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Corresponding author(s):	Dimitrios Spentzos
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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 <u>Editorial Policies</u> and the <u>Editorial Policy Checklist</u>.

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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
	$oxed{x}$ 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>
x	For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings
x	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 <i>d</i> , Pearson's <i>r</i>), 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 No data was collected.

Data analysis The NCI BRB-ArrayTo

The NCI BRB-ArrayTools v4.6.0, R (versions 3.5.1 and 4.1.0), R ggplot2 package, R minfi package, R missMethyl package, R ESTIMATE package, R bumphunter package, R amap package, R pheatmap package, R dendsort package, R hmisc package, R qqman package, R survminer package, SPSS (version 24), CIBERSORTx software were used for data 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 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

The datasets analyzed in this study are available at the TARGET data matrix (https://ocg.cancer.gov/programs/target/data-matrix), the Gene Expression Omnibus repository (GSE125645, GSE97529, GSE59200 and GSE68379), ArrayExpress (E-MTAB-3610), and the PharmacoDB database (https://pharmacodb.pmgenomics.ca/).

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.					
x Life sciences	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				
Life scier	nces study design				
All studies must di	sclose on these points even when the disclosure is negative.				
Sample size	All data used in the study was previously published and sample sizes were chosen by the authors.				
Data exclusions	Three samples were excluded from the TARGET methylation dataset because they did not have clinical follow up (survival times censored at zero months).				
Replication	Where possible we validate findings in independent publicly available datasets.				
Randomization	Not applicable. This is a retrospective study using public data.				
Blinding	Not applicable. This is a retrospective study.				
Reportir	ng for specific materials, systems and methods				
,	ion from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, sted 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	
×	Antibodies	ChIP-seq	
×	Eukaryotic cell lines	Flow cytometry	
×	Palaeontology and archaeology	MRI-based neuroimaging	
×	Animals and other organisms	·	
×	Human research participants		
×	☐ Clinical data		
X	Dual use research of concern		