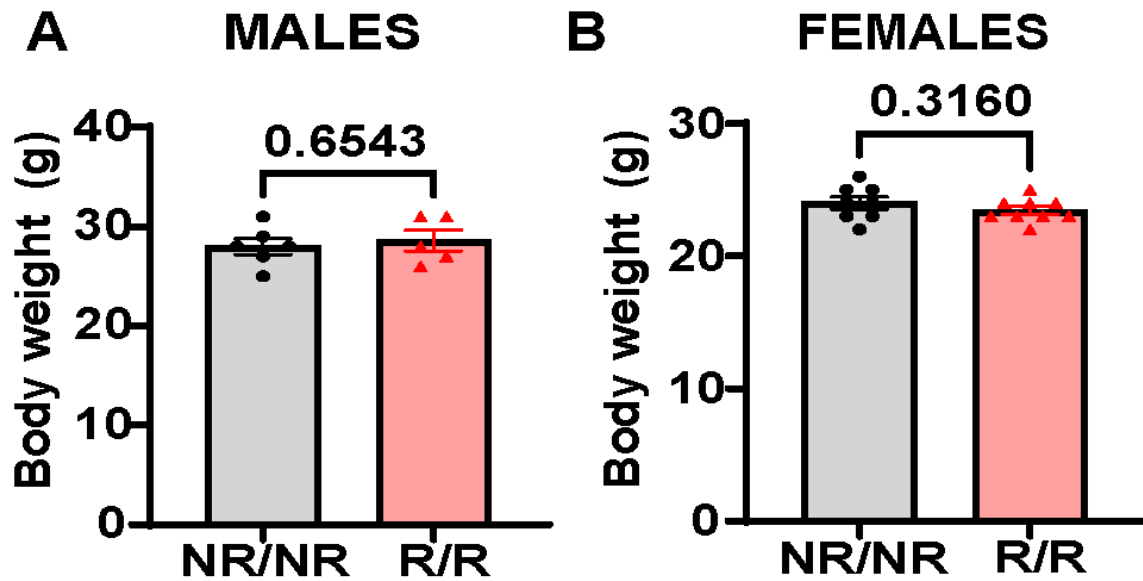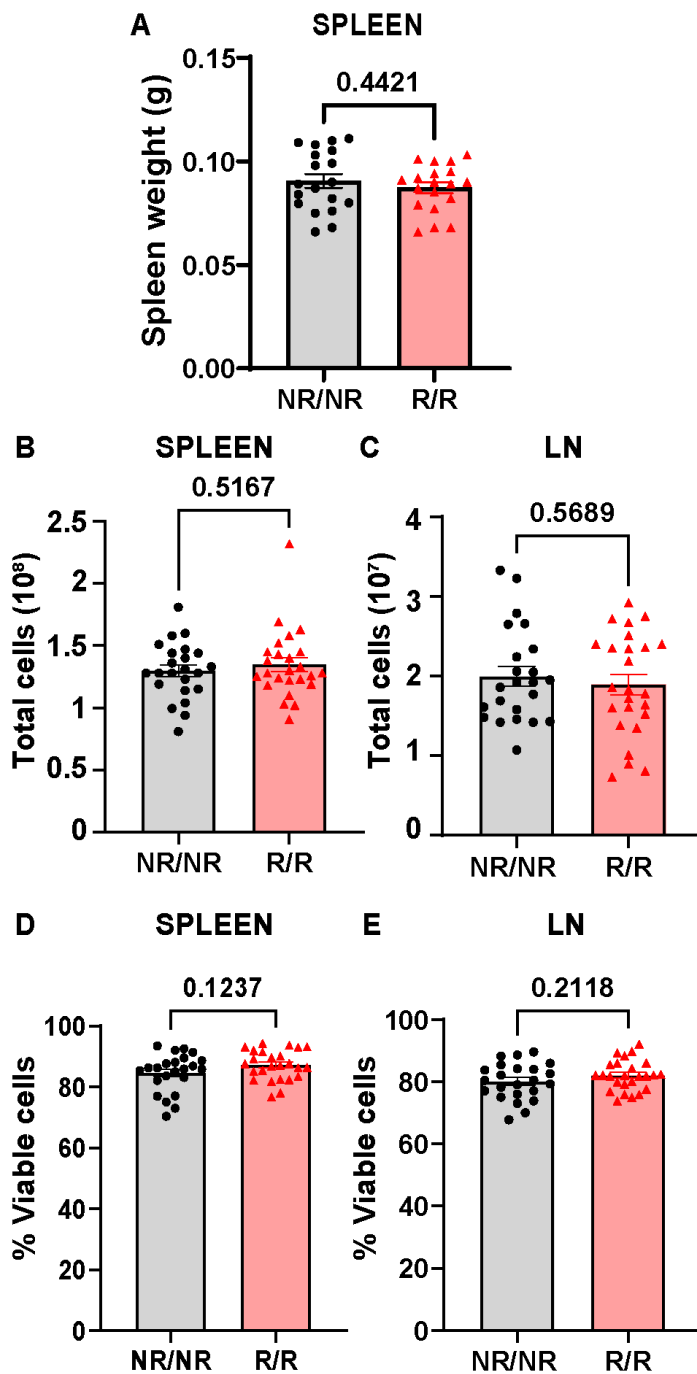Supplementary Figure 1.** *NOD.Ifih1$^{R/R}$* **mice do not display significant changes in *Ifih1* **mRNA in 14-week-old mice.** The relative mRNA expression of *Ifih1* in 14-week-old (**A**) males (*n*=9 *NOD.Ifih1$^{NR/NR}$*, *n*=6 *NOD.Ifih1$^{R/R}$*) and (**B**) females (*n*=11 *NOD.Ifih1$^{NR/NR}$*, *n*=12 *NOD.Ifih1$^{R/R}$*). P values were determined using student's unpaired *t* tests. All data are mean ± SEM.

Supplementary Figure 2. *NOD.Ifih1$^{R/R}$* **mice do not display significant changes in body weight.