

Supplementary Figure 1. Mouse models for genetic marking of lung cells. **A,** Lung photographs and green epifluorescence images (top two rows), as well as fluorescent microscopic images of lung sections (Hoechst 33258 stain, endogenous mT and mG fluorescence, and merged images; bottom four rows) of genetically marked mice employed in these studies at six postnatal weeks (n =5/group). Note absence of mG+ cells in mT/mG and of mT+ cells in mT/mG; Sox 2. Cre mice, mG+ cells in bronchi (b) but not alveoli (a) of mTmG; Scgb1a1. Cre mice, in bronchi and alveoli of mTmG;Sftpc.Cre mice, in alveoli of mTmG;Lyz2.Cre mice, in alveolar capillaries of mTmG; Vav. Cre mice, and in neuroepithelial bodies of mTmG; Nes. Cre mice. ps, pleural space. B, (Left graph), XY plot of mG+ airway versus alveolar cells from A (n = 5/group). Arrows denote the complete and exclusive mG+ marking of airway but not alveolar cells in mTmG; Scgb1a1.Cre mice (green); the exclusive marking of some alveolar but not airway cells in mTmG;Lyz,Cre mice (blue arrows); and the promiscuous marking of alveolar and airway cells in mTmG; Sftpc. Cre mice (olive). (Right graph), data summary from immunostaining of lung sections of lung-marked mice (n = 5/group) for CCSP and SFTPC shown in Figure 1A: XY plot of ratios of mG+ to CCSP+ airway versus mG+ to SFTPC+ alveolar cells. Arrows denote the complete and exclusive mG+ marking of CCSP+ club cells, but not of alveolar cells in mTmG; Scgb1a1. Cre mice (green); the exclusive marking of a fraction of SFTPC+ ATII cells, but not of airway cells in mTmG;Lyz,Cre mice (blue); and the promiscuous marking of mTmG; Sftpc. Cre mice (olive). C, Quantification of genetic/proteinaceous labeling from immunostains of airways of mT/mG; Sftpc. Cre mice (n = 5) for Clara cell secretory protein (CCSP, top two graphs) shows that $75 \pm 24\%$ of mG+ cells are club cells and $66 \pm 11\%$ of club cells are mG+, and from immunostains of distal alveolar regions of mT/mG; Lyz2. Cre mice (n = 5) for SFTPC and LYZ2 (bottom two graphs) that $68 \pm 3\%$ of mG+ cells are ATII cells and 32 \pm 3% alveolar macrophages (AM Φ) and that 47 \pm 4% of ATII cells are mG+, i.e. from the SFTPC+LYZ2+ lineage. **D**, Schematic representation of genetic marking in mTmG;Scgb1a1.Cre, mTmG;Sftpc.Cre, and mTmG;Lyz.Cre mice (left), flow cytometric gating strategy to quantify mG+ and mT+ cells (middle), and data summary from n = 5, 3, and 6

mice/group (right). Numbers above columns are mean \pm SD values of mG+ cell percentage/strain. Data are given as mean \pm SD. Five non-overlapping fields/sample were examined. mG, membranous green fluorescent protein fluorophore; mT, membranous tomato fluorophore.