

Figure S1.  $\gamma$ H2AX foci staining following the exposure of control MCF10A cells to ionizing radiation. Induction of  $\gamma$ H2AX foci was assessed 30 min following exposure to ionizing radiation (2 Gy) or after mock irradiation (0 Gy). Immunostaining was performed by means of a monoclonal anti- $\gamma$ H2AX antibody, as previously described (34). This image gallery shows 4 cells for each experimental condition taken with Metafer 4 (Metasystems).

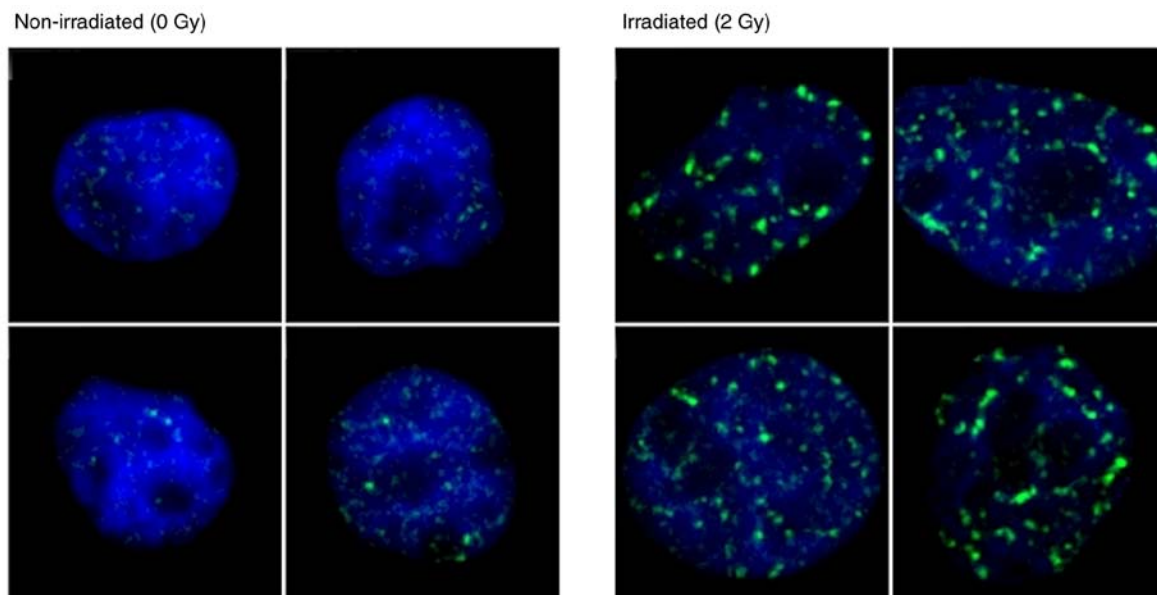


Figure S2. Evaluation of PARP inhibition by olaparib. Immunodetection for PARP activity was performed as previously described by Barazzuol *et al* (35). PARP activation was induced by H<sub>2</sub>O<sub>2</sub> treatment (20 mM, 10 min, left panel). Incubation with olaparib (5 μM) 1 h prior to and during exposure to H<sub>2</sub>O<sub>2</sub>, resulted in the absence of PARP activity (right panel). Images were obtained with Metafer 4 (Metasystems). PARP, poly(ADP-ribose) polymerase.

