# SUPPLEMENTAL MATERIAL

**Supplemental Table 1**. Location and extent of blocks of non-IBD containing genes of the immunoreceptor pathway and effects of ancestral variation on protein coding genes in block. The table indicates only genes with non-synonymous variation from the Brown-Norway (BN) reference genome in each block and omits the immunoglobulin heavy chain (chr 6) because of the extreme variation observed in this gene. The table also indicates confirmation of alleles in whole genome sequence from another SHR-A3 colony (Glasgow) and whether the variant allele is present in either of 2 injury-resistant (SHRSR) lines for which whole genome sequence data is available.

				Variation		Confirm A3	
•	•	Start	End	from BN		genotype in	Present in other
Gene name	Chr	position	position	in	AA change	SHRA3/GIa	SHRSR?
Block Non-IBD	3	158.090.000	161.780.000				
Ndra3	3	158.614.195	158.725.392	A3	Ser17Ara	Y	N
Tam2	3	160.977.905	161.007.260	A3	Asp94Asn	Ŷ	N
Tti1	3	161.113.415	161.140.919	A3	Thr663lle	Ŷ	Y
Tti1	3	161.113.415	161,140,919	A3	Lvs665Phe	Y	Y
		, ,			ý		
Block Non-IBD	6	146,030,000	154,190,000				
Gpr132	6	146,660,012	146,648,353	B2	Asp189Asn	Y	Y
Gpr132	6	146,660,012	146,648,353	A3	Thr26Met	Y	Ν
Gpr132	6	146,660,012	146,648,353	A3	Pro5Ser	Y	Ν
Adam6	6	147,426,645	147,429,132	B2	Val531Met	Y	Y
Wdr60	6	153,061,842	153,007,323	A3	Thr49Ala	Y	Ν
Block Non-IBD	16	17,650,000	23,470,000				
Dydc2	16	18,506,978	18,520,588	A3	Pro123Lys	Y	Ν
Cyp4f18	16	19,269,139	19,385,553	B2	Tyr242His	Y	Y
Myo9b	16	19,529,516	19,614,169	B2	Lys1175Gln	Y	Y
Slc27a1	16	19,859,540	19,876,619	A3	His357Arg	Y	Ν
Fam129c	16	19,887,776	19,899,606	A3	Cys7Ser	Y	Ν
B3gnt3	16	19,984,402	19,993,283	A3	Pro68Arg	Y	Ν
Crlf1	16	20,541,796	20,530,521	B2	Lys32Phe	Y	1 of 2
Gdf1	16	20,712,564	20,698,125	A3	Ala173Thr	Y	Ν
Sugp2	16	20,789,858	20,760,491	B2	Ser298Gly	Y	Y
Armc6	16	20,789,624	20,799,784	B2	Ser263Asn	Y	Y
Tmem161a	16	20,846,231	20,836,630	B2	Arg98His	Y	Y
Rfxank	16	20,879,467	20,886,951	A3	Val39lle	Y	Y
Gatad2a	16	21,093,916	21,184,110	B2	Ser235Gly	Y	Y
Cilp2	16	21,204,960	21,211,953	B2	Thr900Ala	Y	1 of 2
Sh2d4a	16	23,044,750	22,972,707	A3	Gly236Glu	Y	Y
Block Non-IBD	17	6,760,000	16,520,000				
Ntrk2	17	8,156,432	8,464,507	A3	Ala187Val	Y	N
Dok3	17	11,748,066	11,752,826	A3	lle71Val	Y	N
Dok3	17	11,748,066	11,752,826	A3	Ala83Val	Y	N
Dbn1	17	11,788,597	11,802,569	B2	Ala618Val	Y	Y
Slc34a1	17	11,856,946	11,872,100	B2	Ala630Thr	Y	Y
Fgfr4	17	12,099,383	12,113,632	A3	Arg95Gly	Y	N
Hk3	17	12,246,816	12,261,635	B2	Ser770Asn	Y	Y
Ror2	17	14,050,995	14,228,982	A3	Gly744Ser	Y	Ν

Sema4d	17	15,590,977	15,613,214	B2	Gly35Glu	Y	Y
Sema4d	17	15,590,977	15,613,214	B2	Gly617Ser	Y	Y
Nxnl2	17	16,120,216	16,124,963	B2	Ala147Thr	Y	Y

### IgG Fc variants (positions from rn5):

1. Location: chr6:147108547 intron SHR-A3 TAGGAGCTTGGGTCCAGACCTGTCAGACAAAAT SHR-B2 TAGGAGCTTGGGTCCAGACCTGTCAGACAAAAT SHR-B2 different from BN reference

2. Location: chr6:147107456 exon SHR-A3 ACGTTCAGATGCAAGGTCACCAGTGCAGCTTTCCCATCC SHR-B2 ACGTTCAGATGCAAGGTCACTAGTGCAGCTTTCCCATCC SHR-B2 different from BN reference, synonymous

3. Location: chr6:147141845 intron SHR-A3 AGACAAAATGATCACGCATACTTATTCTTGTAGCTGAAACAAC SHR-B2 AGACAAAATGATCACGCATACTTTTTCTTGTAGCTGAAACAAC SHR-A3 different from BN reference

4. Location: chr6: 147141478 intron SHR-A3 TAAGGCATTAGACTGCCTACACAAACCAGGGTGGGCAGAC SHR-B2 TAAGGCATTAGACTGCCTACACAAACCAGGGTGGGCAGAC SHR-A3 different from BN reference

