Questions about the previous material?

We covered the FEL and SLAC selection analyses, site-level methods

- Fixed Effects Likelihood (FEL)
 - A simpler alternative to MEME (looks for pervasive selection)
 - May be more suited for smaller datasets or datasets of low divergence
- Single Likelihood Ancestor Counting (SLAC)
 - A counting-based approach
 - Good for data exploration and visualization

It is not unexpected that site-level positive results can occur when a gene-level test does not yield a positive result

- Lack of power for the global test: if the proportion of sites under selection is very small, a mixture-model test, like BUSTED, will miss it.
- Model violations: MEME supplies much more flexible distributions of dN/dS over sites; compared to alignment-wide k-bin (k=3) BUSTED distribution.
- False positives at site-level: our site-level tests have good statistical properties, but each positive site result could be a false positive; FWER correction would make site-level tests too conservative.
- **Summary**: gene-level selection tests need a minimal proportion of sites to be under selection to be powered; site-level tests should not be used to make inferences about gene-level selection.