

## Fundamental Concepts of Computational Evolution

- ► Given Data: Sequence Alignments
- Markov Chain ruled by a Poisson Process
- ▶ Instantaneous Substitution Rate Matrix, Q
- Obtaining the Transition Probability Matrix, P (t)

$$P(t) = e^{Qt}$$

- Models defined by parameters attached to Substitution Matrix
- Representation of Trees
- Likelihood and its calculation on a tree

#### Sources



Scott V. Edwards.

Natural Selection and Phylogenetic Analysis.

PNAS, 106(22):8799-8800, June 2009.



J. Felsenstein.

Evolutionary Trees from DNA Sequences: A Maximum Likelihood Approach.

Journal of Molecular Evolution, 17:368-376, 1981.

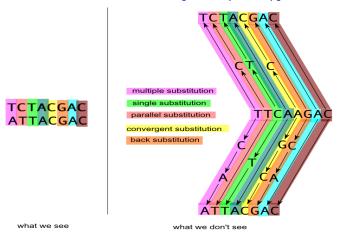


Ziheng Yang.

Computational Molecular Evolution.

Oxford University Press, 2006.

## Alignments: what we don't see [Yang(2006)]



- ▶ multiple hits: we deal with this via Chapman-Kolmogorov.
- convergent substitution, natural selection at work. See [Edwards(2009)].

## Chapman-Kolmogorov Equations

Referred to as **Pulley Principle** by [Felsenstein(1981)].

$$ho_{ij}(t)=\sum_k 
ho_{ik}(t_1)
ho_{kj}(t_2)$$

# Dealing with multiple hits

We get the Poisson distribution by approximating the binomial:

$$\Pr(K = k) = b(k; n, p) = \binom{n}{k} p^k (1 - p)^{n-k}$$

Look at example of K=0, with n large and p small so we way substitute  $\lambda=np$ :

$$b(0; n, p) = (1 - p)^n = \left(1 - \frac{\lambda}{n}\right)^n$$

Taking logs and using the Taylor expansion for log(1-x):

$$\log b(0; n, p) = n \log \left(1 - \frac{\lambda}{n}\right) = -\lambda - \frac{\lambda^2}{2n} - \dots$$

Truncating to first term, and redoing for K = 0, 1, 2, ..., k...:

$$b(0; n, p) \approx e^{-\lambda}; b(1; n, p) \approx \lambda e^{-\lambda}; b(k; n, p) \approx \frac{\lambda^k}{k!} e^{-\lambda};$$