

**Figure S1** Cell annotations. (A) Visualization of UMAP clustering results displaying the aggregation and distribution of cells, with each color representing a cluster; (B) two-dimensional visualization of UMAP clustering results showing the aggregation and distribution of cells from three samples; (C) UMAP distribution plots of eight marker genes; (D) visualization of UMAP clustering results after extraction of epithelial cell population, illustrating the aggregation and distribution of cells, with each color representing a cluster; (E) UMAP distribution plots of eight marker genes after extraction of epithelial cell population. COPD, chronic obstructive pulmonary disease; UMAP, uniform manifold approximation and projection.



**Figure S2** Impact of Nrf2 activator on the SASP of COPD primary cells. (A) Western blot experiment assessing the protein expression of nuclear Nrf2 in each group of cells; (B) Western blot experiment examining the expression of Nrf2, IL-1 $\beta$ , MMP1, and TGF- $\beta$  proteins in primary cells of each group; (C) Western blot experiment analyzing the expression levels of p53, p21, and p16 proteins in primary cells of each group; (D) CCK-8 assay evaluating the proliferative capacity of primary cells in each group; (E) ELISA experiment measuring the levels of TNF- $\alpha$  and IL-6 in the supernatant of primary cell cultures in each group; (F) SA- $\beta$ -Gal staining experiment determining the proportion of SA- $\beta$ -Gal-positive cells in primary cells of each group; (G-I) measurement of enzyme activities of CAT (G), SOD (H) and ROS (I) levels in primary cells of each group. ANOVA was used to compare differences between groups: \*, P<0.05 compared to the model group. Cell experiments were repeated three times. COPD, chronic obstructive pulmonary disease; Nrf2, nuclear factor erythroid 2-related factor 2; SASP, senescence-associated secretory phenotype; IL-1 $\beta$ , interleukin-1 beta; MMP1, matrix metalloproteinase 1; TGF- $\beta$ , transforming growth factor beta; CCK-8, cell counting kit-8; TNF- $\alpha$ , tumor necrosis factor-alpha; IL-6, interleukin-6.