** Body weights measured at a single time point immediately prior to euthanasia in 14-week-old *NOD.Ifih1$^{NR/NR}$* compared to *NOD.Ifih1$^{R/R}$* mice in (**A**) males (*n*=6 *NOD.Ifih1$^{NR/NR}$*, *n*=5 *NOD.Ifih1$^{R/R}$*) and (**B**) females (*n*=8 *NOD.Ifih1$^{NR/NR}$*, *n*=9 *NOD.Ifih1$^{R/R}$*). P values were determined using student's unpaired *t* tests. All data are mean ± SEM.
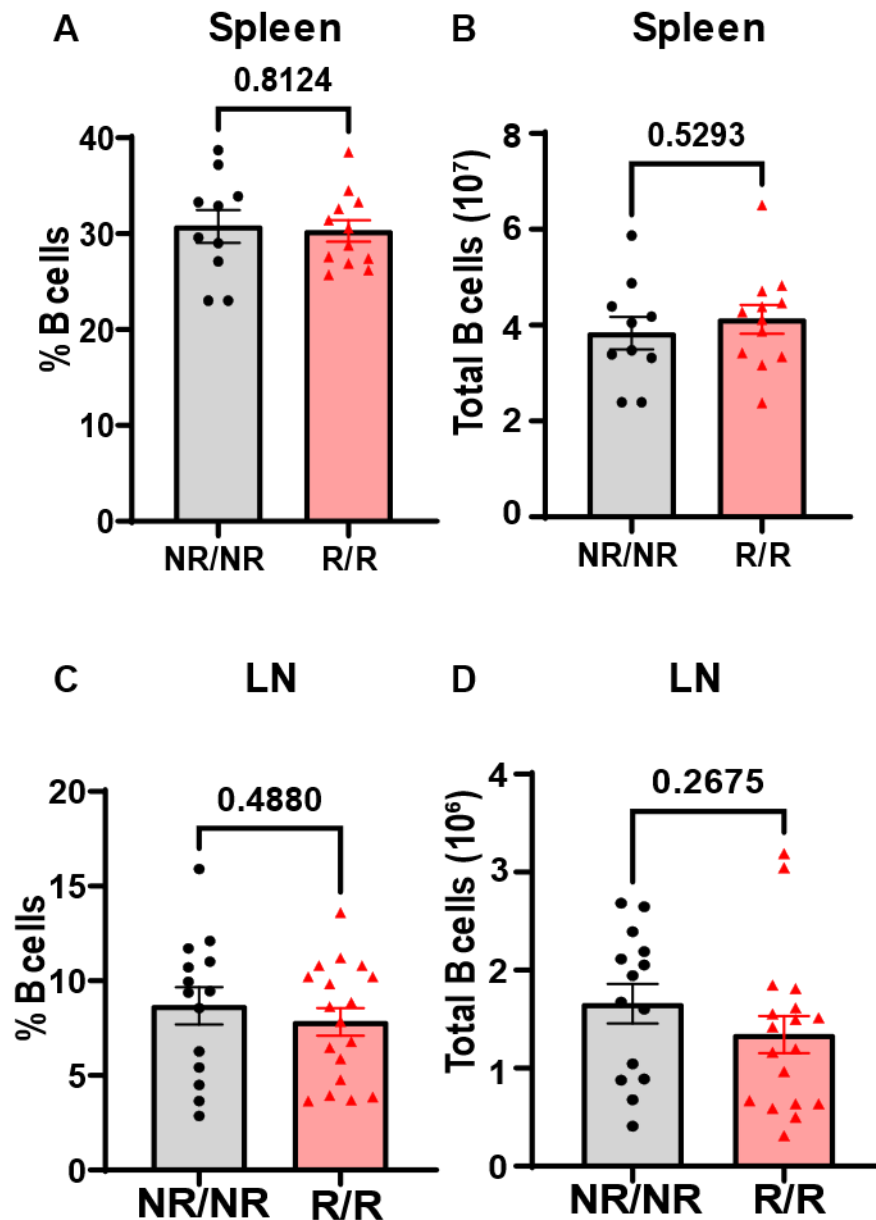
Supplementary Figure 3. *NOD.Ifih1$^{R/R}$* **mice do not exhibit significant changes in splenic weight or cell numbers or viability in spleens or lymph nodes.** (**A**) Spleen weight in grams (g) measured at the time of euthanasia in 14-week-old *NOD.Ifih1$^{NR/NR}$* compared to *NOD.Ifih1$^{R/R}$* mice (*n*=19 mice per genotype). (**B-C**) Total