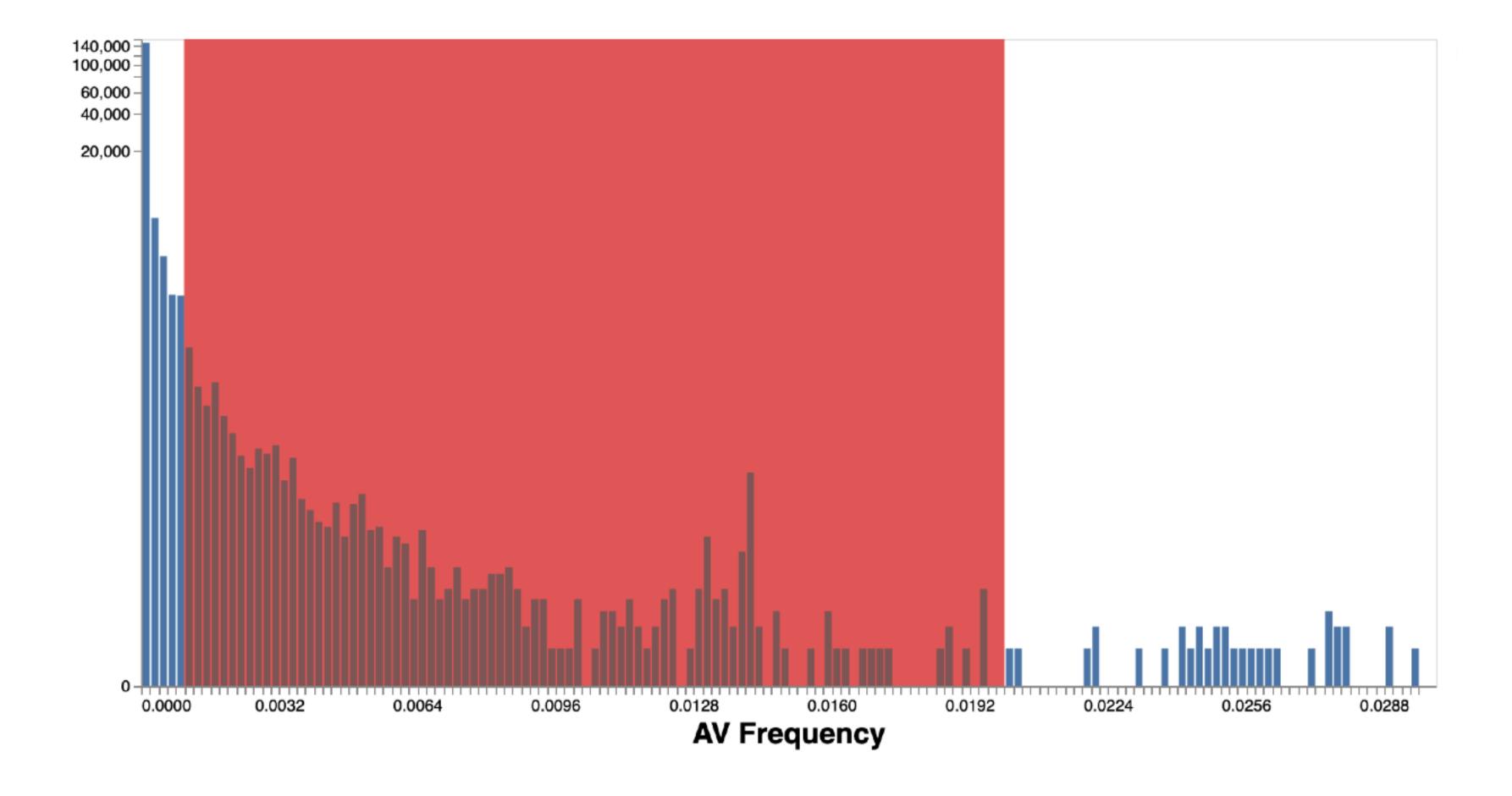
## Which AVs are interesting and non-obvious?

- Less common variants (e.g. 0.1% 2%) are potentially interesting and non-obvious
- Too many (>2000) to test exhaustively



## Analytical framework

- Key mutations have significant phenotypic and epidemiological significance.
- Analysis of substitution patterns in large sets of SARS-CoV-2 data can reveal genomic sites subject to selective pressure => possibly important
- Used a suite of stock and modified dN/dS methods for coding sequence evolution developed over ~2 decades and extensively used in other RNA viruses
- The methods are implemented in the HyPhy software package (www.hyphy.org)
- Our public servers (<u>datamonkey.org</u>) have processed >10,000 complex CoV evolutionary analyses from researchers worldwide.
- Spent considerable effort to scale up analyses and develop data reduction techniques to manage data sizes.
- Collaborative efforts, many other open source tools at covid-19.galaxyproject.org