

Figure S1. Confirmation of increased *RNH1* expression upon induction of the *MET25* promoter. qRT-PCR of *RNH1* transcript levels in *isw1* Δ strains with *pMET25-RNH1* in non-inducing (+ Methionine) and inducing (- Methionine) conditions. *RNH1* transcript level is normalized to the level of *ACT1* transcript and is presented as percent of *ACT1* transcript level. qRT-PCR confirmation of increased *RNH1* expression is from one experiment.

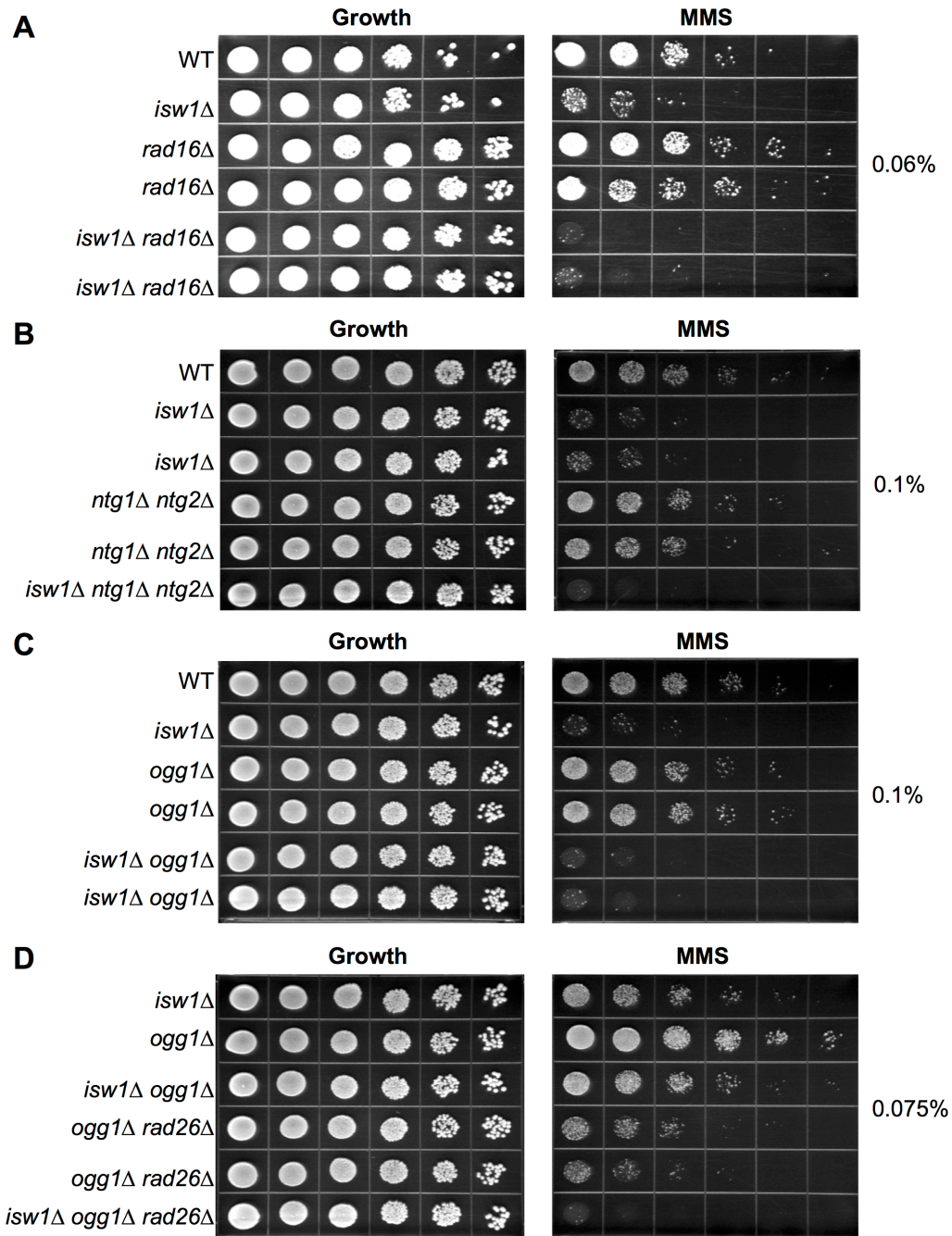


Figure S2. NER and BER gene mutations exhibit a synergistic increase in MMS sensitivity when combined with *isw1Δ*. The indicated strains were plated on growth control plates (YC-Leu-Ura) or MMS plates to test MMS sensitivity of WT and *isw1Δ* strains combined with (A) deletion of the NER gene *RAD16* on plates containing 0.06% MMS, (B) deletion of the BER genes *NTG1* and *NTG2* on plates containing 0.1% MMS, (C) deletion of the BER gene *OGG1* on plates containing 0.1% MMS, and (D) combined deletion of the BER gene *OGG1* and the TCR gene *RAD26* on plates containing 0.075% MMS. Plates were incubated at 30°C for 3 days prior to capturing images.