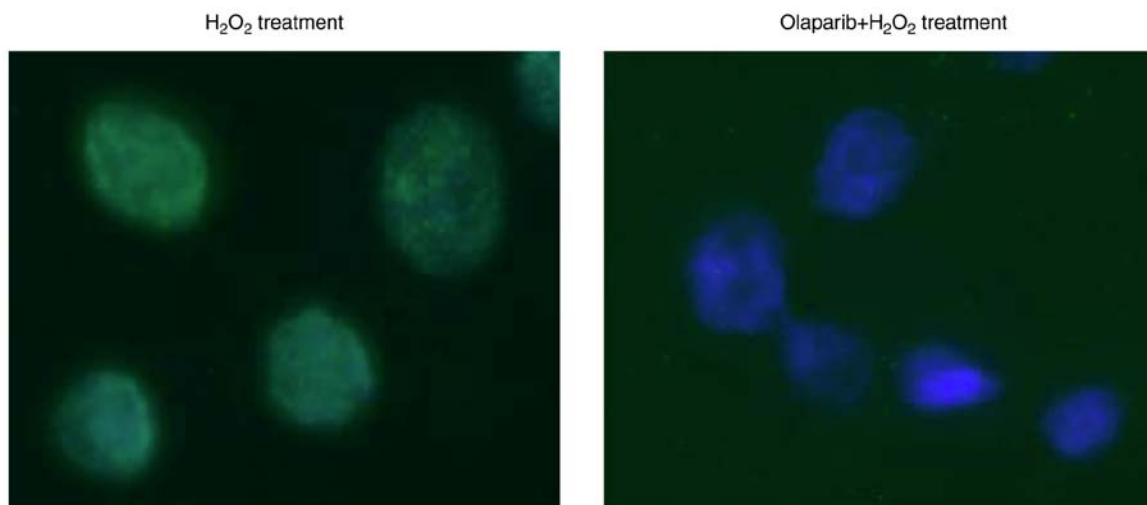


Figure S3. RT-qPCR analysis of *BRCA1* and *BRCA2* mRNA levels ( $\pm$  standard deviation) in control, BRCA1i and BRCA2i cell lines. For an easy comparison, the relative expression in relation to the control sample is shown for each knockdown cell line (n=3). The relative mRNA expression of *BRCA1*, but not that of *BRCA2*, was significantly decreased in the BRCA1i cells, and the relative mRNA expression of *BRCA2*, but not that of *BRCA1*, was significantly decreased in the BRCA2i cells. Statistical analysis was carried out using one-way ANOVA with Tukey's post-hoc test and the detailed results are shown in Table SI.

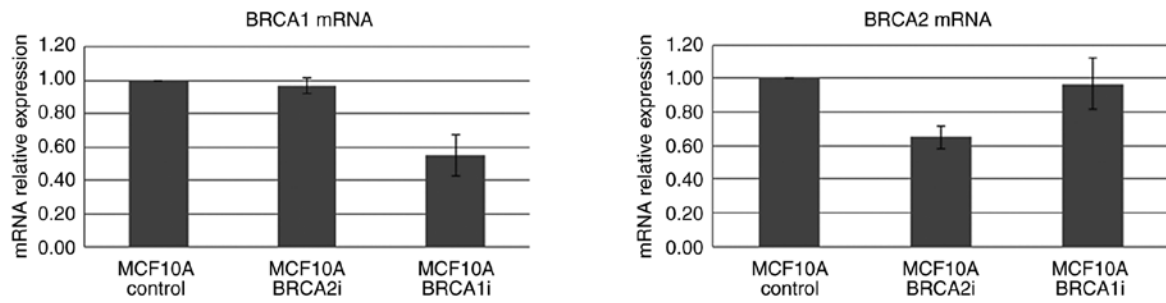


Table SI. Results of statistical analysis with one-way ANOVA and Tukey's test for post hoc significance of multiple comparisons.