5. Location: chr6: 147141081 intron SHR-A3 ACAGCTATTGACCTATTCTACCTTCTTCTTCATCTA SHR-B2 ACAGCTATTGACCTATTTCCACCTTTCTTCTTCATCTA SHR-B2 different from BN reference

6. Location: chr6: 147141059 exon
SHR-A3 TTTCTTCTTCATCTACAGECTCAGAAGTATCATCTGTC
SHR-B2 TTTCTTCTTCATCTACAGECCCAGATATATCATCTGTC
SHR-A3 different from BN reference, non-synonymous, Gly to Val,

7. Location: chr6: 147141057 exon SHR-A3 TCTTCTTCATCTACAGGCTCAGAAGTATCATCTGTCTTCATC SHR-B2 TCTTCTTCATCTACAGTCCCAGATATATCATCTGTCTTCATC SHR-A3 different from BN reference, non-synonymous, Ser to Pro

8. Location: chr6: 147141051-2 exon SHR-A3 TCATCTACAGGCTCAGA<mark>AG</mark>TATCATCTGTCTTCATCTTCCCC SHR-B2 TCATCTACAGTCCCAGA<mark>TA</mark>TATCATCTGTCTTCATCTTCCCC SHR-A3 different from BN reference, non-synonymous, Glu to Asp, Val to Ile

9. Location: chr6: 147140508-9 exon SHR-A3 TCTATCCCCCAGACATTTAT<mark>CT</mark>GGAGTGGAAGATGAACG SHR-B2 TCTATCCCCCAGACATTTAT<mark>AC</mark>GGAGTGGAAGATGAACG SHR-A3 different from BN reference, non-synonymous, Tyr to Val

10. Location: chr6: 147086792 intron SHR-A3 GACAGAAAGGATGCCTACACCCAGAAGGTGTAGGGTAGA SHR-B2 GACAGAAAGGATGCCCACACACAGAAGGTGTAGGGTAGA SHR-A3 different from BN reference

11. Location: chr6: 147086612 intron

SHR-A3 CACTGAAAGATCAAGGCTCCTGGGGCTGGAGAGTTGGCTC SHR-B2 CACTGAAAGATCAAGGCTCCTAGGGCTGGAGAGTTGGCTC SHR-A3 different from BN reference

12. Location: chr6: 147086388 intron SHR-A3 AAGGTCAGAGTGCAGGGTCTAGAGGTGGAAGCTGCTGACACCACTTA SHR-B2 AAGGTCAGAGTGCAGGGTCTAGAAGTGGAAGCTGCTGACACCACTTA SHR-A3 different from BN reference

13. Location: chr6: 147086179 exon SHR-A3 GATACAACCAGCTCCACGGTGACCCTGGGATGCCTGGTCAAGGG SHR-B2 GATACAACCAGCTCCACGGTGACCCTGGGATGCCTGGTCAAGGG SHR-A3 different from BN reference, synonymous

14. Location: chr6: 147086108 exon SHR-A3 CCGTGACCTGGAACTCTGGACCCCTGTCCAGCGATGTG SHR-B2 CCGTGACCTGGAACTCTGGACCCCTGTCCAGCGATGTG SHR-A3 different from BN reference, non-synonymous, Ala to Pro

15. Location: chr6: 147086092 exon SHR-A3 GTCCAGCGATGTGCACACCTTCCCAGCTGTCCTGCAGTCTGGGC SHR-B2 GTCCAGCGATGTGCACACCTTCCCAGCTGTCCTGCAGTCTGGGC SHR-A3 different from BN reference, synonymous

16. Location: chr6: 147086065 exon SHR-A3 CTGTCCTGCAGTCTGGGGCTCTAGACTCTCACCAGCTCAGTGA SHR-B2 CTGTCCTGCAGTCTGGGGCTCTATACTCTCACCAGCTCAGTGA SHR-B2 different from BN reference, synonymous

17. Location: chr6: 147085968 exon SHR-A3 CACCAAGGTGGACAAGAAATTGGTGAGAGGACAACCA SHR-B2 CACCAAGGTGGACAAGAAAGTTGGTGAGAGAACAACCA SHR-A3 different from BN reference, non-synonymous Val to Ile

18. Location: chr6: 147085957 intron SHR-A3 GGACAAGAAAATTGGTGAGAGGACAACCAAGGGACGAGG SHR-B2 GGACAAGAAAGTTGGTGAGAGAGAACCAAGGGGACGAGG SHR-B2 different from BN reference

19. Location: chr6: 147085949 intron SHR-A3 TTGGTGAGAGGAGACAACCAAGGGGACGAGGGGGCTCACTAGAC SHR-B2 TTGGTGAGAGAACAACCAGGGGACGAGGGGGCTCACTAGAC SHR-A3 different from BN reference

20. Location: chr6: 147085908 intron SHR-A3 GTGAGGATAAGGCATTAGAGTGCCTACACCAAGGGT SHR-B2 GTGAGGATAAGGCATTAGAGTGCCTACACCAACCAGGGT SHR-A3 different from BN reference

22. Location: chr6: 147085758 intron SHR-A3 AGACATGATCCTCTGGTATAGATGTCTATGTCATGTATAGGATCATAC SHR-B2 AGACATGATCCTCTGGTATAGATGTCTGTGTCATGCATAGGATCATAC SHR-A3 different from BN reference