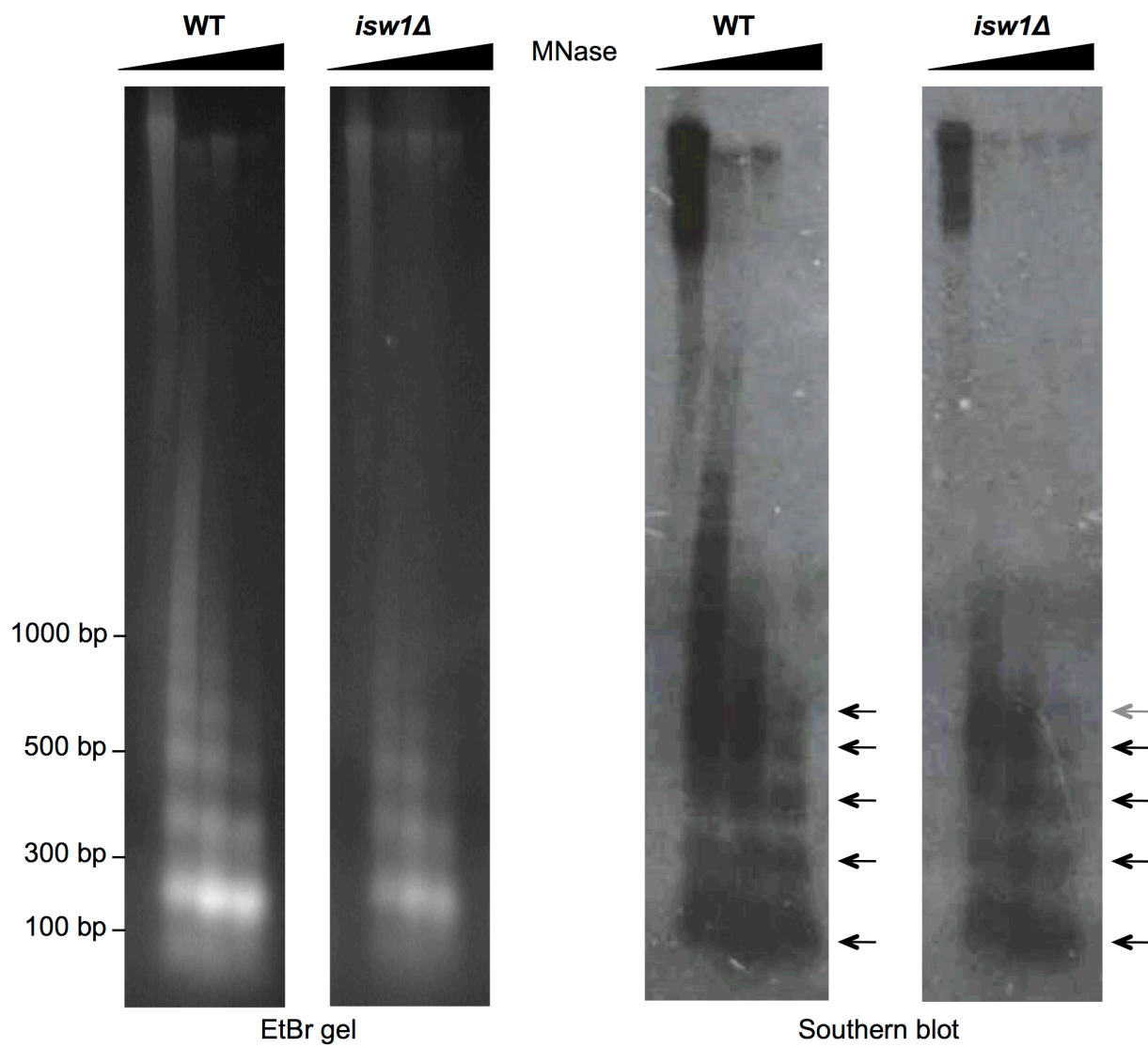


Figure S3. Absence of *Isw1* alters the nucleosome array at the CAG-85 repeat on the *URA3*-YAC. MNase assay of WT and *isw1* Δ strains containing the CAG-85 *URA3*-YAC. The wedge indicates increasing MNase level (0-7.5 U). MNase digested DNA (20-30 μ g) was run on 1.5% agarose with EtBr (left panels) and Southern blotted using a probe ~100bp upstream of the CAG repeat (right panels).

