+$), peritoneal mast cells ($C-kit^+ Fc\epsilon R1\alpha^+$) and peripheral blood eosinophils ($SiglecF^+ Ly6G^-$) from C57BL/6 (black dashed line) and *syk*^{ff} *MRP8-cre*⁺ (black line) mice. Percent GFP positive in the *syk*^{ff} *MRP8-cre*⁺ mice are shown. **C**, GFP expression in various leukocytes as defined above, as well as $CD11b^{lo} F4/80^- CD11c^-$ monocytes/macrophages, basophils ($CD131^+ DX5^+$), and natural killer (NK) cells ($NKp46^+$). Data from *syk*^{+/+} *MRP8-cre*⁺ (black bars, n=3) and *syk*^{ff} *MRP8-cre*⁺ (white bars, n=6-12) mice. Error bars represent SEM.

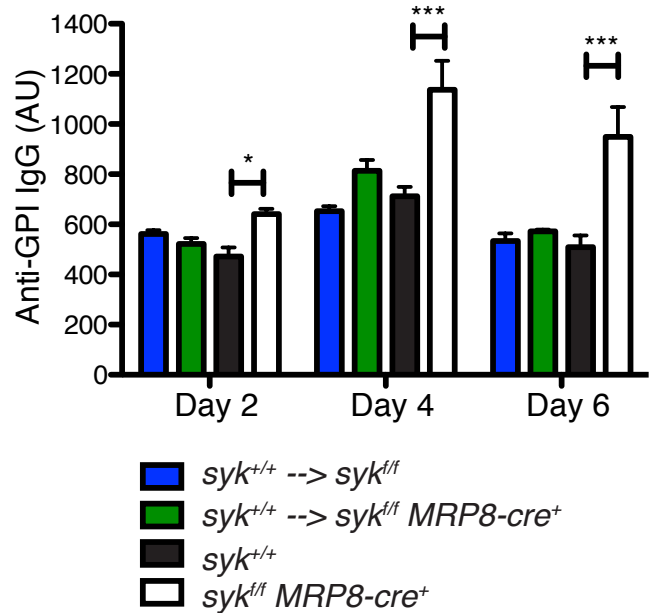
A. Arthritis in *syk^{fl/fl} MRP8-cre⁺* Mice



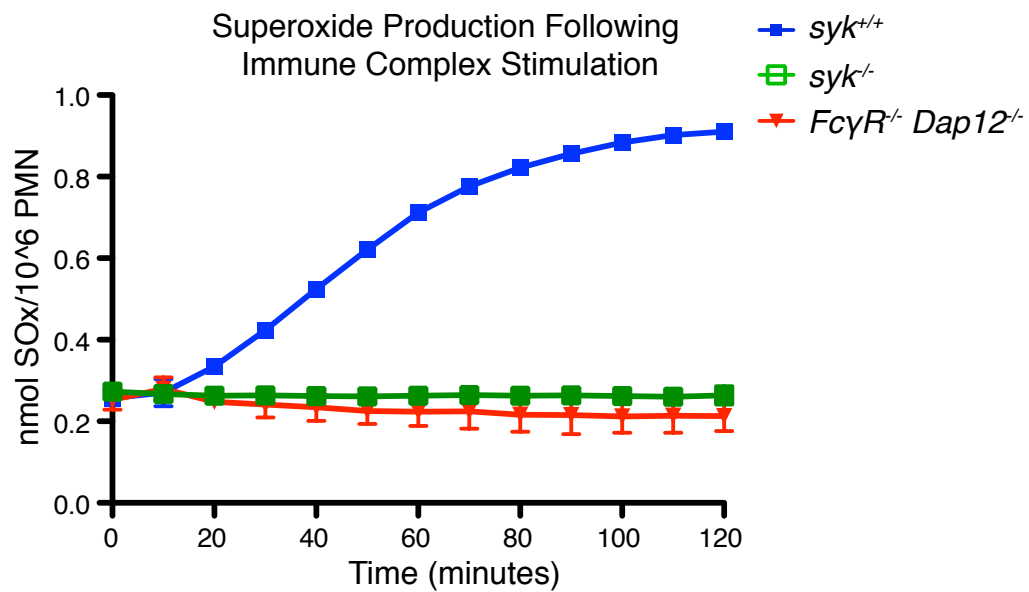
B. Arthritis in Chimeric Mice



C. Serum Anti-IgG Titers



Supplemental Figure 2. Conditional deletion of Syk in neutrophils protects mice from K/BxN serum-induced arthritis. **A**, Clinical score was recorded at the indicated times following K/BxN serum transfer in *syk^{fl/fl}*, *syk^{+/+} MRP8-cre⁺*, and *syk^{fl/fl} MRP8-cre⁺* mice (n=4). Use of these genotypes controls for potential effects of the *syk^{fl/fl}* mutation – showing that *Syk^{fl/fl}* mice respond normally to K/BxN serum. Representative plots from 2 independent experiments. **B**, **C**, *syk^{fl/fl} MRP8-cre⁺* recipients were lethally irradiated and reconstituted with congenically marked (CD45.1) wild type bone marrow to control for the contribution of radioresistant cells to the phenotype of *syk^{fl/fl} MRP8-cre⁺* mice. **B**, Clinical score and **C**, anti-GPI IgG serum titers were recorded at the indicated times following K/BxN serum transfer in *syk^{+/+} --> syk^{fl/fl}* (n=2) and *syk^{+/+} --> syk^{fl/fl} MRP8-cre⁺* (n=3) chimeric mice, and non-irradiated *syk^{+/+}* and *syk^{fl/fl} MRP8-cre⁺* mice (n=3). Error bars represent \pm SEM. * = p < 0.05, *** = p < .001, analyzed by two-way ANOVA.



Supplemental Figure 3. Syk-deficient neutrophils show impaired superoxide production in response to immune complex stimulation *in vitro*. Bone marrow-derived neutrophils from the indicated mice were plated with immune complexes and superoxide (SOx) production was monitored as described (VanZiffle and Lowell, Blood, 144:4871 (2009)). FcγR^{-/-} Dap12^{-/-} cells, lacking all activating FcγRs, served as negative controls.