23. Location: chr6: 147085750 intron SHR-A3 TGGTATAGATGTCTATGTCATGTATAGGATCATACCAGGGACAAA SHR-B2 TGGTATAGATGTCTGTGTCATGTATAGGATCATACCAGGGACAAA SHR-A3 different from BN reference

24. Location: chr6: 147085643 exon SHR-A3 CTCTACAGAGCGCAGAAATGGCGGCAATGGACACAAATG SHR-B2 CTCTACAGAGCGCAGAAATGGCGGCGCA<mark>T</mark>TGGACACAAATG SHR-B2 different from BN reference, non-synonymous, Ile to Asn

25. Location: chr6: 147085580 intron SHR-A3 GCCCAGGTAAGTCACTAGACCTGGGCCCCCAGCTCCACAATGATGGCA SHR-B2 GCCCAGGTAAGTCACTAGACCTGGACCCCAGCTCCACAATGATGGCA SHR-A3 different from BN reference

26. Location: chr6: 147085489 exon SHR-A3 CATTTCTATCTCTCCTCATCAGCTCCTGAACTCTTGGGTGGACCAT SHR-B2 CATTTCTATCTCTCCTCATCAGCTCCTGAACTCTTGGGTGGACCAT SHR-A3 different from BN reference, non-synonymous, Val to Ala

27. Location: chr6: 147085464 exon SHR-A3 TCCTGAACTCTTGGGTGGACCATCGGTCTTCATCTTCCCCCCCAAA SHR-B2 TCCTGAACTCTTGGGTGGACCATCIGTCTTCATCTTCCCGCCAAA SHR-A3 different from BN reference, synonymous

28. Location: chr6: 147085449 exon SHR-A3 CCATCCGTCTTCATCTTCCCCCCAAAGCCCAAGGACATCCTCTTGA SHR-B2 CCATCTGTCTTCATCTTCCCCCCCAAAGCCCAAGGACATCCTCTTGA SHR-A3 different from BN reference, synonymous

29. Location: chr6: 147085379 exon SHR-A3 ACGTGTGTGGTGGTGGTGGATGTGAGGGAGGAGGAGCCAGACGTCCAG SHR-B2 ACGTGTGTGGTGGTGGTGGATGTGAGTGGAGGAGCCGGACGTCCAG SHR-B2 different from BN reference, synonymous

30. Location: chr6: 147085368 exon SHR-A3 TGGATGTGAGCGAGGAGGAGCCAGACGTCCAGTTCAGCTGGTTT SHR-B2 TGGATGTGAGTGAGGAGGAGGAGCCCGGACGTCCAGTTCAGCTGGTTT SHR-A3 different from BN reference, synonymous

31. Location: chr6: 147085102 intron SHR-A3 CTGGGATAGGTATAAGAATGAATGCCCGGGTGGACGGCCATGACCTCTGT SHR-B2 CTGGGATAGGTATAAGAATGAATGCCCGGGTGGACGGCCATGACCTCTGT SHR-A3 different from BN reference

31. Location: chr6: 1470850352 exon SHR-A3 GTCAGAAAACCACAGGTATACGTCATGGGTCCACCGACAGAGCAG SHR-B2 GTCAGAAAACCACAGGTATATGTCATGGGTCCACCGACAGAGCAG SHR-A3 different from BN reference, synonymous 32. Location: chr6: 147084991-5001 exon SHR-A3 GACAGAGCAGTTGACTGAGAAAGCGGTCATTTTGACCTGCTTGA SHR-B2 GACAGAGCAGTTGACTGAGCAAACGGTCAGTTTGACCTGCTTGA SHR-A3 different from BN reference, non-synonymous Gln to Lys

33. Location: chr6: 147084818 exon SHR-A3 GGAGCAGGTGGGATAGCAGAGCGTCCTTCGTCTGCTCCGTGGTCCACGA SHR-B2 GGAGCAGGTGGGATAGCAGAGCGCCCTTCGTCTGCTCCGTGGTCCACGA SHR-A3 different from BN reference, non-synonymous, Pro to Ser

34. Location: chr6: 147084778 exon SHR-A3 GTCCACGAGGGTCTGCACAATCCCCACGTGGAGAAGAGCATCTC SHR-B2 GTCCACGAGGGTCTGCACAATCACCACGTGGAGAAGAGCATCTC SHR-A3 different from BN reference, non-synonymous, His to Arg

35. Location: chr6: 147084600 intron SHR-A3 TTCTTTCCGGGGGTATAGAGCCTAAGTCACGGGCTT SHR-B2 TTCTTTCCGGGGGTATAGAGCCTAGGTCACGGGCTT SHR-A3 different from BN reference

36. Location: chr6: 147189280 exon SHR-A3 AGCCGGTAACCGTAAAATGGAACTGTGGAGCCCTGTCCAGCGGTGTG SHR-B2 AGCCGGTAACCGTAAAATGGAACTATGGAGCCCTGTCCAGCGGTGTG SHR-B2 different from BN reference, non-synonymous Ser to Tyr

37. Location: chr6: 147189135 exon SHR-A3 AGCCCACCCAGCCACCAAAAGCAACTTGATCAAGAGAATTGGTA SHR-B2 AGCCCACCCAGCCACCAAAAGCGACTTGATCAAGAGAATTGGTA SHR-B2 different from BN reference, non-synonymous Asp to Asn

