

Figure S1. Characterization of GBM cell lines and PARP-1 inhibition by NU1025. (A) Representative protein expression of PTEN, AKT, pAKT<sup>(476)</sup> and MGMT in GBM cell lines (T98G, LN18 and U87MG). HeLa cells were used as a positive control for PTEN expression;  $\beta$ -actin was used as an endogenous control. The samples were analyzed by western blot analysis. (B) Effects of NU1025 agent in the reduction of poly-ADP-ribosylation promoted by PARP-1. Cells were treated with NU1025 (NU-100 and NU-200  $\mu$ M) at 20 min prior to H<sub>2</sub>O<sub>2</sub> treatment (20 mM for 10 min), and then analyzed by flow cytometry using PAR-antibody. \*P<0.05. Bars correspond to comparisons between treatments. GBM, glioblastoma; PARP-1, poly[ADP-ribose] polymerase 1; PTEN, phosphatase and tensin homologue deleted on chromosome ten; MGMT, O<sup>6</sup>-methylguanine DNA methyltransferase.

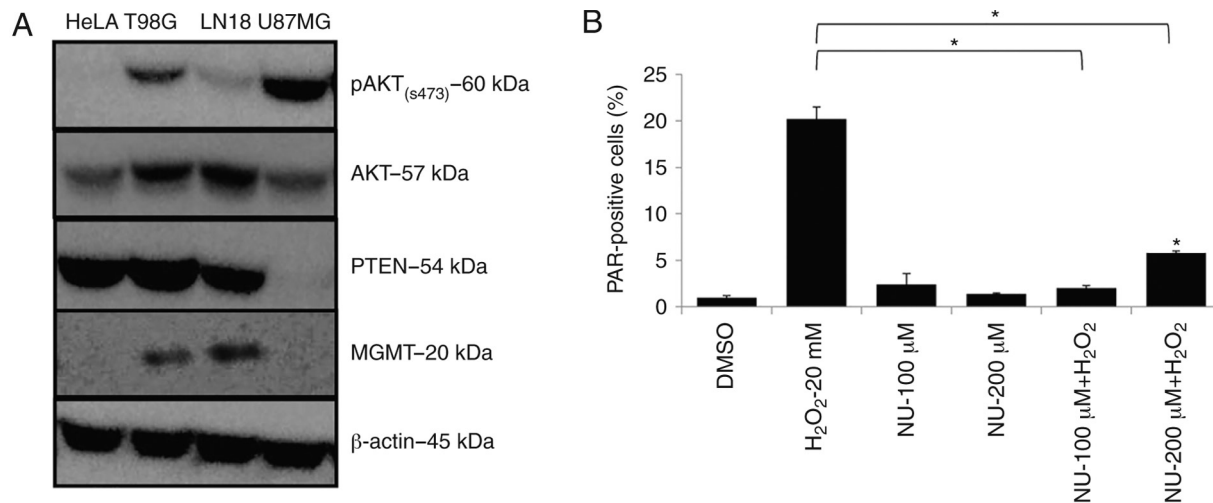


Table SI. List of genes used in expression analysis<sup>a</sup>.

Custom genes TaqMan Array 96 - Well FAST	
Genes	Identification
<i>TBP</i>	Hs00427620_m1
<i>HPRT1</i>	Hs99999909_m1
<i>MGMT</i>	Hs01037698_m1
<i>BRCA1</i>	Hs01556193_m1
<i>BRCA2</i>	Hs01037414_m1
<i>LIG4</i>	Hs00172455_m1
<i>RAD51B</i>	Hs00172522_m1
<i>RAD51C</i>	Hs00427442_m1
<i>RAD51D</i>	Hs00979544_m1
<i>RAD51</i>	Hs00153418_m1
<i>FEN1</i>	Hs00748727_s1
<i>APEX1</i>	Hs00172396_m1
<i>LIG1</i>	Hs01553527_m1
<i>PARP1</i>	Hs00242302_m1
<i>XRCC1</i>	Hs00959834_m1
<i>MSH2</i>	Hs00953523_m1
<i>MSH3</i>	Hs00989003_m1
<i>XPA</i>	Hs00166045_m1
<i>XPC</i>	Hs01104206_m1
<i>XRCC4</i>	Hs00243327_m1
<i>PRKDC</i>	Hs00179161_m1
<i>XRCC6</i>	Hs01922652_g1
<i>XRCC5</i>	Hs00221707_m1

<sup>a</sup>as evaluated by means of custom TaqMan Array 96 - Well FAST plates (Thermo Fischer Scientific, Inc.).

Table SII. Fold change values (log2) evaluated by PCR array (Thermo Fisher Scientific, Inc.) for a set of DNA repair pathway genes in T98G cells submitted to continuous treatments of TMZ (200  $\mu$ M) and NU1025 (200  $\mu$ M), tested alone or in combination.

