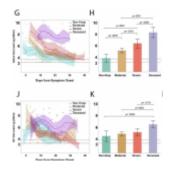
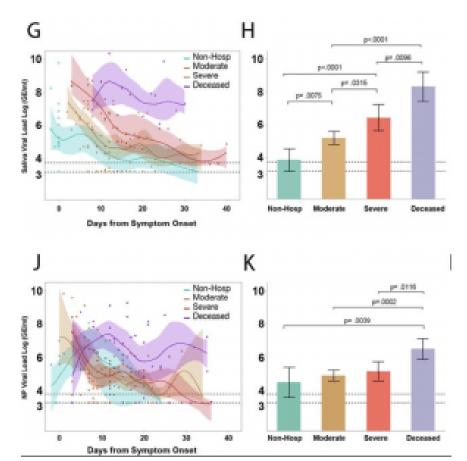
Saliva viral load could be a potential correlate of severe COVID-19





Saliva viral load is strongly associated with spectrum of disease severity throughout the course of illness. (G) Comparison of saliva viral load between non-hospitalized, moderate, severe, and deceased individuals across days from symptom onset. Points represent a unique individual-timepoint and color represents the clinical severity status of that given individual-timepoint as labelled. Moderate and severe classification represents only individual-timepoints of patients who survived. A cubic spline fitted to the mean with a lambda of 0.05 is shown for each severity group. Shading of the lines represent the bootstrap confidence of fit. (H) Least squares Mean analysis comparing mean saliva viral load amongst spectrum of disease over days from symptom onset using Restricted Maximum Likelihood (REML) to account for for the repeated measures same individuals. Significance was adjusted for multiple comparisons using the Tukey method. (J. K) Comparison of nasopharyngeal viral load between spectrums of disease severity a s described for saliva viral load in (G) and (H) respectively. Dotted lines VLP=positivity threshold for viral load 3.75 Log (Genomic Equivalents is [GE]/ml)); VLD=viral load limit of detection is 3.22 Log (GE/ml). (Source: Silva et al., Pre-print)

Determining COVID-19 severity and mortality in patients is an evolving and important topic as it is needed to guide clinical practice. COVID-19 patients' prognosis cannot be simply judged by how they visually appear. In a phenomenon known as "happy hypoxia," patients who do not appear distressed have been found to be extremely hypoxic, with oxygen saturation levels as low as 50%. Patients usually lose consciousness at oxygen saturation levels under 75%.

While blood tests and nasopharyngeal viral load can give some insight on disease severity, the development of other tests such as saliva tests can give clinicians a better picture of patient prognosis.

Silva *et al.*, found that saliva viral load was significantly higher in people with COVID-19 risk factors, correlated with increased disease severity, and was a stronger predictor of mortality over time compared to nasopharyngeal viral load.

The group quantified nasopharyngeal and saliva viral RNA using RT-qPCR in 154 patients admitted to Yale-New Haven Hospital. Plasma cytokines, chemokine, antibodies, and leukocyte profiles were analyzed as well. Patients who were hospitalized were found to have higher salivary viral loads. These patients were also found to have higher levels of known COVID-19 associated inflammatory markers such as IL-6, IL-18, IL-10, CXCL10, and type 1 immune response cytokines, and lower platelet and lymphocyte counts. In patients with fatal cases of COVID-19, the high salivary viral loads correlated with follicular CD4+ T cell depletion and lower anti-spike and anti-receptor binding domain IgG production.

Authors concluded that: "Together these results demonstrated that viral load — as measured by saliva but not nasopharyngeal — is a dynamic unifying correlate of disease presentation, severity, and mortality over time."

Journal Articles:

- <u>Silva et al., (Pre-Print). Saliva viral load is a</u> <u>dynamic unifying correlate of COVID-19 severity and</u> <u>mortality. MedRxiv.</u>
- <u>Couzin-Frankel (2020). The mystery of the pandemic's</u>
 <u>'happy hypoxia'. Science</u>

Summary by Maxwell Chan