



**Supplementary Information for**  
**High throughput functional variant screens *via* in-vivo production of single-stranded DNA**

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**This PDF file includes:**

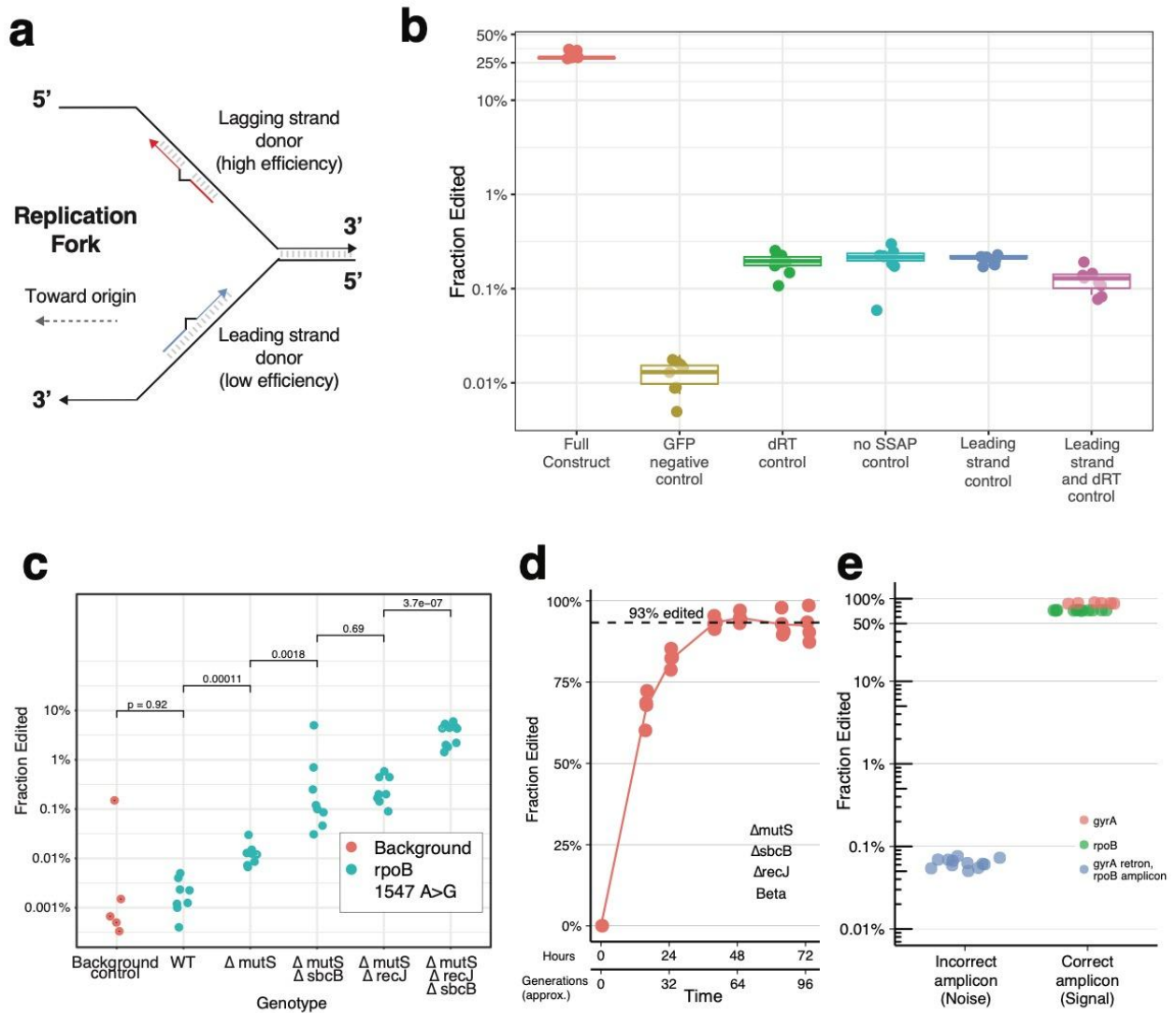
Figures S1 to S6  
Tables S1-S4

**Other supplementary materials for this manuscript include the following:**

All data and scripts necessary to reproduce figures and analysis can be found at <https://github.com/churchlab/rlr> including additional formats of the supplemental files contained here.

**Supplemental\_Dataset\_S1 (supplemental sequence maps):** generalized genbank plasmid maps for Retron Recombineering plasmids used in this study, and a sf.GFP-expressing control plasmid, supplied in Genbank format in a single text file. Please note that some sequence analysis software may require separation into separate files and changing extension to '.gb'.

**Supplemental\_Dataset\_S2:** Filtered mutations observed in whole genome sequencing (summarized in figure S4).

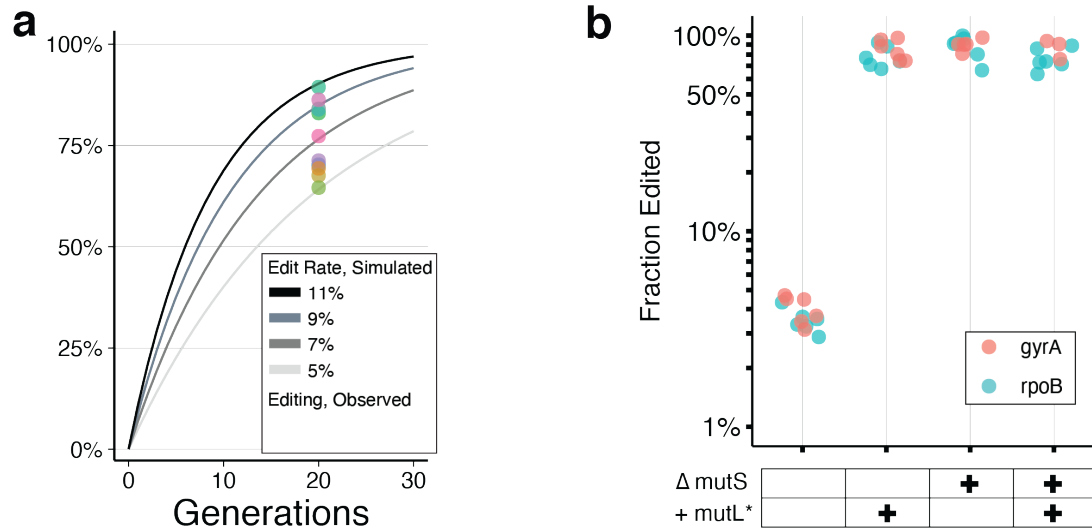


### Supplemental Figure 1:

- (a) A diagram of the replication fork, depicting recombineering donors annealing to either the leading strand of replication, or the lagging strand of replication.
- (b) A confirmation of the mechanism of editing. Editing was compared between the full RLR system expressing the *gyrA* 248 C>T donor using Beta as a Single-stranded annealing protein (SSAP), and control constructs expressing only GFP, having an inactivated retron reverse-transcriptase active site (dRT) and/or reverse-complemented donor DNA annealing to the leading strand, or which do not express an SSAP protein. Editing was performed in the  $\Delta mutS \Delta recJ \Delta sbcB$  background, with Beta as an SSAP. Individual measurements are represented with dots, and the data is summarized by box plots in the style of Tukey.
- (c) The edited fraction of cells, measured phenotypically by resistance to rifampicin at 25ug/mL. These results confirm those shown in Figure 2B, but are measured by plating for CFU, rather than by amplicon sequencing at the edited locus.
- (d) The continuous nature of Retron editing.  $\Delta mutS \Delta recJ \Delta sbcB$  cells expressing the *gyrA* retron with Beta as an SSAP were sampled during continuous growth and induction

in a turbidostat. The edited fraction of the *gyrA* locus was determined by amplicon deep sequencing over time. This data also depicted in figure 2C, but with inferred generations of growth on the X axis rather than time.

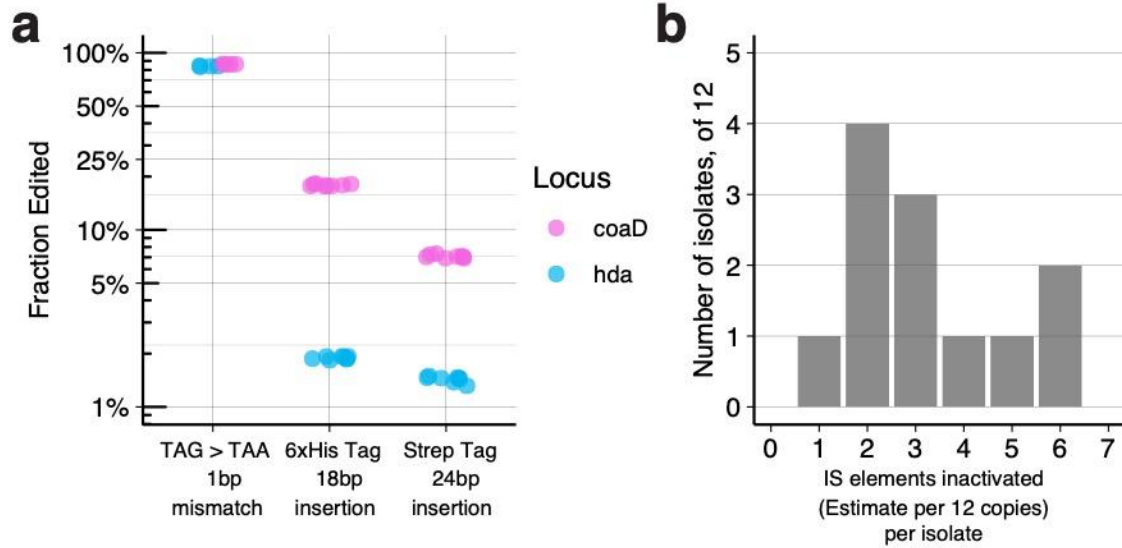
e) Control experiment characterizing error and noise. A culture editing *gyrA* was subjected to amplicon-NGS of the non-edited *rpoB* locus, to determine a background level of editing in the assay, alongside positive controls treated normally. Background editing is most likely due to sequencing error and mis-assigned indices. The mean of this background signal is depicted by a green dotted line in figure 2B, below which



**Supplemental Figure 2:**

a) Mean edited fraction of the loci in figure 2d are shown alongside a simulated editing trajectory of alleles with neutral fitness effect, as per the model introduced in figure S5. This estimate is suggestive of editing rate of 5-11% per generation for these allele.

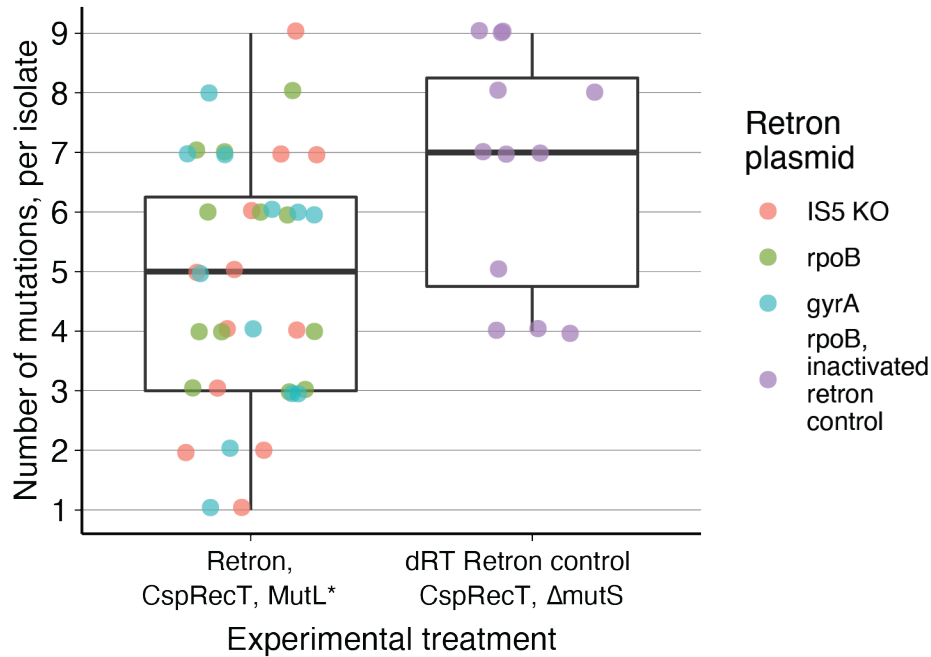
b) Retron plasmids with and without a dominant-negative MutL-E32K variant (mutL\*) were expressed in batch growth, in strains with and without inactivation of mutS ( $\Delta$ mutS). Retron plasmids targeted *gyrA* and *rpoB* loci as per figure 2b and used CspRecT as an SSAP. Four replicate experiments per locus and condition are indicated with dots.



**Supplemental Figure 3:**

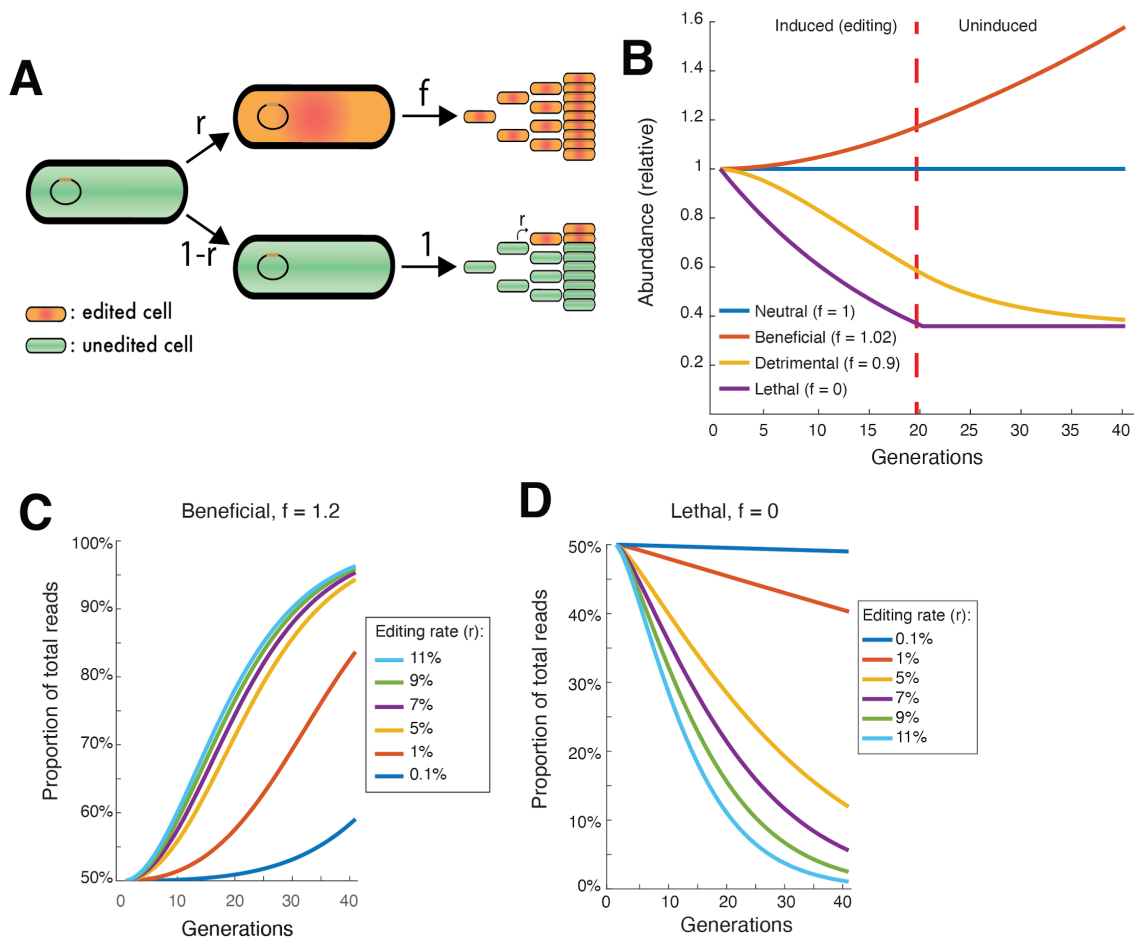
a) For two essential genes, *hda* and *coaD*, retrons were used to incorporate C-terminal tags, to determine the efficiency of incorporating longer insertions. Fraction edited was determined by amplicon-NGS as previously. Results of 8 replicate experiments are depicted with dots, alongside experiments shown in Figure 2D for the same loci, for comparison. Unlike the 1bp mismatch experiment, edited amplicons for insertions are approximately 10% longer than the non-edited amplicons, possibly introducing PCR bias toward un-edited sequences. This makes the edited fractions reported here likely an under-estimate.

b) A retron targeting the 12 copies of the IS5 transposase with a small inactivating insertion was expressed, and whole-genome sequencing was performed on isolates. The number of IS elements inactivated was estimated from the allelic fraction of the inactivation allele observed within reads.