number of cells in (**B**) spleens (*n*=23 *NOD.Ifih1$^{NR/NR}$*, *n*=25 *NOD.Ifih1$^{R/R}$*) and (**C**) lymph nodes (LN, *n*=23 *NOD.Ifih1$^{NR/NR}$*, *n*=25 *NOD.Ifih1$^{R/R}$*). (**D-E**) Percent viable cells in (**D**) spleens (*n*=23 *NOD.Ifih1$^{NR/NR}$*, *n*=25 *NOD.Ifih1$^{R/R}$*) and (**E**) lymph nodes (*n*=23 *NOD.Ifih1$^{NR/NR}$*, *n*=25 *NOD.Ifih1$^{R/R}$*). P values were determined using student's unpaired *t* tests. All data are mean ± SEM.
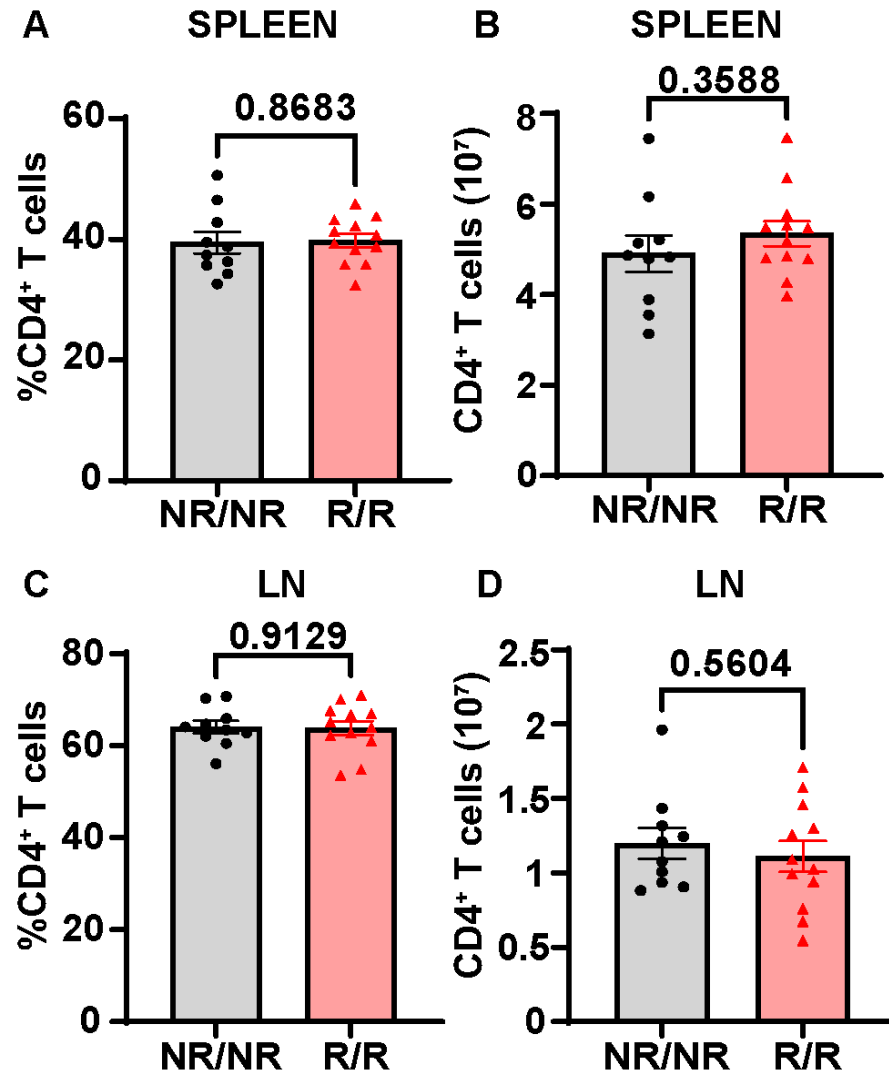
Supplementary Figure 4. *NOD.Ifih1$^{R/R}$* **mice do not exhibit changes in the frequency or total cell numbers of CD4$^+$ T cells.