

Double-blind peer review submissions: write DBPR and your manuscript number here

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Last updated by author(s): YYYY-MM-DD

# **Reporting Summary**

Nature Research 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 Research policies, see Authors & Referees and the Editorial Policy Checklist.

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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
	$\square$ 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. <i>F</i> , <i>t</i> , <i>r</i> ) with confidence intervals, effect sizes, degrees of freedom and <i>P</i> value noted <i>Give P values as exact values whenever suitable.</i>
$\boxtimes$	For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings
$\boxtimes$	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 <u>statistics for biologists</u> contains articles on many of the points above.

### Software and code

Data analysis

Policy information about availability of computer code

Data collection No software was used

Unless otherwise stated, all analysis were done under R v3.3.2 using mainly packages from Bioconductor v 3.4.

Original code used for the analysis of gene neighbors is being further developed in the GeneNeighborhood R package available at https:// github.com/pgpmartin/GeneNeighborhood.

Detailed procedures for the production of each figure are provided in Methods

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/reviewers. We strongly encourage code deposition in a community repository (e.g. GitHub). See the Nature Research 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 list of figures that have associated raw data
- A description of any restrictions on data availability

All raw and processed sequencing data, as well as details of the experimental procedures and data analysis methods are provided in GEO SuperSeries GSE112443

Field-spe	ecific reporting		
Please select the o	ne below that is the best fit for yo	our research. If you are not sure, read the appropriate sections before making your selection.	
X Life sciences	Behavioural & socia	l sciences	
	the document with all sections, see <u>nature</u> .	com/documents/nr-reporting-summary-flat.pdf	
Life scier	nces study desig	ζN	
	sclose on these points even when		
Sample size		culations. Sample sizes for sequencing studies were essentially maximized based on available funding,	
Sample Size		perimental capacity (e.g. ability to handle large number of tubes simultaneously in ChIP-seq experiments)	
Data exclusions	As stated in GEO entry GSE112440	sample WT_12 was removed from the diffenrential expression analysis due to its low sequencing depth	
Data exclasions	compared to the other samples.		
Replication	The RNA-sea study to identify genes	s differentially expressed between hdr1 2.3 triple mutant and wild-type plants was entirely replicated (2)	
Replication	The RNA-seq study to identify genes differentially expressed between bdr1,2,3 triple mutant and wild-type plants was entirely replicated (2 independent studies: GSE112440 and GSE112441). Both studies involved biological triplicates in each group (except after removal of WT_12		
	in GSE112440, see above).  Replicates for ChIP-seg are describe.	d below. ChIP-seg on RNA polymerase II were performed in duplicates (or triplicates for RNAPII ChIP),	
	<u>'</u>	g different phosphorylation states of the largest subunit (RNAPII, S2P and S5P).	
Randomization	Samples/individuals were randomly	allocated to the treatment groups (MPA, 6AU or light experiments)	
DI: I			
Blinding	Except for the personnel of the sequ	uencing facility, investigators were not blinded to group allocation during data collection or analysis.	
We require informat	ion from authors about some types of	aterials, systems and methods materials, experimental systems and methods used in many studies. Here, indicate whether each material, e not sure if a list item applies to your research, read the appropriate section before selecting a response.	
	perimental systems	Methods	
n/a Involved in the	· · · · · · · · · · · · · · · · · · ·	n/a Involved in the study	
Antibodies	·	ChIP-seq	
Eukaryotic	cell lines	Flow cytometry	
Palaeonto	logy	MRI-based neuroimaging	
Animals ar	nd other organisms	_,_	
Human re	search participants		
Clinical da	ta		
—1—			
Antibodies			
Antibodies used	Antibodies are describe	ed (including lot numbers) for each sample in GEO entry GEO112443.	
	. ,	I CTD repeat YSPTSPS antibody [8WG16] - ChIP Grade (abcam, ab817, lot #: GR261055-1, GR261055-2 or	
	GR299078-3 ) Anti-RNA polymerase I	I CTD repeat YSPTSPS (phospho S2) antibody - ChIP Grade (abcam, ab5095, lot #: GR270578-1 or	
	GR295145-1)		
	Anti-RNA polymerase I GR278215-1)	I CTD repeat YSPTSPS (phospho S5) antibody - ChIP Grade (abcam, ab513, lot #: GR118198-2 or	
	Anti-Histone H3 antibo	ody - Nuclear Loading Control and ChIP Grade (abcam, ab1791, lot #: GR312884-1)	
	9 -	.73] Isotype Control- ChIP Grade (abcam, ab18413, lot # GR249563-7) r, clone 4A6 (Millipore-sigma, 05-724, lot #: 2720363).	
Validaki	All 4441	lin this study are nonconsistly available and validated by their second size and second	
Validation	All the antibodies used	in this study are commercially available and validated by their corresponding manufacturers.	

## Eukaryotic cell lines

Policy information about <u>cell lines</u>

Cell line source(s)

State the source of each cell line used.

Authentication

Describe the authentication procedures for each cell line used OR declare that none of the cell lines used were authenticated.

