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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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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	
	$oxed{oxed}$ 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.	
\boxtimes	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 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	
	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 <u>availability of computer code</u>		

Data collection

The raw reads were aligned to the reference human genome (UCSC hg19) with TopHat2(19). Genes were annotated (using NCBI RefSeq) and quantified by HTSeq(20), and DESeq(21) was used to identify differentially expressed genes and significant genes with fold change > 1 and multiple-test adjusted p value < 0.01 were used for interpreting the biological pathways.

Data analysis

Gene Set Enrichment Analysis was performed using stand-alone distribution (http://www.gsea-msigdb.org/gsea/index.jsp)

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 $\underline{\text{availability of data}}$

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

The data supporting the findings of this study are available from the corresponding authors upon reasonable request. Source data for the graphs and charts are provided in the Supplementary Data 1. The RNAseq data has been deposited in the GEO database (accession number GSE163249). Unedited western blot images are available in the Supplementary Information (Supplementary Figures 7–17).

Field-specific reporting				
Please select the or	ne below that	is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.		
∠ Life sciences		Behavioural & social sciences		
For a reference copy of t	he document with	n all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>		
Life scier	ices st	udy design		
		e points even when the disclosure is negative.		
Sample size		animal studies, sample size of tumors/treatment was derived using effect information from previous studies and calculations were based model of unpaired data power =0.8; p<0.05.		
Data exclusions	No data was ex	xcluded		
Replication	All in vitro assa	ays were performed in triplicate and repeated at least three times		
Randomization	Animal groups	were randomized based on tumor volume. All in vitro assays were done in triplicate with no randomization.		
Blinding	All in vitro assays were performed in triplicate and repeated at least three times. Investigator was not blinded. IHC studies are blinded for quantitation by independent evalutions.			
Reportin	g for s	pecific materials, systems and methods		
We require information	on from authors	s about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, by your study. If you are not sure if a list item applies to your research, read the appropriate section before selecting a response.		
Materials & exp				
n/a Involved in th		n/a Involved in the study		
Antibodies	,	ChIP-seq		
Eukaryotic	cell lines	Flow cytometry		
Palaeontolo	ogy and archaed	ology MRI-based neuroimaging		
Animals an	d other organisı	ns		
Human res	earch participar	nts		
Clinical dat	a			
Dual use re	search of conce	rn		
Antibodies				
Antibodies used		odies for GAPDH, p-S6, S6, p-Akt (S473), Akt, p-mTOR (S2448), mTOR, p-STAT3 (Y705), and STAT3 were purchased from Cell		
	_	ling Technology (Danvers MA). LIF and LIFR antibodies were purchased from Santa Cruz Biotechnology (Dallas, TX). β-actin ody was obtained from Sigma-Aldrich (St. Louis, MO). The Ki67 antibody was purchased from Abcam (Cambridge, MA).		
Validation	All the	e antibodies used were commercially validated antibodies and from respectable companies.		
Eukaryotic cell lines				
Policy information about cell lines				
Cell line source(s)		ATCC		
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STR DNA profiling was used to confirm cell identity.

None used

All the model cells utilized were free of mycoplasma contamination

Authentication

Mycoplasma contamination

Commonly misidentified lines (See <u>ICLAC</u> register)

Animals and other organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research		
Laboratory animals	female, athymic nude mice , SCID mice	
Wild animals	None	
Field-collected samples	None	
Ethics oversight	All animal experiments were performed using UTHSA IACUC approved protocol	

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