** **(A-B)** CD4$^+$ T cells in the spleen as **(A)** frequency of live cells and **(B)** total CD4$^+$ cell numbers in *NOD.Ifih1$^{NR/NR}$* compared to *NOD.Ifih1$^{R/R}$* mice (*n*=10 *NOD.Ifih1$^{NR/NR}$*, *n*=12 *NOD.Ifih1$^{R/R}$*). **(C-D)** CD4$^+$ T cells in the lymph nodes (LN) as **(C)** frequency of live cells and **(D)** total CD4$^+$ cell numbers in *NOD.Ifih1$^{NR/NR}$* compared to *NOD.Ifih1$^{R/R}$* mice (*n*=10 *NOD.Ifih1$^{NR/NR}$*, *n*=12 *NOD.Ifih1$^{R/R}$*). P values were determined using student's unpaired *t* tests. All data are mean ± SEM.
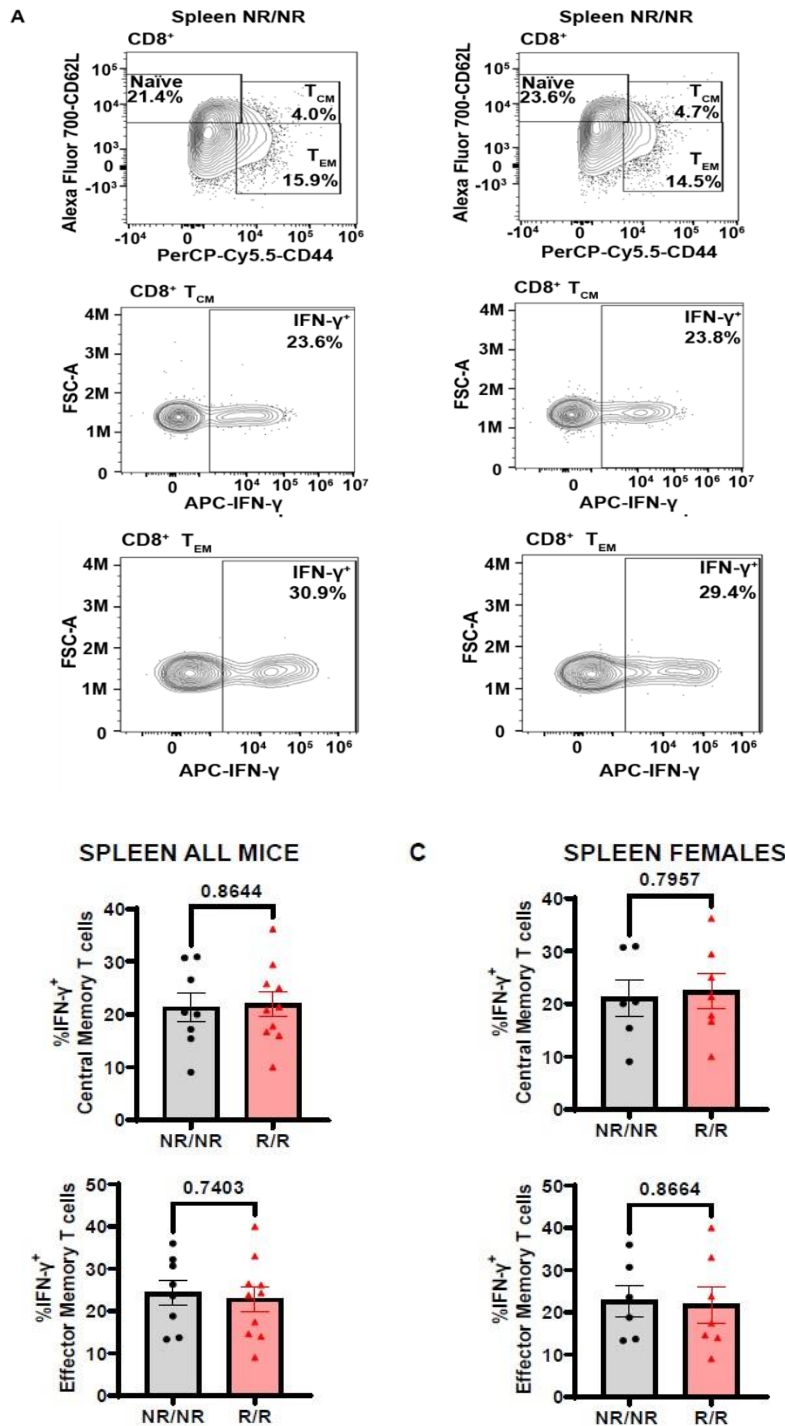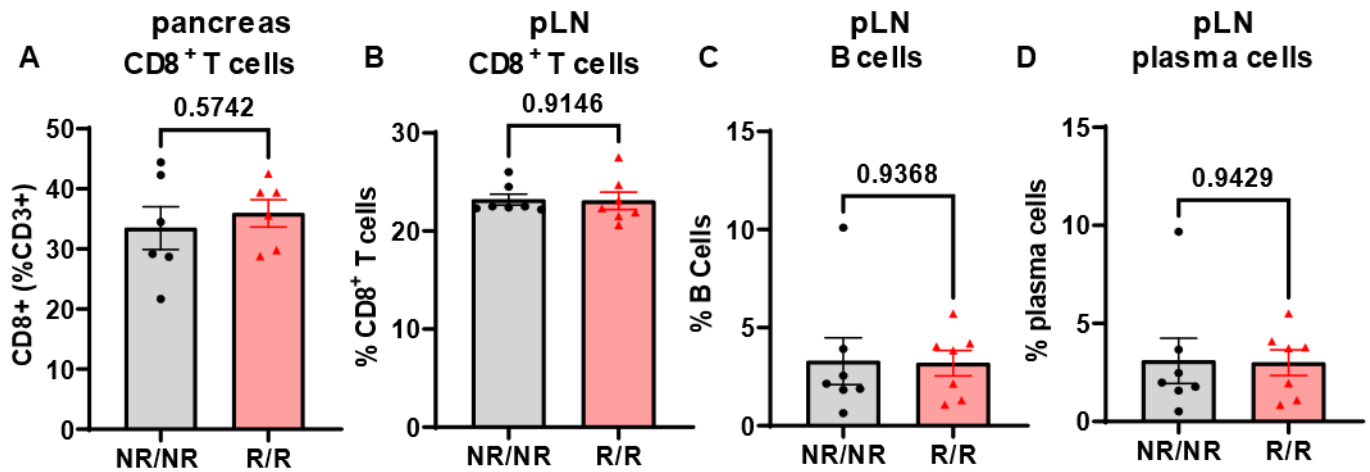
Supplementary Figure 5. *NOD.Ifih1^R* **does not affect the frequency of IFN-γ on splenic CD8$^+$ central memory and effector memory T cells.** (**A**) Representative gating of CD8$^+$ naïve (CD62L$^+$), central memory (CD62L$^+$CD44$^+$, T$_{CM}$), effector memory (CD62L$^-$CD44$^+$, T$_{EM}$), and IFN-γ$^+$ CD8$^+$ central memory and effector memory T cells from PMA/ionomycin-treated spleens. (**B-C**) The frequency of IFN-γ$^+$ CD8$^+$ central memory and effector memory T cells from PMA/ionomycin-treated lymph nodes from (**B**) all mice (*n*=8 *NOD.Ifih1$^{NR/NR}$*, *n*=10 *NOD.Ifih1$^{R/R}$*) and (**C**) females only (*n*=6 *NOD.Ifih1$^{NR/NR}$*, *n*=7 *NOD.Ifih1$^{R/R}$*). P values were determined using student's unpaired *t* tests. All data are mean ± SEM.

