Alignment-wide estimates

- Using standard MLE approaches it is straightforward to obtain point estimates of $dN/dS := \beta/\alpha$
- Can also easily test whether or not dN/dS > 1, or < 1 using the likelihood ratio test (LRT)
- Codon models also support the concepts of synonymous and nonsynonymous distances between sequences using standard properties of Markov processes (exponentially distributed waiting times)

Three example datasets

West Nile Virus NS3 protein

- An interesting case study of how positive selection detection methods lead to testable hypotheses for function discovery
- Brault et al 2007, <u>A single positively selected West Nile viral mutation confers increased virogenesis in American crows</u>

HIV-1 transmission pair

- Partial *env* sequences from two epidemiologically linked individuals
- An example of multiple selective environments (source, recipient, transmission)

SARS-CoV-2 Spike

- Full length spike sequences chosen to represent viral diversity (circa mid 2021)
- Good example for analyzing selection in population samples with many "deadend" intra-host variants