T98G cells-Transcript expression values (fold change log2)							
Genes	DNA repair pathway	Time of treatments (6 h)			Time of treatments (24 h)		
		NU1025 (200 $\mu$ M)	TMZ (200 $\mu$ M)	NU1025 +TMZ	NU1025 (200 $\mu$ M)	TMZ (200 $\mu$ M)	NU1025 +TMZ
<i>PARP1</i>	BER	0.17	0.10	0.38	0.31	<b>1.30</b>	-0.06
<i>APEX1</i>		0.05	0.20	0.50	-0.36	0.34	-0.18
<i>FEN1</i>		-0.61	-0.83	-0.92	0.05	<b>1.43</b>	-0.53
<i>LIG1</i>		0.39	0.20	1.28	-0.11	<b>1.90</b>	<b>1.47</b>
<i>XRCC1</i>		0.37	0.16	0.17	-0.52	0.86	-0.46
<i>BRCA1</i>	HR	-0.06	1.52	0.31	1.18	<b>1.32</b>	0.31
<i>RAD51</i>		0.97	0.20	0.65	0.60	1.16	0.02
<i>RAD51B</i>		0.49	-0.20	-0.21	-0.28	<b>1.63</b>	0.88
<i>RAD51C</i>		0.21	-0.02	-0.76	0.73	<b>1.68</b>	0.89
<i>RAD51D</i>		0.52	0.27	0.57	-0.51	0.49	-0.06
<i>LIG4</i>		<b>-1.98</b>	<b>-2.13</b>	<b>-1.42</b>	0.07	0.93	-0.69
<i>PRKDC</i>	NHEJ	0.61	0.03	0.30	-0.32	<b>1.37</b>	0.37
<i>XRCC5</i>		0.35	-0.44	0.46	0.56	0.43	-0.14
<i>XRCC6</i>		0.31	-0.11	0.53	-0.27	0.02	-0.45
<i>XPA</i>	NER	0.45	0.20	0.0	0.10	0.51	-0.40
<i>XPC</i>		<b>-1.94</b>	<b>-1.60</b>	-1.28	-0.58	-0.01	-1.01
<i>XRCC4</i>		0.27	0.20	0.40	0.13	<b>1.31</b>	0.47
<i>MGMT</i>	MGMT	-0.61	<b>-1.38</b>	-1.01	-0.13	1.03	0.81
<i>MSH2</i>	MMR	0.34	0.36	0.38	0.03	1.05	0.26
<i>MSH3</i>		-0.04	-0.24	-0.09	0.13	0.51	-0.28

The expression was evaluated 6 and 24 h after the start of the treatments using QuantStudio 3 Real-Time PCR Systems and software (Thermo Fisher Scientific, Inc.). *TBP* and *HPRT1* were used as reference genes. The experiments were performed with a pool of cDNA corresponding to the proportional junction of three independent experiments applied in simplicates. Fold change values >|1.3| are represented in bold. BER, base excision repair; HR, homologous recombination; NHEJ, non-homologous end joining; NER, nucleotide excision repair; MGMT, O<sup>6</sup>-methylguanine DNA methyltransferase; MMR, mismatch repair.

Table SIII. Fold change values (log2) evaluated by PCR array (Thermo Fisher Scientific, Inc.) for a set of DNA repair pathway genes in LN18 cells submitted to continuous treatments of TMZ (200  $\mu$ M) and NU1025 (200  $\mu$ M), tested alone or in combination.

LN18 cells-Transcript expression values (fold change log2)							
Genes	DNA repair pathway	Time of treatments (6 h)			Time of treatments (24 h)		
		NU1025 (200 $\mu$ M)	TMZ (200 $\mu$ M)	NU1025 +TMZ	NU1025 (200 $\mu$ M)	TMZ (200 $\mu$ M)	NU1025 +TMZ
<i>PARP1</i>	BER	-0.16	-0.05	-0.06	-0.16	0.21	0.07
<i>APEX1</i>		0.11	0.33	0.05	0.13	0.79	0.64
<i>FEN1</i>		0.16	0.39	0.17	0.09	0.23	0.14
<i>LIG1</i>		-0.16	0.07	-0.08	-1.24	-0.49	-1.03
<i>XRCC1</i>		0.14	0.33	-0.10	-0.33	-0.11	0.22
<i>BRCA1</i>	HR	-0.31	-0.12	-0.33	<b>2.18</b>	0.78	0.52
<i>RAD51</i>		-0.51	0.01	-0.16	0.35	0.97	0.44
<i>RAD51B</i>		-0.18	-0.42	-0.40	0.11	0.10	-0.02
<i>RAD51C</i>		-0.41	0.12	-0.01	0.56	0.67	0.27
<i>RAD51D</i>		0.05	0.09	0.10	0.45	0.50	0.49
<i>LIG4</i>		-0.35	-0.42	-0.10	-0.94	0.06	-0.56
<i>PRKDC</i>	NHEJ	-0.32	-0.27	-0.03	0.25	-0.11	0.25
<i>XRCC5</i>		-0.11	0.05	-0.01	0.01	0.36	0.32
<i>XRCC6</i>		-0.02	0.09	0.18	<b>-1.81</b>	<b>-2.18</b>	<b>-2.18</b>
<i>XPA</i>	NER	-0.09	0.00	0.18	-0.39	0.15	-0.24
<i>XPC</i>		0.23	0.47	0.18	0.11	0.14	0.32
<i>XRCC4</i>		0.01	0.10	-0.04	-0.49	0.63	0.17
<i>MGMT</i>	MGMT	0.40	0.41	0.00	0.98	1.10	0.83
<i>MSH2</i>	MMR	-0.18	0.09	0.09	-0.05	0.61	0.31
<i>MSH3</i>		-0.30	-0.25	-0.20	0.97	0.54	0.71

The expression was evaluated six and 24 h after the start of the treatments using QuantStudio 3 Real-Time PCR Systems and software (Thermo Fisher Scientific). *TBP* and *HPRT1* were used as reference genes. The experiments were performed with a pool of cDNA corresponding to the proportional junction of three independent experiments applied in simplicates. Fold change values >|1.3| are represented in bold. BER, base excision repair; HR, homologous recombination; NHEJ, non-homologous end joining; NER, nucleotide excision repair; MGMT, O<sup>6</sup>-methylguanine DNA methyltransferase; MMR, mismatch repair.