Figure 1







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Supplementary Figure Legends

Supplementary Figure 1: Protein levels of ACE2 and endogenous proteases that contribute to SARS-CoV-2 pathogenesis in CD45⁺ immune cells among groups. A-H, Flow cytometry was used to determine membrane protein levels of ACE2, TMPRSS2, furin, and ADAM17. Fluorescence intensity of a positive cell population was compared to a negative cell population (fluorescence minus one negative control for staining) (Δ MFI). The compared groups were nonsmokers (NS, white), electronic-cigarette vapers (ECIG-vapers, light grey) and tobacco cigarette smokers (TCIG-smokers, light blue). Representative data of percentage of immune (CD45⁺) cells that had positive staining for each protein between compared groups are shown for ACE2 (A), TMPRSS2 (C), ADAM17 (E), furin (G). Summary of data for ACE2 (B) TMPRSS2 (D), ADAM17 (F), furin (H) are shown. I. Representative data of Δ MFI for ACE2 and TMPRSS2 in immune (CD45⁺) cells. Summary of Δ MFI data for ACE2 (J) TMPRSS2 (K), furin (L), ADAM17 (M) are shown. Mann Whitney test was used to compare 2 groups (*p < 0.05, **p < 0.01, ***p < 0.001).

Supplementary Figure 2: Protein levels of ACE2 and endogenous proteases that contribute to SARS-CoV-2 pathogenesis in gut epithelial cells are associated with respective protein levels in CD45⁺ blood immune cells within the same person. A, B. Gut epithelial and immune cells were isolated from colon biopsies of healthy participants who underwent screening colonoscopy (n=11) as described in methods. Flow cytometry was used to determine membrane protein levels of ACE2, TMPRSS2, furin,

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and ADAM17 in peripheral blood mononuclear CD45⁺ blood immune cells (PBMCs), gut CD324⁺ CD45⁻ epithelial and gut CD45⁺ immune cells. Fluorescence intensity of a positive cell population was compared to a negative cell population (fluorescence minus one negative control for staining) (Δ MFI, shown in light grey). **A.** Representative data of Δ MFI for ACE2, TMPRSS2, furin and ADAM17 in gut epithelial and immune (CD45⁺) cells. **B.** Representative data of percentage of gut epithelial and immune (CD45⁺) cells that had positive staining for each protein. C. Scatter plots of protein levels of ACE2 in PBMCs (% of ACE2⁺ PBMCs, x axis) against protein levels of ACE2 in gut epithelial cells (% of ACE2⁺ of CD324⁺CD45⁻ gut cells, y axis)(n = 11). The Spearman correlation coefficient was used for all correlations. D. Scatter plots of protein levels of TMPRSS2 in PBMCs (% of TMPRSS2⁺ PBMCs, x axis) against protein levels of TMPRSS2 in gut epithelial cells (% of TMPRSS2⁺ of CD324⁺CD45⁻ gut cells, y axis)(n = 11). **E.** Scatter plots of protein levels of furin in PBMCs (% of furin⁺ PBMCs, x axis) against protein levels of furin in gut epithelial cells (% of furin⁺ of CD324⁺CD45⁻ gut cells, y axis)(n = 11). **F.** Scatter plots of protein levels of ADAM17 in PBMCs (% of ADAM17⁺ PBMCs, x axis) against protein levels of ADAM17 in gut epithelial cells (% of ADAM17⁺ of CD324⁺CD45⁻ gut cells, y axis)(n = 11).