

1st experiment		NON-SPECIFIC (NS) BAND	HISTONE SIGNAL (HS)	HS/NS	relative to rad53-AID control
<i>rad53-AID</i>	-	8.466.648	26.932.877	3,18	1,00
	NAA	10.735.648	29.300.635	2,73	0,86
<i>rad53-AID lsm1Δ</i>	-	9.438.234	21.198.271	2,25	0,71
	NAA	9.051.527	34.409.999	3,80	1,20

2nd experiment		NON-SPECIFIC (NS) BAND	HISTONE SIGNAL (HS)	HS/NS	relative to rad53-AID control
<i>rad53-AID</i>	-	9.246.062	16.259.371	1,76	1,00
	NAA	9.897.355	17.414.421	1,76	1,00
<i>rad53-AID lsm1Δ</i>	-	10.742.355	14.898.250	1,39	0,79
	NAA	9.068.770	22.048.735	2,43	1,38

t-test (paired, 2 tails)					
		<i>rad53-AID</i>		<i>rad53-AID lsm1Δ</i>	
		-	NAA	-	NAA
<i>rad53-AID</i>	-	-	0.501	0.259	0.026
	NAA	-	-	0.082	0.144
<i>rad53-AID lsm1Δ</i>	-	-	-	-	0.124
	NAA	-	-	-	-

**Source data Figure 3a.** CBR staining for each of the two experiments in which core histones have been purified from *rad53-AID* and *rad53-AID lsm1Δ* cells treated or not with NAA and synchronized with Nocodazole. Quantification and statistics are shown below.

	% of diploid cells			
	replicate 1		replicate 2	
	DMSO	NAA	DMSO	NAA
haploid	100	90	100	70
diploid	0	10	0	5
<i>rad53-AID lsm1Δ</i>	0	10	0	6,66
n	100	100	75	75

t-test (2 tails, unequal variance) | 0,038

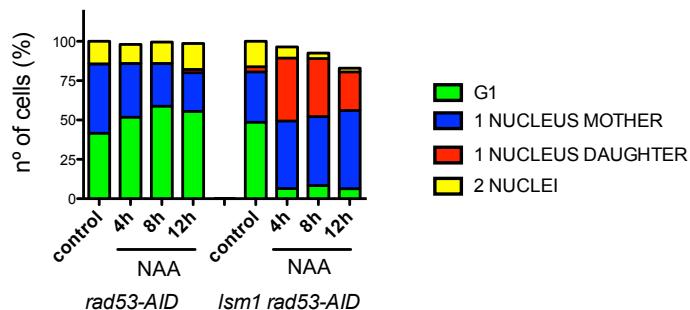
**Source data Figure 3g.** Number of diploids observed by FACS in *rad53-AID lsm1Δ* cells after a 4 hour treatment with DMSO or NAA. This experiment was performed twice. In the first experiment, we counted 100 colonies for each condition. In the second one, we counted 75.

1st experiment		number of colonies		
		expected	counted	survival (NAA/untreated) %
rad53-AID untreated		100	108	80,6
		200	191	91,1
rad53-AID+NAA		100	87	
		200	174	
rad53-AID <i>Ism1Δ</i> untreated		100	68	2,9
		200	101	3,0
rad53-AID <i>Ism1Δ</i> +NAA		100	2	
		200	3	
<b>t-test (paired, two tails)</b>				0,0006
2nd experiment		number of colonies		
		expected	counted	survival (NAA/untreated) %
rad53-AID untreated		100	77	124,7
		200	124	106,5
rad53-AID+NAA		100	96	
		200	132	
rad53-AID <i>Ism1Δ</i> untreated		100	40	17,5
		200	72	12,5
rad53-AID <i>Ism1Δ</i> +NAA		100	7	
		200	9	

**Source data Figure 3-Figure Supplement 3d.** Relative colony formation assay data used to determine the number of viable cells after a four-hour treatment with NAA in *rad53-AID* and *rad53-AID* *Ism1Δ* mutants. % of viable cells is estimated dividing the number of viable cells obtained between in treated extracts divided by the number obtained in non-treated samples.

	biological replicate	RED(viable cells)	GREEN (dead cells) (%)	n	Nº of dead cells (%)		
					AVERAGE	SD	SEM
wild type	vector	1	99,0	306	wild type	vector	1,6
		2	99,2	257			1,2
		3	95,7	300			0,7
	2μΔNEG	1	92,9	240	2μΔNEG	t-test (paired, two tails)	7,7
		2	89,2	204			2,8
		3	94,7	227			1,6
					0,11		

**Source data Figure 3-Figure Supplement 3e.** % of green cells (dead cells, unable to metabolise the compound) compared to red cells (able to metabolise the compound) observed in three independent experiments in wild type cells transformed with an empty vector or the 2μΔNEG vector.



rad53-AID	1st experiment (%)				2nd experiment (%)			
	AS	4H NAA	8H NAA	12H NAA	AS	4H NAA	8H NAA	12H NAA
G1	35,4	38,8	45,8	80,6	41,5	51,8	58,8	55,4
bud, 1 nucleus in mother	45,5	48,8	38,3	12,2	44,0	34,1	27,0	24,6
bud, 1 nucleus in daughter	0,0	0,0	0,0	0,0	0,0	0,0	0,0	2,2
bud, two nuclei	19,2	11,9	15,4	5,9	14,5	12,2	13,7	16,5
n	198	201	201	222	207	164	204	224

rad53-AID <i>Ism1Δ</i>	1st experiment (%)				2nd experiment (%)			
	AS	4H NAA	8H NAA	12H NAA	AS	4H NAA	8H NAA	12H NAA
G1	23,4	5,5	7,0	9,2	48,6	6,4	8,4	6,3
bud, 1 nucleus in mother	44,0	47,6	42,2	44,2	31,9	42,9	43,8	49,7
bud, 1 nucleus in daughter	19,6	46,3	40,6	35,0	3,3	40,0	36,9	24,6
bud, two nuclei	10,5	0,0	0,8	2,5	16,2	7,1	3,4	2,3
n	209	164	128	120	210	140	203	175

**Source data Figure 3-Figure Supplement 3f.** Raw data obtained in the two cell cycle kinetics performed to determine changes in the cell cycle distribution of *rad53-AID* and *rad53-AID* *Ism1Δ* cells after treatment with NAA. Cells were fixed, stained with DAPI and classified according to morphology and number of nuclei using a 100x objective. The image on the top represents the numbers obtained in the second experiment, not included in the main text.