**Supplemental Figure 4:**

Before and after induction and editing, whole genome sequencing was performed on isolates to measure overall mutagenesis, and search for off-target editing. 12 isolates for each of three retron donors expressing mutL\*, and an inactivated retron control in a ΔmutS strain were sequenced, alongside their parent strains, and the number of mutations per isolate are depicted with individual dots, and summarized by box plots per plasmid type.



### Supplemental figure 5:

**(A)** A population genetics model describing the Retron Library Recombining (RLR) process. Each generation, a cell has probability “ $r$ ” of being successfully edited by the retron construct it bears, and if so edited, propagates at rate “ $f$ ” determined by the fitness of this mutation in future generations. If not edited in a given generation, which occurs at a rate  $1-r$ , editing may occur in future generations.

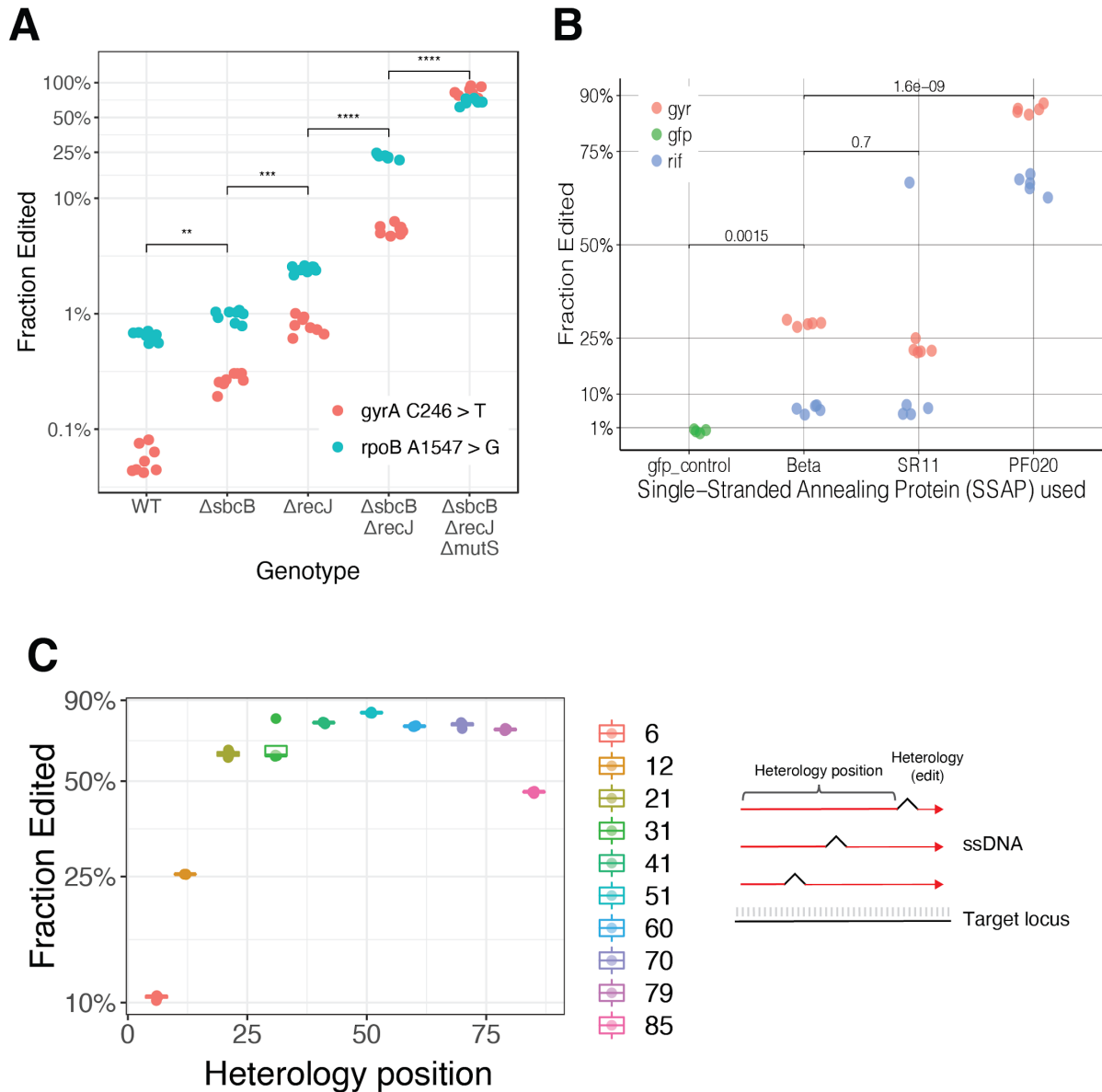
**(B)** The population genetics model predicts that for beneficial alleles, the editing process can be halted (red vertical line), and the lineage of cells marked with the beneficial retron will continue to become more abundant. In contrast, for deleterious or lethal mutations, continued decrease in abundance of a barcoded population is dependent on continued induction and editing.

**(C)** The growth of populations bearing a retron conferring mutations with growth rate 120% that of the parent ( $f = 1.2$ ) are simulated, for different rates of editing ( $r$ ). Population size compared to a simulated neutral parent control, and thus proportion of barcode reads observed, is depicted on the Y axis. The signal observed for beneficial alleles is large, and does not vary substantially if editing frequency varies among alleles. This is compatible with quantitative measurement of beneficial allele fitness, even if editing rate varies somewhat. Additionally, beneficial alleles can be detected even with

inefficient editing.

**(D)** Simulated growth of populations as in Figure S2C, except with a Lethal mutation ( $f = 0$ ). In contrast with Figure S2C, the signal for deleterious or lethal alleles is comparatively small, and scales with editing rate to a greater degree. This indicates that quantification of deleterious phenotypes will be clouded by differences in editing rate among alleles to a greater degree, and that deleterious phenotypes require more efficient editing and/or editing over more generations for robust detection.





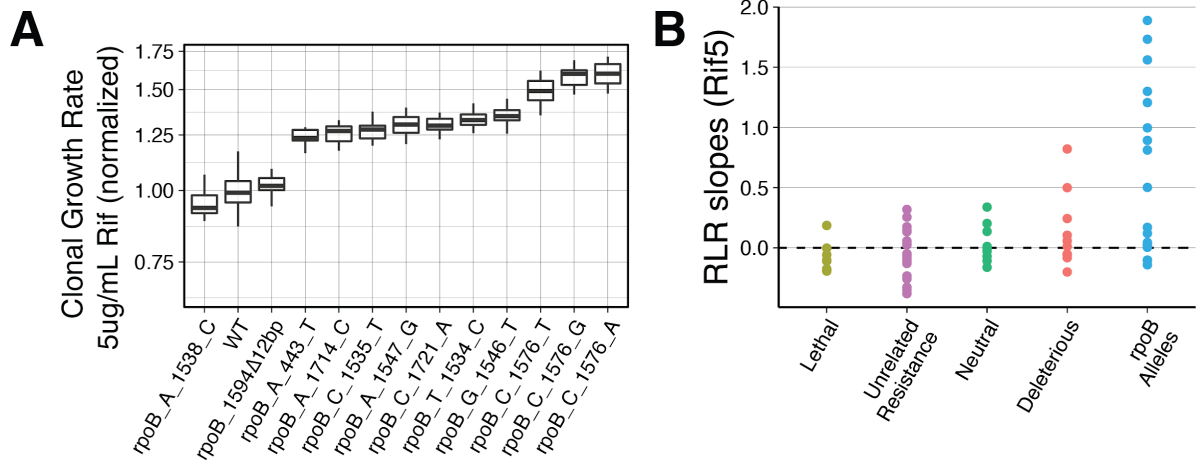
**Supplemental figure 6:**

**(A)** The effect of genotype on editing efficiency when the CspRecT is used rather than Beta as an SSAP. Individual measurements of the edited fraction of cells by amplicon sequencing are depicted by dots, for two loci. \*\*\*\* signifies  $p < 0.0001$ , \*\*\* signifies  $p < 0.001$ , and \*\* signifies  $p < 0.01$ , for two-tailed unpaired t-tests performed for comparisons indicated in brackets.

**(B)** The effect of SSAP protein on editing efficiency. Edited fraction of cells after batch growth was measured by amplicon sequencing for two loci. Retron constructs used Beta, “SR11”, or CspRecT as their SSAP, or expressed only GFP as a negative control. Independent measurements are represented with dots, and p-values of two-tailed, unpaired t-tests are given for the comparisons shown.

**(C)** The effect of heterology position within a 90-mer donor DNA was determined for the *gyrA* 246 C>T mutation, using CspRecT as an SSAP. Donors conferring this

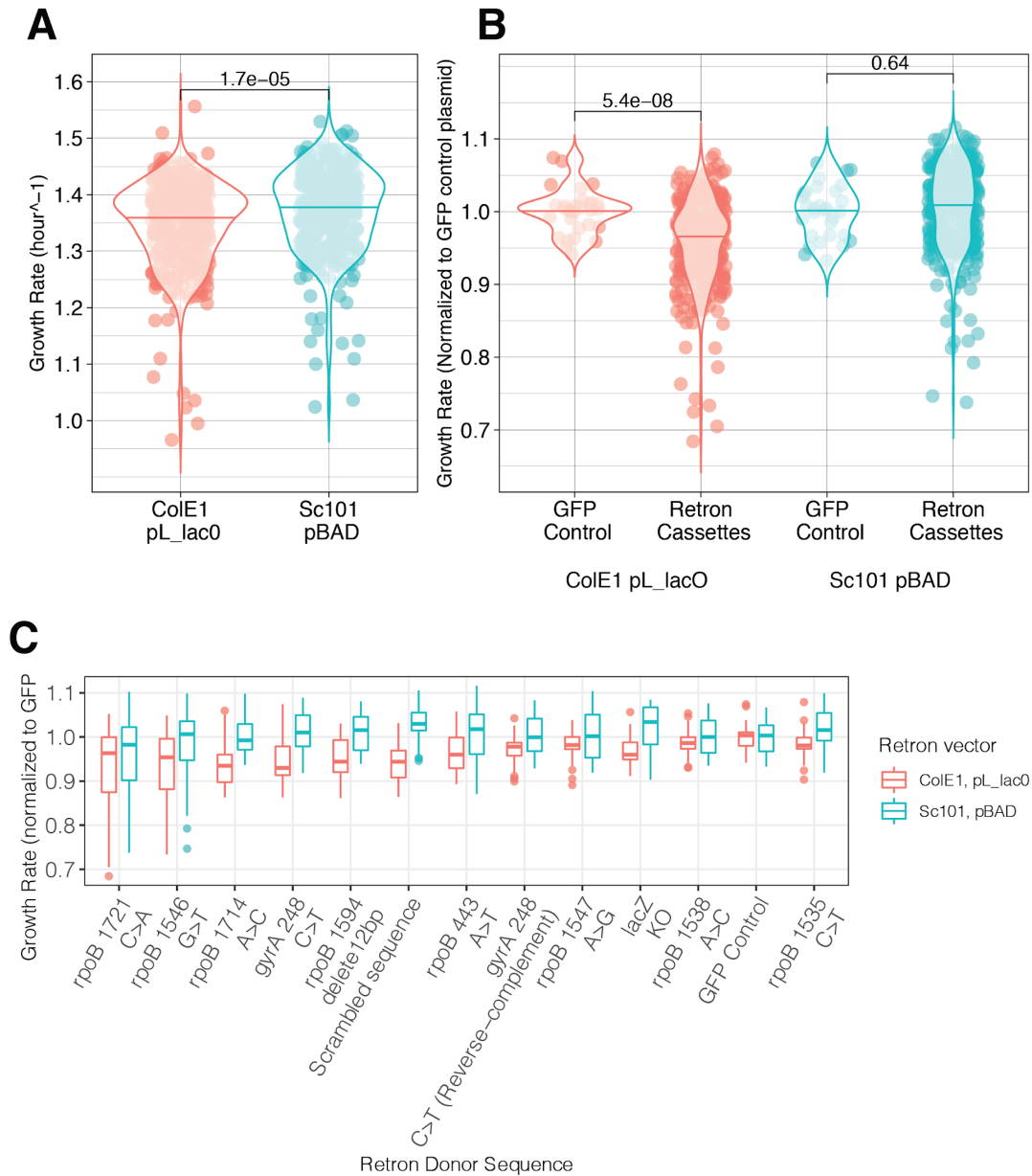
mutation were constructed with the heterologous mutation present in different locations within the donor, as indicated in the diagram. Individual cultures were grown and induced for approximately 20 generations. Box plots in the style of Tukey are shown, and the 4 replicate measurements are additionally represented with dots.



**Supplemental Figure 7:**

**(A)** *rpoB* mutants conferring rifampicin resistance were constructed in the  $\Delta mutS \Delta recJ \Delta sbcB$  background via oligonucleotide recombineering<sup>3</sup>, and growth rates were measured in batch growth with sub-inhibitory rifampicin (5ug/ml). Data is shown normalized to the parental (WT,  $\Delta mutS \Delta recJ \Delta sbcB$ ) growth rate.

**(B)** Summary of RLR enrichment observed in a pooled, quantitative experiment over time. RLR enrichment rate (slope) for all alleles across 3 replicate experiments is indicated by a dot, and different classes of mutants are separated by color.



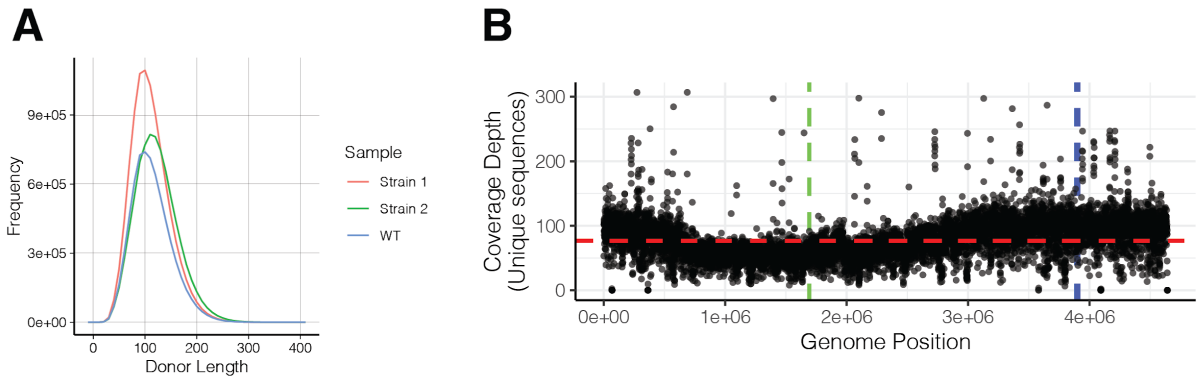
### Supplemental Figure 8:

**(A)** Where previous studies used a medium-copy ColE1 vector and pL\_lacO promoter for expression of the retron components, expression from a low-copy (sc101) vector is less detrimental to growth. Absolute growth rates are pooled here for 12 expressed retron constructs, and violin plot outlines depict the distribution of the data, with the width of outlines indicating the proportion of the data with a given value. Horizontal lines depict the mean growth rate for each vector type. P value for a t-test comparing data for both vector types is given.

**(B)** Retrons are expressed from the ColE1 vector, they are significantly depressed in growth rate compared to sfGFP expressed in the same manner, whereas 12 retons expressed from the SC101 vector cannot be distinguished, as a population. Growth rate is

shown normalized to the GFP control construct for each vector type, and horizontal lines depict the mean growth rate for each condition. P values for t-tests are given for the comparisons indicated with brackets.

(C) Growth rates for 12 induced retron constructs, normalized to the GFP-expressing control plasmid for each vector type. Constructs are ordered by mean growth rate, and box-and-whisker plots are depicted in the style of Tukey, with outlier observations represented by dots.



### Supplemental Figure 9

**(A)** Retron donor size distribution for three representative genomic DNA libraries. Strains 1 and 2 were evolved, Trimethoprim-resistant *E. coli*, and “WT” is *E. coli* MG1655.