38. Location: chr6: 147189069 intron SHR-A3 AATATTTAATCAGGAGGTCAGGCTGGGGGTCAACCCCTTGTATAGAACA SHR-B2 AATATTTAATCAGGAGGTCAGGCTAGGGTCAACCCCTTGTATAGAACA SHR-B2 different from BN reference

39. Location: chr6: 147188923 intron SHR-A3 CAGAAAGTGGTCCTCTGGAATGACTACCAGTAGCTCACATCAGGGACA SHR-B2 CAGAAAGTGGTCCTCTGGAATGCTACCAGTATCTCACATCAGGGACA SHR-B2 different from BN reference

40. Location: chr6: 147188913 intron SHR-A3 CTCTGGAATGACTACCAGTAGCTCACATCAGGGACACACAGA SHR-B2 CTCTGGAATGACTACCAGTATCTCACATCAGGGACACACAGA SHR-B2 different from BN reference

41. Location: chr6: 147188770 intron SHR-A3 TCTCCCTTCCAGGCCCCATTAATGCCCAGTCTTTTCTCTGC SHR-B2 TCTCCCTTCCAGGCCCCATTAATGACCAGTCTTTTCTCTGC SHR-B2 different from BN reference

42. Location: chr6: 147188379 exon SHR-A3 GTCAGTACCCTCCATATCCAGCACCAGGACTGGA SHR-B2 GTCAGTACCCTCCACATCCAGCACCAGGACTGGA SHR-B2 different from BN reference, synonymous

43. Location: chr6: 147188230 intron SHR-A3 CTGGGATGGGCGTAAGAATAAATGCCTGCGTGGACG SHR-B2 CTGGGATGGGCGTAAGAATGAATGCCTGCGTGGACG SHR-B2 different from BN reference

44. Location: chr6: 147188078 exon SHR-A3 ACCAGCTTCTACCCCCCATCCAGTGTGGAGT SHR-B2 ACCAGCTTCTACCCCCCCATCCAGTGTGGAGT SHR-A3 different from BN reference (Ala to Pro)

45. Location: chr6: 147187959 exon SHR-A3 ACAGCAAGCTCAGTGTGGACAC GACAGTTGGATGCGAG SHR-B2 ACAGCAAGCTCAGTGTGGACAC GACAGTTGGATGCGAG SHR-B2 different from BN reference, synonymous Haplotype structure in SHR strains across this region. SHR-A3 and SHRSP/Gla are injury-prone strains. SHR-B2, SHR and SHR/NHsd are injury resistant strains. SHR lines with the same genotype as SHR-A3 at each variant site are colored pink at that site. SHR lines with the same genotype as SHR-B2 are shaded green when this genotype differs from the injury-prone lines. Genome sequence variation for SHRSP/Gla, SHR, SHR/NHsd was provided by the Rat Genome Database.

Variant #	SHR-A3	SHRSP/Gla	SHR-B2	SHR	SHR/NHsd
1	G	G	A	A	A
2	С	С	Т	Т	Т
3	A	A	Т	Т	Т
4	A	A	С	С	С
5	Т	Т	С	С	С
6	G	G	Т	Т	Т
7	Т	Т	С	С	C
8	AG	AG	ТА	ТА	ТА
9	GT	GT	AC	AC	AC
10	G	G	А	A	A
11	G	G	А	А	А
12	G	G	А	А	А
13	C	C	Т	Т	Т
14	С	С	G	G	G
15	C	C	Т	Т	Т
16	C	C	Т	Т	Т
17	A	A	G	G	G
18	G	G	А	A	A
19	А	А	G	G	G
20	С	С	Т	Т	Т
21	A	A	Т	Т	Т
22	A	A	G	G	G
23	Т	Т	С	С	С
24	A	A	Т	Т	Т
25	G	G	А	А	А
26	С	С	Т	Т	Т
27	С	С	Т	Т	Т

28	C	C	G	G	G
29	C	C	Т	Т	Т
30	А	А	C	C	C
31	C	С	Т	Т	Т
32	А	А	C	C	C
33	Т	Т	C	C	C
34	G	G	A	А	А
35	А	А	G	G	G
36	C	C	А	А	А
37	A	А	G	G	G
38	G	G	А	А	А
39	А	А	G	G	G
40	G	G	Т	Т	Т
41	C	C	А	А	А
42	Т	Т	C	C	C
43	A	А	G	G	G
44	C	C	G	G	G
45	G	G	A	А	А

#### Dok3 variants (positions from rn5):

 Location: chr17:11747889 proximal promoter SHR-A3 AAGCTCCCCATAGGCAGGTAGGGTTTCTTGGAACAGCAAGG SHR-B2 AAGCTCCCCATAGGCAGGTAGGTTTCTTGGAACAGCAAGG SHR-B2 differs from BN reference seq

2. Location: chr17:11748271 intron SHR-A3 GTGAAGTGAAAGCTGGAGACGGGAGGTTCTTGGCCCAAGAA SHR-B2 GTGAAGTGAAAGCTGGAGACAGGAGGTTCTTGGCCCAAGAA SHR-A3 differs from BN reference seq

3. Location: chr17:11748474 proximal promoter SHR-A3 CAGGCTGGTTCTCCTGGAGAAGAGGAGATGGGCTGGGAATCTAT SHR-B2 CAGGCTGGTTCTCCTGGAGAAGATGGGCTGGGAATCTAT SHR-A3 differs from BN reference seq

