

Supporting Information

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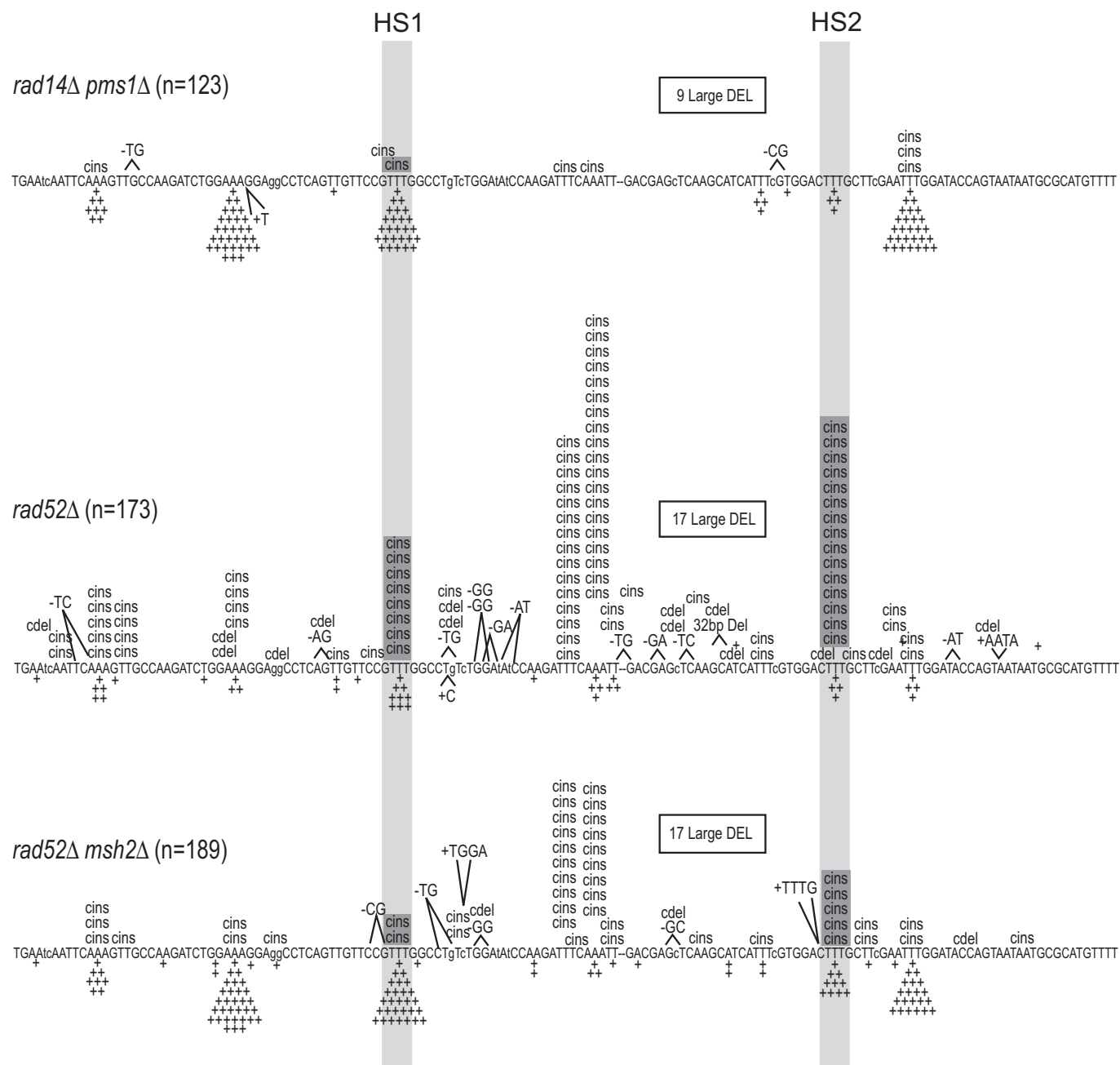


Fig. S1. *lvs2ΔA746-NR* reversion spectra in repair-defective backgrounds. The sequence of the ≈ 150 -bp reversion window is shown. The site of the 1-nt deletion that defines the allele is indicated by a dash and additional changes from the WT sequence are in lowercase. Simple insertions are indicated by “+” below the sequence and other mutation types are indicated above the sequence. The positions of HS1 and HS2 are indicated by light gray shading; complex insertions (“cins”) at these positions are shaded dark gray. “Large DEL” refers to 95-nt deletions with endpoints in 10-nt direct repeats; “cdel” corresponds to -2 events associated with nearby base substitutions. The number (“n”) of independent revertants sequenced from each background is indicated.

Table S1. Yeast strains

Strain	Relevant repair genotype	Construction
SJR1472	<i>msh2Δ::hisG-URA3-hisG</i>	OST of SJR1467 with AatII/XbaI–digested GC1914 (1)
SJR2645	<i>rad14Δ::kan</i>	OST of SJR1467 with PCR fragment amplified from pFA6a–kanMX6 (2)
SJR2672	<i>rad14Δ::kan msh2Δ::hisG-URA3-hisG</i>	OST of SJR2645 with AatII/XbaI–digested GC1914 (1)
SJR2733	<i>rad52Δ::kan</i>	OST of SJR1467 with PCR fragment amplified from pFA6a–kanMX6 (2)
SJR2734	<i>msh2Δ::hisG-URA3-hisG rad52Δ::kan</i>	OST of SJR1472 with PCR fragment amplified from pFA6a–kanMX6 (2)
SJR2773	<i>rad14Δ::kan pms1Δ::hyg</i>	OST of SJR2645 with PCR fragment amplified from hphMX4 (3)
SJR2778	<i>rad14Δ::kan rad52Δ::hyg</i>	OST of SJR2645 with PCR fragment amplified from hphMX4 (3)
SJR2779	<i>rad14Δ::kan msh2Δ::hisG-URA3-hisG rad52Δ::hyg</i>	OST of SJR2672 with PCR fragment amplified from hphMX4 (3)
SJR2808	<i>rad14Δ::kan pms1-G128A,I854M</i>	Sequential TST of SJR2645 with pSR760 [<i>pms1-I854M</i> allele (4)] and pYI–pms1–128 [<i>pms1-G128A</i> allele (5)]

OST, 1-step allele transplacement; TST, 2-step allele transplacement. All strains were derived from SJR1467 (*MAT α ade2-101_{oc} his3 Δ 200 ura3 Δ Nco lys2 Δ A746-NR*) by transformation.