Table S1. Yeast strains used in this study^a

Strain	Genotype	YAC	Reference
VPS105	<i>MATα ade2 ade3 can1 leu2-3,112 ura3Δ0 trp1Δ lys2-801</i>	none	Schultz and Zakian 1994
BY4705	<i>MATα ade2Δ::hisG his3Δ200 leu2Δ0 lys2Δ0 met15Δ0 trp1Δ63 ura3Δ0</i>	none	Brachmann et al. 1998
BY4742	<i>MATα his3Δ1 leu2Δ0 lys2Δ0 ura3Δ0</i>	none	Brachmann et al. 1998
CFY809, 810	BY4705	CAG-85 <i>LEU2 URA3</i>	Yang and Freudenreich 2007
CFY967, 968	BY4742, <i>isw1Δ::kanMX4</i>	CAG-85 <i>LEU2 URA3</i>	
CFY1359, 1360	BY4742, <i>cbf1Δ::kanMX6</i>	CAG-85 <i>LEU2 URA3</i>	
CFY1683	BY4705, <i>isw1Δ::kanMX4</i>	CAG-85 <i>LEU2 URA3</i>	
CFY2001, 2002	BY4742, <i>isw1Δ::kanMX4 rad52Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>	
CFY2050	BY4705, <i>esa1-1851-kanMX6</i>	CAG-85 <i>LEU2 URA3</i>	House et al. 2014
CFY2097, 2098	BY4705, <i>isw2Δ::kanMX4</i>	CAG-85 <i>LEU2 URA3</i>	
CFY2101, 2102	BY4705, <i>apn1Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>	
CFY2103, 2104	BY4742, <i>apn1Δ::HIS3MX6 isw1Δ::kanMX4</i>	CAG-85 <i>LEU2 URA3</i>	
CFY2105, 2106	BY4705, <i>rad14Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>	
CFY2107, 2108	BY4742, <i>isw1Δ::kanMX4 rad14Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>	
CFY2181, 2182	BY4705, <i>ioc4Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>	
CFY2190, 2191	BY4705, <i>ioc3Δ::TRP1 ioc4Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>	
CFY2205, 2206	BY4705, <i>ioc2Δ::kanMX4</i>	CAG-85 <i>LEU2 URA3</i>	
CFY2207, 2208	BY4705, <i>ioc3Δ::TRP1</i>	CAG-85 <i>LEU2 URA3</i>	
CFY2209, 2210	BY4705, <i>ioc2Δ::kanMX4 ioc4Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>	

CFY2211, 2212	BY4705, <i>ioc2Δ::kanMX4 ioc3Δ::TRP1 ioc4Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2213, 2214	BY4705, <i>rad26Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2216	BY4742, <i>isw1Δ::kanMX4 rad26Δ::HISMX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2319, 2320	BY4705, <i>rad16Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2321, 2322	BY4742, <i>isw1Δ::kanMX4 rad16Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2323, 2324	BY4705, <i>ogg1Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2325, 2326	BY4742, <i>isw1Δ::kanMX4 ogg1Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2335, 2336	BY4705, <i>set1Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2339, 2340	BY4705, <i>sas3Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2341, 2342	BY4742, <i>isw1Δ::kanMX4 sas3Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2345, 2346	BY4705, <i>esa1-1851-kanMX6 isw1Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2423, 2424	BY4705, <i>apn1Δ::HIS3MX6 rad26Δ::HPH</i>	CAG-85 <i>LEU2 URA3</i>
CFY2425, 2426	BY4742, <i>apn1Δ::HIS3MX6 isw1Δ::kanMX4 rad26Δ::HPH</i>	CAG-85 <i>LEU2 URA3</i>
CFY2427, 2428	BY4705, <i>ogg1Δ::HIS3MX6 rad26Δ::HPH</i>	CAG-85 <i>LEU2 URA3</i>
CFY2429	BY4742, <i>isw1Δ::kanMX4 ogg1Δ::HIS3MX6 rad26Δ::HPH</i>	CAG-85 <i>LEU2 URA3</i>
CFY2441	BY4705, <i>rad52Δ::kanMX4</i>	CAG-85 <i>LEU2 URA3</i>
CFY2447, 2448	BY4705, <i>ntg1Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2449, 2450	BY4742, <i>isw1Δ::kanMX4 ntg1Δ::HIS3MX6</i>	CAG-85 <i>LEU2 URA3</i>
CFY2451, 2452	BY4705, <i>ntg2Δ::HPH</i>	CAG-85 <i>LEU2 URA3</i>
CFY2453, 2454	BY4742, <i>isw1Δ::kanMX4 ntg2Δ::HPH</i>	CAG-85 <i>LEU2 URA3</i>
CFY2455, 2456	BY4705, <i>ntg1Δ::HIS3MX6 ntg2Δ::HPH</i>	CAG-85 <i>LEU2 URA3</i>
CFY2457	BY4742, <i>isw1Δ::kanMX4 ntg1Δ::HIS3MX6 ntg2Δ::HPH</i>	CAG-85 <i>LEU2 URA3</i>

