

Reporting Summary

Nature Portfolio wishes to improve the reproducibility of the work that we publish. This form provides structure for consistency and transparency in reporting. For further information on Nature Portfolio policies, see our [Editorial Policies](#) and the [Editorial Policy Checklist](#).

Statistics

For all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.

n/a Confirmed

- The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement
- A statement on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly
- The statistical test(s) used AND whether they are one- or two-sided
Only common tests should be described solely by name; describe more complex techniques in the Methods section.
- A description of all covariates tested
- A description of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons
- A full description of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) AND variation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)
- For null hypothesis testing, the test statistic (e.g. F , t , r) with confidence intervals, effect sizes, degrees of freedom and P value noted
Give P values as exact values whenever suitable.
- For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings
- For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes
- Estimates of effect sizes (e.g. Cohen's d , Pearson's r), indicating how they were calculated

Our web collection on [statistics for biologists](#) contains articles on many of the points above.

Software and code

Policy information about [availability of computer code](#)

Data collection Microscopy images were acquired using a Zeiss AxioImager M2 or D2 widefield fluorescence microscope and ZEN 2012 software (blue edition, version 1.1.0.0). Western blot images were acquired using a Odyssey CLx with Image studio lite software (v5.2) or for western blot with ECL a Amersham Imager 680.

Data analysis Microscopy images were analyzed in Image J (1.48v). Mass spectrometry data was analyzed using MaxQuant software (v1.6.14) and analysis output was further processed in the Perseus (v1.6.14) computational platform and further processed in Microsoft Excel 365 for comprehensive visualization. Graphs were plotted and analyzed using Graphpad Prism 8 (v8.4.2).

For manuscripts utilizing custom algorithms or software that are central to the research but not yet described in published literature, software must be made available to editors and reviewers. We strongly encourage code deposition in a community repository (e.g. GitHub). See the Nature Portfolio [guidelines for submitting code & software](#) for further information.

Data

Policy information about [availability of data](#)

All manuscripts must include a [data availability statement](#). This statement should provide the following information, where applicable:

- Accession codes, unique identifiers, or web links for publicly available datasets
- A description of any restrictions on data availability
- For clinical datasets or third party data, please ensure that the statement adheres to our [policy](#)

Mass spectrometry proteomics data are presented in main Figures 1a, b and Fig 2a, b, c, d and Fig 7a, b, and have been deposited to the ProteomeXchange Consortium via the PRIDE partner repository (<https://www.ebi.ac.uk/pride/>) with the dataset identifier PXD025226 (Perez-Riverol et al., 2019). The data is also included as Excel in the Source Data file

Field-specific reporting

Please select the one below that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.

Life sciences Behavioural & social sciences Ecological, evolutionary & environmental sciences

For a reference copy of the document with all sections, see [nature.com/documents/nr-reporting-summary-flat.pdf](https://www.nature.com/documents/nr-reporting-summary-flat.pdf)

Life sciences study design

All studies must disclose on these points even when the disclosure is negative.

| | |
|-----------------|--|
| Sample size | No statistical method was used to predetermine sample size. Sample sizes were chosen for the different experimental approaches based on the technical difficulty and throughput of the individual assays, the chosen sample sizes are consistent with previous publications. |
| Data exclusions | No data was excluded. |
| Replication | All replication attempts were successful. The number of replicate experiments are indicated in the figure legends of the manuscript and at least two replicates were performed for each individual approach. Most results were confirmed in multiple cell lines and using complementary approaches. - Mass spec findings were confirmed by reciprocal immunoprecipitation and western blot analyses - Effects of knock-out of proteins of interest were confirmed by rescue experiments - Global UV irradiation experiments were confirmed by local UV irradiation experiment |
| Randomization | There was no allocation of test subjects for any experiments, thus randomization was not applicable to our study |
| Blinding | Data analyses were performed by unbiased software programs/algorithms blinding was therefore not applicable to our study |

Reporting for specific materials, systems and methods

We require information from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, system or method listed is relevant to your study. If you are not sure if a list item applies to your research, read the appropriate section before selecting a response.

