Neutral expectation

- A random mutation is ~3 times more likely to be non-synonymous that synonymous, depending on the variety of factors, such as codon composition, transition/transversion ratios, etc.
- We need to **estimate** the proportion of <u>random</u> mutations that are synonymous, and use it as a reference to compute **dS**.
- In early literature, these quantities were codified as synonymous and non-synonymous "sites" and/or mutational opportunity.
- As a very crude approximation (assuming that third positions ~ synonymous), each codon has 1 synonymous and 2 non-synonymous sites.

Evolutionary Modes

Positive Selection (Diversifying)

dS < dN or $\omega := dN/dS > 1$

Negative Selection

 $dS > dN \text{ or } \omega < 1$

Neutral Evolution

 $dS \simeq dN \text{ or } \omega \simeq 1$