4. Location: chr17:11749072 exon 2
SHR-A3 GCCGTAGAGGGGAACGCCGGGTCATACGCTTGGCTGACTGT
SHR-B2 GCCGTAGAGGGGAACGCCGGGATCATACGCTTGGCTGACTGT
SHR-A3 differs from BN reference seq, non-synon Ile to Val (I71V)

6. Location: chr17:11749790 intron SHR-A3 TCCTCACCAAGTTGCTCATGCAGCAGCTGGAGAGGTGCTGC SHR-B2 TCCTCACCAAGTTGCTCATGCTGCAGCTGGAGAGGTGCTGC SHR-B2 differs from BN reference seq

7. Location: chr17:11750490 intron SHR-A3 CTAAAACCCACTGAACTGGGTACCTTAAAATAGTAATGTCC SHR-B2 CTAAAACCCACTGAACTGGG<mark>A</mark>ACCTTAAAATAGTAATGTCC SHR-B2 differs from BN reference seq

8. Location: chr17:11750514 intron SHR-A3 TTAAAATAGTAATGTCCAGGCCTTGAGAAATGACTCCGTAG SHR-B2 TTAAAATAGTAATGTCCAGGACTTGAGAAATGACTCCGTAG SHR-B2 differs from BN reference seq

9. Location: chr17:11750578 intron SHR-A3 AGAGTAGTAGAGTTCAATTCCCAGCAACCACATGGCAGCTC SHR-B2 AGAGTAGTAGAGTTCAATTCCCAGCAACCACATGGCAGCTC SHR-B2 differs from BN reference seq

10. Location: chr17:11750642 intron SHR-A3 AGAGGACCTGACACCCTCACCCCAATCCAATACACACAAAA SHR-B2 AGAGGACCTGACACCCTCACCCCAATCCAATACACAAAAA SHR-B2 differs from BN reference seq

11. Location: chr17:11750740 intron

SHR-A3 TGTAGACAAGGTTTCTCCAT<mark>A</mark>TAGCCCTGGCTGTTCTAGAA SHR-B2 TGTAGACAAGGTTTCTCCAT<mark>G</mark>TAGCCCTGGCTGTTCTAGAA SHR-B2 differs from BN reference seq

12. Location: chr17:11751049 intron SHR-A3 CTGTCTTGAAAACAAAAAGGAAAAAAAAGTGCAGGTAG SHR-B2 CTGTCTTGAAAACAAAAAGGAAAAAAAGTGCAGGTAG SHR-B2 differs from BN reference seq

13. Location: chr17:11751393 intron SHR-A3 CCCTGGAGTGTGGGGTCAGAGTGAGAGTGATGGCGGAGAGTG SHR-B2 CCCTGGAGTGTGGGGTCAGAGTGGAGACTGATGGCGGAGAGTG SHR-B2 differs from BN reference seq

14. Location: chr17:11752923 3'-UTR SHR-A3 TGCTCAGTGTAGCAAACGCCGGGCCTCGGCTGGACCCTGAG SHR-B2 TGCTCAGTGTAGCAAACGCCAGGCCTCGGCTGGACCCTGAG SHR-A3 differs from BN reference seq

Haplotype structure in SHR strains across this region. SHR-A3 and SHRSP/Gla are injury-prone strains. SHR-B2, SHR and SHR/NHsd are injury resistant strains. SHR lines with the same genotype as SHR-A3 at each variant site are colored pink at that site. SHR lines with the same genotype as SHR-B2 are shaded green when this genotype differs from the injury-prone lines. Genome sequence variation for SHRSP/Gla, SHR, SHR/NHsd was provided by the Rat Genome Database.

Variant #	SHR-A3	SHRSP/Gla	SHR-B2	SHR	SHR/NHsd
1	С	С	Т	Т	Т
2	G	G	A	А	А
3	А	А	G	G	G
4	G	G	A	А	А
5	Т	Т	C	С	С
б	GA	GA	TT	TT	TT
7	Т	Т	A	A	A
8	G	G	А	А	А
9	С	C	Т	Т	Т
10	G	G	A	А	A
11	A	A	G	G	G
12	А	А	-	-	-
13	Т	Т	C	С	С
14	G	G	A	А	А

### Src variants (positions are rn5):

 Location: chr3:158138524 Promoter SHR-A3 CAGGACCTTAGTCTGCCTATCTGCTTAGTGGGGTAGGGTGTCTTGTCTGTT SHR-B2 CAGGACCTTAGTCTGCCTATCTGCTTAGTGGGTAGGGTGTCTTGTCTGTT SHR-A3 differs from BN reference sequence
 Location: chr3:158138450 Promoter Kozak consensus sequence SHR-A3 GGGATTTCTGTCGCCCCCAGCCAGGCCATGGGCAGCAACAAGAGCAAGC SHR-B2 GGGATTTCTGTCGCCCCCAGCCAGGACCATGGGCAGCAACAAGAGCAAGC SHR-B2 GGGATTTCTGTCGCCCCCAGCCAGGACCATGGGCAGCAACAAGAGCAAGC SHR-A3 differs from BN reference sequence
 Location: chr3:158136014 Intron SHR-A3 TATTCACCCCTCAAGATCCAGGGTGGGGCTGCCCTCCCTGCAAGCAGGT SHR-B2 TATTCACCCCTCAAGATCCAGGGTGGGGCTGCCCTCCCTGCAAGCAGGT SHR-A3 differs from BN reference sequence
 Location: chr3:158134950 Intron

