Adaptive dating and fast proposals: revisiting the phylogenetic relaxed clock model

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## S1 Appendix: Rate quantiles

## 1 Piecewise linear approximation

In this article we introduced a linear piecewise approximation of the i-CDF (inverse cumulative distribution function) to improve the computational performance of the quant parameterisation. Let  $\hat{F}^{-1}(\mathcal{R}_i)$  be the piecewise approximation of the i-CDF  $F^{-1}(\mathcal{R}_i)$ . The approximation consists of n pieces (where n = 100 is fixed). Due to the nonlinear nature of small and large quantiles in a Log-normal distribution, the first and last pieces are not linear approximations but rather equal to the underlying distribution itself.

$$\hat{F}^{-1}(q) = \begin{cases} F^{-1}(q) & \text{if } q \leq \frac{1}{n} \text{ or } q \geq \frac{n-1}{n} \\ F^{-1}(\lfloor v \rfloor) + \left( F^{-1}(\lfloor v \rfloor + 1) - F^{-1}(\lfloor v \rfloor) \right) \left( v - \lfloor v \rfloor \right) & \text{otherwise.} \end{cases}$$
(1)

where v = q(n-1) indexes quantile q into piece number  $\lfloor v \rfloor$ . Values from the underlying function  $F^{-1}$  are cached, enabling rapid computation.

## 2 Tree operators for rate quantiles

Zhang and Drummond 2020 introduced several tree operators for the *real* parameterisation – including ConstantDistance, SimpleDistance, and SmallPulley [1]. In this appendix, these three operators are extended to the *quant* parameterisation. Following the notation presented in the main article, let  $t_i$  be the time of node i, let  $0 < q_i < 1$  be the rate quantile of node i, and let  $r_i = \hat{F}^{-1}(q_i)$  be the real rate of node i where  $\hat{F}^{-1}$  is the linear approximation of the i-CDF.

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#### Constant Distance

Let  $\mathcal{X}$  be a uniformly-at-random sampled internal node on tree  $\mathcal{T}$ . Let  $\mathcal{L}$  and  $\mathcal{R}$  be the left and right child of  $\mathcal{X}$ , respectively, and let  $\mathcal{P}$  be the parent of  $\mathcal{X}$ . Under the *quant* parameterisation, the ConstantDistance operator works as follows:

Step 1. Propose a new height for  $t_{\chi}$ :

$$t_{\chi}' \leftarrow t_{\chi} + s\Sigma \tag{2}$$

where  $\Sigma$  is drawn from a proposal transition distribution (Uniform or Bactrian), and s is a tunable step size. Ensure that  $\max\{t_{\mathcal{L}}, t_{\mathcal{R}}\} < t_{\mathcal{X}}' < t_{\mathcal{P}}$ , and if the constraint is broken then reject the proposal.

Step 2. Recalculate  $q_{\mathcal{X}}$  as:

$$q_{\mathcal{X}'} \leftarrow \hat{F}\left(r_{\mathcal{X}'}\right)$$

$$\leftarrow \hat{F}\left(\frac{t_{\mathcal{P}} - t_{\mathcal{X}}}{t_{\mathcal{P}} - t_{\mathcal{X}'}}r_{\mathcal{X}}\right)$$

$$\leftarrow \hat{F}\left(\frac{t_{\mathcal{P}} - t_{\mathcal{X}}}{t_{\mathcal{P}} - t_{\mathcal{X}'}}F^{-1}(q_{\mathcal{X}})\right).$$

$$(3)$$

This ensures that the genetic distance between  $\mathcal{X}$  and P remains constant after the operation by enforcing the constraint:

$$r_{\mathcal{X}}(t_{\mathcal{P}} - t_{\mathcal{X}}) = r_{\mathcal{X}}'(t_{\mathcal{P}} - t_{\mathcal{X}}'). \tag{4}$$

Step 3. Similarly, propose new rate quantiles for the two children  $\mathcal{C} \in \{\mathcal{L}, \mathcal{R}\}$ :

$$q_{\mathcal{C}'} \leftarrow \hat{F}\left(r_{\mathcal{C}'}\right)$$

$$\leftarrow \hat{F}\left(\frac{t_{\mathcal{X}} - t_{\mathcal{C}}}{t_{\mathcal{X}'} - t_{\mathcal{C}}} \times r_{\mathcal{C}}\right)$$

$$\leftarrow \hat{F}\left(\frac{t_{\mathcal{X}} - t_{\mathcal{C}}}{t_{\mathcal{X}'} - t_{\mathcal{C}}} \times \hat{F}^{-1}(q_{\mathcal{C}})\right).$$

$$(5)$$

Ensure that  $0 < q_i' < 1$  for all proposed nodes  $i \in \{\mathcal{X}, L, R\}$ , and if the constraint is broken then reject the proposal. This constraint can only be broken from numerical issues.