Experiment	Group(s)	One-way ANOVA P-value	Tukey's post-hoc test Comparison, P-value
Mean number of RAD51 foci per cell (Fig. 4)	Control, BRCA1i, BRCA2i no radiation, no olaparib	0.12	Control vs. BRCA1, P=0.7987 Control vs. BRCA2i, P=0.1154 BRCA1i vs. BRCA2i, P=0.3339
	Control, BRCA1i, BRCA2i no radiation, yes olaparib	0.53	Control vs. BRCA1i, P=0.5058 Control vs. BRCA2i, P=0.9159 BRCA1i vs. BRCA2i, P=0.7477
	Control, BRCA1i, BRCA2i yes radiation, no olaparib	<0.00001	Control vs. BRCA1i, P<0.00001 Control vs. BRCA2i, P<0.00001 BRCA1i vs. BRCA2i, P=0.7216
	Control, BRCA1i, BRCA2i yes radiation, yes olaparib	<0.00001	Control vs. BRCA1i, P=0.0001 Control vs. BRCA2i, P<0.00001 BRCA1i vs. BRCA2i, P=0.4456
	Controls, all experimental conditions (yes/no radiation, yes/no olaparib)	<0.00001	no IR no olaparib vs. no IR yes olaparib, P=0.9777 no IR no olaparib vs. yes IR no olaparib, P<0.00001 no IR no olaparib vs. yes IR yes olaparib, P<0.00001 no IR yes olaparib vs. yes IR no olaparib, P=0.0001 no IR yes olaparib vs. yes IR yes olaparib, P<0.00001 yes IR no olaparib vs. yes IR yes olaparib, P=0.0487 no IR no olaparib vs. no IR yes olaparib, P=0.9943 no IR no olaparib vs. yes IR no olaparib, P=0.9620 no IR no olaparib vs. yes IR yes olaparib, P=0.4949 no IR yes olaparib vs. yes IR no olaparib, P=0.9954 no IR yes olaparib vs. yes IR yes olaparib, P=0.3546 yes IR no olaparib vs. yes IR yes olaparib, P=0.2472 no IR no olaparib vs. no IR yes olaparib, P=0.3271 no IR no olaparib vs. yes IR no olaparib, P=0.9994 no IR no olaparib vs. yes IR yes olaparib, P=0.5897 no IR yes olaparib vs. yes IR no olaparib, P=0.2724 no IR yes olaparib vs. yes IR yes olaparib, P=0.9666 yes IR no olaparib vs. yes IR yes olaparib, P=0.5176
	BRCA1i, all experimental conditions (yes/no radiation, yes/no olaparib)	0.24	Control vs. BRCA1i, P=0.0001 Control vs. BRCA2i, P=0.0001 BRCA1i vs. BRCA2i, P=0.4456
	BRCA2i, all experimental conditions (yes/no radiation, yes/no olaparib)	0.19	Control vs. BRCA1i, P=0.0001 Control vs. BRCA2i, P=0.0001 BRCA1i vs. BRCA2i, P=0.4456
	Dose: 0 Gy	0.015	Control vs. BRCA1i, P=0.1262 Control vs. BRCA2i, P=0.4615 BRCA1i vs. BRCA2i, P=0.0126
	Dose: 0.25 Gy	<0.00001	Control vs. BRCA1i, P<0.00001 Control vs. BRCA2i, P=0.1454 BRCA1i vs. BRCA2i, P<0.00001
	Dose: 0.5 Gy	<0.00001	Control vs. BRCA1i, P<0.00001 Control vs. BRCA2i, P<0.00001
	Radiation dose vs. mean number of micronuclei per 1000 binucleated cells (Fig. 5)		

Table SI. Continued.

Experiment	Group(s)	One-way ANOVA P-value	Tukey's post-hoc test Comparison, P-value
Radiation dose vs. fraction of surviving cells (Fig. 6)	Dose: 1 Gy	<0.00001	Control vs. BRCA2i, P=0.0002 BRCA1i vs. BRCA2i, P=0.0002 Control vs. BRCA1i, P<0.0001 Control vs. BRCA2i, P=0.0007 BRCA1i vs. BRCA2i, P<0.0001 Control vs. BRCA1i, P<0.0001 Control vs. BRCA2i, P=0.0002 BRCA1i vs. BRCA2i, P<0.0001 Control vs. BRCA1i, P<0.0001 Control vs. BRCA2i, P=0.0002 BRCA1i vs. BRCA2i, P=0.0018
	Dose: 2 Gy	<0.00001	Reference value
	Dose: 4 Gy	<0.00001	Control vs. BRCA1i, P=0.0080 Control vs. BRCA2i, P=0.0011 BRCA1i vs. BRCA2i, P<0.0001 Control vs. BRCA1i, P<0.00001 Control vs. BRCA2i, P=0.0001 BRCA1i vs. BRCA2i, P<0.00001 Control vs. BRCA1i, P<0.00001 Control vs. BRCA2i, P=0.0609 BRCA1i vs. BRCA2i, P<0.00001 Control vs. BRCA1i, P<0.00001 Control vs. BRCA2i, P=0.0146 BRCA1i vs. BRCA2i, P<0.00001 Control vs. BRCA1i, P<0.00001 Control vs. BRCA2i, P=0.0440 BRCA1i vs. BRCA2i, P=0.0007 Control vs. BRCA2i, P=0.0421 Control vs. BRCA2i, P=0.1886 Control vs. BRCA1i, P=0.0001 Control vs. BRCA2i, P=0.9172 BRCA1i vs. BRCA2i, P=0.0002 Control vs. BRCA2i, P=0.0061 Control vs. BRCA1i, P=0.9472 BRCA1i vs. BRCA2i, P=0.0114
	Dose: 0 Gy Dose: 0.5 Gy	Reference <0.00001	
	Dose: 1 Gy	<0.00001	
	Dose: 2 Gy	<0.00001	
	Dose: 3 Gy	<0.00001	
	Dose: 4 Gy	<0.00001	
	Dose: 6 Gy Dose: 8 Gy	0.0421 0.1886 <0.00001	
BRCA1 mRNA relative expression (Fig. S3)			
BRCA2 mRNA relative expression (Fig. S3)			