**(B)** Depth of coverage of a representative genomic retron library is depicted. Dots indicate the mean coverage of unique retron donor sequences across 500bp windows of the genome. The mean coverage across all windows is 78, and is indicated with a dashed red line. Dashed gray lines mark the genomic positions of the origin and terminus of the genome, helping to clarify increased coverage as DNA content increases with proximity to the origin.

**Supplemental Table S1 (Supplemental sequences):** annotated list of selected oligonucleotides used in the study

oligonucleotide name	oligonucleotide sequence	oligonucleotide description
TetA_K O_tetR _bla_F	CTCGCAGAAAGAAACATATCCATGAAATCCCGC CCCGAATTCATatgaccgatgatgaagaaaagaatttgaaactagtt a	integrate bla at $\lambda$ :tetR of ECNR1, Fwd
TetA_K O_tetR _bla_R	CATTATTTGCCGACTACCTTGGTGATCTCGCCTT TCACGTAGTGGACAAAActgtataaaaaaggatcaattttgaact ctctccc	integrate bla at $\lambda$ :tetR of ECNR1, Rev
araBAD fwd (kanfrt)	GAGCGTGACCAGGTTGTGTTTTGGCAGCGCCAG GTTGGCTATTCCGGGGATCCGTCGACC	inactivate araBAD, Fwd
arabad rev	CTGACGCTTTTTATCGCAACTCTCTACTGTTTCTC CATACTGTAGGCTGGAGCTGCTTCG	inactivate araBAD, Rev
mutSka n_total_ del_fwd	CTAACAATTCCCGATAGTCTTTTGCTATCGGGAA TATTAACGACGAATTCAGGAAACCCG	mutS inactivate, Fwd
mutSka n_total_ del_rev	GGTTATGTCCGGTTCCTGATATTAATGGGGTG TGATGGGAATTCAGGAAAACAGACAG	mutS inactivate, Rev
oMS_0 178	g*a*ggcaattcagcgggcaagtctgccgttcatcgacttcagTTATC Aaagttgatctgttgttcacgcaattattaccgctggt	recJ inactivation oligo
oMS_0 179	g*g*cagacagcagaataacggatttaacctaataatgatgaatgacTAATG Acaacaatctaccttttgttcacgattacgaaacctttgg	sbcB inactivation oligo
oMS_7 33	CCTACACGACGCTCTTCCGATCTNGTTACCCTGA CCGTCTACCAG	gyrA fwd genomic PCR1
oMS_7 34	CCTACACGACGCTCTTCCGATCTNNGTTACCCTG ACCGTCTACCAG	
oMS_7 44	CCTACACGACGCTCTTCCGATCTNNGTTACCCT GACCGTCTACCAG	
oMS_7 39	GAGTTCAGACGTGTGCTCTTCCGATCTNNCTGGA ACAAAGCCTATAAAAAATCTGCC	gyrA rev genomic PCR1

oMS_7 40	GAGTTCAGACGTGTGCTCTTCCGATCTNCTGGAA CAAAGCCTATAAAAAATCTGCC	
oMS_7 45	GAGTTCAGACGTGTGCTCTTCCGATCTNNNCTGG AACAAAGCCTATAAAAAATCTGCC	
oMS_7 29	CCTACACGACGCTCTTCCGATCTNCTGATGCCAC AGGATATGATCAACG	rpoB fwd genomic PCR1
oMS_7 30	CCTACACGACGCTCTTCCGATCTNNCTGATGCCA CAGGATATGATCAACG	
oMS_7 35	GAGTTCAGACGTGTGCTCTTCCGATCTNNGAGT GCGGAGATACGACGTTTG	rpoB rev genomic PCR1
oMS_7 36	GAGTTCAGACGTGTGCTCTTCCGATCTNGAGTGC GGAGATACGACGTTTG	
retron_ donor_f wd_1	cctacacgacgcttccgatctNNTCTGAGTTACTGTCTGTT TTCCTGAATTC	for amplification of retron donors from plasmid, Fwd
retron_ donor_f wd_2	cctacacgacgcttccgatctNNBTCTGAGTTACTGTCTGT TTCCTGAATTC	
retron_ donor_f wd_3	cctacacgacgcttccgatctNNNHTCTGAGTTACTGTCTG TTTTCTGAATTC	
retron_ donor_f wd_4	cctacacgacgcttccgatctNNNNBTCTGAGTTACTGTCT GTTTTCTGAATTC	
retron_ donor_f wd_5	cctacacgacgcttccgatctTCTGAGTTACTGTCTGTTTT CCTGAATTC	
retron_ donor_r ev_1	gagttcagacgtgtgcttccgatctNAAGTGTTCAAATATTC AGCGGATTTTCATG	for amplification of retron donors from plasmid, Rev
retron_ donor_r ev_2	gagttcagacgtgtgcttccgatctNVAAGTGTTCAAATATT CAGCGGATTTTCATG	
retron_ donor_r ev_3	gagttcagacgtgtgcttccgatctNNVAAGTGTTCAAATAT TCAGCGGATTTTCATG	



retron_ donor_r ev_4	gagttcagacgtgtgcttccgatctNNNDAAGTGTTCAAATA TTCAGCGGATTCATG	
murF fwd	cctacacgacgcttccgatctACGATTTTAGTTAAGGGTTC ACGTAG	for amplification of TAG>TAA loci from genome
yafF fwd	cctacacgacgcttccgatctTGCGAAAAGCAGCCATGGA C	
pgpA fwd	cctacacgacgcttccgatctCGCAGGCATCCTGTATTTTA TCG	
sucB fwd	cctacacgacgcttccgatctGCGAATCCGTGGGCTTCCTG	
lolA fwd	cctacacgacgcttccgatctCTGTGGATGCAGCGAAATTT AC	
lpxK fwd	cctacacgacgcttccgatctGTAGACGCACAGCTTTCAG G	
fabH fwd	cctacacgacgcttccgatctTAAGCCGGGGCAGTTGGTT C	
hemA fwd	cctacacgacgcttccgatctCCAACGAAATCACTTCAAC AGG	
fliN fwd	cctacacgacgcttccgatctGTTGCCGATAAATATGGCGT G	
hda fwd	cctacacgacgcttccgatctccGGCCTACGAATACTATGG ATTAAC	
mreC fwd	cctacacgacgcttccgatctGGAAAACAATCAGGTTATC CGGC	
coaD fwd	cctacacgacgcttccgatctCTCTTCATCGTTGGTGAAAG AGG	
murF rev	gagttcagacgtgtgcttccgatctCGATGGCGCGAAACGTC AG	
yafF rev	gagttcagacgtgtgcttccgatctAAAGTTTCCGGCATGGC AGG	
pgpA rev	gagttcagacgtgtgcttccgatctCGAACTGGAAACGCCGT ATAAAC	

sucB rev	gagttcagacgtgtgctcttccgatctCTTCAATTGCCGGAGAA GGC	
lolA rev	gagttcagacgtgtgctcttccgatctCTAAATTTTCTGGCCGCA TACG	
lpxK rev	gagttcagacgtgtgctcttccgatctATTACGCGCATCAGCAA GG	
fabH rev	gagttcagacgtgtgctcttccgatctGCCAGCATTCCAACGGT TTG	
hemA rev	gagttcagacgtgtgctcttccgatctGCTTCCAGTTTGGCAAC GATAG	
fliN rev	gagttcagacgtgtgctcttccgatctGGGCAATAATGGCGATG AGTG	
hda rev	gagttcagacgtgtgctcttccgatctATCAGTTGGATCGTGCG TCG	
mreC rev	gagttcagacgtgtgctcttccgatctTCTCCACAAAGGGCTAC GCC	
coaD rev	gagttcagacgtgtgctcttccgatcttGCCAGAAGTAATTCATG CGC	
oMS_8 90	/5Phos/GAATTCAGGAAAACAGAGAGACCgAACT CCAGTCUACACTCTACACGctGGTCTCGTTTCCTG AATTC*T	Custom Hairpin adapter, ligates to TA-tailed genomic fragments
oMS_8 88	GACTGGAGTTcGGTCTCTCTGT	5prime fwd primer for amplifying genomic fragments
oMS_8 89	ACACTCTACACGctGGTCTCG	3prime rev primer for amplifying genomic fragments
oMS_8 91	ATTCAGGTCTCGGAAACCCGTTTTTTCTGACG	genomic Vector Fwd, for genomic insert retron vector
oMS_8 92	TATAGGTCTCAACAGACAGTAACTCAGATTCAA TGC	genomic Vector Rev, for genomic insert retron vector
oMS_9 57	T*A*AAAAAAATTGTCGCCACTATACGTAAAGCG TAAACCGTCGTCaACTGGTGCGAGGATGATGTTG AGGAAAATTTTATATTCTGCTGGC	MAGE oligo for introduction of [C- >t].49764 mutation

(folA region)

oMS_9 58	C*A*GGTATGGCGGCCATAATCACGGGTTTATT TAAGGTGTTGCGTTTAAACCAGGCGAGcTCGGC AGGCAGGTTCCACGGCATGGCGTTT	MAGE oligo for introduction of [T- >g].49902 mutation (folA region)
oMS_9 59	A*A*TTCTATACAAAATTACCGCCGCTCCAGAAT CTCAcAGCAATAGCTGTGAGAGTTCTGCGCATCA GCATCGTGGAATTCGCTGAATACC	MAGE oligo for introduction of [T- >g].50279 mutation (folA region)
oMS_9 60	G*T*GCGTAATCTCAGACAGCGGGTTGTTCTGGT CCATAAACTGAGGCAGCTGGCTGGAACCGAAGA ACTCTTTCACTGCTGCGGAAATCGG	MAGE oligo for introduction of rpoB_1547A.G mutation
oMS_9 61	A*G*ATACGACGTTTGTGCGTAATCTCAGACAGC GGGTTGTTCTGGCCATAAACTGAGACAGCTGG CTGGAACCGAAGAACTCTTTCACTG	MAGE oligo for introduction of rpoB_1534T.C mutation
oMS_9 62	C*C*GTAGTGAGTCGGGTGTACGTCTCGAACTTC GAAGCCTGCACGATCACGGGTCAGACCGCCTGG GCCGAGTGCGGAGATACGACGTTTG	MAGE oligo for introduction of rpoB_A_1623_T mutation
oMS_9 63	C*G*GACCTTCAGGGGTTTCGATTGGACATACGC GACCGTAGTGAGCCGGGTGTACGTCTCGAACTT CGAAGCCTGCACGTTACGGGTCAG	MAGE oligo for introduction of rpoB_A_1657_G mutation
oMS_9 64	C*G*GACCTTCAGGGGTTTCGATTGGACATACGC GACCGTAGTGAGACGGGTGTACGTCTCGAACTT CGAAGCCTGCACGTTACGGGTCAG	MAGE oligo for introduction of rpoB_A_1657_T mutation
oMS_9 65	A*C*GGGTCAGACCGCCTGGGCCGAGTGCGGAG ATACGACGTTTGTTCGTAATCTCAGACAGCGGG TTGTTCTGGTCCATAAACTGAGACAG	MAGE oligo for introduction of rpoB_C_1576_A mutation
oMS_9 66	A*C*GGGTCAGACCGCCTGGGCCGAGTGCGGAG ATACGACGTTTGTCCGTAATCTCAGACAGCGGG TTGTTCTGGTCCATAAACTGAGACAG	MAGE oligo for introduction of rpoB_C_1576_G mutation
oMS_9 67	C*G*GGTCAGACCGCCTGGGCCGAGTGCGGAGA TACGACGTTTGTACGTAATCTCAGACAGCGGGT TGTTCTGGTCCATAAACTGAGACAGC	MAGE oligo for introduction of rpoB_C_1576_T

		mutation
oMS_9 68	C*G*GGTCAGACCGCCTGGGCCGAGTGCGGAGA TACGACGTTTGTGAATAATCTCAGACAGCGGGT TGTTCTGGTCCATAAACTGAGACAGC	MAGE oligo for introduction of rpoB_CG_1574_TT mutation
oMS_9 69	A*T*ACGACGTTTGTGCGTAATCTCAGACAGCGG GTTGTTCTGGTCTATAAACTGAGACAGCTGGCTG GAACCGAAGAACTCTTTCCTGCT	MAGE oligo for introduction of rpoB_G_1545_A mutation
oMS_9 70	A*T*ACGACGTTTGTGCGTAATCTCAGACAGCGG GTTGTTCTGGTCAATAAACTGAGACAGCTGGCT GGAACCGAAGAACTCTTTCCTGCT	MAGE oligo for introduction of rpoB_G_1545_T mutation
i5_2_1	aatgatacggcgaccaccgagatctacacaagccattacactctttccctacac gacgctctccgat	Custom Illumina i5 index 2_1
i5_2_2	aatgatacggcgaccaccgagatctacacttcggtaaacactctttccctacacg acgctctccgat	Custom Illumina i5 index 2_2
i5_2_3	aatgatacggcgaccaccgagatctacacTATAGCCTacactctttccct acacgacgctctccgat	Custom Illumina i5 index 2_3
i5_2_4	aatgatacggcgaccaccgagatctacacATAGAGGCacactctttccc tacacgacgctctccgat	Custom Illumina i5 index 2_4
i5_2_5	aatgatacggcgaccaccgagatctacacCCTATCCTacactctttccct acacgacgctctccgat	Custom Illumina i5 index 2_5
i5_2_6	aatgatacggcgaccaccgagatctacacGGCTCTGAacactctttccct acacgacgctctccgat	Custom Illumina i5 index 2_6
i5_2_7	aatgatacggcgaccaccgagatctacacAGGCGAAGacactctttccc tacacgacgctctccgat	Custom Illumina i5 index 2_7
i5_2_8	aatgatacggcgaccaccgagatctacacTAATCTTAacactctttccct acacgacgctctccgat	Custom Illumina i5 index 2_8
i5_2_9	aatgatacggcgaccaccgagatctacacCAGGACGTacactctttccc tacacgacgctctccgat	Custom Illumina i5 index 2_9
i5_2_10	aatgatacggcgaccaccgagatctacacGTACTIONGACacactctttccct acacgacgctctccgat	Custom Illumina i5 index 2_10
PCR2_r 01_8me	CAAGCAGAAGACGGCATAACGAGATCCGTGAGA	Custom Illumina i7 index 1 (A1)

r	GTGACTGGAGTTCAGACGTGTGCTCTT	
PCR2_r 02_8me r	CAAGCAGAAGACGGCATAACGAGATCATCAAGTG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 2 (A2)
PCR2_r 03_8me r	CAAGCAGAAGACGGCATAACGAGATAAAGACGGA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 3 (A3)
PCR2_r 04_8me r	CAAGCAGAAGACGGCATAACGAGATCGAACTTAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 4 (A4)
PCR2_r 05_8me r	CAAGCAGAAGACGGCATAACGAGATGATAGACA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 5 (A5)
PCR2_r 06_8me r	CAAGCAGAAGACGGCATAACGAGATGGTGCGAA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 6 (A6)
PCR2_r 07_8me r	CAAGCAGAAGACGGCATAACGAGATCACCTTACG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 7 (A7)
PCR2_r 08_8me r	CAAGCAGAAGACGGCATAACGAGATCCTAATCCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 8 (A8)
PCR2_r 09_8me r	CAAGCAGAAGACGGCATAACGAGATAACGTGAT GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 9 (A9)
PCR2_r 10_8me r	CAAGCAGAAGACGGCATAACGAGATAGTACAAG GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 10 (A10)
PCR2_r 11_8me r	CAAGCAGAAGACGGCATAACGAGATAACCGAGA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 11 (A11)
PCR2_r 12_8me r	CAAGCAGAAGACGGCATAACGAGATAACGCTTAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 12 (A12)
PCR2_r 13_8me	CAAGCAGAAGACGGCATAACGAGATAAGGTACA	Custom Illumina i7 index 13 (B1)