SHR-A3 ACAGTGTTTCTCCAGGAACTACTGAGCTCTGGGGGGGCCGGGGCAGTACAG SHR-B2 ACAGTGTTTCTCCAGGAACTACTGAGCTCTGGGGGGGCCGGGGCAGTACAG SHR-A3 differs from BN reference sequence

5. Location: chr3:158131765 Intron SHR-A3 ATGGCAACCAAGCCAGGCTGCTTCCTGTGGGGCCTGGGGGCTCAGCAGGAGA SHR-B2 ATGGCAACCAAGCCAGGCTGCTTCCTTGTGGGGCCTGGGGGCTCAGCAGGAGA SHR-A3 differs from BN reference sequence

6. Location: chr3:158129031 Intron SHR-A3 TGTGGCCAGGTCTCACACCAAGCCTCAGCCTCCTGATGTGAACAC SHR-B2 TGTGGCCAGGTCTCACACCACGCCTCAGCCTCCTGATGTGAACAC SHR-A3 differs from BN reference sequence

7. Location: chr3:158128929 Intron SHR-A3 AGTGCTGTGCGGAGGGAGGAGTAGTGGGAAGTAGGTGTCCGGTCTCCCG SHR-B2 AGTGCTGTGCGGAGGGAGGAGTAATGGGAAGTAGGTGTCCCGGTCTCCCG SHR-A3 differs from BN reference sequence

8. Location: chr3:158127846 Intron SHR-A3 CAGGCTGACCTCTGCTCAGATGCCCACACGGCAGACTGAGGGTAAC SHR-B2 CAGGCTGACCTCTGCTCAGATGCCCCCACAGGCAGACTGAGGGTAAC SHR-A3 differs from BN reference sequence

9. Location: chr3:158127694 Intron SHR-A3 TCTTTTGTCCCTTACCACTGACAAGATCCCTGTTGTTGCTGGTGTG SHR-B2 TCTTTTGTCCCTTACCACTGACAGGATCCCTGTTGTTGCTGGTGTG SHR-A3 differs from BN reference sequence

10. Location: chr3:158127366 Intron SHR-A3 GCCTCCGTTGCACCTTGAGTTGGTTTAGGTGAGACCCTGTCTCAGAACAA SHR-B2 GCCTCCGTTGCACCTTGAGTTGGTTTAGGTGAGACCCTGTCTCAGAACAA SHR-A3 differs from BN reference sequence

11. Location: chr3:158126261 Intron SHR-A3 TACACAGCTCGGCAAGGTGGGAGTTCCTGGCGGTTATCTGATGGTATCGG SHR-B2 TACACAGCTCGGCAAGGTGGGAGTGCCTGGCGGTTATCTGATGGTATCGG SHR-A3 differs from BN reference sequence

### 12. Location: chr3:158125852 Intron SHR-A3 TGCCTCTGGTCAGGACTCTTAGCCTCCAAGACAGTGTTCTGTACAGCATA SHR-B2 TGCCTCTGGTCAGGACTCTTAGCCTCCAAGGCAGTGTTCTGTACAGCATA SHR-A3 differs from BN reference sequence

Haplotype structure in SHR strains across this region. SHR-A3 and SHRSP/Gla are injury-prone strains. SHR-B2, SHR and SHR/NHsd are injury resistant strains. SHR lines with the same genotype as SHR-A3 at each variant site are colored pink at that site. SHR lines with the same genotype as SHR-B2 are shaded green when this genotype differs from the injury-prone lines. Genome sequence variation for SHRSP/Gla, SHR, SHR/NHsd was provided by the Rat Genome Database.

Variant #	SHR-A3	SHRSP/Gla	SHR-B2	SHR	SHR/NHsd
1	С	С	Т	Т	Т
2	G	G	А	А	А
3	G	G	А	G	G
4	G	G	C	С	С
5	С	C	Т	Т	Т
6	А	А	C	С	C
7	G	G	А	А	А
8	А	А	G	А	А
9	А	А	G	G	G
10	С	С	Т	Т	Т
11	Т	Т	G	Т	Т
12	А	А	G	G	G

### Syk variants (positions from rn5):

1. Location: chr17:14759592 proximal promoter SHR-A3 GTTTCCTATTTTTAGCTTCAGAGTTTTTTTTTTTCATGTTACT SHR-B2 GTTTCCTATTTTTTAGCTTCAGAGTTTTTTTTTTTTCATGTTACT SHR-A3 differs from BN reference seq

2. Location: chr17:14730906 Intron SHR-A3 GTGTTCAATCCCAGCATTACAAAGGCAGAGTAGGGAATCTGTTC SHR-B2 GTGTTCAATCCCAGCATTACGAAGGCAGAGTAGGGAATCTGTTC SHR-B2 differs from BN reference seq

3. Location: chr17:14728977 Intron SHR-A3 CTGAGCCTCGCTGCCCCTGCACAGCGCATGCCTGCTCCGGTTGTG SHR-B2 CTGAGCCTCGCTGCCCCTGCACACCGCATGCCTGCTCCGGTTGTG SHR-B2 differs from BN reference seq