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CFY3012	BY4705	pGAL1 CAG-100 LEU2 URA3	
CFY3013	BY4705, <i>isw1Δ::kanMX4</i>	pGAL1 CAG-100 LEU2 URA3	
CFY3273, 3274	VPS105	CAG-85 LEU2 ADE2-URA3	
CFY3331, 3332	VPS105, <i>isw1Δ::kanMX4</i>	CAG-85 LEU2 ADE2-URA3	
CFY3399, 3411	BY4742, <i>isw1Δ::kanMX4 MET17 pMET25-RNH1-natMX</i>	CAG-85 LEU2 URA3	
CFY3580	BY4705	CAG-70 LEU2 URA3 2T	Su and Freudenreich 2017
CFY3694, 3695	BY4705, <i>set2Δ::HIS3MX6</i>	CAG-85 LEU2 URA3	
CFY3817, 3818	BY4705, <i>isw1Δ::kanMX4</i>	CAG-70 LEU2 URA3 2T	
CFY4035, 4036	BY4705, <i>chd1Δ::HPH</i>	CAG-85 LEU2 URA3	
CFY4037, 4038	BY4742, <i>chd1Δ::HPH isw1Δ::kanMX4</i>	CAG-85 LEU2 URA3	

^aUnless otherwise noted, strains were constructed during the course of this study or are part of the standard lab collection.

Table S2. CAG repeat stability assay data

WT, CAG-85 URA3-YAC	Total Rxns	Contractions				Expansions				
		#	%	Fold/WT	p-value ^b to WT	#	%	Fold/WT	p-value ^b to WT	p-value ^b to <i>isw1Δ</i>
Wild-type	299	33	11.0	--	--	4	1.3	--	--	--
Chromatin remodeling										
<i>isw1Δ</i>	307	42	13.7	1.2	0.33	25	8.1	6.1	7.0 x10⁻⁵	--
<i>isw2Δ</i>	158	24	15.2	1.4	0.23	3	1.9	1.4	0.07	--
<i>chd1Δ</i>	104	11	10.6	0.96	1	7	6.7	5.0	8.0 x10⁻³	--
<i>isw1Δ chd1Δ</i>	104	14	13.5	1.2	0.48	5	4.8	3.6	0.053	0.38
Isw1 complexes										
<i>ioc3Δ</i>	156	15	9.6	0.87	0.75	6	3.9	2.9	0.099	--
<i>ioc2Δ</i>	156	23	14.7	1.3	0.29	6	3.9	2.9	0.099	--
<i>ioc4Δ</i>	156	19	12.2	1.1	0.76	8	5.1	3.8	0.027	--
<i>ioc2Δ ioc4Δ</i>	156	18	11.5	1.1	0.88	8	5.1	3.8	0.027	--
<i>ioc3Δ ioc4Δ</i>	156	16	10.3	0.93	0.87	7	4.5	3.4	0.052	--
<i>ioc2Δ ioc3Δ ioc4Δ</i>	156	13	8.3	0.76	0.42	5	3.2	2.4	0.29	--
CBF1 and histone methyltransferases (HMTs)										
<i>set1Δ</i>	149	19	12.8	1.2	0.16	1	0.67	0.50	1	--
<i>set2Δ</i>	180	30	16.7	1.5	0.094	5	2.8	2.1	0.31	--
<i>cbf1Δ</i>	104	5	4.8	0.44	0.08	4	3.8	2.9	0.2	--
Histone acetyltransferases (HATs)										
<i>sas3Δ</i>	154	23	14.9	1.4	0.23	4	2.6	1.9	0.45	--
<i>esa1-1851</i>	125	25	20.0	1.8	0.054	10	8.0	6.0	5.0 x10⁻³	--
<i>isw1Δ sas3Δ</i>	156	19	12.2	1.1	0.76	12	7.7	5.8	8.7 x10⁻⁴	1
<i>isw1Δ esa1-1851</i>	104	28	26.9	2.4	2.2 x 10⁻⁴	13	12.5	9.3	1.1 x10⁻⁵	0.24
CAG-85 ADE2-URA3-YAC										
WT	148	27	18.2	--	--	3	2.0	--	--	--
<i>isw1Δ</i>	200	34	17.0	0.93	0.78	3	1.5	0.74	0.70	--

