## **Intersection Matrix**

Deep sequenced gDNA samples: 53 convalescent samples 25 follow-up samples



Selected in frame templates



Template n>2



Create a variable 'CAS\_V\_J'

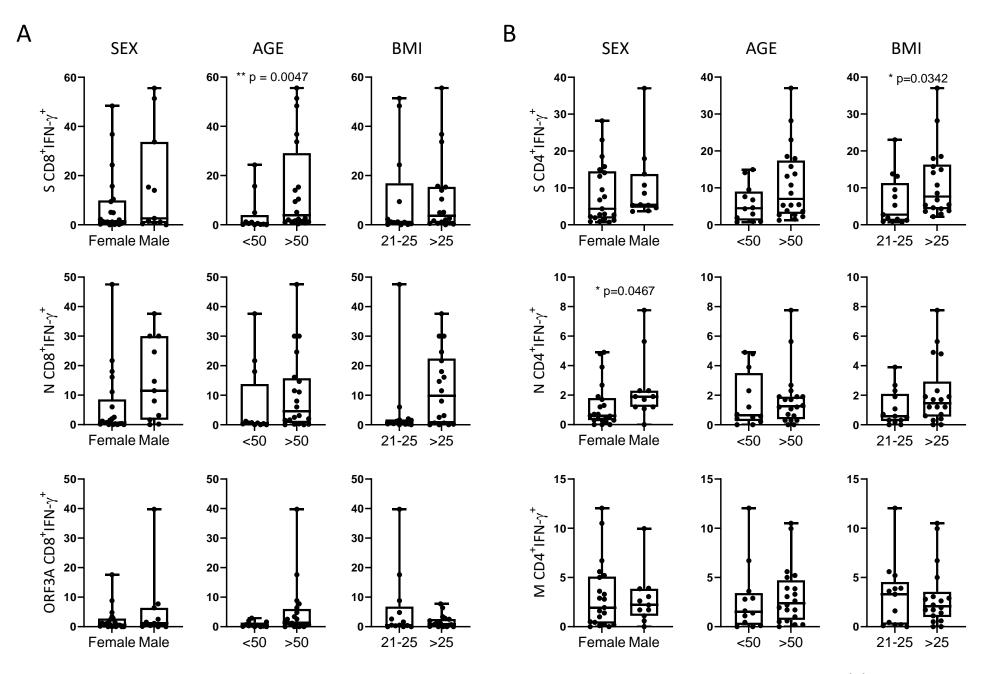


Summed the productive frequencies of individual CAS\_V\_J within a patient

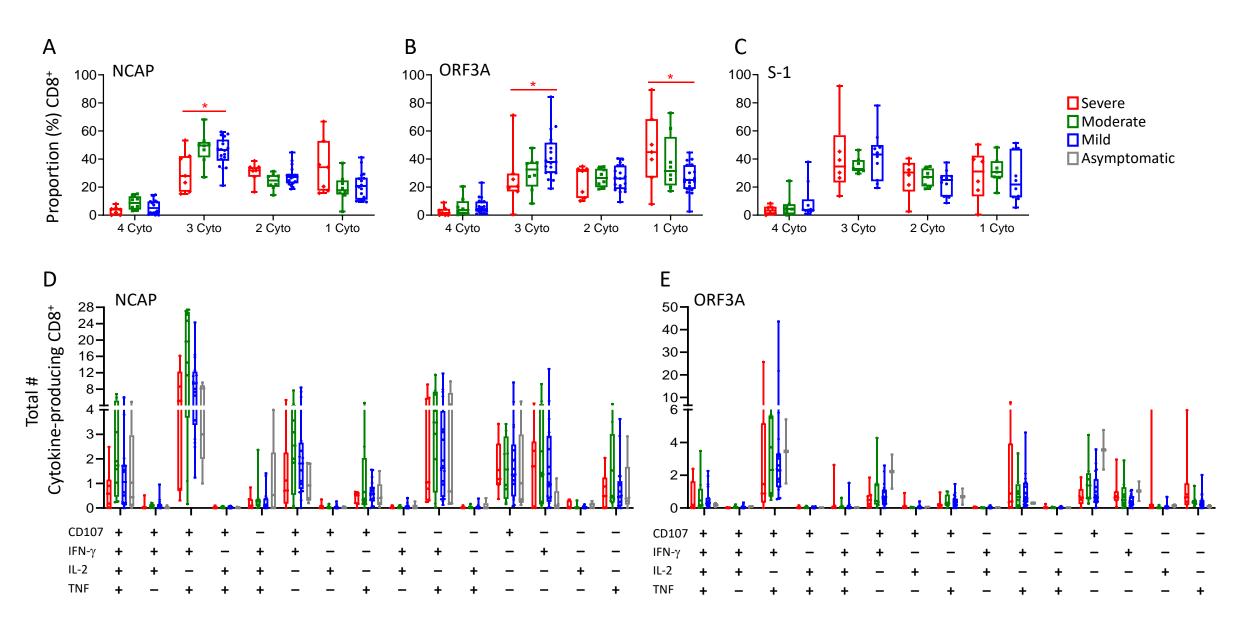


Intersection Matrix to determine shared CAS\_V\_J templates between donors

