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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 <u>Authors & Referees</u> and the <u>Editorial Policy Checklist</u>.

Sta	atis	tics		
For	all st	atistical analy:	ses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.	
n/a	Cor	nfirmed		
	\boxtimes	The exact sar	mple size (n) for each experimental group/condition, given as a discrete number and unit of measurement	
	\boxtimes	A statement	on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly	
	\boxtimes	The statistica	ll test(s) used AND whether they are one- or two-sided tests should be described solely by name; describe more complex techniques in the Methods section.	
	\boxtimes	A description	of all covariates tested	
\boxtimes		A description	of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons	
	\boxtimes	A full description	tion of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) n (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)	
\boxtimes		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 Give <i>P</i> values as exact values whenever suitable.		
\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			
\boxtimes	Estimates of effect sizes (e.g. Cohen's d , Pearson's r), indicating how they were calculated			
	ı		Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.	
So	ftw	vare and o	code	
Poli	cy in	formation abo	out <u>availability of computer code</u>	
Data collection		ollection	BD FACSDiva™ software v6.1.3; CCP4 Suite, XDS package, GROMACS, Modoller 9.20	
Data analysis		nalysis	Excel 2007, Prism 7, FlowJo v7.6.1, origin 7	

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 <u>data availability statement</u>. 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

A reporting summary for this article is available as a Supplementary Information file. The source data underlying Figure 1b, 1d, 1e, 1g, 3c, 3d, 3e, 4b, 4c, 4d, 4e, 4f, 4g, Supplementary Figures 2, 3a, 3b, 7, 8a, 9a, 9b, 10 and Table 2 can be found in the Source Data files.

The structure of VanR complexed with vanillic acid has been deposited in the Protein Data Bank [www.pdb.org], with a PDB ID code of 6LG2.

Field-spe	cific reporting			
Please select the or	e below that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.			
\times Life sciences	Behavioural & social sciences Ecological, evolutionary & environmental sciences			
For a reference copy of t	ne document with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>			
Life scier	ces study design			
All studies must dis	All studies must disclose on these points even when the disclosure is negative.			
Sample size	The data are from three replicates.			
Data exclusions	No data were excluded.			
Replication	The data are from three replicates and all attempts at replication were successful.			
Randomization	Samples were tested in one group. There is no group allocation.			
Blinding	Samples were tested in one group. There is no group allocation.			
Reportin	g for specific materials, systems and methods			
We require information	on from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each materi			
•	ed 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			
n/a Involved in th	perimental systems Methods n/a Involved in the study			
Antibodies	ChIP-seq			
Eukaryotic cell lines Flow cytometry				
Palaeontology MRI-based neuroimaging				
Animals and other organisms				
Human research participants				
Clinical dat				
Flow Cytome	etry			
Plots				
Confirm that:				
The axis labels state the marker and fluorochrome used (e.g. CD4-FITC).				
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 c	All plots are contour plots with outliers or pseudocolor plots.			
🔀 A numerical v	alue for number of cells or percentage (with statistics) is provided.			
Methodology				
Sample preparati	cells were precultured overnight in LB medium and then diluted to OD600=0.2 in the same medium in the presence of corresponding inducer. The cells were then grown for 12 h and FACS sorting was performed.			
Instrument	Instrument FACSAria II sorter (BD, San Jose, USA)			
Software	Software BD FACSDiva™ software v6.1.3; FlowJo v7.6.1			
Cell population a	Cell population abundance A total of 350000 cells were used for sorting.			
Gating strategy The least fluorescent 80% cells (negative sorting) and most fluorescent 1% cells (positive sorting) were sorted.				