Supplementary Figure 6. *NOD.Ifih1^R* **does not affect the frequency or total number of B cells. (A-B)** Splenic B cells from 14-week-old mice (*n=10 NOD.Ifih1$^{NR/NR}$*, *n=12 NOD.Ifih1$^{R/R}$*) as **(A)** frequency of live cells and **(B)** total number of splenic B cells. **(C-D)** Lymph node (LN)-derived B cells from 14-week-old mice (*n=14 NOD.Ifih1$^{NR/NR}$*, *n=18 NOD.Ifih1$^{R/R}$*) as **(C)** frequency of live cells and **(D)** total number of lymph node B cells. P values were determined using student's unpaired *t* tests. All data are mean ± SEM.
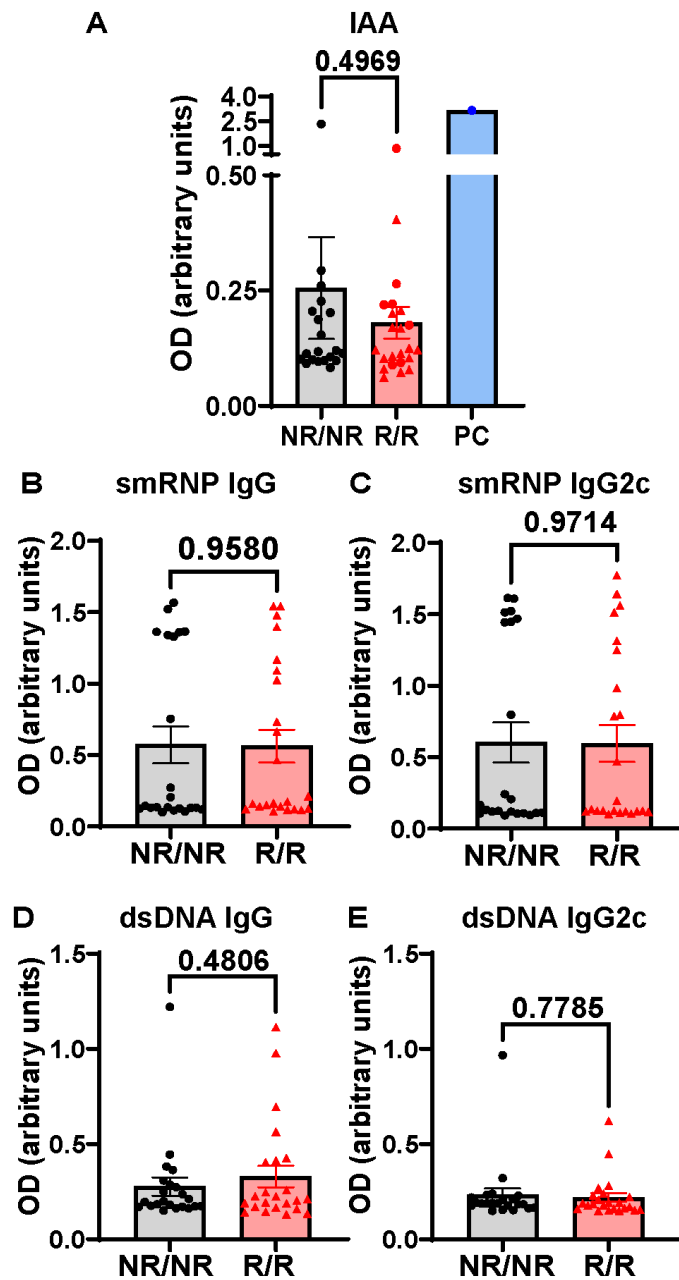
Supplementary Figure 7. *NOD.Ifih1^{R/R}* **mice do not display significant changes in insulin, smRNP, or dsDNA autoantibody production.** **(A)** OD measurements from insulin autoantibody (IAA) ELISAs indicative of the levels of IAA in serum from 14-week-old mice (*n*=20 *NOD.Ifih1^{NR/NR}*, *n*=23 *NOD.Ifih1^{R/R}*). PC: positive control provided by manufacturer. **(B-C)** OD measurements from smRNP **(B)** IgG and **(C)** IgG2c autoantibody ELISAs indicative of the levels of smRNP autoantibody in serum from 14-week-old mice (*n*=22 *NOD.Ifih1^{NR/NR}*, *n*=23 *NOD.Ifih1^{R/R}*). **(D-E)** OD measurements from dsDNA **(D)** IgG and **(E)** IgG2c autoantibody ELISAs indicative of the levels of dsDNA autoantibody in serum from 14-week-old mice (*n*=22 *NOD.Ifih1^{NR/NR}*, *n*=23 *NOD.Ifih1^{R/R}*). P values were determined using student's unpaired *t* tests. All data are mean ± SEM.