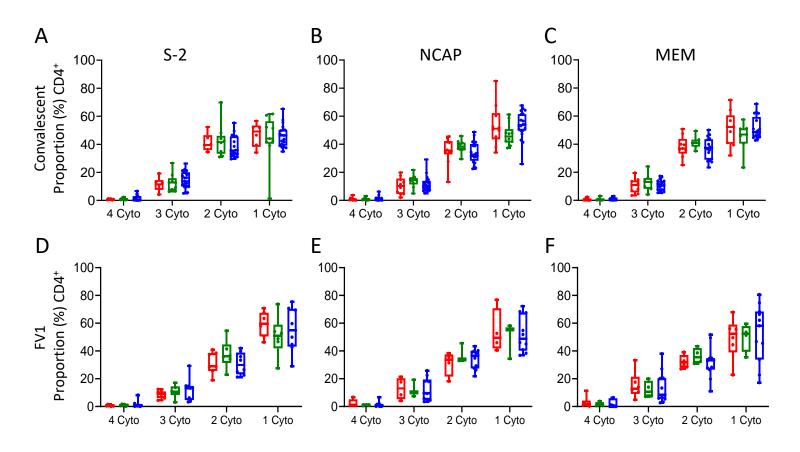
**Supplementary Figure 1**. Flow chart of the intersection matrix analysis.



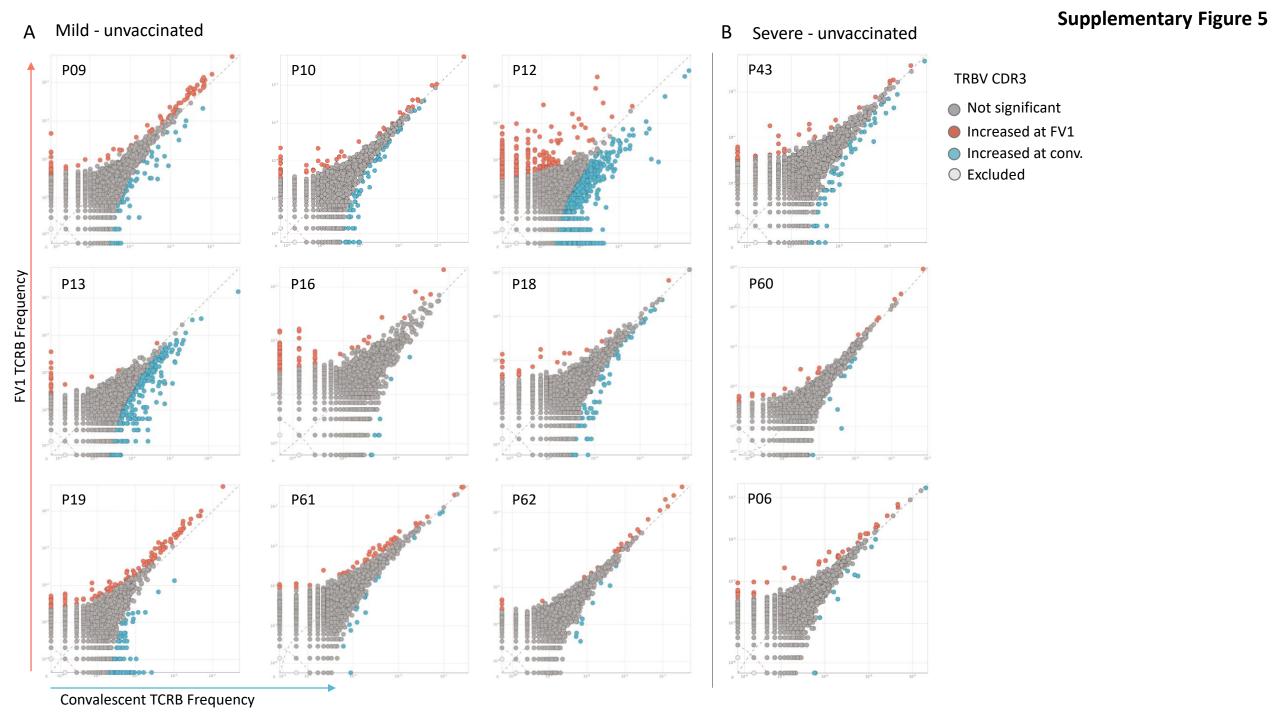
**Supplementary Figure 2**. Impact of Sex, Age and BMI on the SARS-CoV-2 T cell responses. Data represents the frequency of **(A)** CD8<sup>+</sup>IFN- $\gamma$ <sup>+</sup> T cells and **(B)** CD4+IFN- $\gamma$ <sup>+</sup> T cells



