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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>.

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For	statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.	
n/a	onfirmed	
	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 coeffi AND variation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)	icient
	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>	ł
\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	
	$\overline{\Box}$ 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

Policy information about availability of computer code

Data collection

No software used for data collection. All raw data used in this analysis are from TCGA and METABRIC public repositories. All processed data used in the modeling process are deposited in our GitHub account: https://github.com/xyouli/DNA-based-predictors-of-nongenetic-cancer-phenotypes

Data analysis

All code used for data analysis in this manuscript are deposited in GitHub: https://github.com/xyouli/DNA-based-predictors-of-non-genetic-cancer-phenotypes

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

The TCGA data referenced during the study are available from the TCGA website (the Broad Institute TCGA GDAC Firehose: https://gdac.broadinstitute.org/). The METABRIC data is available from the European Genome-phenome Archive at the European Bioinformatics Institute (https://www.ebi.ac.uk/ega/). All the other data sets supporting the findings of this study are available within the article, the supplementary information tables and our GitHub repository and from the corresponding author upon reasonable request.

Field-spe	ecific reporting			
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All studies must dis	sclose on these points even when the disclosure is negative.			
Sample size	NA			
Data exclusions	NA			
Replication	NA			
Randomization	NA			
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