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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 our Editorial Policies and the Editorial Policy Checklist.

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1016	ali StatiSticai ali	aryses, commit that the following items are present in the right elegend, table legend, main text, or interious section.
n/a	Confirmed	
	🗶 The exact	sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement
x	A stateme	ent on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly
×		tical test(s) used AND whether they are one- or two-sided on tests should be described solely by name; describe more complex techniques in the Methods section.
×	A descript	cion of all covariates tested
	🗶 A descript	cion of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons
×		cription of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) tion (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)
×		ypothesis testing, the test statistic (e.g. F , t , r) with confidence intervals, effect sizes, degrees of freedom and P value noted es as exact values whenever suitable.
×	For Bayes	ian analysis, information on the choice of priors and Markov chain Monte Carlo settings
x	For hierar	chical 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.
Sof	ftware an	d code
Polic	cy information	about <u>availability of computer code</u>
Da	ta collection	The data was downloaded manually via a web-browser. The ICGC PANCANCER ANALYSIS OF WHOLE GENOMES data portal (https://

Data analysis

For the ROC-curve analysis the ROCR package was used and referenced in the manuscript.

dcc.icgc.org/pcawg) was used to ensure correctness of sample mapping.

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 Research guidelines for submitting code & software for further information.

Data

Policy information about <u>availability of data</u>

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

Only data referenced in Nonneville and Reddel has been used in this study.

Life sciences study design

Data exclusions No data was excluded. Replication Not applicable. Randomization No randomization was necessary.	Sample size	This formal reply was a reanalysis of the data in the "Matters arising" article.
	Data exclusions	No data was excluded.
Randomization No randomization was necessary.	Replication	Not applicable.
	Randomization	No randomization was necessary.
Blinding Not applicable.	Blinding	Not applicable.

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	Involved in the study	n/a	Involved in the study
X	Antibodies	×	ChIP-seq
×	Eukaryotic cell lines	×	Flow cytometry
X	Palaeontology and archaeology	X	MRI-based neuroimaging
×	Animals and other organisms		
×	Human research participants		
×	☐ Clinical data		
x	Dual use research of concern		