- Greene CN, Jinks-Robertson S (1997) Frameshift intermediates in homopolymer runs are removed efficiently by yeast mismatch repair proteins. *Mol Cell Biol* 17:2844–2850.
- Longtine MS, et al. (1998) Additional modules for versatile and economical PCR–based gene deletion and modification in *Saccharomyces cerevisiae*. *Yeast* 14:953–961.
- Goldstein AL, McCusker JH (1999) Three new dominant drug resistance cassettes for gene disruption in *Saccharomyces cerevisiae*. *Yeast* 15:1541–1553.
- Welz-Voegele C, et al. (2002) Alleles of the yeast *PMS1* mismatch-repair gene that differentially affect recombination- and replication-related processes. *Genetics* 162:1131–1145.
- Tran PT, Liskay M (2000) Functional studies on the candidate ATPase domains of *Saccharomyces cerevisiae* MutL α . *Mol Cell Biol* 20:6390–6398.

Table S2. Distributions of hotspot 1 complex events in different strain backgrounds

Hotspot 1 position		Number of events													Total			
C	C	T	T	T	G	G	C	WT	<i>msh2Δ</i>	<i>rad14Δ</i>	<i>rad14Δ msh2Δ</i>	<i>rad14Δ rad52Δ</i>	<i>rad14Δ rad52Δ msh2Δ</i>	<i>rad14Δ pms1Δ</i>	<i>rad14Δ pms1Δ msh2Δ</i>	<i>rad52Δ</i>	<i>rad52Δ msh2Δ</i>	Total
-	-	+	-	-	-	T	-	2	11	3	3	1	1	1	1	1	1	21
-	-	+	-	-	-	T	-	1	8	2	1	1	1	1	1	1	1	14
-	-	+	-	-	-	-	1							1		2	1	6
-	-	+	-	-	-	T	-			2	1	1	1	1	1		1	5
-	-	+	-	-	-	-	A		3	1								4
-	-	+	-	-	-	T	-	1	1	1	1	1	1	1	1	1	1	3
T	-	+	-	-	-	-	-			2	1							2
-	-	+	-	-	-	A	-			2								2
-	-	+	-	-	-	T	T			2								3
-	-	+	-	-	-	-	C		1	1				1				1
T	G	+	-	-	-	-	-	1		1								1
-	-	+	-	-	-	-	-			1						1		1
*	-	+	-	-	-	-	-			1								1
-	-	+	-	-	-	-	-											1
-	-	+	-	-	-	-	-											1
-	A	+	-	-	-	-	-											1
-	-	+	-	-	-	-	-											1
-	-	+	-	-	-	-	A	T										1
T	-	+	-	-	-	-	-	G						1				1
-	-	+	-	-	-	T	A	-								1		1
-	-	+	-	-	-	T	-	G								1		1
Total								3	24	0	0	15	11	1	6	8	2	70

The 3T run where the selected frameshift occurs is italicized and bold; the G where base substitutions most frequently occur is bold and underlined. A base substitution outside of the core hotspot sequence is indicated by an asterisk. *pms1-GA,IM=pms1-GT28A,1854M*.

Table S3. Distributions of hotspot 2 complex events in different strain backgrounds

Hotspot 2 position		Number of events												Total											
G	A	C	T	T	T	G	C	T	WT	msh2Δ	rad14Δ	rad52Δ	rad14Δ	rad52Δ	rad14Δ	rad52Δ	msh2Δ	rad14Δ	pms1Δ	rad14Δ	pms1-GA,IM	rad52Δ	msh2Δ	Total	
-	-	-	-	+	T	-	-	-	1		7		11		5				1		9		3		37
-	-	T	-	+	T	-	-	-	2		2		3		1				1		1				7
-	-	-	-	+	-	T	-	-			2		1		1				1		1				4
-	-	-	-	+	A	-	-	-	2		2												1		3
-	-	G	T	+	A	-	-	-	1		1								1						2
-	-	-	-	+	C	-	-	-					1								1				2
-	-	-	T	+	-	-	-	-							1								1		2
-	-	A	-	+	T	-	-	-	1		1										1				2
*	-	A	+	+	T	-	-	-																	1
*	-	A	+	+	-	-	-	-					1												1
-	-	G	T	+	-	-	-	-																	1
-	-	-	-	+	T	A	-	-					1												1
*	-	G	T	+	A	A	-	-					1												1
*	-	-	A	+	T	-	C	-					1												1
-	-	G	-	+	T	-	-	-					1												1
-	-	-	A	+	+	T	-	-					1												1
-	-	-	-	+	+	T	T	-							1										1
-	-	T	T	+	-	-	-	-							1										1
-	-	G	T	+	T	-	-	-							1							1			1
-	-	-	-	+	T	-	C	-																	1
Total									1	0	13		24		10				0	3		15		5	72

The 3T run where the selected frameshift occurs is italicized and bold; the G where base substitutions most frequently occur is bold and underlined. A base substitution outside the core hotspot sequence is indicated by an asterisk.