4. Location: chr17:14721420 Exon SHR-A3 GCCTGCCATGGACACCGAGGTATATGAGAGTCCTTACGCTG SHR-B2 GCCTGCCATGGACACTGAGGTATATGAGAGTCCTTACGCTG Synonymous, codon is Thr, third base of codon affected SHR-A3 differs from BN reference seq

5. Location: chr17:14706170 3'UTR SHR-A3 GGTCACGTGGTATCCACACAGAGGGAAGCGAAGAATGAACAGGAA SHR-B2 GGTCACGTGGTATCCACACTGAGGAAGCGAAGAATGAACAGGAA SHR-A3 differs from BN reference seq

6. Location: chr17:14704598 3'UTR SHR-A3 GGGTTGGGACTGACCATTCGTTATCGGGATGTGACCAAGCAGAACGGAGG SHR-B2 GGGTTGGGACTGACCATTCGTTATCAGGATGTGACCAAGCAGAACGGAGG SHR-B2 differs from BN reference seq

7. Location: chr17:14704464 3'UTR SHR-A3 TCTGTTGAAAACTGTGTCTTGCTGTCAAGATTAATTCTCTCCCCTG SHR-B2 TCTGTTGAAAACTGTGTCTTGCTGTTTAAGATTAATTCTCTCCCCCTG SHR-B2 differs from BN reference seq

9. Location: chr17:14704196 3'UTR SHR-A3 CCCTAATCTGCCTCGCCACATCCTACTGACCTCGCCACACCACATACC SHR-B2 CCCTAATCTGCCTCGCCACATACCACTGACCTCGCCACACCACATACC SHR-A3 differs from BN reference seq Haplotype structure in SHR strains across this region. SHR-A3 and SHRSP/Gla are injury-prone strains. SHR-B2, SHR and SHR/NHsd are injury resistant strains. SHR lines with the same genotype as SHR-A3 at each variant site are colored pink at that site. SHR lines with the same genotype as SHR-B2 are shaded green when this genotype differs from the injury-prone lines. Uncolored boxes indicate no genomic sequence variation data for SHRSP/Gla, SHR or SHR/NHsd. Genome sequence variation for SHRSP/Gla, SHR, SHR/NHsd was provided by the Rat Genome Database.

Variant #	SHR-A3	SHRSP/Gla	SHR-B2	SHR	SHR/NHsd
1	-		Т		
2	А	А	G	G	G
3	G		-		
4	С	C	Т	Т	Т
5	А	А	Т	Т	Т
6	G	G	А	А	А
7	С	C	Т	Т	Т
8	С	C	А	А	А
9	Т	Т	C	C	C

### JunD variants (positions from rn5):

1. Location: chr16:20344584 proximal promoter SHR-A3 TGAGATCTTGTCTTAAAATAACCCAAAACAAAGATTAAA SHR-B2 TGAGATCTTGTCTTAAAACAAACAAAGATTAAA SHR-B2 differs from BN reference seq

3. Location: chr16:20344265 proximal promoter SHR-A3 CTATGGTGGGCGTGACGCGGGGCGTGGCTTCTCAGAGG SHR-B2 CTATGGTGGGCGTGACGC-GGGCGTGGCTTCTCAGAGG SHR-B2 differs from BN reference seq

6. Location: chr16:20343878 proximal promoter SHR-A3 CCCGCCCCTTTATGCAAATCCACGACGTCACCTCGAGCC SHR-B2 CCCGCCCCTTTATGCAAATCCACGACGTCACCTCGAGCC SHR-B2 differs from BN reference seq

7. Location: chr16:20342252-4 3'-UTR SHR-A3 TTTTAAAGAAACGGGAAG AAAAAAAAAAAACCTCCC SHR-B2 TTTTAAAGAAACGGGAAGAAGAAAAAAAAAACCTCCC SHR-A3 differs from BN reference seq

Haplotype structure in SHR strains across this region. SHR-A3 and SHRSP/Gla are injury-prone strains. SHR-B2, SHR and SHR/NHsd are injury resistant strains. SHR lines with the same genotype as SHR-A3 at each variant site are colored pink at that site. SHR lines with the same genotype as SHR-B2 are shaded green when this genotype differs from the injury-prone lines. Uncolored boxes indicate no genomic sequence variation data for SHRSP/Gla, SHR or SHR/NHsd. Genome sequence variation for SHRSP/Gla, SHR, SHR/NHsd was provided by the Rat Genome Database.

Variant #	SHR-A3	SHRSP/Gla	SHR-B2	SHR	SHR/NHsd
1	Т	Т	С	С	С
2	Т	Т	C	C	C
3	G		-		
4	GA				
5			ТА		

6	С	С	Т	Т	Т
7			AAG		

# Supplemental statistical analysis for Figure 6.

### SHR-A3 ctrl BP vs time

Call:

Im(formula = sample1\$BP ~ poly(sample1\$Week, 2, raw = TRUE))

Residuals: Min -24.5763	1Q -7.7596	Median -0.4215	3Q 7.9176	Max 24.1168	
Coefficients:					
		Estimate	Std. Error	t value	Pr(> t )
(Intercept)		198.1606	5.0170	39.498	< 2e-16 ***
poly(sample1\$Wee	k, 2, raw = TRUE	E)1 8.6372	2.5579	3.377	0.00115 **
poly(sample1\$Wee	k, 2, raw = TRUE	)2 -0.6258	0.2774	-2.256	0.02693 *