r	GTGACTGGAGTTCAGACGTGTGCTCTT	
PCR2_r 14_8me r	CAAGCAGAAGACGGCATAACGAGATACAGCAGA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 14 (B2)
PCR2_r 15_8me r	CAAGCAGAAGACGGCATAACGAGATACCTCCAAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 15 (B3)
PCR2_r 16_8me r	CAAGCAGAAGACGGCATAACGAGATACGTATCAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 16 (B4)
PCR2_r 17_8me r	CAAGCAGAAGACGGCATAACGAGATACTATGCAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 17 (B5)
PCR2_r 18_8me r	CAAGCAGAAGACGGCATAACGAGATAGAGTCAA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 18 (B6)
PCR2_r 19_8me r	CAAGCAGAAGACGGCATAACGAGATAGCAGGAA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 19 (B7)
PCR2_r 20_8me r	CAAGCAGAAGACGGCATAACGAGATCAACCACA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 20 (B8)
PCR2_r 21_8me r	CAAGCAGAAGACGGCATAACGAGATCAAGACTA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 21 (B9)
PCR2_r 22_8me r	CAAGCAGAAGACGGCATAACGAGATCAATGGAA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 22 (B10)
PCR2_r 23_8me r	CAAGCAGAAGACGGCATAACGAGATCACTTCGAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 23 (B11)
PCR2_r 24_8me r	CAAGCAGAAGACGGCATAACGAGATCAGCGTTAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 24 (B12)
PCR2_r 25_8me	CAAGCAGAAGACGGCATAACGAGATCATAACAA	Custom Illumina i7 index 25 (C1)

r	GTGACTGGAGTTCAGACGTGTGCTCTT	
PCR2_r 26_8me r	CAAGCAGAAGACGGCATAACGAGATCCAGTTCAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 26 (C2)
PCR2_r 27_8me r	CAAGCAGAAGACGGCATAACGAGATCCGAAGTA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 27 (C3)
PCR2_r 28_8me r	CAAGCAGAAGACGGCATAACGAGATCCTCCTGAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 28 (C4)
PCR2_r 29_8me r	CAAGCAGAAGACGGCATAACGAGATCGCATAACA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 29 (C5)
PCR2_r 30_8me r	CAAGCAGAAGACGGCATAACGAGATCTGAGCCA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 30 (C6)
PCR2_r 31_8me r	CAAGCAGAAGACGGCATAACGAGATCTGGCATAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 31 (C7)
PCR2_r 32_8me r	CAAGCAGAAGACGGCATAACGAGATGAATCTGA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 32 (C8)
PCR2_r 33_8me r	CAAGCAGAAGACGGCATAACGAGATGAGCTGAA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 33 (C9)
PCR2_r 34_8me r	CAAGCAGAAGACGGCATAACGAGATGCCACATA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 34 (C10)
PCR2_r 35_8me r	CAAGCAGAAGACGGCATAACGAGATGCGAGTAA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 35 (C11)
PCR2_r 36_8me r	CAAGCAGAAGACGGCATAACGAGATGCTAACGA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 36 (C12)
PCR2_r 37_8me	CAAGCAGAAGACGGCATAACGAGATGCTCGGTAG	Custom Illumina i7 index 37 (D1)

r	TGACTGGAGTTCAGACGTGTGCTCTT	
PCR2_r 38_8me r	CAAGCAGAAGACGGCATAACGAGATGGAGAACA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 38 (D2)
PCR2_r 39_8me r	CAAGCAGAAGACGGCATAACGAGATGTACGCAA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 39 (D3)
PCR2_r 40_8me r	CAAGCAGAAGACGGCATAACGAGATGTCGTAGA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 40 (D4)
PCR2_r 41_8me r	CAAGCAGAAGACGGCATAACGAGATGTCTGTCAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 41 (D5)
PCR2_r 42_8me r	CAAGCAGAAGACGGCATAACGAGATTAGGATGA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 42 (D6)
PCR2_r 43_8me r	CAAGCAGAAGACGGCATAACGAGATTATCAGCAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 43 (D7)
PCR2_r 44_8me r	CAAGCAGAAGACGGCATAACGAGATTCCGTCTAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 44 (D8)
PCR2_r 45_8me r	CAAGCAGAAGACGGCATAACGAGATTCTTCACAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 45 (D9)
PCR2_r 46_8me r	CAAGCAGAAGACGGCATAACGAGATTGAAGAGA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 46 (D10)
PCR2_r 47_8me r	CAAGCAGAAGACGGCATAACGAGATTGGAACAA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 47 (D11)
PCR2_r 48_8me r	CAAGCAGAAGACGGCATAACGAGATTGGCTTCAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 48 (D12)
PCR2_r 49_8me	CAAGCAGAAGACGGCATAACGAGATTGGTGGTAG	Custom Illumina i7 index 49 (E1)



r	TGACTGGAGTTCAGACGTGTGCTCTT	
PCR2_r 50_8me r	CAAGCAGAAGACGGCATAACGAGATTTACGCAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 50 (E2)
PCR2_r 51_8me r	CAAGCAGAAGACGGCATAACGAGATAATCCGTCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 51 (E3)
PCR2_r 52_8me r	CAAGCAGAAGACGGCATAACGAGATAATGTTGCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 52 (E4)
PCR2_r 53_8me r	CAAGCAGAAGACGGCATAACGAGATACACGACC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 53 (E5)
PCR2_r 54_8me r	CAAGCAGAAGACGGCATAACGAGATACAGATTCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 54 (E6)
PCR2_r 55_8me r	CAAGCAGAAGACGGCATAACGAGATAGATGTAC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 55 (E7)
PCR2_r 56_8me r	CAAGCAGAAGACGGCATAACGAGATCAAGGAGC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 56 (E8)
PCR2_r 57_8me r	CAAGCAGAAGACGGCATAACGAGATCCATCCTCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 57 (E9)
PCR2_r 58_8me r	CAAGCAGAAGACGGCATAACGAGATCCGACAAC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 58 (E10)
PCR2_r 59_8me r	CAAGCAGAAGACGGCATAACGAGATCCTCTATCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 59 (E11)
PCR2_r 60_8me r	CAAGCAGAAGACGGCATAACGAGATCGACACAC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 60 (E12)
PCR2_r 61_8me	CAAGCAGAAGACGGCATAACGAGATCGGATTGCG	Custom Illumina i7 index 61 (F1)

r	TGACTGGAGTTCAGACGTGTGCTCTT	
PCR2_r 62_8me r	CAAGCAGAAGACGGCATAACGAGATGAACAGGC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 62 (F2)
PCR2_r 63_8me r	CAAGCAGAAGACGGCATAACGAGATGACAGTGC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 63 (F3)
PCR2_r 64_8me r	CAAGCAGAAGACGGCATAACGAGATGAGTTAGC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 64 (F4)
PCR2_r 65_8me r	CAAGCAGAAGACGGCATAACGAGATGATGAATC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 65 (F5)
PCR2_r 66_8me r	CAAGCAGAAGACGGCATAACGAGATGCCAAGAC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 66 (F6)
PCR2_r 67_8me r	CAAGCAGAAGACGGCATAACGAGATAACAAGCTA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 67 (F7)
PCR2_r 68_8me r	CAAGCAGAAGACGGCATAACGAGATAAACATCG GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 68 (F8)
PCR2_r 69_8me r	CAAGCAGAAGACGGCATAACGAGATACATTGGCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 69 (F9)
PCR2_r 70_8me r	CAAGCAGAAGACGGCATAACGAGATACCACTGTG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 70 (F10)
PCR2_r 71_8me r	CAAGCAGAAGACGGCATAACGAGATCGCTGATCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 71 (F11)
PCR2_r 72_8me r	CAAGCAGAAGACGGCATAACGAGATCAGATCTGG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 72 (F12)
PCR2_r 73_8me	CAAGCAGAAGACGGCATAACGAGATCTGTAGCCG	Custom Illumina i7 index 73 (G1)

r	TGACTGGAGTTCAGACGTGTGCTCTT	
PCR2_r 74_8me r	CAAGCAGAAGACGGCATAACGAGATAGTGGTCA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 74 (G2)
PCR2_r 75_8me r	CAAGCAGAAGACGGCATAACGAGATAACAACCA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 75 (G3)
PCR2_r 76_8me r	CAAGCAGAAGACGGCATAACGAGATACACAGAA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 76 (G4)
PCR2_r 77_8me r	CAAGCAGAAGACGGCATAACGAGATACGCTCGA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 77 (G5)
PCR2_r 78_8me r	CAAGCAGAAGACGGCATAACGAGATAGATCGCA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 78 (G6)
PCR2_r 79_8me r	CAAGCAGAAGACGGCATAACGAGATAGTCACTAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 79 (G7)
PCR2_r 80_8me r	CAAGCAGAAGACGGCATAACGAGATCGACTGGA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 80 (G8)
PCR2_r 81_8me r	CAAGCAGAAGACGGCATAACGAGATCTCAATGAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 81 (G9)
PCR2_r 82_8me r	CAAGCAGAAGACGGCATAACGAGATGACTAGTA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 82 (G10)
PCR2_r 83_8me r	CAAGCAGAAGACGGCATAACGAGATGTGTTCTAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 83 (G11)
PCR2_r 84_8me r	CAAGCAGAAGACGGCATAACGAGATAAATCACCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 84 (G12)
PCR2_r 85_8me	CAAGCAGAAGACGGCATAACGAGATAAGGACAC	Custom Illumina i7 index 85 (H1)

r	GTGACTGGAGTTCAGACGTGTGCTCTT	
PCR2_r 86_8me r	CAAGCAGAAGACGGCATAACGAGATAGCACCTCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 86 (H2)
PCR2_r 87_8me r	CAAGCAGAAGACGGCATAACGAGATAGCCATGC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 87 (H3)
PCR2_r 88_8me r	CAAGCAGAAGACGGCATAACGAGATAGGCTAAC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 88 (H4)
PCR2_r 89_8me r	CAAGCAGAAGACGGCATAACGAGATATCATTCCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 89 (H5)
PCR2_r 90_8me r	CAAGCAGAAGACGGCATAACGAGATCTAAGGTCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 90 (H6)
PCR2_r 91_8me r	CAAGCAGAAGACGGCATAACGAGATATGCCTAAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 91 (H7)
PCR2_r 92_8me r	CAAGCAGAAGACGGCATAACGAGATATTGAGGA GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 92 (H8)
PCR2_r 93_8me r	CAAGCAGAAGACGGCATAACGAGATAAGAGATC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 93 (H9)
PCR2_r 94_8me r	CAAGCAGAAGACGGCATAACGAGATATAGCGAC GTGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 94 (H10)
PCR2_r 95_8me r	CAAGCAGAAGACGGCATAACGAGATATTGGCTCG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 95 (H11)
PCR2_r 96_8me r	CAAGCAGAAGACGGCATAACGAGATATCCTGTAG TGACTGGAGTTCAGACGTGTGCTCTT	Custom Illumina i7 index 96 (H12)

**Supplemental table S2**: summarized data and source data for Figure 2B. Edited fraction across genotype and SSAP.

genotype	locus	fraction_edited	ssap	well
WT	rpoB	9.32E-05	Beta	A1
WT	rpoB	0.00016102	Beta	A2
WT	rpoB	0.0001508	Beta	A3
WT	rpoB	0.0001307	Beta	A4
WT	rpoB	0.00017252	Beta	A5
WT	rpoB	7.12E-05	Beta	A6
WT	rpoB	0.00011803	Beta	A7
WT	rpoB	0.00014323	Beta	A8
WT	gyrA	3.75E-05	Beta	E1
WT	gyrA	1.58E-05	Beta	E2
WT	gyrA	5.26E-06	Beta	E3
WT	gyrA	1.06E-05	Beta	E4
WT	gyrA	2.66E-05	Beta	E5
WT	gyrA	5.29E-06	Beta	E6

WT	<i>gyrA</i>	5.48E-06	Beta	E7
WT	<i>gyrA</i>	1.13E-05	Beta	E8
$\Delta$ mutS	<i>rpoB</i>	0.00029032	Beta	A9
$\Delta$ mutS	<i>rpoB</i>	0.00019444	Beta	A10
$\Delta$ mutS	<i>rpoB</i>	0.00031214	Beta	A11
$\Delta$ mutS	<i>rpoB</i>	0.00020312	Beta	A12
$\Delta$ mutS	<i>rpoB</i>	0.00031395	Beta	B9
$\Delta$ mutS	<i>rpoB</i>	0.00027199	Beta	B10
$\Delta$ mutS	<i>rpoB</i>	0.00025501	Beta	B11
$\Delta$ mutS	<i>rpoB</i>	0.00037342	Beta	B12
$\Delta$ mutS	<i>gyrA</i>	0.00167809	Beta	E9
$\Delta$ mutS	<i>gyrA</i>	0.00145778	Beta	E10
$\Delta$ mutS	<i>gyrA</i>	0.00176807	Beta	E11
$\Delta$ mutS	<i>gyrA</i>	0.0017596	Beta	E12
$\Delta$ mutS	<i>gyrA</i>	0.00165089	Beta	F9
$\Delta$ mutS	<i>gyrA</i>	0.00616607	Beta	F10
$\Delta$ mutS	<i>gyrA</i>	0.00164878	Beta	F11
$\Delta$ mutS	<i>gyrA</i>	0.00152824	Beta	F12

$\Delta\text{mutS}, \Delta\text{recJ}$	rpoB	0.00322439	Beta	B1
$\Delta\text{mutS}, \Delta\text{recJ}$	rpoB	0.00291499	Beta	B2
$\Delta\text{mutS}, \Delta\text{recJ}$	rpoB	0.00331901	Beta	B3
$\Delta\text{mutS}, \Delta\text{recJ}$	rpoB	0.00294365	Beta	B4
$\Delta\text{mutS}, \Delta\text{recJ}$	rpoB	0.00329234	Beta	B5
$\Delta\text{mutS}, \Delta\text{recJ}$	rpoB	0.0032829	Beta	B6
$\Delta\text{mutS}, \Delta\text{recJ}$	rpoB	0.00312337	Beta	B7
$\Delta\text{mutS}, \Delta\text{recJ}$	rpoB	0.00352062	Beta	B8
$\Delta\text{mutS}, \Delta\text{recJ}$	gyrA	0.03687721	Beta	F1
$\Delta\text{mutS}, \Delta\text{recJ}$	gyrA	0.03631908	Beta	F2
$\Delta\text{mutS}, \Delta\text{recJ}$	gyrA	0.04317863	Beta	F3
$\Delta\text{mutS}, \Delta\text{recJ}$	gyrA	0.0344794	Beta	F4
$\Delta\text{mutS}, \Delta\text{recJ}$	gyrA	0.03569975	Beta	F5
$\Delta\text{mutS}, \Delta\text{recJ}$	gyrA	0.03528977	Beta	F6
$\Delta\text{mutS}, \Delta\text{recJ}$	gyrA	0.03638852	Beta	F7
$\Delta\text{mutS}, \Delta\text{recJ}$	gyrA	0.03654135	Beta	F8
$\Delta\text{mutS}, \Delta\text{recJ}, \Delta\text{sbcB}$	rpoB	0.06061889	Beta	C9
$\Delta\text{mutS}, \Delta\text{recJ}, \Delta\text{sbcB}$	rpoB	0.05995104	Beta	C10

$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	rpoB	0.01053677	Beta	C11
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	rpoB	0.05843985	Beta	C12
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	rpoB	0.05923514	Beta	D1
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	rpoB	0.06256935	Beta	D2
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	rpoB	0.06053654	Beta	D3
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	rpoB	0.06256011	Beta	D4
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	rpoB	0.05899817	Beta	D5
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	rpoB	0.06520138	Beta	D6
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	gyrA	0.28783597	Beta	G9
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	gyrA	0.28481606	Beta	G10
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	gyrA	0.2888569	Beta	G11
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	gyrA	0.26955921	Beta	G12
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	gyrA	0.29316805	Beta	H1
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	gyrA	0.29879149	Beta	H2
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	gyrA	0.29285045	Beta	H3
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	gyrA	0.29838652	Beta	H4
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	gyrA	0.29406375	Beta	H5
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$	gyrA	0.29958465	Beta	H6



$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>gyrA</i>	0.84498444	CspRecT	G1
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>gyrA</i>	0.8508817	CspRecT	G2
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>gyrA</i>	0.82264361	CspRecT	G3
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>gyrA</i>	0.8452907	CspRecT	G4
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>gyrA</i>	0.81406457	CspRecT	G5
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>gyrA</i>	0.85009398	CspRecT	G6
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>gyrA</i>	0.84937543	CspRecT	G7
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>gyrA</i>	0.84486369	CspRecT	G8
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>rpoB</i>	0.70608883	CspRecT	H1
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>rpoB</i>	0.68284453	CspRecT	H2
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>rpoB</i>	0.70397889	CspRecT	H3
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>rpoB</i>	0.69858698	CspRecT	H4
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>rpoB</i>	0.70239808	CspRecT	H5
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>rpoB</i>	0.70515084	CspRecT	H6
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>rpoB</i>	0.70141649	CspRecT	H7
$\Delta\text{mutS}$ , $\Delta\text{recJ}$ , $\Delta\text{sbcB}$ , + CspRecT	<i>rpoB</i>	0.70163794	CspRecT	H8
$\Delta\text{mutS}$ , $\Delta\text{sbcB}$	<i>rpoB</i>	0.06134563	Beta	C1
$\Delta\text{mutS}$ , $\Delta\text{sbcB}$	<i>rpoB</i>	0.00136385	Beta	C2

