# nature portfolio

Raw single cell RNA sequencing and Seurat object will be available before publication.

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## **Reporting Summary**

Nature Portfolio 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 Portfolio 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	1.		
(a Confirmed			
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 repeate	dly		
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 Give <i>P</i> values as exact values whenever suitable.			
For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings			
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 <u>availability of computer code</u>			
Data collection Chromium Controller Firmware (10X Genomics).			
Data analysis Cell Ranger (Version 6.1.2); R software (Version 4.1.2); Seurat package (Version 4.1.1).			
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 ecreviewers. We strongly encourage code deposition in a community repository (e.g. GitHub). See the Nature Portfolio guidelines for submitting code & software for further in			
Data			
Policy information about <u>availability of data</u> 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 description of any restrictions on data availability  - For clinical datasets or third party data, please ensure that the statement adheres to our <u>policy</u>			

Research involving human participants, their data, or biological material		
Policy information about studies with <a href="https://human.participants.or">human data</a> . See also policy information about		

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

Academy of Medical Sciences.

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Ethics oversight

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.

Life sciences

Behavioural & social sciences

Ecological, evolutionary & environmental sciences

The use of human lung tissue in this study was approved by the Human Ethics Committee of Fuwai Hospital, Chinese

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 No statistical methods were used to pre-determine sample size.

Data exclusions No data were excluded from the analysis.

Replication We performed single cell RNAseq on 10 lungs, ), with comparable results among the donors in the same group. The same samples were used for the histological validation experiments.

Randomization Randomization was not relevant due to the study design.

Blinding Blinding was not relevant due to the study design.

### 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 Involve	ed in the study	n/a	Involved in the study	
Ant	tibodies	$\boxtimes$	ChIP-seq	
Euk	karyotic cell lines	$\boxtimes$	Flow cytometry	
∑ Pal	aeontology and archaeology	$\boxtimes$	MRI-based neuroimaging	
Ani	mals and other organisms			
Clir	nical data			
Dua Dua	al use research of concern			
∑ □ Pla	nts			

#### **Antibodies**

Antibodies used

Rabbit anti-FTL (Proteintech, #10727-1-AP, 1:200) Mouse anti-SFTPC (Abcam, #ab90716,  $1 \mu g/ml$ )

Rabbit anti-EEF1A1 (Proteintech, #11402-1-AP, 1:200) Rabbit anti-AGER (Abcam, #ab216329, 1:1000) Mouse anti-VIM (Abcam, #ab8978, 2µg/ml) Rabbit anti-p21 (Proteintech, #10355-1-AP, 1:1000)

Validation

Validation statements are available from manufacturers' websites:

Rabbit anti-FTL (https://www.ptgcn.com/products/FTL-Antibody-10727-1-AP.htm)

Mouse anti-SFTPC (https://www.abcam.cn/prosurfactant-protein-c-antibody-ab90716.html) Rabbit anti-EEF1A1 (https://www.ptgcn.com/products/EEF1A1-Antibody-11402-1-AP.htm)

Rabbit anti-AGER (https://www.abcam.cn/rage-antibody-epr21171-ab216329.html)
Mouse anti-VIM (https://www.abcam.cn/vimentin-antibody-rv202-cytoskeleton-marker-ab8978.html)

Rabbit anti-p21 (https://www.ptgcn.com/products/P21-Antibody-10355-1-AP.htm)

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Seed stocks	N/A
Novel plant genotypes	N/A
Authentication	N/A