TT CAG-70 TT URA3-YAC (2T-YAC)

WT	326	17	5.2	--	--	3	0.92	--	--	--
<i>isw1Δ</i>	88	4	4.6	0.87	1	2	2.3	2.5	0.29	--

pGAL1 CAG-100 URA3-YAC

WT, glucose	200	16	8.0	--	--	2	1.0	--	--	--
<i>isw1Δ</i> , glucose	199	14	7.0	0.88	0.85	4	2.0	2.0	0.45	--
WT, galactose	200	8	4.0	--	--	4	2.0	--	--	--
<i>isw1Δ</i> , galactose	200	14	7.0	1.8	0.27	9	4.5	2.3	0.26	--

RNase H overexpression

<i>isw1Δ</i> , RNase H not induced	103	8	7.8	0.71	0.45	7	6.8	5.2	7.9 x10⁻³	--
<i>isw1Δ</i> , RNase H induced	104	10	9.6	0.87	0.85	8	7.7	5.9	3.0 x10⁻³	--

HR pathway

<i>rad52Δ</i>	199	45	22.6	2.1	6.4 x 10⁻⁴	4	2.0	1.5	1	--
<i>isw1Δ rad52Δ</i>	256	24	9.4	0.85	0.58	20	7.8	5.8	2.1 x10⁻⁴	1

NER pathway

<i>rad14Δ</i>	204	25	12.3	1.1	0.67	11	5.4	4.0	0.014	--
<i>rad26Δ</i>	143	20	14.0	1.3	0.43	2	1.4	1.1	1	--
<i>rad16Δ</i>	156	23	14.7	1.3	0.29	9	5.8	4.3	0.014	--
<i>isw1Δ rad14Δ</i>	208	15	7.2	0.65	0.17	5	2.4	1.8	0.50	6.6 x10⁻³
<i>isw1Δ rad26Δ</i>	123	12	9.8	0.88	0.86	4	3.3	2.4	0.24	0.087
<i>isw1Δ rad16Δ</i>	156	22	14.1	1.3	0.37	11	7.1	5.3	3.6 x10⁻³	0.86

BER pathway

<i>apn1Δ</i>	208	16	7.7	0.70	0.23	1	0.48	0.36	0.65	--
<i>ogg1Δ</i>	156	23	14.7	1.3	0.29	4	2.6	1.9	0.46	--
<i>ntg1Δ</i>	208	22	10.6	0.96	1	4	1.9	1.4	0.72	--
<i>ntg2Δ</i>	208	20	9.6	0.87	0.66	6	2.9	2.2	0.33	--
<i>ntg1Δ ntg2Δ</i>	207	26	12.7	1.1	0.67	2	0.97	0.72	1	--

<i>isw1Δ apn1Δ</i>	208	15	7.2	0.65	0.17	3	1.4	1.1	1	6.1 x10⁻⁴
<i>isw1Δ ogg1Δ</i>	156	13	8.3	0.76	0.42	3	1.9	1.4	0.70	6.8 x10⁻³
<i>isw1Δ ntg1Δ</i>	208	23	11.1	1.0	1	4	1.9	1.4	0.72	2.8 x10⁻³
<i>isw1Δ ntg2Δ</i>	208	23	11.1	1.0	1	8	3.9	2.9	0.08	0.066
<i>isw1Δ ntg1Δ ntg2Δ</i>	208	23	11.1	1.0	1	10	4.8	3.6	0.026	0.16
NER and BER pathway										
<i>apn1Δ rad26Δ</i>	102	17	16.7	1.5	0.16	2	2.0	1.5	0.65	--
<i>isw1Δ apn1Δ rad26Δ</i>	96	18	18.8	1.7	0.057	2	2.1	1.6	0.64	0.037

^bp-values in bold font show significant differences of $p \leq 0.05$.

Supporting Information References

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