Mycoplasma contamination

Confirm that all cell lines tested negative for mycoplasma contamination OR describe the results of the testing for mycoplasma contamination OR declare that the cell lines were not tested for mycoplasma contamination.

Commonly misidentified lines (See ICLAC register)

Name any commonly misidentified cell lines used in the study and provide a rationale for their use.

## Palaeontology

Specimen provenance Provide provenance information for specimens and describe permits that were obtained for the work (including the name of the issuing authority, the date of issue, and any identifying information).

Dating methods

If new dates are provided, describe how they were obtained (e.g. collection, storage, sample pretreatment and measurement), where they were obtained (i.e. lab name), the calibration program and the protocol for quality assurance OR state that no new

dates are provided.

Tick this box to confirm that the raw and calibrated dates are available in the paper or in Supplementary Information.

## Animals and other organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research

Laboratory animals For laboratory animals, report species, strain, sex and age OR state that the study did not involve laboratory animals.

Wild animals

Provide details on animals observed in or captured in the field; report species, sex and age where possible. Describe how animals were caught and transported and what happened to captive animals after the study (if killed, explain why and describe method; if

released, say where and when) OR state that the study did not involve wild animals.

Field-collected samples

For laboratory work with field-collected samples, describe all relevant parameters such as housing, maintenance, temperature, photoperiod and end-of-experiment protocol OR state that the study did not involve samples collected from the field.

Ethics oversight Identify the organization(s) that approved or provided guidance on the study protocol, OR state that no ethical approval or quidance was required and explain why not.

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

## Human research participants

Policy information about studies involving human research participants

Describe the covariate-relevant population characteristics of the human research participants (e.g. age, gender, genotypic information, past and current diagnosis and treatment categories). If you filled out the behavioural & social sciences study design questions and have nothing to add here, write "See above."

**Recruitment**Describe how participants were recruited. Outline any potential self-selection bias or other biases that may be present and how these are likely to impact results.

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

### Clinical data

Policy information about clinical studies

All manuscripts should comply with the ICMJE guidelines for publication of clinical research and a completed CONSORT checklist must be included with all submissions.

Clinical trial registration

Provide the trial registration number from ClinicalTrials.gov or an equivalent agency.

Study protocol

Note where the full trial protocol can be accessed OR if not available, explain why.

Outcomes Describe how you pre-defined primary and secondary outcome measures and how you assessed these measures.

## ChIP-sea

#### 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.

To review GEO accession GSE112443:

go to https://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE112443

Files in database submission

ChIP-seq files related to this study are in GEO entries GSE113059 (Myc-BDR1 and Myc-BDR2 ChIP-seq) and GSE113078 (RNA polymerase II ChIP-seq). Both Series are part of GSE112443 SuperSeries.

Fastq, bigwig andbed files (BDR1 and BDR2 peaks) are provided for the following samples in GSE113059:

- Myc\_BDR1
- Myc BDR2
- Myc\_WT
- input\_BDR2\_lib1\_seq1
- input\_BDR2\_lib1\_seq2
- input\_BDR2\_lib2
- -input BDR1
- input\_WT

Fastq and bigwig files are provided for the following samples in GSE113078:

- RNAPII\_WT\_rep1
- RNAPII\_bdrs\_rep1
- S2P\_WT\_rep1
- S2P\_bdrs\_rep1
- S5P\_WT\_rep1
- S5P\_bdrs\_rep1
- input\_WT\_rep1\_lib1
- input\_bdrs\_rep1\_lib1
- RNAPII\_WT\_rep2\_seq1
- RNAPII\_bdrs\_rep2\_seq1
- S2P\_WT\_rep2
- S2P\_bdrs\_rep2 S5P\_WT\_rep2
- S5P bdrs rep2
- RNAPII\_WT\_rep2\_seq2
- RNAPII\_bdrs\_rep2\_seq2
- input\_WT\_rep1\_seq2
- RNAPII\_bdrs\_rep2\_seq2
- input\_WT\_rep1\_lib2 input\_bdrs\_rep1\_lib2
- IgG\_WT\_rep3
- -input WT rep3
- RNAPII\_WT\_rep3

Genome browser session (e.g. UCSC)

http://jbrowseiu.pgpmartin.fr

#### Methodology

Replicates

In the file names above \_rep1, 2 or 3 designates independent biological replicates. \_seq1 or 2 indicates independent sequencing of the same library and \_lib1 or 2 designates independent technical replicates at the level of library preparation (and sequencing). The results presented here were supported by all replictates.

Sequencing depth

Describe the sequencing depth for each experiment, providing the total number of reads, uniquely mapped reads, length of reads and whether they were paired- or single-end.

Antibodies

Antibodies used for ChIP-seq are described in the Antibody section above and for each sample in GEO entry GEO112443 (including lot numbers).