Signif. codes: 0 "\*\*\*" 0.001 "\*\*" 0.01 "\*" 0.05 ".' 0.1 " 1

Residual standard error: 11.37 on 77 degrees of freedom Multiple R-squared: 0.3089, Adjusted R-squared: 0.2909 F-statistic: 17.21 on 2 and 77 DF, p-value: 6.644e-07

#### SHR-A3 +MMF BP vs time

Call: Im(formula = sample2\$BP ~ poly(sample2\$Week, 2, raw = TRUE))

Residuals:           Min         1Q           -30.8104         -3.299	Median 1 0.8181	3Q 6.3479	Max 28.5903	
Coefficients:				
	Estimate	Std. Error	t value	Pr(> t )
(Intercept)	190.2577	4.9032	38.803	<2e-16 ***
poly(sample2\$Week, 2, ra	w = TRUE)1 3.0319	2.4999	1.213	0.229
poly(sample2\$Week, 2, ra	w = TRUE)2 -0.2889	0.2711	-1.065	0.290
Y				

Signif. codes: 0 '\*\*\*' 0.001 '\*\*' 0.01 '\*' 0.05 '.' 0.1 ' ' 1

Residual standard error: 11.11 on 77 degrees of freedom Multiple R-squared: 0.02247, Adjusted R-squared: -0.002918 F-statistic: 0.8851 on 2 and 77 DF, p-value: 0.4168

### SHR-B2 ctrl BP vs time

Call:

Im(formula = sample3\$BP ~ poly(sample3\$Week, 2, raw = TRUE))

Residuals: Min -24.0196	1Q -5.0269	Median -0.6765	3Q 5.2464	Max 13.5330	
Coefficients:					
		Estimate	Std. Error	t value	Pr(> t )
(Intercept)		164.9447	3.4117	48.346	<2e-16 ***
poly(sample3\$	Week, 2, raw = TRL	JE)1 4.1114	1.7395	2.364	0.0213 *
poly(sample3\$	Week, 2, raw = TRL	JE)2 -0.3412	0.1887	-1.809	0.0754 .

Signif. codes: 0 '\*\*\*' 0.001 '\*\*' 0.01 '\*' 0.05 '.' 0.1 ' ' 1

Residual standard error: 6.917 on 61 degrees of freedom Multiple R-squared: 0.1513, Adjusted R-squared: 0.1235 F-statistic: 5.437 on 2 and 61 DF, p-value: 0.006719

### SHR-B2 +MMF BP vs time

Call:
Im(formula = sample4\$BP ~ poly(sample4\$Week, 2, raw = TRUE))

Residuals:

Min	1Q	Median	3Q	Max
-16.149	-7.277	-3.458	7.755	19.859
0 11 1				
Coefficients:				
		Estimate	Std. Error	t value Pr(> t )
(Intercept)		164.9861	4.5248	36.462 <2e-16 ***
poly(sample4\$Weel	k, 2, raw = TRUE)1	2.7700	2.3527	1.177 0.243
poly(sample4\$Weel	k, 2, raw = TRUE)2	-0.3237	0.2573	-1.258 0.213

Signif. codes: 0 '\*\*\*' 0.001 '\*\*' 0.01 '\*' 0.05 '.' 0.1 ' ' 1

Residual standard error: 9.597 on 63 degrees of freedom Multiple R-squared: 0.02533, Adjusted R-squared: -0.005608 F-statistic: 0.8188 on 2 and 63 DF, p-value: 0.4456 Supplemental Figure 1.

Renal JunD expression determined by Affymetrix array in F2 rats inheriting contrasting JunD alleles, is determined in cis by the JunD allele inherited.



Supplemental Figure 2.

Representative photomicrographs of CD43 immunohistochemistry in SHR-A3 and SHR-B2 animals at 25 wks of age with (+ MMF) and without (CTRL) 8 weeks of MMF treatment



# Supplemental Figure 3.

Divergence at the nucleotide level between the rat reference genome sequence derived from the Brown Norway strain and SHR-A3 and SHR-B2 in the haplotype block on chromosome 3 that contains Src (chr3:148,157,256-148,170,524). The vertical axis indicates the number of bases that differ in each rat strain compared to the reference. This value was calculated by dividing the region into bins of 100,000 bases and summing the number of variants for each bin across the entire region. In regions of identity by descent SHR-A3 and SHR-B2 are descended form a common ancestor and share the same divergence from the rat reference genome. In this block of non-identity by descent SHR-A3 is more divergent from the rat reference strain than SHR-B2 and differences in divergence vary across this region.



Chr%%%Mbases)%

# Supplemental Figure 4.

Divergence at the nucleotide level between the rat reference genome sequence derived from the Brown Norway strain and SHR-A3 and SHR-B2 in the haplotype block on chromosome 16 that contains JunD (chr16:19,239,694-19,241,529). In this block of non-identity by descent the degree of divergence is less between SHR-A3 and SHR-B2, possibly indicating that this region of the chromosome is derived from a more recent common ancestor that is shared by the two lines than is suggested by Supplemental Figure 3 where the nucleotide divergence is greater.



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Supplemental Figure 5.

Divergence at the nucleotide level between the rat reference genome sequence derived from the Brown Norway strain and SHR-A3 and SHR-B2 in the haplotype block on chromosome 17 that contains Dok3 (chr17: 15,154,414-15,159,609) and Syk (chr17: 18,440,672-18,498,001).



Chr%17%2Mbases)%