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SARS-CoV-2–infected primary human airway epithelia illustrate mucus hypersecretion



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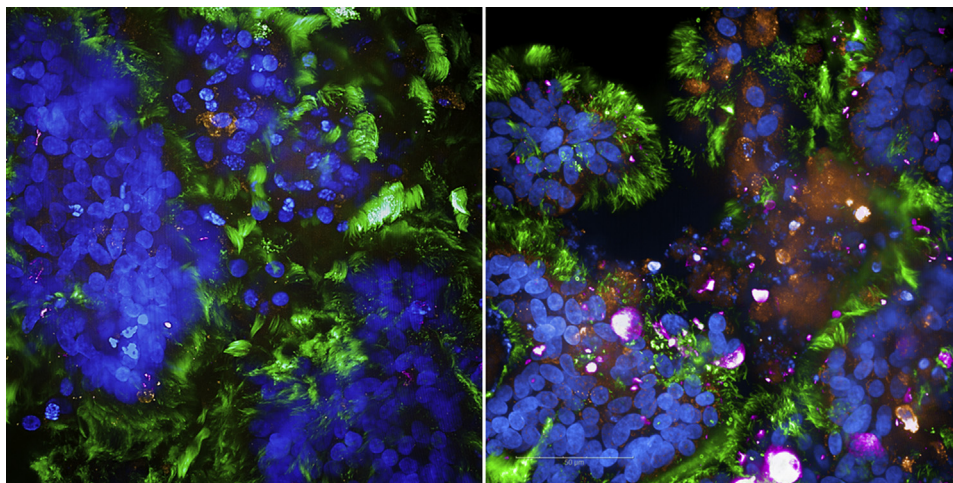


FIG 1. SARS-CoV-2 causes exacerberated mucus hypersecretion and plug formation. Visualization of mock-treated, uninfected or SARS-CoV-2 infected human airway tissues were analyzed after 2 days by immunofluorescence analyses (nuclei [blue], ciliated cells [green], mucous-producing cells [orange], SARS-CoV-2 [pink]).

Key word: SARS-CoV-2

To monitor first SARS-CoV-2 interactions with primary, fully differentiated, ciliated, and mucous-producing epithelial tissue models, infection with clinical isolates derived from patients with SARS-CoV-2 was performed and monitored after 2 days by using confocal microscopy (Fig 1 [right]). Mock-treated, uninfected (UI) cultures served as negative controls (left). Immunofluorescence analyses revealed a significant infection of the tissue model with SARS-CoV-2 (pink [right]) and significant destruction, as recently described by us¹ and as observed by high fragmentation of nuclei (blue) in particular within mucous areas (orange [right]). Infection with SARS-CoV-2 of both ciliated (green [right]) and mucous-producing cells (orange [right]) was detected. These analyses, which were performed by using the MUC5AC antibody

for goblet cell staining in SARS-CoV-2–infected and UI tissues, revealed that infected human airway tissues showed an exacerberated mucus hypersecretion and mucus plugs (orange [right]), whereas in UI tissues no such plugs were monitored (left). UI tissues illustrated goblet cell staining (orange [left]) and intact, multilayered epithelia (blue [left]). Mucus plug formation was described when critically ill COVID-19 patients with airway obstruction and respiratory failure were analyzed (reviewed in Khan et al²), and mucus hypersecretion in combination with proinflammatory cytokine activation, which we recently illustrated in our model,¹ and reduced mucociliary clearance³ shape a vicious circle resulting in airway tissue destruction. Of note, these clinical manifestations were not only reported in COVID-19 but also during other viral infections, colonization by pathogenic opportunistic bacteria, or asthma (reviewed in Khan et al²).

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