Supplementary Figure 1. Weight loss of different Ace2TM mice infected with SARS-CoV-2 WA-1. (A) 8-14-week-old WT mice, Ace2H353K mice, Ace2S82M/F83Y mice, and Ace2TM mice were infected with WA-1 (2.34x10^5 PFU). Graph shows percent change in baseline body weight. (B) qRT-PCR of WA-1 Spike RNA loads in the lungs of mice at 3 dpi, shown as fold-change over RNA levels in WT infected mice. Total RNA input was normalized prior to qRT-PCR analysis.
**Supplementary Figure 2.** Histopathology scoring sheet of lung specimens from WA-1 and Omicron infected Ace2™ and WT mice. Tissue pathology was assessed based on the presence of the indicated features in distinct areas from each lung specimen in (A) WT non-infected, (B) WA-1 infected WT and Ace2™ mice and (C) Omicron infected WT and Ace2™ mice.