# **Supporting Information**

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Fig. S1. IgG polyclonal and autoAbs from 14- to 15-month-old B6-*Fas<sup>(pr</sup>* mice. IgG concentrations in serum were determined by ELISA in wild-type (*wt/wt*) and mutant (*3d/3d*) mice. IgM RF was anti-IgG1.

## Table S1. ANA in B6-Fas<sup>lpr</sup> 3d mice

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Unc93b1	Age (mo)	ANA
Group A		
wt/wt	5.5	4 + Homog, Meta +
wt/wt	5.5	3 + Fine Spk, Meta +
wt/wt	6.7	3 + Fine Spk, Meta +
wt/wt	7.4	3 + Fine Spk, Meta +
wt/wt	7.4	3–4 + Find Spk, Meta +
wt/3d	6.9	2 + Fine Spk, Meta +
wt/3d	7.5	3–4 $+$ Fine Spk, Meta $+$
wt/3d	7.5	2–3 + Fine Spk, Meta +
wt/3d	7.6	4 + Homog, Meta +
3d/3d	6.1	Neg
3d/3d	7.2	Neg
3d/3d	7.6	Neg
3d/3d	7.7	Neg
Group B		
wt/wt	14.5	3–4 + Homogenous, Meta +
wt/wt	14.5	3–4 + Homogenous, Meta +
wt/wt	15.1	1–2 $+$ Fine Spk, Meta $+$
wt/wt	15.1	3 + Clumpy Spk, Meta +
wt/wt	15.1	2–3 + Homogenous, Meta +
3d/3d	14.5	Neg
3d/3d	14.5	Trace Fine Spk, Meta $+*$
3d/3d	14.6	1 + Spk in some cells <sup>†</sup>
3d/3d	14.6	Neg <sup>‡</sup>
3d/3d	14.6	Neg
3d/3d	14.8	Neg
3d/3d	14.8	Neg

ANA staining was performed with 1/100 serum dilution and anti-IgG-FITC. All positive nuclear staining patterns are consistent with anti-chromatin. Trace amounts of staining are considered negative. Homog, homogeneous; Meta, metaphase; Spk, speckled; Neg, negative. Unusual staining patterns for mice are shown in footnotes below. \*Metaphase chromosomes positive with only faint staining of other nuclear components. <sup>†</sup>Human anti-PCNA-like pattern.

<sup>+</sup>1 + Golgi apparatus, but no nuclear staining.

#### Table S2. Lymph node and spleen T cell populations in young and old B6-Fas<sup>lpr</sup> 3d mice

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Unc93b1	Body (g)	Organ (mg)	Cell no. (× 10 <sup>6</sup> )	T cells (%)	CD4 (%)	CD8 (%)	DN (%)	CD8 CD62L <sup>+</sup> (%)	CD8 CD44 <sup>hi</sup> (%)
Spleen									
Young wt	11.0 ± 0.7	49 ± 5	33.3 ± 5.2	$\textbf{22.0} \pm \textbf{1.0}$	$\textbf{52.9} \pm \textbf{0.6}$	$32.1 \pm 0.1$	$13.6\pm0.6$	92.7 ± 0.4	41.1 ± 2.8
Young 3d	13.4 ± 0.3	46 ± 3	25.3 ± 3.7	25.7 ± 1.9	$54.5 \pm 0.5$	$30.3\pm0.6$	$14.0\pm0.6$	93.7 ± 0.4	$28.9 \pm 0.7$
P-value	0.02					0.01			< 0.0001
Old <i>wt</i>	23.9 ± 1.1	340 ± 90	103.4 ± 32.2	36.8 ± 2.	51.6 ± 3.3	23.2 ± 3.4	23.4 ± 1.4	18.0 ± 4.3	85.8 ± 3.3
Old <i>3d</i>	29.7 ± 1.5	90 ± 10	$48.9 \pm 4.5$	18.8 ± 1.3	46.2 ± 3.1	$24.2 \pm 3.4$	$\textbf{28.0} \pm \textbf{2.1}$	$34.7 \pm 4.9$	$80.7\pm6.0$
P-value	0.007	0.01	0.07	< 0.0001				0.03	
Lymph nodes									
Young wt		19 ± 3	$4.3 \pm 0.9$	69.8 ± 1.7	56.1 ± 1.0	$40.2\pm0.9$	$2.4\pm0.2$	95.3 ± 0.4	22.0 ± 2.7
Young 3d		21 ± 2	8.4 ± 1.6	64.1 ± 2.1	54.6 ± 0.4	38.3 ± 3.2	$3.2\pm0.2$	96.5 ± 0.3	$14.3\pm0.9$
P-value									0.0005
Old <i>wt</i>		596 ± 175	105 ± 32	71.1 ± 4.1	37.6 ± 2.3	19.6 ± 3.1	40.3 ± 5.1	41.3 ± 6.8	76.5 ± 3.8
Old 3d		310 ± 68	113 ± 27	66.4 ± 3.4	12.9 ± 1.4	10.4 ± 1.9	74.1 ± 2.7	68.2 ± 6.1	66.4 ± 4.2
P-value					< 0.0001	0.02	< 0.0001	0.01	

