

Figure S1: Effects of aging and *JunD* deficiency on the number of lung β-galactosidase positive cells in mice treated by NAC or vehicle.

(A) The number of  $\beta$ -galactosidase positive cells was higher in aged than in young JunD<sup>-/-</sup> mice. (B) Representative micrographs showing cells stained for  $\beta$ -galactosidase positive cells activity at pH 6. Results are individual values and means. \*\**P*<0.01, \**P*<0.05.



**Figure S2: Muc5AC and Muc5B expression in lung tissue from** *JunD<sup>-/-</sup>* **and WT mice: effects of NAC treatment.** Lung Muc5AC (left panel) and Muc5B (right panel) mRNA levels did not differ between aged and young mice and were not affected by *JunD* inactivation.

![](_page_2_Figure_1.jpeg)

### Figure S3: NAC treatment decreases p53 protein expression in control and JunD<sup>-/-</sup> mouse lungs

p53 expression assessed by Western blotting in mouse lung homogenates (left panel). Representative Western blots (right panel). Results are individual values and means. \*\*P<0.01, and \*P<0.05.

![](_page_3_Figure_1.jpeg)

### Figure S4: NAC treatment decreases p16 mRNA expression in control and *JunD<sup>-/-</sup>* mouse lungs

Lung p16 mRNA levels were assessed by quantitative PCR after reverse transcription. Results are individual values and means for 8 animals per group. \*\**P*<0.01, and \**P*<0.05.

![](_page_4_Figure_1.jpeg)

# Figure S5: NAC treatment stimulates growth of mouse embryonic fibroblasts (MEFs) previously subjected to a 3T3 immortalization protocol.

Primary mouse embryonic fibroblasts from control and  $JunD^{-/-}$  mice grown initially in a low-oxygen environment to increase their survival rate and previously subjected to a 3T3 immortalization protocol (13) were studied for assessment of cell proliferation in response to NAC (1mM) or vehicle. (A) Cells were fixed and stained using crystal violet. (B) NAC treatment slightly increased BrdU incorporation. Results are individual values and means for 8 animals per group. \*\*P<0.01, and \*P<0.05.

## Table S1

BMI

, ,	·			
	Control		COPD	
	Mean	sem	Mean	sem
n	8		8	
Females/males	4/4		4/4	
Age, yr	63.2	±5.41	61.1	±2.40
FEV %	98.9	±7.34	72.4*	±6.34
FVC%	100.2	±4.44	87.2	±6.42
FEV/FVC,%	91.6	±4.55	63**	±1.88
Pack-years	12.5	±6.41	34.4*	±3.46

**Table S1:** Clinical features and pathological variables in all the patients with chronic obstructive pulmonary disease and control smokers from whom lung samples were obtained.

Abbreviations: FEV, forced expiratory volume; FVC, forced vital capacity; BMI, body mass index.

24.1

±1.44

±0.98

24.4

![](_page_6_Figure_0.jpeg)

### Fig1: actin

![](_page_6_Figure_2.jpeg)

![](_page_7_Figure_0.jpeg)

![](_page_7_Figure_1.jpeg)

![](_page_7_Figure_2.jpeg)

![](_page_7_Figure_3.jpeg)

![](_page_8_Figure_0.jpeg)

![](_page_8_Figure_1.jpeg)

![](_page_8_Figure_2.jpeg)

### Fig3: gH2.AX young

![](_page_9_Figure_1.jpeg)

#### Fig3: actin young

![](_page_9_Figure_3.jpeg)

### Fig3: gH2.AX aged

![](_page_9_Figure_5.jpeg)

![](_page_9_Figure_6.jpeg)

![](_page_9_Figure_7.jpeg)

Fig S3: p53

![](_page_10_Figure_1.jpeg)

Fig S3: actin

![](_page_10_Figure_3.jpeg)