Supplementary Figure 8. *NOD.Ifih1^R* **does not affect the frequency of CD8$^+$ T cells in the pancreas or the frequency of immune cells in the draining lymph nodes of 14-week-old mice. (A-B)** The frequency of CD8$^+$ T cells in the **(A)** pancreas (*n*=6 per genotype) and **(B)** pancreatic lymph nodes (pLN, *n*=7 per genotype). **(C)** The frequency of B cells (B220$^+$) in the pancreatic lymph nodes (*n*=7 per genotype). **(D)** The frequency of plasma cells (CD19$^+$B220$^-$CD138$^+$) in the pancreatic lymph nodes (*n*=7 per genotype). All data are from 14-week-old mice. P values were determined using student's unpaired *t* tests. All data are mean ± SEM.

Supplementary Table 1. PCR and qRT-PCR Primers

Primers for PCR/sequencing	
A946 F1	5'- TTCAAGTGAGTGCAGCAAGA -3'
A946 R1	5'- AGGCAATTCCACCCACTTCT -3'
A946 R2	5'- GTGCACCATCATTGTTCCCC -3'
Primers for qRT-PCR	
Gene/primer	**Sequence**
Hprt-forward	5'-TTG CTG ACC TGC TGG ATT ACA-3'
Hprt-reverse	5'-CCC CGT TGA CTG ACT GAT CAT TAC A-3'
Ifih1-forward	5'-AGA TCA ACA CCT GTG GTA ACA CC-3'
Ifih1-reverse	5'-CTC TAG GGC CTC CAC GAA CA-3'
Ifit1-forward	5'-TGC TGA GAT GGA CTG TGA GG-3'
Ifit1-reverse	5'-CTC CAC TTT CAG AGC CTT CG-3'
Mx1-forward	5'-GGC AGA CAC CAC ATA CAA CC-3'
Mx1-reverse	5'-CCT CAG GCT AGA TGG CAA G-3'
Oas1-forward	5' CTT AGC ATG GAG CAC GGA CT-3'
Oas1-reverse	5' AGC ACC TTG GAA GCA TCT CTC-3'

Supplementary Table 2. Flow Cytometry Antibodies

Antibody	Company	Catalog #
BD Horizon™ BUV661 Rat Anti-Mouse CD19	BD Biosciences	612971
Brilliant Violet 711™ anti-mouse CD138 (Syndecan-1)	Biolegend	142519
Near IR fluorescent reactive dye	Thermo Fisher	L34994 A
Pacific Blue™ anti-mouse/human CD45R/B220	Biolegend	103227
Alexa Fluor® 700 anti-mouse/human CD45R/B220	Biolegend	103232
PE anti-mouse CD80	Biolegend	104707
PE/Cy7 anti-mouse CD3ε	Biolegend	100320
PerCP/Cyanine5.5 anti-mouse CD8a	Biolegend	100734
Brilliant Violet 711™ anti-mouse CD8	Biolegend	100759
Spark NIR™ 685 anti-mouse CD69	Biolegend	104557
Spark Violet™ 538 anti-mouse CD4	Biolegend	100485
APC anti-mouse IFN-γ	Biolegend	505810
PE-Dazzle 594 anti-mouse IL-10	Biolegend	505034
PerCP/Cyanine5.5 anti-mouse/human CD44	Biolegend	103032
Alexa Fluor® 700 anti-mouse CD62L	Biolegend	104426
BD Horizon™ BUV395 Rat Anti-Mouse CD45	BD Biosciences	564279