$\Delta\text{mutS}, \Delta\text{sbcB}$	rpoB	0.00131544	Beta	C3
$\Delta\text{mutS}, \Delta\text{sbcB}$	rpoB	0.00154202	Beta	C4
$\Delta\text{mutS}, \Delta\text{sbcB}$	rpoB	0.00135054	Beta	C5
$\Delta\text{mutS}, \Delta\text{sbcB}$	rpoB	0.00145458	Beta	C6
$\Delta\text{mutS}, \Delta\text{sbcB}$	rpoB	0.00154701	Beta	C7
$\Delta\text{mutS}, \Delta\text{sbcB}$	rpoB	0.00158267	Beta	C8
$\Delta\text{mutS}, \Delta\text{sbcB}$	gyrA	0.0145603	Beta	G1
$\Delta\text{mutS}, \Delta\text{sbcB}$	gyrA	0.01433665	Beta	G2
$\Delta\text{mutS}, \Delta\text{sbcB}$	gyrA	0.01419199	Beta	G3
$\Delta\text{mutS}, \Delta\text{sbcB}$	gyrA	0.01401869	Beta	G4
$\Delta\text{mutS}, \Delta\text{sbcB}$	gyrA	0.01334759	Beta	G5
$\Delta\text{mutS}, \Delta\text{sbcB}$	gyrA	0.01271887	Beta	G6
$\Delta\text{mutS}, \Delta\text{sbcB}$	gyrA	0.01318843	Beta	G7
$\Delta\text{mutS}, \Delta\text{sbcB}$	gyrA	0.01315153	Beta	G8

**Supplemental table S3:** summarized data from Figure 3C, showing the enrichment scores for all alleles

SampleNum	rif	Count	nickname	seq_freq	enrichment.post.rif
S65	12.5	0	wecA_ko	4.53E-07	6.57E-05
S66	12.5	1	wecA_ko	2.58E-07	3.41E-05
S67	12.5	0	wecA_ko	3.38E-07	4.01E-05
S68	25	0	wecA_ko	4.03E-07	5.84E-05
S69	25	0	wecA_ko	4.26E-07	5.63E-05
S70	25	0	wecA_ko	5.30E-07	6.29E-05
S71	50	0	wecA_ko	6.00E-07	8.69E-05
S72	50	0	wecA_ko	6.00E-07	7.93E-05
S73	50	0	wecA_ko	6.99E-07	8.30E-05
S74	100	0	wecA_ko	7.82E-07	0.00011328
S75	100	0	wecA_ko	4.63E-07	6.12E-05
S76	100	0	wecA_ko	7.10E-07	8.43E-05
S65	12.5	0	typA_ko	4.53E-07	5.02E-05
S66	12.5	0	typA_ko	2.58E-07	2.98E-05
S67	12.5	0	typA_ko	3.38E-07	3.73E-05

S68	25	0	typA_ko	4.03E-07	4.46E-05
S69	25	0	typA_ko	4.26E-07	4.93E-05
S70	25	0	typA_ko	5.30E-07	5.85E-05
S71	50	0	typA_ko	6.00E-07	6.64E-05
S72	50	0	typA_ko	6.00E-07	6.95E-05
S73	50	0	typA_ko	6.99E-07	7.71E-05
S74	100	0	typA_ko	7.82E-07	8.66E-05
S75	100	0	typA_ko	4.63E-07	5.36E-05
S76	100	0	typA_ko	7.10E-07	7.84E-05
S65	12. 5	1	yciW_ko	4.53E-07	3.12E-05
S66	12. 5	0	yciW_ko	2.58E-07	1.79E-05
S67	12. 5	3	yciW_ko	1.01E-06	6.99E-05
S68	25	485	yciW_ko	0.0001953 6	0.01347102
S69	25	0	yciW_ko	4.26E-07	2.96E-05
S70	25	0	yciW_ko	5.30E-07	3.65E-05
S71	50	0	yciW_ko	6.00E-07	4.14E-05
S72	50	1	yciW_ko	6.00E-07	4.18E-05

S73	50	0	yciW_ko	6.99E-07	4.82E-05
S74	100	1	yciW_ko	7.82E-07	5.39E-05
S75	100	0	yciW_ko	4.63E-07	3.22E-05
S76	100	0	yciW_ko	7.10E-07	4.90E-05
S65	12. 5	0	ptsP_ko	4.53E-07	4.76E-05
S66	12. 5	0	ptsP_ko	2.58E-07	2.57E-05
S67	12. 5	0	ptsP_ko	3.38E-07	3.54E-05
S68	25	1	ptsP_ko	4.03E-07	4.23E-05
S69	25	1680	ptsP_ko	0.0007152 6	0.07139618
S70	25	0	ptsP_ko	5.30E-07	5.55E-05
S71	50	0	ptsP_ko	6.00E-07	6.30E-05
S72	50	0	ptsP_ko	6.00E-07	5.99E-05
S73	50	0	ptsP_ko	6.99E-07	7.32E-05
S74	100	0	ptsP_ko	7.82E-07	8.21E-05
S75	100	0	ptsP_ko	4.63E-07	4.62E-05
S76	100	0	ptsP_ko	7.10E-07	7.43E-05
S65	12. 5	0	evgA_ko	4.53E-07	6.68E-05

S66	12. 5	0	evgA_ko	2.58E-07	3.52E-05
S67	12. 5	1	evgA_ko	3.38E-07	4.31E-05
S68	25	0	evgA_ko	4.03E-07	5.94E-05
S69	25	0	evgA_ko	4.26E-07	5.82E-05
S70	25	0	evgA_ko	5.30E-07	6.76E-05
S71	50	0	evgA_ko	6.00E-07	8.85E-05
S72	50	0	evgA_ko	6.00E-07	8.20E-05
S73	50	0	evgA_ko	6.99E-07	8.92E-05
S74	100	0	evgA_ko	7.82E-07	0.00011525
S75	100	0	evgA_ko	4.63E-07	6.32E-05
S76	100	0	evgA_ko	7.10E-07	9.06E-05
S65	12. 5	1	ackA_ko	4.53E-07	3.14E-05
S66	12. 5	0	ackA_ko	2.58E-07	1.73E-05
S67	12. 5	1	ackA_ko	3.38E-07	2.36E-05
S68	25	0	ackA_ko	4.03E-07	2.79E-05
S69	25	1	ackA_ko	4.26E-07	2.85E-05
S70	25	0	ackA_ko	5.30E-07	3.71E-05

S71	50	0	ackA_ko	6.00E-07	4.15E-05
S72	50	0	ackA_ko	6.00E-07	4.02E-05
S73	50	0	ackA_ko	6.99E-07	4.89E-05
S74	100	0	ackA_ko	7.82E-07	5.41E-05
S75	100	0	ackA_ko	4.63E-07	3.10E-05
S76	100	864	ackA_ko	0.0006134 4	0.04294498
S65	12. 5	0	yobF_ko	4.53E-07	6.82E-05
S66	12. 5	0	yobF_ko	2.58E-07	3.66E-05
S67	12. 5	1	yobF_ko	3.38E-07	4.56E-05
S68	25	0	yobF_ko	4.03E-07	6.07E-05
S69	25	0	yobF_ko	4.26E-07	6.05E-05
S70	25	0	yobF_ko	5.30E-07	7.15E-05
S71	50	0	yobF_ko	6.00E-07	9.03E-05
S72	50	0	yobF_ko	6.00E-07	8.52E-05
S73	50	0	yobF_ko	6.99E-07	9.44E-05
S74	100	0	yobF_ko	7.82E-07	0.00011771
S75	100	0	yobF_ko	4.63E-07	6.57E-05

S76	100	0	yobF_ko	7.10E-07	9.59E-05
S65	12. 5	1	dapA_ko	4.53E-07	4.95E-05
S66	12. 5	0	dapA_ko	2.58E-07	2.69E-05
S67	12. 5	0	dapA_ko	3.38E-07	3.67E-05
S68	25	0	dapA_ko	4.03E-07	4.40E-05
S69	25	0	dapA_ko	4.26E-07	4.44E-05
S70	25	0	dapA_ko	5.30E-07	5.76E-05
S71	50	1	dapA_ko	6.00E-07	6.55E-05
S72	50	0	dapA_ko	6.00E-07	6.26E-05
S73	50	0	dapA_ko	6.99E-07	7.60E-05
S74	100	0	dapA_ko	7.82E-07	8.54E-05
S75	100	0	dapA_ko	4.63E-07	4.83E-05
S76	100	0	dapA_ko	7.10E-07	7.72E-05
S65	12. 5	0	dapD_ko	4.53E-07	9.63E-05
S66	12. 5	0	dapD_ko	2.58E-07	5.07E-05
S67	12. 5	0	dapD_ko	3.38E-07	7.15E-05
S68	25	0	dapD_ko	4.03E-07	8.56E-05



S69	25	0	dapD_ko	4.26E-07	8.38E-05
S70	25	0	dapD_ko	5.30E-07	0.0001121
S71	50	0	dapD_ko	6.00E-07	0.00012746
S72	50	0	dapD_ko	6.00E-07	0.00011814
S73	50	0	dapD_ko	6.99E-07	0.00014787
S74	100	0	dapD_ko	7.82E-07	0.00016607
S75	100	0	dapD_ko	4.63E-07	9.11E-05
S76	100	0	dapD_ko	7.10E-07	0.00015023
S65	12. 5	0	dnaJ_ko	4.53E-07	4.29E-05
S66	12. 5	0	dnaJ_ko	2.58E-07	2.47E-05
S67	12. 5	0	dnaJ_ko	3.38E-07	3.16E-05
S68	25	0	dnaJ_ko	4.03E-07	3.81E-05
S69	25	0	dnaJ_ko	4.26E-07	4.07E-05
S70	25	0	dnaJ_ko	5.30E-07	4.96E-05
S71	50	0	dnaJ_ko	6.00E-07	5.68E-05
S72	50	2428	dnaJ_ko	0.0014570 3	0.13943068
S73	50	0	dnaJ_ko	6.99E-07	6.54E-05

S74	100	0	dnaJ_ko	7.82E-07	7.40E-05
S75	100	0	dnaJ_ko	4.63E-07	4.43E-05
S76	100	0	dnaJ_ko	7.10E-07	6.64E-05
S65	12.	1	endA_ko	4.53E-07	3.76E-05
	5				
S66	12.	2652	endA_ko	0.0006831	0.05636256
	5			4	
S67	12.	0	endA_ko	3.38E-07	2.74E-05
	5				
S68	25	0	endA_ko	4.03E-07	3.34E-05
S69	25	1008	endA_ko	0.0004291	0.03540789
				6	
S70	25	0	endA_ko	5.30E-07	4.30E-05
S71	50	0	endA_ko	6.00E-07	4.97E-05
S72	50	259	endA_ko	0.0001554	0.01282343
				2	
S73	50	1	endA_ko	6.99E-07	5.68E-05
S74	100	0	endA_ko	7.82E-07	6.48E-05
S75	100	0	endA_ko	4.63E-07	3.82E-05
S76	100	0	endA_ko	7.10E-07	5.77E-05
S65	12.	0	recJ_ko	4.53E-07	2.54E-05
	5				
S66	12.	1639	recJ_ko	0.0004221	0.023002
	5			9	

S67	12.	0	recJ_ko	3.38E-07	1.84E-05
	5				
S68	25	0	recJ_ko	4.03E-07	2.26E-05
S69	25	0	recJ_ko	4.26E-07	2.32E-05
S70	25	0	recJ_ko	5.30E-07	2.88E-05
S71	50	0	recJ_ko	6.00E-07	3.36E-05
S72	50	0	recJ_ko	6.00E-07	3.27E-05
S73	50	0	recJ_ko	6.99E-07	3.80E-05
S74	100	0	recJ_ko	7.82E-07	4.38E-05
S75	100	3918	recJ_ko	0.0018126	0.09875871
				9	
S76	100	2	recJ_ko	1.42E-06	7.73E-05
S65	12.	0	sbcB_ko	4.53E-07	2.94E-05
	5				
S66	12.	0	sbcB_ko	2.58E-07	1.62E-05
	5				
S67	12.	0	sbcB_ko	3.38E-07	2.18E-05
	5				
S68	25	1	sbcB_ko	4.03E-07	2.61E-05
S69	25	0	sbcB_ko	4.26E-07	2.68E-05
S70	25	0	sbcB_ko	5.30E-07	3.42E-05
S71	50	1	sbcB_ko	6.00E-07	3.89E-05

S72	50	0	sbcB_ko	6.00E-07	3.78E-05
S73	50	0	sbcB_ko	6.99E-07	4.51E-05
S74	100	0	sbcB_ko	7.82E-07	5.07E-05
S75	100	2	sbcB_ko	9.25E-07	5.83E-05
S76	100	0	sbcB_ko	7.10E-07	4.58E-05
S65	12. 5	0	dnaA_ko	4.53E-07	4.50E-05
S66	12. 5	0	dnaA_ko	2.58E-07	2.70E-05
S67	12. 5	0	dnaA_ko	3.38E-07	3.65E-05
S68	25	0	dnaA_ko	4.03E-07	4.00E-05
S69	25	0	dnaA_ko	4.26E-07	4.46E-05
S70	25	0	dnaA_ko	5.30E-07	5.72E-05
S71	50	1	dnaA_ko	6.00E-07	5.95E-05
S72	50	0	dnaA_ko	6.00E-07	6.29E-05
S73	50	0	dnaA_ko	6.99E-07	7.55E-05
S74	100	0	dnaA_ko	7.82E-07	7.76E-05
S75	100	0	dnaA_ko	4.63E-07	4.85E-05
S76	100	0	dnaA_ko	7.10E-07	7.67E-05

S65	12. 5	0	galE_ko	4.53E-07	7.44E-05
S66	12. 5	0	galE_ko	2.58E-07	4.08E-05
S67	12. 5	0	galE_ko	3.38E-07	5.41E-05
S68	25	0	galE_ko	4.03E-07	6.62E-05
S69	25	1	galE_ko	4.26E-07	6.74E-05
S70	25	272	galE_ko	0.0001441 1	0.0230831
S71	50	0	galE_ko	6.00E-07	9.85E-05
S72	50	0	galE_ko	6.00E-07	9.49E-05
S73	50	0	galE_ko	6.99E-07	0.00011195
S74	100	0	galE_ko	7.82E-07	0.00012835
S75	100	0	galE_ko	4.63E-07	7.32E-05
S76	100	0	galE_ko	7.10E-07	0.00011373
S65	12. 5	0	mutS_ko	4.53E-07	6.58E-05
S66	12. 5	1	mutS_ko	2.58E-07	4.10E-05
S67	12. 5	0	mutS_ko	3.38E-07	5.61E-05
S68	25	0	mutS_ko	4.03E-07	5.86E-05
S69	25	0	mutS_ko	4.26E-07	6.78E-05

S70	25	0	mutS_ko	5.30E-07	8.80E-05
S71	50	0	mutS_ko	6.00E-07	8.72E-05
S72	50	0	mutS_ko	6.00E-07	9.55E-05
S73	50	0	mutS_ko	6.99E-07	0.00011609
S74	100	1	mutS_ko	7.82E-07	0.0001136
S75	100	0	mutS_ko	4.63E-07	7.37E-05
S76	100	0	mutS_ko	7.10E-07	0.00011794
S65	12. 5	0	mutL_ko	4.53E-07	3.33E-05
S66	12. 5	0	mutL_ko	2.58E-07	1.85E-05
S67	12. 5	0	mutL_ko	3.38E-07	2.50E-05
S68	25	0	mutL_ko	4.03E-07	2.96E-05
S69	25	0	mutL_ko	4.26E-07	3.06E-05
S70	25	0	mutL_ko	5.30E-07	3.93E-05
S71	50	0	mutL_ko	6.00E-07	4.41E-05
S72	50	0	mutL_ko	6.00E-07	4.31E-05
S73	50	0	mutL_ko	6.99E-07	5.18E-05
S74	100	1	mutL_ko	7.82E-07	5.75E-05

