

Reporting Summary

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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 Genomic sequencing data was collected using HiSeq-2500/4000, Nextseq550 and Novaseq 6000 system. Proteomics data was collected using Orbitrap Fusion Lumas platform.

Data analysis NGS data was analyzed using bowtie2-2.1.0, samtools-1.2, MACS2-2.1.1, HOMER v4.0 and Genomics Suite 7.18. Proteomics data was analyzed by Peak Studio X. AzureSpot Analysis Software 2.0 was used to perform western blot densitometry. Sigmaplot 13.0 was used for statistical analysis.

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](#)

High throughput sequencing data were deposited in the Gene Expression Omnibus database under accession number GSE133655. Proteomics data were deposited in PRIDE database under accession number PXD027143 and PXD017881.

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	Sample sizes were chosen according to the standards of the field (at least three independent biological replicates for proteomics, at least two independent biological replicates for high throughput sequencing and other molecular and cellular experiments)
Data exclusions	No data exclusion was performed in the study
Replication	We performed recycling assay three times. For Proteomics data, 9 replicates were examined. For ChIP-seq, 4 replicates were sequenced. All attempts on replication were successful.
Randomization	N/A No animal subjects were used in the study.
Blinding	N/A Investigators and samples were not blinded. Blinding is not used in the field.

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

n/a	Involved 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 checked="" type="checkbox"/>	<input 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

Methods

n/a	Involved in the study
<input type="checkbox"/>	<input checked="" 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

Pol II (N-20) Santa Cruz Cat#sc-899
 Pol II (8WG16) Abcam Cat#ab817
 Pol II (phospho S2) Abcam Cat#ab5095
 PAF1 Abcam Cat#ab20662
 PAF1 Abcam Cat#ab137519
 PAF1 Novus Biologicals Cat#NB600-273
 LEO1 Bethyl Laboratories Cat#A300-175A
 WDR61 Novus Biologicals Cat#NBP2-59231
 WDR61 Novus Biologicals Cat#NBP2-46359
 Rtf1 Abcam, Cat#ab99362
 H3 Abcam Cat#ab209023
 TBP Abcam Cat#ab197874
 MED6 Thermo Cat# PA5-40846
 SPT6H (H-227) Santa Cruz Biotechnology Cat# sc-367017
 TOP2B (H-286) Santa Cruz Biotechnology Cat# sc-13059
 TAF2 (6B3) Santa Cruz Biotechnology Cat# sc-735
 XRN2 (H-300) Santa Cruz Biotechnology Cat# sc-99237
 Goat anti-mouse IgG (H+L) secondary antibody, HRP Azure biosystems Cat#AC2115
 Goat anti-rabbit IgG (H+L) Secondary antibody, HRP Azure biosystems Cat#AC2114
 Donkey anti-goat IgG-HRP Santa Cruz Biotechnology Cat# sc-2020

Validation

The validation of all primary antibodies except for PAF1 antibody for the species and application, noting any validation statements on the manufacturer's website. The validation of PAF1 antibody for the species and application, noting any validation statements or data provided in the manuscript.

Eukaryotic cell lines

Policy information about [cell lines](#)

Cell line source(s)

The human prostate cancer cell line abl was provided by Dr. Zoran Culig (Innsbruck Medical University, Austria). The human female HeLa cell line was obtained from ATCC. The human benign prostatic hyperplasia cell line was obtained from Sigma.

Authentication

Cells were authenticated by short tandem repeat (STR) profiling and karyotyping or as described previously (Culig, et al. Br J Cancer 1999;81(2):242-51).

Mycoplasma contamination

The cell lines will be routinely tested to ensure they are free of mycoplasma contamination (VenorTMGeM Mycoplasma Detection Kit, Sigma-Aldrich). All cell lines tested negative for mycoplasma contamination.

Commonly misidentified lines
(See [ICLAC](#) register)

No commonly misidentified cell lines were used in the study.

ChIP-seq

Data deposition

- Confirm that both raw and final processed data have been deposited in a public database such as [GEO](#).
- Confirm that you have deposited or provided access to graph files (e.g. BED files) for the called peaks.

Data access links

May remain private before publication.

<https://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?token=cbonqaqkntyfnot&acc=GSE133655>

Files in database submission

GSM3914212 siCon-FP1h-0-PAF
 GSM3914213 siCon-FP1h-10-PAF
 GSM3914214 siCon-FP1h-20-PAF
 GSM3914215 siCon-FP1h-30-PAF
 GSM3914216 siCon-FP1h-40-PAF
 GSM3914217 siPAF-FP-0-5095
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 GSM5430247 Abl_RTF1_rep4

Genome browser session
 (e.g. [UCSC](#))

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Methodology

Replicates	We performed experiments using four to five biological replicates.
Sequencing depth	All data are 50-bp single-end. Each data was normalized to the same sequencing depth (50 million).
Antibodies	PAF1 (Abcam, ab20662) phosphorylated RNA pol II (Abcam, phospho S2) leo1 (Bethyl Laboratories, A300-175A) Rtf1 (Abcam, ab99362)
Peak calling parameters	Default parameters were used
Data quality	All peaks are at 1% FDR
Software	NGS data was analyzed using bowtie2-2.1.0, samtools-1.2, MACS2-2.1.1, HOMER v4.0 and Genomics Suite 7.0.