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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>.

Statistics

For all statistical analyses, 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 sample 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	
		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.	
\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 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)	
\boxtimes		For null hypothesis testing, the test statistic (e.g. F, t, r) with confidence intervals, effect sizes, degrees of freedom and P value noted Give P 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	
		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 Software was used	
Data analysis	Natural abundance calculation script available at: https://github.com/wencgu/nac	

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

Previously published data is indicated. Non-published, raw data is included in the supplemental excel file.

Field-specific reporting

Life sciences

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.

Behavioural & social sciences Ecological, evolutionary & environmental sciences

For a reference copy of the document with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>

Life sciences study design

All studies must disclose on these points even when the disclosure is negative.

Sample size	N/A
Data exclusions	N/A
Replication	N/A
Randomization	N/A
Blinding	N/A

Reporting for specific materials, systems and methods

Methods

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

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

Eukaryotic cell lines

Policy information about <u>cell lines</u>			
Cell line source(s)	ATCC		
Authentication	HCC15 cells (non-small cell lung cancer) were identified using DNA fingerprinting.		
Mycoplasma contamination	Cells were confirmed to be mycoplasma free using the e-Myco kit (Boca Scientific).		
Commonly misidentified lines (See I <u>CLAC</u> register)	Name any commonly misidentified cell lines used in the study and provide a rationale for their use.		

Animals and other organisms

Policy information about <u>studies involving animals; ARRIVE guidelines</u> recommended for reporting animal research				
Laboratory animals	NOD.Cg-Prkdc <scid> Il2rg<tm1wjl>/SzJ (NSG) Mice</tm1wjl></scid>			
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

All procedures were approved by UT Southwestern's Animal Care and Use Committee in accordance with the Guide for the Care and Use of Laboratory Animals.

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				
Population characteristics	All patients with thoracic tumors are eligible			
Recruitment	All patients with thoracic tumors are eligible			
Ethics oversight	Patients with NSCLC were enrolled in an IRB-approved protocol after obtaining informed consent (NCT02095808)			

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