S75	100	1	mutL_ko	4.63E-07	3.32E-05
S76	100	0	mutL_ko	7.10E-07	5.26E-05
S65	12. 5	5	mcrA_ko	2.26E-06	0.00040154
S66	12. 5	0	mcrA_ko	2.58E-07	4.29E-05
S67	12. 5	0	mcrA_ko	3.38E-07	5.43E-05
S68	25	0	mcrA_ko	4.03E-07	7.14E-05
S69	25	0	mcrA_ko	4.26E-07	7.08E-05
S70	25	0	mcrA_ko	5.30E-07	8.52E-05
S71	50	0	mcrA_ko	6.00E-07	0.00010634
S72	50	0	mcrA_ko	6.00E-07	9.98E-05
S73	50	0	mcrA_ko	6.99E-07	0.00011244
S74	100	0	mcrA_ko	7.82E-07	0.00013855
S75	100	0	mcrA_ko	4.63E-07	7.70E-05
S76	100	0	mcrA_ko	7.10E-07	0.00011423
S65	12. 5	10	marR_mut_V84E	4.53E-06	0.00018641
S66	12. 5	0	marR_mut_V84E	2.58E-07	1.07E-05
S67	12. 5	1	marR_mut_V84E	3.38E-07	1.49E-05

S68	25	0	marR_mut_V84E	4.03E-07	1.66E-05
S69	25	0	marR_mut_V84E	4.26E-07	1.78E-05
S70	25	1	marR_mut_V84E	5.30E-07	2.34E-05
S71	50	0	marR_mut_V84E	6.00E-07	2.47E-05
S72	50	0	marR_mut_V84E	6.00E-07	2.50E-05
S73	50	0	marR_mut_V84E	6.99E-07	3.09E-05
S74	100	0	marR_mut_V84E	7.82E-07	3.22E-05
S75	100	1	marR_mut_V84E	4.63E-07	1.93E-05
S76	100	0	marR_mut_V84E	7.10E-07	3.14E-05
S65	12. 5	1	gyrA_mut_S83L	4.53E-07	2.28E-05
S66	12. 5	0	gyrA_mut_S83L	2.58E-07	1.29E-05
S67	12. 5	1	gyrA_mut_S83L	3.38E-07	1.69E-05
S68	25	0	gyrA_mut_S83L	4.03E-07	2.03E-05
S69	25	0	gyrA_mut_S83L	4.26E-07	2.14E-05
S70	25	0	gyrA_mut_S83L	5.30E-07	2.66E-05
S71	50	0	gyrA_mut_S83L	6.00E-07	3.02E-05
S72	50	0	gyrA_mut_S83L	6.00E-07	3.02E-05



S73	50	0	gyrA_mut_S83L	6.99E-07	3.50E-05
S74	100	0	gyrA_mut_S83L	7.82E-07	3.94E-05
S75	100	0	gyrA_mut_S83L	4.63E-07	2.33E-05
S76	100	0	gyrA_mut_S83L	7.10E-07	3.56E-05
S65	12. 5	1	gyrA_mut_D87G	4.53E-07	1.86E-05
S66	12. 5	1	gyrA_mut_D87G	2.58E-07	1.02E-05
S67	12. 5	0	gyrA_mut_D87G	3.38E-07	1.39E-05
S68	25	0	gyrA_mut_D87G	4.03E-07	1.66E-05
S69	25	0	gyrA_mut_D87G	4.26E-07	1.68E-05
S70	25	0	gyrA_mut_D87G	5.30E-07	2.18E-05
S71	50	0	gyrA_mut_D87G	6.00E-07	2.46E-05
S72	50	1	gyrA_mut_D87G	6.00E-07	2.37E-05
S73	50	0	gyrA_mut_D87G	6.99E-07	2.87E-05
S74	100	0	gyrA_mut_D87G	7.82E-07	3.21E-05
S75	100	1	gyrA_mut_D87G	4.63E-07	1.83E-05
S76	100	0	gyrA_mut_D87G	7.10E-07	2.92E-05
S65	12. 5	0	envZ_mut_A396T	4.53E-07	1.90E-05

S66	12. 5	1	envZ_mut_A396T	2.58E-07	1.09E-05
S67	12. 5	1	envZ_mut_A396T	3.38E-07	1.47E-05
S68	25	0	envZ_mut_A396T	4.03E-07	1.69E-05
S69	25	1	envZ_mut_A396T	4.26E-07	1.79E-05
S70	25	0	envZ_mut_A396T	5.30E-07	2.31E-05
S71	50	0	envZ_mut_A396T	6.00E-07	2.51E-05
S72	50	0	envZ_mut_A396T	6.00E-07	2.53E-05
S73	50	0	envZ_mut_A396T	6.99E-07	3.05E-05
S74	100	0	envZ_mut_A396T	7.82E-07	3.27E-05
S75	100	0	envZ_mut_A396T	4.63E-07	1.95E-05
S76	100	0	envZ_mut_A396T	7.10E-07	3.10E-05
S65	12. 5	0	trkH_mut_T350K	4.53E-07	2.09E-05
S66	12. 5	3	trkH_mut_T350K	7.73E-07	3.47E-05
S67	12. 5	1	trkH_mut_T350K	3.38E-07	1.48E-05
S68	25	1	trkH_mut_T350K	4.03E-07	1.85E-05
S69	25	0	trkH_mut_T350K	4.26E-07	1.91E-05
S70	25	0	trkH_mut_T350K	5.30E-07	2.32E-05

S71	50	0	trkH_mut_T350K	6.00E-07	2.76E-05
S72	50	0	trkH_mut_T350K	6.00E-07	2.70E-05
S73	50	0	trkH_mut_T350K	6.99E-07	3.05E-05
S74	100	1	trkH_mut_T350K	7.82E-07	3.60E-05
S75	100	1	trkH_mut_T350K	4.63E-07	2.08E-05
S76	100	1	trkH_mut_T350K	7.10E-07	3.10E-05
S65	12.5	4	soxR_mut_D137E	1.81E-06	7.65E-05
S66	12.5	0	soxR_mut_D137E	2.58E-07	1.10E-05
S67	12.5	0	soxR_mut_D137E	3.38E-07	1.43E-05
S68	25	0	soxR_mut_D137E	4.03E-07	1.70E-05
S69	25	290	soxR_mut_D137E	0.00012347	0.00526505
S70	25	0	soxR_mut_D137E	5.30E-07	2.24E-05
S71	50	0	soxR_mut_D137E	6.00E-07	2.53E-05
S72	50	0	soxR_mut_D137E	6.00E-07	2.56E-05
S73	50	0	soxR_mut_D137E	6.99E-07	2.95E-05
S74	100	0	soxR_mut_D137E	7.82E-07	3.30E-05
S75	100	1	soxR_mut_D137E	4.63E-07	1.97E-05

S76	100	0	soxR_mut_D137E	7.10E-07	3.00E-05
S65	12.5	1	ycbZ_mut_S438R	4.53E-07	2.74E-05
S66	12.5	0	ycbZ_mut_S438R	2.58E-07	1.54E-05
S67	12.5	0	ycbZ_mut_S438R	3.38E-07	1.95E-05
S68	25	49	ycbZ_mut_S438R	1.97E-05	0.00119298
S69	25	0	ycbZ_mut_S438R	4.26E-07	2.54E-05
S70	25	0	ycbZ_mut_S438R	5.30E-07	3.06E-05
S71	50	0	ycbZ_mut_S438R	6.00E-07	3.63E-05
S72	50	0	ycbZ_mut_S438R	6.00E-07	3.58E-05
S73	50	0	ycbZ_mut_S438R	6.99E-07	4.04E-05
S74	100	0	ycbZ_mut_S438R	7.82E-07	4.72E-05
S75	100	1	ycbZ_mut_S438R	4.63E-07	2.76E-05
S76	100	1	ycbZ_mut_S438R	7.10E-07	4.10E-05
S65	12.5	3	folA_mut_P21L	1.36E-06	6.23E-05
S66	12.5	0	folA_mut_P21L	2.58E-07	1.20E-05
S67	12.5	1	folA_mut_P21L	3.38E-07	1.58E-05
S68	25	0	folA_mut_P21L	4.03E-07	1.85E-05

S69	25	0	folA_mut_P21L	4.26E-07	1.98E-05
S70	25	0	folA_mut_P21L	5.30E-07	2.47E-05
S71	50	1	folA_mut_P21L	6.00E-07	2.75E-05
S72	50	0	folA_mut_P21L	6.00E-07	2.79E-05
S73	50	0	folA_mut_P21L	6.99E-07	3.26E-05
S74	100	0	folA_mut_P21L	7.82E-07	3.58E-05
S75	100	0	folA_mut_P21L	4.63E-07	2.15E-05
S76	100	0	folA_mut_P21L	7.10E-07	3.31E-05
S65	12. 5	2	folA_mut_W30C	9.06E-07	6.56E-05
S66	12. 5	0	folA_mut_W30C	2.58E-07	1.94E-05
S67	12. 5	1	folA_mut_W30C	3.38E-07	2.62E-05
S68	25	1	folA_mut_W30C	4.03E-07	2.92E-05
S69	25	2	folA_mut_W30C	8.52E-07	6.41E-05
S70	25	0	folA_mut_W30C	5.30E-07	4.10E-05
S71	50	0	folA_mut_W30C	6.00E-07	4.34E-05
S72	50	0	folA_mut_W30C	6.00E-07	4.52E-05
S73	50	0	folA_mut_W30C	6.99E-07	5.41E-05

S74	100	0	folA_mut_W30C	7.82E-07	5.66E-05
S75	100	1	folA_mut_W30C	4.63E-07	3.48E-05
S76	100	0	folA_mut_W30C	7.10E-07	5.50E-05
S65	12. 5	66	folA_mut_W30G	2.99E-05	0.00109125
S66	12. 5	2	folA_mut_W30G	5.15E-07	1.89E-05
S67	12. 5	0	folA_mut_W30G	3.38E-07	1.22E-05
S68	25	0	folA_mut_W30G	4.03E-07	1.47E-05
S69	25	0	folA_mut_W30G	4.26E-07	1.56E-05
S70	25	0	folA_mut_W30G	5.30E-07	1.91E-05
S71	50	0	folA_mut_W30G	6.00E-07	2.19E-05
S72	50	0	folA_mut_W30G	6.00E-07	2.20E-05
S73	50	0	folA_mut_W30G	6.99E-07	2.52E-05
S74	100	0	folA_mut_W30G	7.82E-07	2.85E-05
S75	100	0	folA_mut_W30G	4.63E-07	1.70E-05
S76	100	0	folA_mut_W30G	7.10E-07	2.56E-05
S65	12. 5	0	folA_mut_I94L	4.53E-07	1.69E-05
S66	12. 5	1	folA_mut_I94L	2.58E-07	9.78E-06

S67	12. 5	3	folA_mut_I94L	1.01E-06	3.96E-05
S68	25	0	folA_mut_I94L	4.03E-07	1.51E-05
S69	25	0	folA_mut_I94L	4.26E-07	1.62E-05
S70	25	0	folA_mut_I94L	5.30E-07	2.07E-05
S71	50	0	folA_mut_I94L	6.00E-07	2.24E-05
S72	50	0	folA_mut_I94L	6.00E-07	2.28E-05
S73	50	0	folA_mut_I94L	6.99E-07	2.73E-05
S74	100	1	folA_mut_I94L	7.82E-07	2.92E-05
S75	100	0	folA_mut_I94L	4.63E-07	1.76E-05
S76	100	0	folA_mut_I94L	7.10E-07	2.77E-05
S65	12. 5	2	folA_mut_F153S	9.06E-07	3.35E-05
S66	12. 5	0	folA_mut_F153S	2.58E-07	9.72E-06
S67	12. 5	0	folA_mut_F153S	3.38E-07	1.28E-05
S68	25	0	folA_mut_F153S	4.03E-07	1.49E-05
S69	25	0	folA_mut_F153S	4.26E-07	1.61E-05
S70	25	223	folA_mut_F153S	0.0001181 5	0.00448491
S71	50	2	folA_mut_F153S	1.20E-06	4.43E-05

S72	50	1	folA_mut_F153S	6.00E-07	2.26E-05
S73	50	1	folA_mut_F153S	6.99E-07	2.65E-05
S74	100	0	folA_mut_F153S	7.82E-07	2.89E-05
S75	100	0	folA_mut_F153S	4.63E-07	1.74E-05
S76	100	1	folA_mut_F153S	7.10E-07	2.70E-05
S65	12. 5	1	folA_mut_D27E	4.53E-07	2.73E-05
S66	12. 5	1	folA_mut_D27E	2.58E-07	1.56E-05
S67	12. 5	1	folA_mut_D27E	3.38E-07	2.16E-05
S68	25	0	folA_mut_D27E	4.03E-07	2.43E-05
S69	25	1	folA_mut_D27E	4.26E-07	2.58E-05
S70	25	0	folA_mut_D27E	5.30E-07	3.39E-05
S71	50	1	folA_mut_D27E	6.00E-07	3.61E-05
S72	50	1	folA_mut_D27E	6.00E-07	3.64E-05
S73	50	0	folA_mut_D27E	6.99E-07	4.47E-05
S74	100	0	folA_mut_D27E	7.82E-07	4.71E-05
S75	100	0	folA_mut_D27E	4.63E-07	2.81E-05
S76	100	0	folA_mut_D27E	7.10E-07	4.54E-05



S65	12. 5	0	folA_mut_M20V	4.53E-07	3.69E-05
S66	12. 5	2	folA_mut_M20V	5.15E-07	4.22E-05
S67	12. 5	2	folA_mut_M20V	6.76E-07	5.57E-05
S68	25	0	folA_mut_M20V	4.03E-07	3.28E-05
S69	25	0	folA_mut_M20V	4.26E-07	3.49E-05
S70	25	0	folA_mut_M20V	5.30E-07	4.37E-05
S71	50	0	folA_mut_M20V	6.00E-07	4.89E-05
S72	50	2	folA_mut_M20V	1.20E-06	9.83E-05
S73	50	0	folA_mut_M20V	6.99E-07	5.77E-05
S74	100	0	folA_mut_M20V	7.82E-07	6.37E-05
S75	100	0	folA_mut_M20V	4.63E-07	3.79E-05
S76	100	0	folA_mut_M20V	7.10E-07	5.86E-05
S65	12. 5	0	trkH_mut_G84D	4.53E-07	3.38E-05
S66	12. 5	0	trkH_mut_G84D	2.58E-07	1.91E-05
S67	12. 5	0	trkH_mut_G84D	3.38E-07	2.70E-05
S68	25	1	trkH_mut_G84D	4.03E-07	3.00E-05
S69	25	0	trkH_mut_G84D	4.26E-07	3.15E-05

S70	25	0	trkH_mut_G84D	5.30E-07	4.24E-05
S71	50	0	trkH_mut_G84D	6.00E-07	4.47E-05
S72	50	0	trkH_mut_G84D	6.00E-07	4.44E-05
S73	50	0	trkH_mut_G84D	6.99E-07	5.60E-05
S74	100	0	trkH_mut_G84D	7.82E-07	5.83E-05
S75	100	0	trkH_mut_G84D	4.63E-07	3.42E-05
S76	100	0	trkH_mut_G84D	7.10E-07	5.69E-05
S65	12. 5	1	trkH_mut_I143S	4.53E-07	2.84E-05
S66	12. 5	1	trkH_mut_I143S	2.58E-07	1.64E-05
S67	12. 5	0	trkH_mut_I143S	3.38E-07	2.07E-05
S68	25	0	trkH_mut_I143S	4.03E-07	2.52E-05
S69	25	0	trkH_mut_I143S	4.26E-07	2.72E-05
S70	25	3	trkH_mut_I143S	1.59E-06	9.75E-05
S71	50	0	trkH_mut_I143S	6.00E-07	3.75E-05
S72	50	0	trkH_mut_I143S	6.00E-07	3.83E-05
S73	50	0	trkH_mut_I143S	6.99E-07	4.29E-05
S74	100	0	trkH_mut_I143S	7.82E-07	4.89E-05