Materials & experimental systems

Methods

- | | |
|-------------------------------------|---|
| n/a | Included in the study |
| <input type="checkbox"/> | <input checked="" type="checkbox"/> Antibodies |
| <input type="checkbox"/> | <input checked="" type="checkbox"/> Eukaryotic cell lines |
| <input checked="" type="checkbox"/> | <input type="checkbox"/> Palaeontology and archaeology |
| <input type="checkbox"/> | <input checked="" type="checkbox"/> Animals and other organisms |
| <input checked="" type="checkbox"/> | <input type="checkbox"/> Human research participants |
| <input checked="" type="checkbox"/> | <input type="checkbox"/> Clinical data |
| <input checked="" type="checkbox"/> | <input type="checkbox"/> Dual use research of concern |

- | | |
|-------------------------------------|---|
| n/a | Included in the study |
| <input checked="" type="checkbox"/> | <input type="checkbox"/> ChIP-seq |
| <input checked="" type="checkbox"/> | <input type="checkbox"/> Flow cytometry |
| <input checked="" type="checkbox"/> | <input type="checkbox"/> MRI-based neuroimaging |

Antibodies

Antibodies used

6-4PP, Mouse, Cosmo bio, NM-DND-002, Immunoblot: 1:2000, N/A
 ALC1, Rabbit, Homemade, WB: 1:1000, aML#144
 Alexa 488 anti-rabbit IgG, Goat, Thermo fisher Scientific A-11034, IF: 1:1000, aML#012
 Alexa 488 anti-mouse IgG, Goat, Thermo fisher Scientific A-11029, IF: 1:1000, aML#013
 Alexa 555 anti-rabbit IgG, Goat, Thermo fisher Scientific A-21429, IF: 1:1000, aML#014
 Alexa 555 anti-mouse IgG, Goat, Thermo fisher Scientific A-21424, IF: 1:1000, aML#015
 Alexa 555 anti-mouse IgG, Donkey, Thermo fisher Scientific A-31570, IF: 1:1000, aML#171
 Alexa 647 anti-rabbit IgG, Goat, Thermo fisher Scientific A-21245, IF: 1:1000, aML#016
 Alexa 647 anti-mouse IgG, Goat, Thermo fisher Scientific A-21235, IF: 1:1000, aML#017
 Alexa 647 anti-goat IgG, Donkey, Thermo fisher Scientific A32849, IF: 1:1000, aML#176
 CF680 anti-rabbit IgG Goat, Biotium, VWR #20067, WB: 1:10000, aML#010
 CF770 anti-mouse IgG Goat, Biotium, VWR #20077, WB: 1:10000, aML#009
 CPD, Mouse, Cosmo Bio (TDM2 clone); CAC-NM-DND-001, IF: 1:1000 Immunoblot: 1:4000, N/A
 DDB2, Goat, R&D Systems Netherlands; AF3297-SP, WB: 1:1000, aML#107
 GFP, Goat, Homemade, WB: 1:2000, N/A
 GFP, Mouse, Roche, 11814460001, WB: 1:1000, aML#011
 PAR, Mouse, Mouse monoclonal 10H (ascites) (Homemade), WB: 1:500, N/A
 PAR, Mouse, Trevigen, 4335-MC-100, IF: 1:1000, aML#174

PAR-binding reagent, Rabbit, Millipore; MABE1031, IF: 1:500, N/A
 PAR-binding reagent, Rabbit, Millipore; MABE1016, WB: 1:1000, N/A
 PARP1, Rabbit, Cell signalling; #9542S, WB: 1:1000, aML#060
 PARP1, Rabbit, Homemade, WB: 1:10,000, N/A
 PARP1, Mouse, C2-10: Enzo: BML-SA250-0050, WB: 1:2000, N/A
 PARP2, Mouse, Enzo; clone: 4G8 (ALX-804-639-L001), WB: 1:200, aML#126
 PARP2, Rabbit, Active Motif: Cat# 39743, WB: 1:1000, N/A
 Tubulin, Mouse, Sigma; T6199, WB: 1:1000, aML#008
 XPA, Rabbit, Gift from Rick Wood (CJ1), WB: 1 in 10.000, aML#079
 XPB (ERCC3, p89), Mouse, Millipore, MABE1123, WB: 1 in 2000, aML#101
 XPC, Rabbit, Novus Biologicals: NB100-58801, WB: 1:1000 IF: 1:500, aML#077
 XPC, Rabbit, Gene Tex: GTX70309, WB: 1:1000, N/A

Validation

The following antibodies were validated in knockout cells:

ALC1, Rabbit Homemade WB: 1:1000 aML#144
 DDB2, Goat R&D Systems Netherlands; AF3297-SP WB: 1:1000 aML#107
 PARP1, Rabbit Cell signalling; #9542S WB: 1:1000 aML#060
 PARP1, Rabbit Homemade WB: 1:10,000 N/A
 PARP1, Mouse C2-10: Enzo: BML-SA250-0050 WB: 1:2000 N/A
 PARP2, Mouse Enzo; clone: 4G8 (ALX-804-639-L001) WB: 1:200 aML#126
 PARP2, Rabbit Active Motif: Cat# 39743 WB: 1:1000 N/A
 XPA, Rabbit Gift from Rick Wood (CJ1) WB: 1 in 10.000 aML#079
 XPC, Rabbit Novus Biologicals: NB100-58801 WB: 1:1000, IF: 1:500 aML#077
 XPC, Rabbit Gene Tex: GTX70309 WB: 1:1000 N/A

The following antibodies showed a reduced signal in PARP1-KO cells demonstrating specificity

PAR, Mouse Homemade (10H) WB: 1:500 N/A
 PAR, Mouse Trevigen, 4335-MC-100 IF: 1:1000 aML#174
 Poly-ADP-ribose binding reagent, Rabbit Millipore; MABE1031 IF: 1:500 N/A

UV-specific staining that disappears with expected kinetics and persists in XPC-KO cells

6-4PP, Mouse Cosmo bio, NM-DND-002 Immunoblot: 1:2000 N/A
 CPD, Mouse Cosmo Bio (TDM2 clone); CAC-NM-DND-001 IF: 1:1000, Immunoblot: 1:4000 N/A

The following antibodies were validated in Co-IP experiments:

XPB, (ERCC3, p89) Mouse Millipore, MABE1123 WB: 1 in 2000 aML#101
 GFP, Goat Homemade WB: 1:2000 N/A
 GFP, Mouse Roche, 11814460001 WB: 1:1000 aML#011

This antibody is a commonly used loading control:

Tubulin, Mouse Sigma; T6199 WB: 1:1000 aML#008

Eukaryotic cell lines

Policy information about [cell lines](#)

Cell line source(s)

U2OS Nicholas D Lakin (Ronson et al., 2018)
 U2OS 2-6-3 Susan Janicki (Janicki et al., 2004)
 U2OS PARP1-KO Nicholas D Lakin (Ronson et al., 2018)
 U2OS PARP2-KO Nicholas D Lakin (Ronson et al., 2018)
 U2OS(FRT) Daniel Durocher (Panier et al., 2012)
 U2OS(FRT) ALC1-KO This study
 U2OS(FRT) ALC1-KO + GFP-ALC1E175Q This study
 U2OS(FRT) ALC1-KO + GFP-ALC1WT This study
 U2OS(FRT) ALC1-KO + GFP-ALC1ΔMACRO This study
 U2OS(FRT) CSA-KO (van der Weegen et al., 2020)
 U2OS(FRT) DDB2 ALC1-dKO + GFP-DDB2 This study
 U2OS(FRT) DDB2-KO This study
 U2OS(FRT) DDB2-KO + GFP-DDB2 This study
 U2OS(FRT) DDB2-KO + GFP-ALC1 This study
 U2OS(FRT) GFP-ALC1 This study
 U2OS(FRT) GFP-NLS Haico van Attikum (Luijsterburg et al., 2017)
 U2OS(FRT) PARP1-GFP This study
 U2OS(FRT) GFP-PARP2 This study
 U2OS(FRT) XPC ALC1-dKO + XPC-GFP This study
 U2OS(FRT) XPC-KO This study
 U2OS(FRT) XPC-KO + GFP-PARP2 This study
 U2OS(FRT) XPC-KO + PARP1-GFP This study
 U2OS(FRT) XPC-KO + XPC-GFP This study

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|--|--|
| | U2OS(FRT) XPC-KO + GFP-ALC1 This study |
| Authentication | Cells were authenticated by STR profiling. All knockout cells were validated by Western blot analysis and DNA sequencing |
| Mycoplasma contamination | All cell lines were routinely tested for mycoplasma and were nested negative |
| Commonly misidentified lines (See ICLAC register) | No commonly misidentified cell lines were used in this study |

Animals and other organisms

Policy information about [studies involving animals](#); [ARRIVE guidelines](#) recommended for reporting animal research

| | |
|-------------------------|--|
| Laboratory animals | C. elegans strains wild type Bristol B2 and animals carrying alleles xpc-1 (tm3886), parp-1 (ok988) and parp-2 (ok344) |
| Wild animals | This study did not involve wild animals |
| Field-collected samples | This study did not involve samples collected from the field |
| Ethics oversight | No ethics oversight is required for studies using C. elegans |

Note that full information on the approval of the study protocol must also be provided in the manuscript.