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Corresponding author(s):	Sven Diederichs	
Initial submission	Revised version	Final submission

Life Sciences Reporting Summary

Nature Research wishes to improve the reproducibility of the work that we publish. This form is intended for publication with all accepted life science papers and provides structure for consistency and transparency in reporting. Every life science submission will use this form; some list items might not apply to an individual manuscript, but all fields must be completed for clarity.

For further information on the points included in this form, see Reporting Life Sciences Research. For further information on Nature Research policies, including our data availability policy, see Authors & Referees and the Editorial Policy Checklist.

pol	policies, including our data availability policy, see Authors & Referees and the Editorial Policy Checklist.			
•	Experimental design			
1.	Sample size			
	Describe how sample size was determined.	n.a.		
2.	Data exclusions			
	Describe any data exclusions.	Synonymous mutations found in COSMIC were excluded if: 1) they were not derived from whole genome sequencing studies, 2) they were duplicates of the same mutation in the same sample, 3) if they were misannotated.		
3.	Replication			
	Describe whether the experimental findings were reliably reproduced.	Each experiment was reproduced at least three times. Means and SEM are provided for all figures where applicable.		
4.	Randomization			
	Describe how samples/organisms/participants were allocated into experimental groups.	n.a.		
5.	Blinding			
	Describe whether the investigators were blinded to group allocation during data collection and/or analysis.	n.a.		
	Note: all studies involving animals and/or human research partici	pants must disclose whether blinding and randomization were used.		
6.	Statistical parameters			
	For all figures and tables that use statistical methods, con Methods section if additional space is needed).	firm that the following items are present in relevant figure legends (or in the		
n/a	Confirmed			
	The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement (animals, litters, cultures, etc.			
	A description of how samples were collected, noting whether measurements were taken from distinct samples or whether the same sample was measured repeatedly			
	A statement indicating how many times each experiment was replicated			
	The statistical test(s) used and whether they are one- or two-sided (note: only common tests should be described solely by name; more complex techniques should be described in the Methods section)			
\times	A description of any assumptions or corrections, such as an adjustment for multiple comparisons			
	The test results (e.g. <i>P</i> values) given as exact values whenever possible and with confidence intervals noted			
	A clear description of statistics including central tendency (e.g. median, mean) and variation (e.g. standard deviation, interquartile range)			
	Clearly defined error bars			

See the web collection on statistics for biologists for further resources and guidance.

Software

Policy information about availability of computer code

7. Software

Describe the software used to analyze the data in this study.

The data was analyzed using Excel, Galaxy, RNAsnp, remuRNA, Vienna RNA package. The database was built in shinyR.

For manuscripts utilizing custom algorithms or software that are central to the paper but not yet described in the published literature, software must be made available to editors and reviewers upon request. We strongly encourage code deposition in a community repository (e.g. GitHub). *Nature Methods* guidance for providing algorithms and software for publication provides further information on this topic.

Materials and reagents

Policy information about availability of materials

8. Materials availability

Indicate whether there are restrictions on availability of unique materials or if these materials are only available for distribution by a for-profit company.

All material is available.

9. Antibodies

Describe the antibodies used and how they were validated for use in the system under study (i.e. assay and species).

β-actin (A2228, Sigma-Aldrich, 1:20000), anti-V5 (1:1000-1:2500, R96025, Thermo Scientific), anti-GAPDH (1:10000, #MAB374, Millipore).

10. Eukaryotic cell lines

- a. State the source of each eukaryotic cell line used.
- b. Describe the method of cell line authentication used.
- c. Report whether the cell lines were tested for mycoplasma contamination.
- d. If any of the cell lines used are listed in the database of commonly misidentified cell lines maintained by ICLAC, provide a scientific rationale for their use.

HEK293, HeLa

STR profiling

Cell lines were regularly tested for mycoplasma (every 3 months) and always found negative.

n.a.

▶ Animals and human research participants

Policy information about studies involving animals; when reporting animal research, follow the ARRIVE guidelines

11. Description of research animals

Provide details on animals and/or animal-derived materials used in the study.

n.a.

Policy information about studies involving human research participants

12. Description of human research participants

Describe the covariate-relevant population characteristics of the human research participants.

n.a.