S75	100	0	trkH_mut_I143S	4.63E-07	2.95E-05
S76	100	0	trkH_mut_I143S	7.10E-07	4.36E-05
S65	12. 5	0	trkH_mut_L80Q	4.53E-07	3.00E-05
S66	12. 5	0	trkH_mut_L80Q	2.58E-07	1.79E-05
S67	12. 5	1	trkH_mut_L80Q	3.38E-07	2.28E-05
S68	25	0	trkH_mut_L80Q	4.03E-07	2.66E-05
S69	25	0	trkH_mut_L80Q	4.26E-07	2.95E-05
S70	25	0	trkH_mut_L80Q	5.30E-07	3.58E-05
S71	50	2	trkH_mut_L80Q	1.20E-06	7.94E-05
S72	50	0	trkH_mut_L80Q	6.00E-07	4.16E-05
S73	50	0	trkH_mut_L80Q	6.99E-07	4.72E-05
S74	100	0	trkH_mut_L80Q	7.82E-07	5.17E-05
S75	100	1	trkH_mut_L80Q	4.63E-07	3.21E-05
S76	100	0	trkH_mut_L80Q	7.10E-07	4.80E-05
S65	12. 5	1	trkH_mut_L80R	4.53E-07	3.07E-05
S66	12. 5	0	trkH_mut_L80R	2.58E-07	1.72E-05
S67	12. 5	4047	trkH_mut_L80R	0.0013669 3	0.09322508

S68	25	0	trkH_mut_L80R	4.03E-07	2.73E-05
S69	25	0	trkH_mut_L80R	4.26E-07	2.84E-05
S70	25	0	trkH_mut_L80R	5.30E-07	3.61E-05
S71	50	1	trkH_mut_L80R	6.00E-07	4.06E-05
S72	50	0	trkH_mut_L80R	6.00E-07	4.00E-05
S73	50	0	trkH_mut_L80R	6.99E-07	4.77E-05
S74	100	0	trkH_mut_L80R	7.82E-07	5.29E-05
S75	100	0	trkH_mut_L80R	4.63E-07	3.08E-05
S76	100	0	trkH_mut_L80R	7.10E-07	4.84E-05
S65	12. 5	0	marR_ko	4.53E-07	2.74E-05
S66	12. 5	1	marR_ko	2.58E-07	1.59E-05
S67	12. 5	1	marR_ko	3.38E-07	2.04E-05
S68	25	0	marR_ko	4.03E-07	2.44E-05
S69	25	0	marR_ko	4.26E-07	2.62E-05
S70	25	0	marR_ko	5.30E-07	3.20E-05
S71	50	2	marR_ko	1.20E-06	7.26E-05
S72	50	0	marR_ko	6.00E-07	3.69E-05

S73	50	0	marR_ko	6.99E-07	4.22E-05
S74	100	0	marR_ko	7.82E-07	4.73E-05
S75	100	0	marR_ko	4.63E-07	2.85E-05
S76	100	0	marR_ko	7.10E-07	4.29E-05
S65	12. 5	0	envZ_ko	4.53E-07	3.05E-05
S66	12. 5	0	envZ_ko	2.58E-07	1.75E-05
S67	12. 5	0	envZ_ko	3.38E-07	2.32E-05
S68	25	1	envZ_ko	4.03E-07	2.71E-05
S69	25	1	envZ_ko	4.26E-07	2.89E-05
S70	25	0	envZ_ko	5.30E-07	3.63E-05
S71	50	0	envZ_ko	6.00E-07	4.03E-05
S72	50	0	envZ_ko	6.00E-07	4.08E-05
S73	50	0	envZ_ko	6.99E-07	4.79E-05
S74	100	0	envZ_ko	7.82E-07	5.26E-05
S75	100	0	envZ_ko	4.63E-07	3.14E-05
S76	100	2	envZ_ko	1.42E-06	9.74E-05
S65	12. 5	1	trkH_ko	4.53E-07	2.57E-05

S66	12. 5	0	trkH_ko	2.58E-07	1.42E-05
S67	12. 5	3	trkH_ko	1.01E-06	5.42E-05
S68	25	0	trkH_ko	4.03E-07	2.29E-05
S69	25	0	trkH_ko	4.26E-07	2.35E-05
S70	25	0	trkH_ko	5.30E-07	2.83E-05
S71	50	1	trkH_ko	6.00E-07	3.40E-05
S72	50	0	trkH_ko	6.00E-07	3.31E-05
S73	50	0	trkH_ko	6.99E-07	3.74E-05
S74	100	1	trkH_ko	7.82E-07	4.43E-05
S75	100	0	trkH_ko	4.63E-07	2.55E-05
S76	100	0	trkH_ko	7.10E-07	3.80E-05
S65	12. 5	1	folA_ko	4.53E-07	2.48E-05
S66	12. 5	2	folA_ko	5.15E-07	2.86E-05
S67	12. 5	0	folA_ko	3.38E-07	1.91E-05
S68	25	0	folA_ko	4.03E-07	2.21E-05
S69	25	0	folA_ko	4.26E-07	2.36E-05
S70	25	0	folA_ko	5.30E-07	2.99E-05

S71	50	0	folA_ko	6.00E-07	3.29E-05
S72	50	0	folA_ko	6.00E-07	3.33E-05
S73	50	0	folA_ko	6.99E-07	3.95E-05
S74	100	0	folA_ko	7.82E-07	4.28E-05
S75	100	1	folA_ko	4.63E-07	2.57E-05
S76	100	0	folA_ko	7.10E-07	4.01E-05
S65	12. 5	0	pgi_ko	4.53E-07	2.87E-05
S66	12. 5	0	pgi_ko	2.58E-07	1.59E-05
S67	12. 5	0	pgi_ko	3.38E-07	2.00E-05
S68	25	0	pgi_ko	4.03E-07	2.55E-05
S69	25	0	pgi_ko	4.26E-07	2.63E-05
S70	25	1	pgi_ko	5.30E-07	3.14E-05
S71	50	0	pgi_ko	6.00E-07	3.80E-05
S72	50	0	pgi_ko	6.00E-07	3.71E-05
S73	50	0	pgi_ko	6.99E-07	4.15E-05
S74	100	0	pgi_ko	7.82E-07	4.95E-05
S75	100	0	pgi_ko	4.63E-07	2.86E-05

S76	100	1	pgi_ko	7.10E-07	4.21E-05
S65	12.5	0	galK_ko	4.53E-07	2.98E-05
S66	12.5	0	galK_ko	2.58E-07	1.82E-05
S67	12.5	0	galK_ko	3.38E-07	2.30E-05
S68	25	0	galK_ko	4.03E-07	2.65E-05
S69	25	0	galK_ko	4.26E-07	3.02E-05
S70	25	0	galK_ko	5.30E-07	3.60E-05
S71	50	0	galK_ko	6.00E-07	3.94E-05
S72	50	0	galK_ko	6.00E-07	4.25E-05
S73	50	0	galK_ko	6.99E-07	4.75E-05
S74	100	0	galK_ko	7.82E-07	5.14E-05
S75	100	0	galK_ko	4.63E-07	3.28E-05
S76	100	0	galK_ko	7.10E-07	4.83E-05
S65	12.5	3	fabA_KO	1.36E-06	7.26E-05
S66	12.5	0	fabA_KO	2.58E-07	1.39E-05
S67	12.5	0	fabA_KO	3.38E-07	1.78E-05
S68	25	0	fabA_KO	4.03E-07	2.15E-05



S69	25	1	fabA_KO	4.26E-07	2.29E-05
S70	25	0	fabA_KO	5.30E-07	2.79E-05
S71	50	0	fabA_KO	6.00E-07	3.21E-05
S72	50	0	fabA_KO	6.00E-07	3.23E-05
S73	50	0	fabA_KO	6.99E-07	3.68E-05
S74	100	0	fabA_KO	7.82E-07	4.18E-05
S75	100	0	fabA_KO	4.63E-07	2.49E-05
S76	100	0	fabA_KO	7.10E-07	3.73E-05
S65	12. 5	0	gyrA_KO	4.53E-07	3.52E-05
S66	12. 5	0	gyrA_KO	2.58E-07	2.06E-05
S67	12. 5	1	gyrA_KO	3.38E-07	2.75E-05
S68	25	0	gyrA_KO	4.03E-07	3.13E-05
S69	25	0	gyrA_KO	4.26E-07	3.40E-05
S70	25	0	gyrA_KO	5.30E-07	4.32E-05
S71	50	0	gyrA_KO	6.00E-07	4.66E-05
S72	50	0	gyrA_KO	6.00E-07	4.80E-05
S73	50	2	gyrA_KO	1.40E-06	0.0001139

S74	100	0	gyrA_KO	7.82E-07	6.07E-05
S75	100	0	gyrA_KO	4.63E-07	3.70E-05
S76	100	0	gyrA_KO	7.10E-07	5.79E-05
S65	12. 5	0	lacZ_KO	4.53E-07	3.78E-05
S66	12. 5	0	lacZ_KO	2.58E-07	2.17E-05
S67	12. 5	0	lacZ_KO	3.38E-07	2.84E-05
S68	25	0	lacZ_KO	4.03E-07	3.37E-05
S69	25	0	lacZ_KO	4.26E-07	3.58E-05
S70	25	0	lacZ_KO	5.30E-07	4.46E-05
S71	50	0	lacZ_KO	6.00E-07	5.01E-05
S72	50	0	lacZ_KO	6.00E-07	5.05E-05
S73	50	0	lacZ_KO	6.99E-07	5.88E-05
S74	100	0	lacZ_KO	7.82E-07	6.53E-05
S75	100	0	lacZ_KO	4.63E-07	3.90E-05
S76	100	0	lacZ_KO	7.10E-07	5.98E-05
S65	12. 5	45	lacZ_KO2	2.04E-05	0.00185809
S66	12. 5	0	lacZ_KO2	2.58E-07	2.37E-05

S67	12. 5	2	lacZ_KO2	6.76E-07	5.81E-05
S68	25	0	lacZ_KO2	4.03E-07	3.67E-05
S69	25	0	lacZ_KO2	4.26E-07	3.92E-05
S70	25	0	lacZ_KO2	5.30E-07	4.56E-05
S71	50	0	lacZ_KO2	6.00E-07	5.47E-05
S72	50	0	lacZ_KO2	6.00E-07	5.52E-05
S73	50	0	lacZ_KO2	6.99E-07	6.01E-05
S74	100	0	lacZ_KO2	7.82E-07	7.12E-05
S75	100	0	lacZ_KO2	4.63E-07	4.26E-05
S76	100	0	lacZ_KO2	7.10E-07	6.11E-05
S65	12. 5	0	recA_KO	4.53E-07	3.09E-05
S66	12. 5	0	recA_KO	2.58E-07	1.83E-05
S67	12. 5	1	recA_KO	3.38E-07	2.37E-05
S68	25	0	recA_KO	4.03E-07	2.75E-05
S69	25	0	recA_KO	4.26E-07	3.03E-05
S70	25	0	recA_KO	5.30E-07	3.71E-05
S71	50	2196	recA_KO	0.0013172	0.08987631

S72	50	0	recA_KO	6.00E-07	4.27E-05
S73	50	0	recA_KO	6.99E-07	4.90E-05
S74	100	1	recA_KO	7.82E-07	5.33E-05
S75	100	0	recA_KO	4.63E-07	3.30E-05
S76	100	0	recA_KO	7.10E-07	4.98E-05
S65	12. 5	46	rpoB_A_1538_C	2.08E-05	0.0013945
S66	12. 5	138	rpoB_A_1538_C	3.55E-05	0.00237748
S67	12. 5	74	rpoB_A_1538_C	2.50E-05	0.00166847
S68	25	81	rpoB_A_1538_C	3.26E-05	0.00218348
S69	25	55	rpoB_A_1538_C	2.34E-05	0.00156611
S70	25	122	rpoB_A_1538_C	6.46E-05	0.00431463
S71	50	77	rpoB_A_1538_C	4.62E-05	0.00309086
S72	50	72	rpoB_A_1538_C	4.32E-05	0.00288973
S73	50	85	rpoB_A_1538_C	5.94E-05	0.00396541
S74	100	53	rpoB_A_1538_C	4.14E-05	0.00277194
S75	100	52	rpoB_A_1538_C	2.41E-05	0.00160903
S76	100	137	rpoB_A_1538_C	9.73E-05	0.00649304

S65	12. 5	72350	rpoB_A_1547_G	0.0327741 9	1.74348159
S66	12. 5	257867	rpoB_A_1547_G	0.0664246 7	3.73486669
S67	12. 5	194295	rpoB_A_1547_G	0.0656260 1	3.49098096
S68	25	28324	rpoB_A_1547_G	0.0114091	0.60692769
S69	25	30207	rpoB_A_1547_G	0.0128606 7	0.72311823
S70	25	33034	rpoB_A_1547_G	0.0175014 1	0.93098892
S71	50	227	rpoB_A_1547_G	0.0001361 6	0.00724322
S72	50	495	rpoB_A_1547_G	0.0002970 5	0.01670213
S73	50	2695	rpoB_A_1547_G	0.0018834 6	0.10019068
S74	100	21	rpoB_A_1547_G	1.64E-05	0.00087306
S75	100	1285	rpoB_A_1547_G	0.0005945 1	0.03342773
S76	100	4	rpoB_A_1547_G	2.84E-06	0.00015107
S65	12. 5	0	rpoB_A_1623_T	4.53E-07	2.82E-05
S66	12. 5	0	rpoB_A_1623_T	2.58E-07	1.60E-05
S67	12. 5	0	rpoB_A_1623_T	3.38E-07	2.11E-05
S68	25	68	rpoB_A_1623_T	2.74E-05	0.00170487
S69	25	1	rpoB_A_1623_T	4.26E-07	2.64E-05

S70	25	0	rpoB_A_1623_T	5.30E-07	3.30E-05
S71	50	0	rpoB_A_1623_T	6.00E-07	3.73E-05
S72	50	0	rpoB_A_1623_T	6.00E-07	3.72E-05
S73	50	0	rpoB_A_1623_T	6.99E-07	4.36E-05
S74	100	0	rpoB_A_1623_T	7.82E-07	4.86E-05
S75	100	1	rpoB_A_1623_T	4.63E-07	2.87E-05
S76	100	0	rpoB_A_1623_T	7.10E-07	4.43E-05
S65	12.5	1	rpoB_A_1657_G	4.53E-07	3.05E-05
S66	12.5	1	rpoB_A_1657_G	2.58E-07	1.81E-05
S67	12.5	0	rpoB_A_1657_G	3.38E-07	2.38E-05
S68	25	0	rpoB_A_1657_G	4.03E-07	2.72E-05
S69	25	0	rpoB_A_1657_G	4.26E-07	2.99E-05
S70	25	0	rpoB_A_1657_G	5.30E-07	3.74E-05
S71	50	0	rpoB_A_1657_G	6.00E-07	4.04E-05
S72	50	2208	rpoB_A_1657_G	0.00132501	0.09319431
S73	50	0	rpoB_A_1657_G	6.99E-07	4.93E-05
S74	100	0	rpoB_A_1657_G	7.82E-07	5.27E-05

S75	100	0	rpoB_A_1657_G	4.63E-07	3.25E-05
S76	100	0	rpoB_A_1657_G	7.10E-07	5.01E-05
S65	12. 5	0	rpoB_A_1657_T	4.53E-07	3.10E-05
S66	12. 5	0	rpoB_A_1657_T	2.58E-07	1.68E-05
S67	12. 5	1	rpoB_A_1657_T	3.38E-07	2.30E-05
S68	25	0	rpoB_A_1657_T	4.03E-07	2.76E-05
S69	25	0	rpoB_A_1657_T	4.26E-07	2.78E-05
S70	25	0	rpoB_A_1657_T	5.30E-07	3.60E-05
S71	50	0	rpoB_A_1657_T	6.00E-07	4.11E-05
S72	50	1	rpoB_A_1657_T	6.00E-07	3.92E-05
S73	50	0	rpoB_A_1657_T	6.99E-07	4.75E-05
S74	100	1	rpoB_A_1657_T	7.82E-07	5.35E-05
S75	100	0	rpoB_A_1657_T	4.63E-07	3.02E-05
S76	100	0	rpoB_A_1657_T	7.10E-07	4.83E-05
S65	12. 5	800	rpoB_A_1714_C	0.0003624	0.01765444
S66	12. 5	1222	rpoB_A_1714_C	0.0003147 8	0.01541766
S67	12. 5	1809	rpoB_A_1714_C	0.0006110 2	0.02906726