LN includes cervical, axillary, inguinal, and mesenteric. CD4, CD8, and DN T cell data are expressed as a percentage of T cells; CD62L<sup>+</sup> and CD44<sup>hi</sup> subsets are the percentage of CD8 T cells. Young mice were 25–29 days old, n = 4/group. Old mice were 14.5–15.1 months old, n = 6-7/group. *P*-values < 0.05 are shown

#### Table S3. Splenic B cell populations in young and old B6-Fas<sup>lpr</sup> 3d mice

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Unc93b1	B cells (%)	B cells (no.)	Follicular CD21 <sup>Io</sup> CD23 <sup>hi</sup> (%)	Marginal zone CD21 <sup>hi</sup> CD23 <sup>lo</sup> (%)	CD21 <sup>1</sup> °CD23 <sup>1</sup> ° total (%)	CD21 <sup>I</sup> °CD23 <sup>I°</sup> IgM— subset (%)	Plasma cells CD138+ B220– (%)
Young wt	$58.2 \pm 0.7$	19.4 ± 3.2	$64.9\pm0.8$	$3.8\pm0.2$	$\textbf{27.2} \pm \textbf{0.3}$	17.1 ± 1.1	$\textbf{2.4}\pm\textbf{0.2}$
Young 3d	57.8 ± 1.1	$14.6 \pm 2.2$	72.9 ± 1.2	$4.3\pm0.2$	$19.9 \pm 1.3$	$17.5 \pm 0.1$	$2.6\pm0.2$
<i>P</i> -value			0.0007		0.0006		
Old wt	19.6 ± 5.0	24.1 ± 11.6	32.5 ± 7.7	15.3 ± 5.5	45.4 ± 13.8	51.9 ± 7.4	8.6 ± 2.8
Old 3d	$63.2 \pm 4.1$	$31.6 \pm 4.1$	$66.3 \pm 3.1$	$15.9 \pm 2.2$	$10.6 \pm 2.8$	$37.2 \pm 3.9$	$3.0\pm0.8$
P-value	0.00002		0.0005		0.01	0.06	0.03

Follicular, marginal zone, and CD21<sup>Io</sup>CD23<sup>Io</sup> subsets are expressed as a percentage of B cells; IgM- cells are shown as a percentage of the CD21<sup>Io</sup>CD23<sup>Io</sup> subset. For young mice, calculated numbers of CD21<sup>Io</sup>CD23<sup>Io</sup> B cells per spleen were significantly greater in wt mice (5.3  $\pm$  0.9 versus 2.9  $\pm$  0.3  $\times$  10<sup>6</sup> cells/spleen, *P* = 0.025), while numbers of follicular B cells were not significantly different (12.5  $\pm$  1.9 versus 10.7  $\pm$  1.7  $\times$  10<sup>6</sup> cells/spleen, *P* = 0.5). See Table S2 legend for ages and number of mice.

#### Table S4. ANA in male BXSB 3d mice

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Unc93b1	Age (mo)	ANA	
wt/wt	2.1	2 + Fine Spk, Meta +	
wt/wt	2.1	2–3 + Fine Spk, Meta +	
wt/wt	2.6	1 + Fine Spk	
wt/wt	2.6	1–2 + Fine Spk	
3d/3d	2.6	Neg	
3d/3d	3.0	Neg	
3d/3d	3.3	Neg	
3d/3d	3.6	Neg	
3d/3d	4.3	Neg	
3d/3d	4.5	Neg	
3d/3d	6.0	Neg	
3d/3d	7.1	Neg	

ANA staining was performed with 1/100 serum dilution and anti-IgG-FITC. All positive nuclear staining patterns are consistent with anti-chromatin. Meta, metaphase; Spk, speckled; Neg, negative.

### Table S5. ANA in B6-Fas<sup>Ipr</sup> 3d mice treated with lipid A

Lipid A treated	Age (mo)	ANA		
wt/wt	3.2	1–2 + Fine Spk, Meta +*		
wt/wt	4.1	2–3 + Fine Spk, Meta +*		
wt/wt	5.5	4 + Homog (strong), Meta +		
wt/wt	5.5	4 + Homog (strong), Meta +		
3d/3d	6.0	Neg		
3d/3d	6.0	Neg		
3d/3d	6.1	Neg		

ANA staining was performed with 1/100 serum dilution and anti-IgG-FITC. All positive nuclear staining patterns are consistent with anti-chromatin. Lipid A was administered twice weekly for 2 months. Homog, homogeneous; Meta, metaphase; Spk, speckled; Neg, negative.

\*Two lipid A-treated wt mice had early mortality requiring measurement of serum samples drawn at earlier time points.

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