SUPPLEMENTARY TABLES AND FIGURES

| Name of cell line | Histology | Origin |
|-------------------|----------------------------|---|
| HBEC3-KT | Human bronchial epithelial | Established from bronchial epithelial biopsy ¹ |
| A549 | ADC | Established in 1972 from carcinomatous tissue |
| H1299 | ADC | Established from metastatic lymph node |
| HCC827 | ADC | Established from lung tissue |
| H1975 | ADC | Established in 1988 |
| H460 | LCC | Established in 1982 from pleural fluid |
| SKMES-1 | SCC | Pleural fluid of metastatic squamous cell |
| | | carcinoma |
| EBC-1 | SCC | Established from squamous cell carcinoma |
| HTB182 | SCC | Established in 1982 from squamous cell |
| | | carcinoma |
| CRL5889 | SCC | Stage 1 squamous cell carcinoma from smoker |
| | | donor tissue |
| H226 | SCC | Established from pleural fluid |

Supplementary Table 1. Histology and origin of cell lines used.

ADC; adenocarcinoma, LCC; large cell carcinoma, SCC; squamous cell carcinoma.

1. Ramirez, R.D., Sheridan, S., Girard, L., Sato, M., Kim, Y., Pollack, J. et al. Immortalization of human bronchial epithelial cells in the absence of viral oncoproteins. *Cancer Res.* **64**, 9027-9034 (2004).



Supplementary Figure 1. Prognostic value of COMMD4 in ADC and SCC. A and B. Univariate Kaplan-Meier analysis of overall survival in ADC (A) and SCC (B) cases.HR; hazard ratio, CI; confidence interval, ADC; adenocarcinoma, SCC; squamous cell carcinoma.



Supplementary Figure 2. COMMD4 expression. A and B. Representative images of negative (0), weak (1) and moderate (2) scores for COMMD4 nuclear and cytoplasmic staining of adenocarcinoma (A) and squamous cell carcinoma (B) TMAs. N; nuclear, C; cytoplasmic. C. Graph showing COMMD4 nuclear and cytoplasmic staining intensity of adenocarcinomas and squamous carcinomas across all TMA cores. D. Fluorescence staining of the negative control (secondary antibody only) and the DAPI nuclear staining of the TMAs.



В

Α



Supplementary Figure 3. COMMD4 expression and COMMD4-depletion causes mitotic catastrophe. A. An immunoblot showing the depletion of COMMD4 using control siRNA or COMMD4 siRNA #2 or #3 in the H1299 NSCLC cell line. β -actin shows the loading. B. α -tubulin staining (green) in control HBEC3-KT and H460, H1975, H1299, and CRL5889 NSCLC cell lines. DAPI shows the nucleus. Scale bar denotes 15 μ m.

4



В

Α



Supplementary Figure 4. Hypersensitivity to irradiation in COMMD4-depleted cells and complementation of the hypersensitivity with the overexpression of COMMD4. A. Clonogenic cell viability assay in H1299 cells transfected with control or COMMD4 siRNA (#2 or #3) and treated with varying doses of irradiation (IR). B. Clonogenic assay demonstrating the correction of the IR defect in cells depleted of COMMD4 (with siRNA #2) using a COMMD4 siRNA-resistant plasmid and overexpression of COMMD4-FLAG in the H460 cells. *; P < 0.05 from three independent experiments. Error bars represent mean \pm SD.



Supplementary Figure 5. Cell cycle analysis in COMMD4-depleted cells. A. Flow cytometry analyses of the DNA content profile for control siRNA and siRNA #2 and siRNA #3 treated HBEC3-KT, H460, H1975 and CRL5889 cell lines. Live cells were stained with Annexin V-488 and propidium iodide and assessed using a CytoFLEX flow cytometer. The DNA content was determined by staining the fixed cells with propidium iodide prior to flow cytometry. B. Quantification of (A) using FlowJo v10 software to quantify the proportion of cells within the G/M phase of the cell cycle in the cell lines analysed. *p < 0.05, n.s; not significant. Error bars represent mean \pm S.D from three independent experiments. HBEC3-KT, control cell line and H460, H1975 and CRL5889 are NSCLC cells. sub G10; cells in sub G0/G1 phase, G10; cells in G0/G1 phase, S; cells in S phase, G2M; cells in the G2/M phase and >4N; aneuploid cells > 4N.