

Supplementary Figure 1

 $\mathrm{SW}_{\mathrm{HEL}}$ frequency B220^{lo} Representative flow cytometry plots and of cells identified A) as CD45.2⁺CD45.1⁻CD3⁻B220⁺HEL^{WT+} day 5 after immunization. B) Representative flow cytometry plots, frequency, and absolute cell counts of GL7+Fas+ GCBs at the indicated time points after immunization identified as CD45.2+CD45.1-CD3-B220+HELWT+. C) gMFIs of relevant receptors on donor-specific GCBs, identified as CD45.2⁺CD45.1⁻CD3⁻B220⁺HEL^{WT+}CD38^{lo}Fas⁺. D) In vitro analysis of IRF4 expression levels between *Irf4^{+/+}* and *Irf4^{+/-}* B cells. Equivalent numbers of CD45.2⁺CD45.1⁺ Irf4^{+/+} or CD45.2⁺CD45.1⁻ Irf4^{+/-} B cells were stimulated in cocultures with algD-Dextran and analyzed at 72 or 96 hours for expression of the CD45 allotypic markers and IRF4 by flow cytometry. E) BCR expression; HEL^{WT+} gMFI on total SW_{HEL}, IgG1⁺ SW_{HEL}, or SW_{HEL} GCBs. Panel on right is IgG1 gMFI on total SW_{HEL} on day 10 and 15. We note that, in some panels, dots that cluster together represent mice from separate experiments. Experiment in (A) is from 15 mice in 4 experiments performed. Experiment in (B) is from 16 mice in 4 experiments performed, while (C) is from 5 mice in 1 experiment performed; contour plots are concatenated files from all mice of a given group in a given experiment. Experiments in (E) are from 12 mice in 3 experiments performed. Unpaired t tests were performed to determine significance: all ns, not significant (p > 0.05), unless noted. ** $p \le 0.01$, **** $p \le 0.0001$.



Supplementary Figure 2

A) Frequency of mutated "C" in either $Irf4^{+/+}$ or $Irf4^{+/-}$ total or high affinity clones within "WRC" AID hotspot motif sequences over the VDJ. B) Frequency of $Irf4^{+/+}$ and $Irf4^{+/-}$ total SW_{HEL} cells or total SW_{HEL} GCBs at indicated time points, 1:3 initial cotransfer ($Irf4^{+/+} : Irf4^{+/-}$), transfer and immunization as in **Fig. 2A**. C) Frequency of high affinity SW_{HEL} or IgG1⁺ SW_{HEL} cells, 1:3 initial cotransfer ($Irf4^{+/+} : Irf4^{+/-}$). Lines linking dots denote measurements from a single mouse. D) Equal numbers of $Irf4^{+/+}$ and $Irf4^{-/-}$ SW_{HEL} cells were cultured in vitro with indicated amounts of HEL^{2X}-IE α or HEL, and MHCII presentation of IE α was measured by Y-Ae frequency and gMFI. $Irf4^{+/+}$ cells were distinguished from $Irf4^{-/-}$ by CFSE labeling. Flow cytometry plots shown are representative of 3 separate co-culture experiments and lines linking dots denote measurements from a single well. Paired t tests were performed to determine significance: ** p ≤ 0.01, ns, not significant (p > 0.05). E) Frequency of cells in the scRNA-seq experiment having a given number of sequences encoded by the anti-CD45.1 CITE-seq approach; line represents gating for genotype calling of single cells. A. Cluster:

0 2 tSNE

-30

-60

-50

-25

25

0 tSNE_1

50



Supplementary Figure 3

A) Heat map displaying relative expression of genes that most distinguished identified clusters. B) Distribution of cells expressing signature genes of the G1, G2/M, or S phase of the cell cycle in the t-SNE reduction projection.

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