S68	25	3	rpoB_A_1714_C	1.21E-06	5.89E-05
S69	25	0	rpoB_A_1714_C	4.26E-07	2.09E-05
S70	25	0	rpoB_A_1714_C	5.30E-07	2.52E-05
S71	50	0	rpoB_A_1714_C	6.00E-07	2.92E-05
S72	50	0	rpoB_A_1714_C	6.00E-07	2.94E-05
S73	50	2	rpoB_A_1714_C	1.40E-06	6.65E-05
S74	100	2	rpoB_A_1714_C	1.56E-06	7.61E-05
S75	100	0	rpoB_A_1714_C	4.63E-07	2.27E-05
S76	100	0	rpoB_A_1714_C	7.10E-07	3.38E-05
S65	12. 5	220	rpoB_A_443_T	9.97E-05	0.00769318
S66	12. 5	1052	rpoB_A_443_T	0.0002709 9	0.01971743
S67	12. 5	1404	rpoB_A_443_T	0.0004742 2	0.03413992
S68	25	0	rpoB_A_443_T	4.03E-07	3.11E-05
S69	25	12	rpoB_A_443_T	5.11E-06	0.00037174
S70	25	2	rpoB_A_443_T	1.06E-06	7.63E-05
S71	50	0	rpoB_A_443_T	6.00E-07	4.63E-05
S72	50	2089	rpoB_A_443_T	0.0012536	0.09121379



S73	50	0	rpoB_A_443_T	6.99E-07	5.03E-05
S74	100	0	rpoB_A_443_T	7.82E-07	6.03E-05
S75	100	1	rpoB_A_443_T	4.63E-07	3.37E-05
S76	100	3	rpoB_A_443_T	2.13E-06	0.00015334
S65	12.5	39596	rpoB_C_1535_T	0.01793679	1.19574727
S66	12.5	126592	rpoB_C_1535_T	0.03260918	2.19744792
S67	12.5	69145	rpoB_C_1535_T	0.02335475	1.56192447
S68	25	18550	rpoB_C_1535_T	0.00747207	0.49812178
S69	25	14853	rpoB_C_1535_T	0.00632368	0.42613665
S70	25	21118	rpoB_C_1535_T	0.01118831	0.74825482
S71	50	1825	rpoB_C_1535_T	0.00109467	0.0729756
S72	50	445	rpoB_C_1535_T	0.00026704	0.01799533
S73	50	343	rpoB_C_1535_T	0.00023971	0.01603157
S74	100	87	rpoB_C_1535_T	6.80E-05	0.00453268
S75	100	0	rpoB_C_1535_T	4.63E-07	3.12E-05
S76	100	1	rpoB_C_1535_T	7.10E-07	4.75E-05
S65	12.5	483233	rpoB_C_1576_A	0.21890212	15.6722414

S66	12. 5	784842	rpoB_C_1576_A	0.2021696	14.6145837
S67	12. 5	638289	rpoB_C_1576_A	0.2155915 6	15.4743802
S68	25	578819	rpoB_C_1576_A	0.2331523	16.6924792
S69	25	520496	rpoB_C_1576_A	0.2216018 6	16.0193168
S70	25	357000	rpoB_C_1576_A	0.1891385 7	13.5756806
S71	50	238142	rpoB_C_1576_A	0.1428420 6	10.2267409
S72	50	274695	rpoB_C_1576_A	0.1648434 7	11.9163254
S73	50	258562	rpoB_C_1576_A	0.1807015 1	12.9700991
S74	100	110423	rpoB_C_1576_A	0.0862979	6.17847632
S75	100	135162	rpoB_C_1576_A	0.0625334 8	4.52046621
S76	100	130502	rpoB_C_1576_A	0.0926557 4	6.65049309
S65	12. 5	240194	rpoB_C_1576_G	0.1088066 8	8.08910455
S66	12. 5	505311	rpoB_C_1576_G	0.1301644 4	9.17993732
S67	12. 5	326547	rpoB_C_1576_G	0.1102960 8	7.79220043
S68	25	289143	rpoB_C_1576_G	0.1164688	8.65873631
S69	25	233316	rpoB_C_1576_G	0.0993345 9	7.00564077
S70	25	227468	rpoB_C_1576_G	0.1205125 3	8.513972

S71	50	261766	rpoB_C_1576_G	0.1570121	11.6728864
				8	
S72	50	231843	rpoB_C_1576_G	0.1391281	9.8121084
				4	
S73	50	197965	rpoB_C_1576_G	0.1383520	9.77429682
				2	
S74	100	201990	rpoB_C_1576_G	0.1578594	11.7358751
				4	
S75	100	387174	rpoB_C_1576_G	0.1791283	12.6331471
S76	100	219810	rpoB_C_1576_G	0.1560639	11.0256103
				6	
S65	12.	766570	rpoB_C_1576_T	0.3472523	29.106547
	5			6	
S66	12.	124688	rpoB_C_1576_T	0.3211890	24.9071387
	5	7		4	
S67	12.	924816	rpoB_C_1576_T	0.3123703	24.9977827
	5				
S68	25	840428	rpoB_C_1576_T	0.3385302	28.3754604
				1	
S69	25	885610	rpoB_C_1576_T	0.3770496	29.2389406
				2	
S70	25	715070	rpoB_C_1576_T	0.3788440	30.3174175
				3	
S71	50	684565	rpoB_C_1576_T	0.4106149	34.4175765
				9	
S72	50	776184	rpoB_C_1576_T	0.4657852	36.1200893
				1	
S73	50	569662	rpoB_C_1576_T	0.3981203	31.8600239
				2	
S74	100	656197	rpoB_C_1576_T	0.5128317	42.9853458
				9	
S75	100	110225	rpoB_C_1576_T	0.5099660	39.5461679
		8		7	

S76	100	680647	rpoB_C_1576_T	0.48325584	38.6730892
S65	12.5	0	rpoB_C_1721_A	4.53E-07	2.90E-05
S66	12.5	2087	rpoB_C_1721_A	0.0005376	0.03514962
S67	12.5	26	rpoB_C_1721_A	8.78E-06	0.00057105
S68	25	1	rpoB_C_1721_A	4.03E-07	2.58E-05
S69	25	15	rpoB_C_1721_A	6.39E-06	0.00041755
S70	25	0	rpoB_C_1721_A	5.30E-07	3.45E-05
S71	50	0	rpoB_C_1721_A	6.00E-07	3.85E-05
S72	50	2	rpoB_C_1721_A	1.20E-06	7.85E-05
S73	50	1	rpoB_C_1721_A	6.99E-07	4.54E-05
S74	100	3	rpoB_C_1721_A	2.34E-06	0.00015032
S75	100	1	rpoB_C_1721_A	4.63E-07	3.02E-05
S76	100	0	rpoB_C_1721_A	7.10E-07	4.62E-05
S65	12.5	1968	rpoB_CACTCGGCCAG_1595_DELETE	0.00089149	0.08260284
S66	12.5	13372	rpoB_CACTCGGCCAG_1595_DELETE	0.00344453	0.31176625
S67	12.5	5430	rpoB_CACTCGGCCAG_1595_DELETE	0.00183406	0.16741736
S68	25	5066	rpoB_CACTCGGCCAG_1595_DELETE	0.00204062	0.18907695

S69	25	6077	rpoB_CACTCGGCCAG_1595_DELETE	0.00258729	0.23417706
S70	25	5170	rpoB_CACTCGGCCAG_1595_DELETE	0.00273907	0.25002803
S71	50	927	rpoB_CACTCGGCCAG_1595_DELETE	0.00055603	0.05152006
S72	50	1585	rpoB_CACTCGGCCAG_1595_DELETE	0.00095115	0.08608934
S73	50	1379	rpoB_CACTCGGCCAG_1595_DELETE	0.00096374	0.08797264
S74	100	150	rpoB_CACTCGGCCAG_1595_DELETE	0.00011723	0.01086197
S75	100	791	rpoB_CACTCGGCCAG_1595_DELETE	0.00036596	0.0331233
S76	100	82	rpoB_CACTCGGCCAG_1595_DELETE	5.82E-05	0.00531441
S65	12.5	4	rpoB_CG_1574_TT	1.81E-06	0.00012073
S66	12.5	1	rpoB_CG_1574_TT	2.58E-07	1.71E-05
S67	12.5	2	rpoB_CG_1574_TT	6.76E-07	4.43E-05
S68	25	0	rpoB_CG_1574_TT	4.03E-07	2.68E-05
S69	25	0	rpoB_CG_1574_TT	4.26E-07	2.83E-05
S70	25	0	rpoB_CG_1574_TT	5.30E-07	3.48E-05
S71	50	1	rpoB_CG_1574_TT	6.00E-07	4.00E-05
S72	50	1	rpoB_CG_1574_TT	6.00E-07	3.99E-05
S73	50	1	rpoB_CG_1574_TT	6.99E-07	4.59E-05

S74	100	0	rpoB_CG_1574_TT	7.82E-07	5.21E-05
S75	100	0	rpoB_CG_1574_TT	4.63E-07	3.08E-05
S76	100	0	rpoB_CG_1574_TT	7.10E-07	4.66E-05
S65	12. 5	0	rpoB_G_1545_A	4.53E-07	2.08E-05
S66	12. 5	4	rpoB_G_1545_A	1.03E-06	4.73E-05
S67	12. 5	0	rpoB_G_1545_A	3.38E-07	1.56E-05
S68	25	0	rpoB_G_1545_A	4.03E-07	1.85E-05
S69	25	1	rpoB_G_1545_A	4.26E-07	1.96E-05
S70	25	0	rpoB_G_1545_A	5.30E-07	2.44E-05
S71	50	0	rpoB_G_1545_A	6.00E-07	2.75E-05
S72	50	2631	rpoB_G_1545_A	0.0015788 5	0.07253659
S73	50	0	rpoB_G_1545_A	6.99E-07	3.22E-05
S74	100	1	rpoB_G_1545_A	7.82E-07	3.58E-05
S75	100	0	rpoB_G_1545_A	4.63E-07	2.13E-05
S76	100	0	rpoB_G_1545_A	7.10E-07	3.28E-05
S65	12. 5	2	rpoB_G_1545_T	9.06E-07	4.51E-05
S66	12. 5	5663	rpoB_G_1545_T	0.0014587 5	0.07367214

S67	12. 5	6	rpoB_G_1545_T	2.03E-06	0.00010244
S68	25	0	rpoB_G_1545_T	4.03E-07	2.00E-05
S69	25	0	rpoB_G_1545_T	4.26E-07	2.15E-05
S70	25	0	rpoB_G_1545_T	5.30E-07	2.68E-05
S71	50	0	rpoB_G_1545_T	6.00E-07	2.98E-05
S72	50	0	rpoB_G_1545_T	6.00E-07	3.03E-05
S73	50	0	rpoB_G_1545_T	6.99E-07	3.53E-05
S74	100	1	rpoB_G_1545_T	7.82E-07	3.89E-05
S75	100	0	rpoB_G_1545_T	4.63E-07	2.34E-05
S76	100	1	rpoB_G_1545_T	7.10E-07	3.59E-05
S65	12. 5	600970	rpoB_G_1546_T	0.2722363 9	20.5576473
S66	12. 5	932750	rpoB_G_1546_T	0.2402696 3	17.6607245
S67	12. 5	794721	rpoB_G_1546_T	0.2684287 9	19.0230411
S68	25	721553	rpoB_G_1546_T	0.2906465 4	21.9478706
S69	25	655159	rpoB_G_1546_T	0.2789348	20.5027609
S70	25	528019	rpoB_G_1546_T	0.2797444 2	19.824959
S71	50	477431	rpoB_G_1546_T	0.2863721 2	21.6250918

S72	50	371455	rpoB_G_1546_T	0.2229088	16.3846381
S73	50	400180	rpoB_G_1546_T	0.2796742 4	19.8199853
S74	100	310620	rpoB_G_1546_T	0.2427560 8	18.3314724
S75	100	530780	rpoB_G_1546_T	0.2455684 5	18.0502081
S76	100	376402	rpoB_G_1546_T	0.2672434 7	18.9390397
S65	12. 5	1424	rpoB_KO	0.0006450 6	0.0359368
S66	12. 5	0	rpoB_KO	2.58E-07	1.46E-05
S67	12. 5	1	rpoB_KO	3.38E-07	1.87E-05
S68	25	3	rpoB_KO	1.21E-06	6.73E-05
S69	25	1	rpoB_KO	4.26E-07	2.41E-05
S70	25	2	rpoB_KO	1.06E-06	5.87E-05
S71	50	0	rpoB_KO	6.00E-07	3.34E-05
S72	50	0	rpoB_KO	6.00E-07	3.40E-05
S73	50	0	rpoB_KO	6.99E-07	3.87E-05
S74	100	0	rpoB_KO	7.82E-07	4.35E-05
S75	100	0	rpoB_KO	4.63E-07	2.62E-05
S76	100	0	rpoB_KO	7.10E-07	3.94E-05



S65	12. 5	0	rpsL_KO	4.53E-07	3.82E-05
S66	12. 5	0	rpsL_KO	2.58E-07	2.28E-05
S67	12. 5	0	rpsL_KO	3.38E-07	3.14E-05
S68	25	0	rpsL_KO	4.03E-07	3.39E-05
S69	25	0	rpsL_KO	4.26E-07	3.76E-05
S70	25	0	rpsL_KO	5.30E-07	4.93E-05
S71	50	0	rpsL_KO	6.00E-07	5.05E-05
S72	50	0	rpsL_KO	6.00E-07	5.30E-05
S73	50	0	rpsL_KO	6.99E-07	6.50E-05
S74	100	0	rpsL_KO	7.82E-07	6.58E-05
S75	100	0	rpsL_KO	4.63E-07	4.09E-05
S76	100	0	rpsL_KO	7.10E-07	6.60E-05

**Supplemental table S4:** summary of aberrantly low and high genomic coverage obtained in the RLR library, resulting from alignment artifacts.

window	mean_window_count	notes
224500	228.046	ribosomal RNA
228500	235.416	ribosomal RNA
274000	306.606	mobile element IS5A
283500	228.31	adjacent to mobile elements IS1B, IS1C, IS30
381500	250.26	mobile element IS2A
574500	284.334	mobile element IS5B
688000	306.722	mobile element IS5D
1395000	296.984	mobile element IS5F
1466500	244.556	adjacent to mobile elements IS2B, IS30C
1649500	243.912	adjacent to mobile element IS2E
2067500	233.486	adjacent to mobile element IS5H
2100500	297.728	adjacent to mobile element IS5I
2287500	235.154	adjacent to mobile element IS5K
2288000	275.48	adjacent to mobile element IS5K
2725000	225.774	adjacent to ribosomal RNA

2726000	233.762	ribosomal RNA
2726500	245.828	ribosomal RNA
2995000	243.252	adjacent to mobile element IS2H
3129000	297.266	adjacent to mobile element IS5LO
3185000	233.872	adjacent to mobile element IS2I
3364500	281.456	adjacent to mobile element IS5R
3424000	256.484	ribosomal RNA
3426000	226.362	ribosomal RNA
3651000	286.762	adjacent to mobile element IS5T
3943000	246.63	ribosomal RNA
4034500	230.372	ribosomal RNA
4035000	228.734	ribosomal RNA
4165500	240.72	ribosomal RNA
4166000	230.952	ribosomal RNA
4168000	246.918	ribosomal RNA
4169500	235.474	ribosomal RNA
4207000	231.986	ribosomal RNA
4207500	230.008	ribosomal RNA

4209500 246.766 ribosomal RNA

window	mean_window _count	notes
67500	0	DE(araD-araB)567, this region absent in BW25113( $\Delta$ lacA) parent strain evolved in Baym et al. 2016
68000	0	
68500	0	
69000	0	
69500	0	
70000	0	
361000	0	lacZ4787(del)(::rrnB-3) mutation and $\Delta$ lacA alterations in BW25113(lacA) parent strain. No lacZ,Y,A
364500	0	
365000	0	
365500	0	
3581000	0	
4093500	0	DE(rhaD-rhaB)568 mutation in BW25113( $\Delta$ lacA) parent strain evolved in Baym et al. 2016
4094000	0	
4094500	0	
4095000	0	
4095000	0	

4640500 0 yjjY and yjtD loci, end of reference genome sequence. Decreased coverage likely an artifact of BWA-MEM alignment of reads from a circular genome to a linear reference sequence.

4641000 0

4641500 0

4642000 0