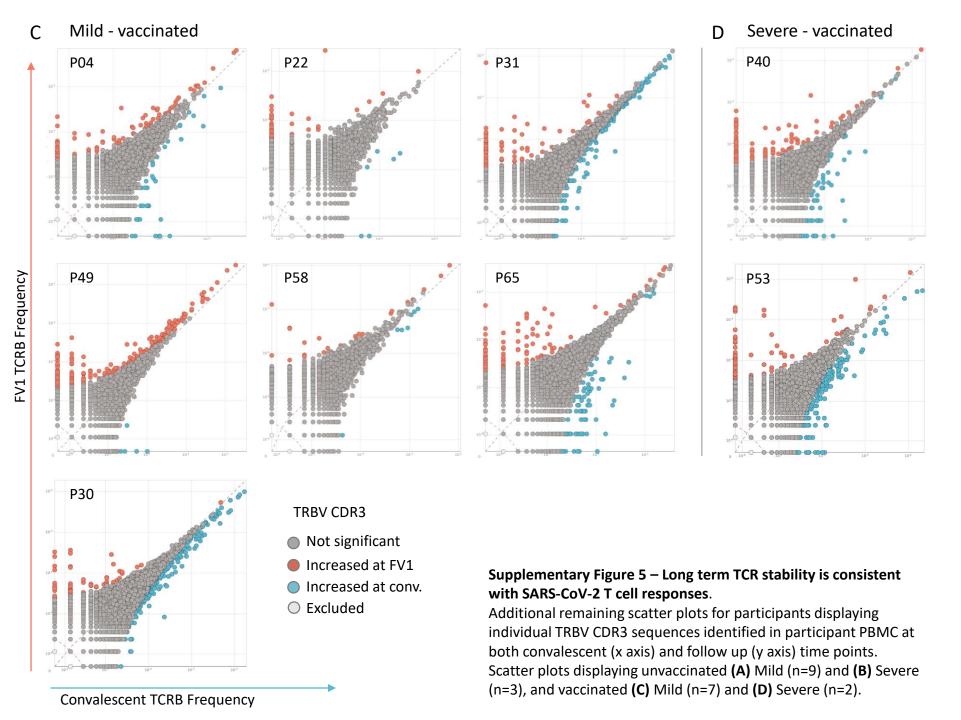
Supplementary Figure 3. SARS-CoV-2 convalescent CD8<sup>+</sup> T cell responses show increased polyfunctionality. Data represents the polyfunctional cytokine (cyto) profile of (A) NCAP-specific, (B) ORF3A-specific and (C) S-1 specific T cells. Data in (F) NCAP-specific and (E) ORF3A-specifi, represents the total number of cytokine-producing CD8<sup>+</sup> T cells



