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Last updated by author(s):	Jun 9, 2020

Reporting Summary

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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
	\mathbf{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
×	\square 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 was used.

GraphPad Prism 8.0 (GraphPad Software, Inc., San Diego, Ca, USA).

ANOVA was performed using SPSS (Version 12.0, Chicago, IL, USA).

Multivariate statistical analyses, such as principal component analysis (PCA) and principal component analysis (PLSR), were performed with SIMCA-P+ (Version 11.0 Umetrics, Umeå, Sweden).

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

Data analysis

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 $% \left(1\right) =\left(1\right) \left(1\right) \left($
- A description of any restrictions on data availability

We have provided a full data availability statement in the manuscript.

Field-specific reporting

Life sciences study design

All studies must disclose on these points even when the disclosure is negative.		
Sample size	This study selected sample sizes based on previous studies. In addition, the accuracy of the research hypothesis and results was verified by repeated experiments.	
Data exclusions	Data were not excluded from this study.	
Replication	All experiments were replicated independently to verify the reproducibility of the results, and all relevant attempts were successful.	
Randomization	Mice were randomly assigned to groups.	
Blinding	In this experiment, human subjects are not involved and blinding was not possible.	

Reporting for specific materials, systems and 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	Methods	
n/a Involved in the study	n/a Involved in the study	
x Antibodies	X ChIP-seq	
Eukaryotic cell lines	Flow cytometry	
Palaeontology	MRI-based neuroimaging	
Animals and other organisms	•	
Human research participants		
Clinical data		

Antibodies

Antibodies used

PDK4 (Santa Cruz, CA, USA, sc-14495); PDH1a (Abcam, Cambridge, UK, ab10330); p-PDH1a (Abcam, Cambridge, UK, ab92696); b-Actin (Santa Cruz, CA, USA, sc-47778)

Validation

PDK4 (Santa Cruz, CA, USA, sc-14495) is reacted with mouse, rat, and human origin by Western Blotting (goat polyclonal antibody, 47kDa).

PDH1a (Abcam, Cambridge, UK, ab10330) is reacted with mouse, rat, and human origin by Western Blotting (mouse monoclonal antibody, 43kDa).

p-PDH1a (Abcam, Cambridge, UK, ab92696) is reacted with mouse and human origin by Western Blotting (rabbit polyclonal antibody, 43kDa).

b-Actin (Santa Cruz, CA, USA, sc-47778) is reacted with mouse, rat, and human origin by Western Blotting (mouse monoclonal antibody, 43kDa).

Eukaryotic cell lines

Policy information about <u>cell lines</u>	
Cell line source(s)	HepG2, Human hepatocellular carcinoma C2C12, Mouse myoblast
Authentication	HepG2 (Human hepatocellular carcinoma, Korean Cell Line Bank, KCLB NO. 88065) Cell line STR Profile (D3S1358: 15,16 vWA: 17 FGA: 22,25 Amelogenin: X,Y TH01: 9 TPOX: 8,9 CSF1P0: 10,11 D5S818: 11,12 D13S317: 9,13 D7S820: 10) C2C12 (Mouse myoblast, ATCC, CRL-1772)
Mycoplasma contamination	Cells were not contaminated with mycoplasma.
Commonly misidentified lines (See ICLAC register)	As a result of ICLAC verification, no cell lines were misidentified in this study.

Animals and other organisms

Policy information about <u>studies involving animals</u>; <u>ARRIVE guidelines</u> recommended for reporting animal research

Laboratory animals C57BL/6J male mice were purchased from Samtako Co. (Kyunggido, Korea), and PPARα-deficient male mice were purchased from Taconic (Hudson, NY, USA).

Wild animals This study did not involve wild animals.

Field-collected samples All mice were maintained under a 12-h light/12-h dark cycle at a temperature of 21-25°C and a relative humidity of 50-60%. At

the end of the experimental period, the mice were fasted for 12 h and sacrificed at the same atmosphere.

Ethics oversight All animal experiments were performed according to a protocol approved by the Animal Experiment Committee of Korea

University (Protocol No. KUIACUC-20090420-4).

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