Peak calling parameters

Reads were mapped to the TAIR10 genome using Bowtie2 (v2.2.6, Langmead & Salzberg, Nature Methods, 2012) with the -dovetail parameter and and maximum insert size of 1kb

Peaks for BDR1 and BDR2 were called using MACS2 (v2.1.0, Zhang et al., Genome Biology, 2008) in paired-end mode using Myc\_WT as control. Peaks with a p-value<0.01 were kept. Peaks located in blacklisted regions were removed (see blacklist bed file provided as supplemental file).

Data quality

Read quality was evaluated with fastgc (v 0.11.14)

Adapters were trimmed with Trimmomatic (v0.33, Bolger, Lohse & Usabel, Bioinformatics, 2014) in paired end mode After alignment, duplicate reads were identified with Picard (v2.2.4) MarkDuplicates and removed. Samtools v 1.3 was used to keep only reads mapped in proper pairs with mapping quality (MapQ) above 2.

Software

In addition to the software above, ChIP-seq data were analyzed with custom R scripts under R (v3.3.2) and Bioconductor (v3.4)

## Flow Cytometry

Plots						
Confirm that:						
The axis labels state the marker and fluorochrome used (e.g. CD4-FITC).						
The axis scales are clearly	The axis scales are clearly visible. Include numbers along axes only for bottom left plot of group (a 'group' is an analysis of identical markers).					
All plots are contour plots with outliers or pseudocolor plots.						
A numerical value for number of cells or percentage (with statistics) is provided.						
Methodology						
Sample preparation	Describe the sample preparation, detailing the biological source of the cells and any tissue processing steps used.					
Instrument	Identify the instrument used for data collection, specifying make and model number.					
Software	Describe the software used to collect and analyze the flow cytometry data. For custom code that has been deposited into a community repository, provide accession details.					
Cell population abundance	Describe the abundance of the relevant cell populations within post-sort fractions, providing details on the purity of the samples and how it was determined.					
Gating strategy	Describe the gating strategy used for all relevant experiments, specifying the preliminary FSC/SSC gates of the starting cell population, indicating where boundaries between "positive" and "negative" staining cell populations are defined.					
Tick this box to confirm the	nat a figure exemplifying the gating strategy is provided in the Supplementary Information.					

## Magnetic resonance imaging

## Experimental design

Aperimental design

Indicate task or resting state; event-related or block design.

Design specifications

Design type

Specify the number of blocks, trials or experimental units per session and/or subject, and specify the length of each trial or block (if trials are blocked) and interval between trials.

Behavioral performance measures

State number and/or type of variables recorded (e.g. correct button press, response time) and what statistics were used to establish that the subjects were performing the task as expected (e.g. mean, range, and/or standard deviation across subjects).

### Acquisition

Imaging type(s)

Specify: functional, structural, diffusion, perfusion.

Field strength

Specify in Tesla

Sequence & imaging parameters

Specify the pulse sequence type (gradient echo, spin echo, etc.), imaging type (EPI, spiral, etc.), field of view, matrix size, slice thickness, orientation and TE/TR/flip angle.

Area of acquisition

State whether a whole brain scan was used OR define the area of acquisition, describing how the region was determined.

Diffusion MRI Used

Not used

#### Preprocessing

Preprocessing software

Provide detail on software version and revision number and on specific parameters (model/functions, brain extraction, segmentation, smoothing kernel size, etc.).

Normalization

If data were normalized/standardized, describe the approach(es): specify linear or non-linear and define image types used for transformation OR indicate that data were not normalized and explain rationale for lack of normalization.

Normalization template

Describe the template used for normalization/transformation, specifying subject space or group standardized space (e.g. original Talairach, MNI305, ICBM152) OR indicate that the data were not normalized.

Noise and artifact removal	Describe your procedure(s) for artifact and structured noise removal, specifying motion parameters, tissue signals and physiological signals (heart rate, respiration).			
Volume censoring	Define your software and/or method and criteria for volume censoring, and state the extent of such censoring.			
Statistical modeling & inference				
Model type and settings	Specify type (mass univariate, multivariate, RSA, predictive, etc.) and describe essential details of the model at the first and second levels (e.g. fixed, random or mixed effects; drift or auto-correlation).			
Effect(s) tested	Define precise effect in terms of the task or stimulus conditions instead of psychological concepts and indicate whether ANOVA or factorial designs were used.			
Specify type of analysis: Whole brain ROI-based Both				
Statistic type for inference (See <u>Eklund et al. 2016</u> )	Specify voxel-wise or cluster-wise and report all relevant parameters for cluster-wise methods.			
Correction	Describe the type of correction and how it is obtained for multiple comparisons (e.g. FWE, FDR, permutation or Monte Carlo).			
Models & analysis				
n/a   Involved in the study   Functional and/or effective connectivity   Graph analysis   Multivariate modeling or predictive analysis				
Functional and/or effective connective	Report the measures of dependence used and the model details (e.g. Pearson correlation, partial correlation, mutual information).			

Multivariate modeling and predictive analysis

etc.).

Graph analysis

Specify independent variables, features extraction and dimension reduction, model, training and evaluation metrics.

Report the dependent variable and connectivity measure, specifying weighted graph or binarized graph, subject- or group-level, and the global and/or node summaries used (e.g. clustering coefficient, efficiency,