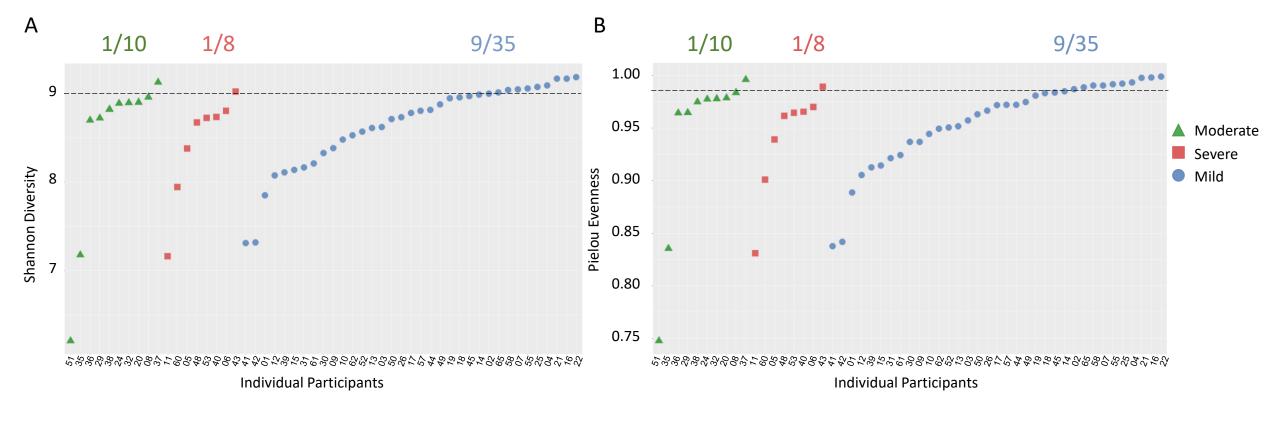
**Supplementary Figure 4**. SARS-CoV-2 CD4<sup>+</sup> T cell responses show reduced polyfunctionality that persists long-term. Data represents the polyfunctional cytokine (cyto) profile of **(A&D)** S-2 specific T cells, **(B&E)** NCAP-specific T cells and **(C&F)** Mem-specific T cells at the convalescent (top panels) and follow-up (bottom panels) timepoints.



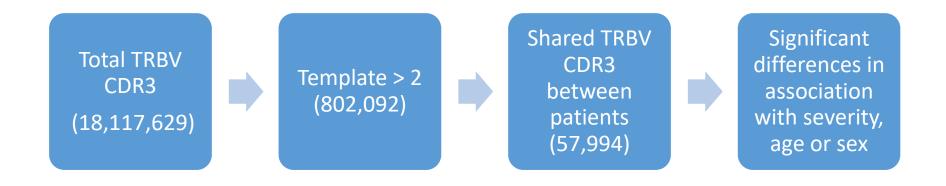
**Supplementary Figure 5 continued.** 



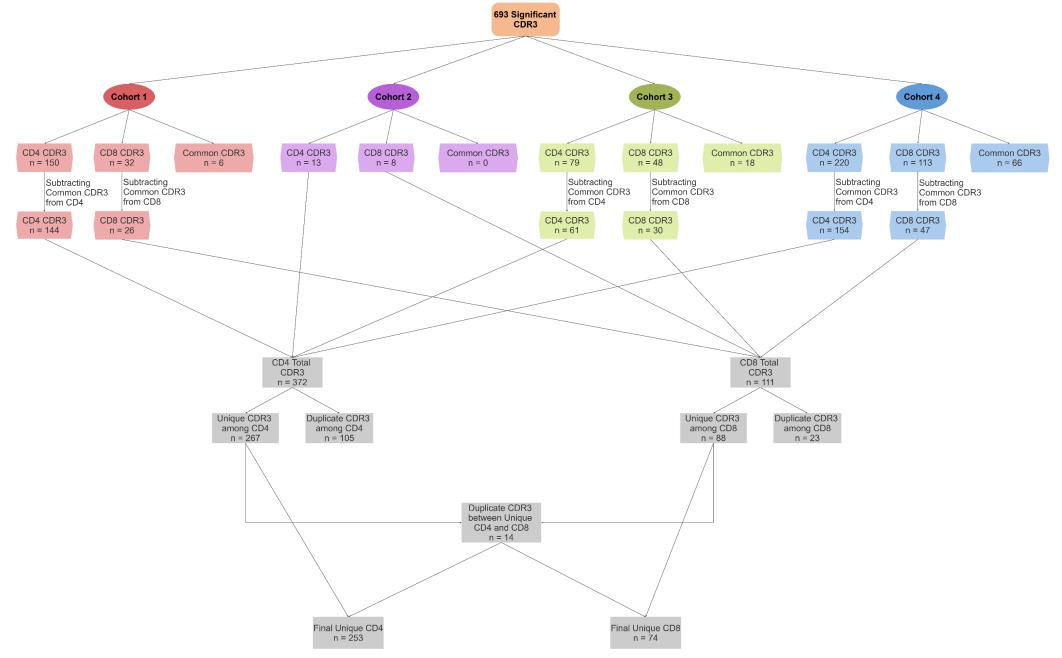
## **Supplementary Figure 6**



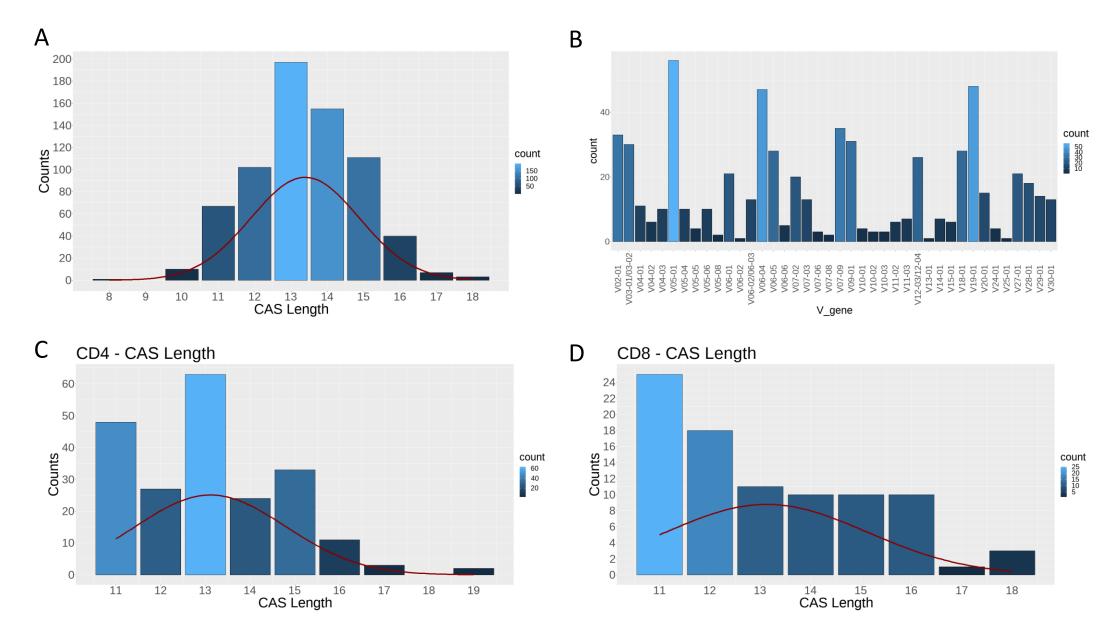
**Supplementary Figure 6**: Scatter plots comparing the Shannon Diversity **(A)** and Pielou Evenness **(B)** between moderate (green, n=10), severe (red, n=8), and mild (blue, n=35) COVID-19 severity cohorts. The dotted black line indicates the 80<sup>th</sup> percentile in each dataset. Graphs were prepared using R software.



**Supplementary Figure 7**. Flowchart of the total number of TRBV CDR3 sequences defined during the intersectional matrix analysis used to identify shared sequences between participants.



**Supplementary Figure 8. Prevalence of severity associated TRBV CDR3** in the CD4+ or CD8+ T cell compartment. TRBV CDR3 sequences identified in four different cohorts contained separate CD4+ and CD8+ population analysis were assessed for the presence of the severity-associated TRBV sequences. Common TRBV CDR3 sequences in both CD4+ and CD8+ populations were subtracted from the final analysis.



**Supplementary Figure 9**. Bar graphs displaying the **(A)** standard length distribution and **(B)** distribution of V-gene usage for the TRBV CDR3 (n=693) identified to be significantly increased in association with disease severity. And, the standard length distribution for **(C)** SARS-CoV-2-specific CD4 TRBV CRD3 and **(D)** SARS-CoV-2-specific CD8 TRBV CDR3 sequences.