 $\underline{Step\ 4}$ . Finally, in order to calculate the Metropolis-Hastings-Green ratio, return the determinant of the Jacobian matrix:

$$J = \begin{bmatrix} \frac{\partial t_{\chi'}}{\partial t_{\chi}} & \frac{\partial t_{\chi'}}{\partial q_{\chi}} & \frac{\partial t_{\chi'}}{\partial q_{\zeta}} & \frac{\partial t_{\chi'}}{\partial q_{R}} \\ \frac{\partial q_{\chi'}}{\partial q_{\chi'}} & \frac{\partial q_{\chi'}}{\partial q_{\chi}} & \frac{\partial q_{\chi'}}{\partial q_{\zeta}} & \frac{\partial q_{\chi'}}{\partial q_{R}} \\ \frac{\partial q_{\zeta'}}{\partial t_{\chi}} & \frac{\partial q_{\zeta'}}{\partial q_{\chi}} & \frac{\partial q_{\zeta'}}{\partial q_{\zeta}} & \frac{\partial q_{\zeta'}}{\partial q_{R}} \\ \frac{\partial q_{R'}}{\partial t_{\chi}} & \frac{\partial q_{R'}}{\partial q_{\chi}} & \frac{\partial q_{R'}}{\partial q_{\zeta}} & \frac{\partial q_{R'}}{\partial q_{R}} \end{bmatrix}$$

$$= \begin{bmatrix} \frac{\partial t_{\chi'}}{\partial t_{\chi}} & 0 & 0 & 0 \\ \frac{\partial q_{\chi'}}{\partial t_{\chi}} & \frac{\partial q_{\chi'}}{\partial q_{\chi}} & 0 & 0 \\ \frac{\partial q_{\chi'}}{\partial t_{\chi}} & \frac{\partial q_{\chi'}}{\partial q_{\chi}} & 0 & 0 \\ \frac{\partial q_{\zeta'}}{\partial t_{\chi}} & 0 & \frac{\partial q_{\zeta'}}{\partial q_{\zeta}} & 0 \\ \frac{\partial q_{\zeta'}}{\partial t_{\chi}} & 0 & 0 & \frac{\partial q_{\zeta'}}{\partial q_{\zeta}} \end{bmatrix}.$$

$$(6)$$

As J is triangular, its determinant |J| is equal to the product of diagonal elements:

$$\ln|J| = \ln\left\{\frac{\partial t_{\mathcal{X}'}}{\partial t_{\mathcal{X}}} \times \frac{\partial q_{\mathcal{X}'}}{\partial q_{\mathcal{X}}} \times \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{L}}} \times \frac{\partial q_{\mathcal{R}'}}{\partial q_{\mathcal{R}}}\right\} 
= \ln 1 + \ln D\hat{F}\left(\frac{t_{\mathcal{P}} - t_{\mathcal{X}}}{t_{\mathcal{P}} - t_{\mathcal{X}'}} \times \hat{F}^{-1}(q_{\mathcal{X}})\right) + \ln \frac{\partial}{\partial q_{\mathcal{X}}} \frac{t_{\mathcal{P}} - t_{\mathcal{X}}}{t_{\mathcal{P}} - t_{\mathcal{X}'}} \hat{F}^{-1}(q_{\mathcal{X}}) 
+ \ln D\hat{F}\left(\frac{t_{\mathcal{X}} - t_{\mathcal{L}}}{t_{\mathcal{X}'} - t_{\mathcal{L}}} \times \hat{F}^{-1}(q_{\mathcal{L}})\right) + \ln \frac{\partial}{\partial q_{\mathcal{L}}} \frac{t_{\mathcal{X}} - t_{\mathcal{L}}}{t_{\mathcal{X}'} - t_{\mathcal{L}}} \hat{F}^{-1}(q_{\mathcal{L}}) 
+ \ln D\hat{F}\left(\frac{t_{\mathcal{X}} - t_{\mathcal{R}}}{t_{\mathcal{X}'} - t_{\mathcal{R}}} \times \hat{F}^{-1}(q_{\mathcal{R}})\right) + \ln \frac{\partial}{\partial q_{\mathcal{R}}} \frac{t_{\mathcal{X}} - t_{\mathcal{R}}}{t_{\mathcal{X}'} - t_{\mathcal{R}}} \hat{F}^{-1}(q_{\mathcal{R}}) 
= \ln D\hat{F}\left(\frac{t_{\mathcal{Y}} - t_{\mathcal{X}}}{t_{\mathcal{P}} - t_{\mathcal{X}'}} \times \hat{F}^{-1}(q_{\mathcal{X}})\right) + \ln D\hat{F}^{-1}(q_{\mathcal{X}}) + \ln \frac{t_{\mathcal{P}} - t_{\mathcal{X}'}}{t_{\mathcal{P}} - t_{\mathcal{X}'}} 
+ \ln D\hat{F}\left(\frac{t_{\mathcal{X}} - t_{\mathcal{L}}}{t_{\mathcal{X}'} - t_{\mathcal{L}}} \times \hat{F}^{-1}(q_{\mathcal{R}})\right) + \ln D\hat{F}^{-1}(q_{\mathcal{L}}) + \ln \frac{t_{\mathcal{X}} - t_{\mathcal{L}}}{t_{\mathcal{X}'} - t_{\mathcal{L}}} 
+ \ln D\hat{F}\left(\frac{t_{\mathcal{X}} - t_{\mathcal{R}}}{t_{\mathcal{X}'} - t_{\mathcal{R}}} \times \hat{F}^{-1}(q_{\mathcal{R}})\right) + \ln D\hat{F}^{-1}(q_{\mathcal{R}}) + \ln \frac{t_{\mathcal{X}} - t_{\mathcal{R}}}{t_{\mathcal{X}'} - t_{\mathcal{R}}}. \tag{7}$$

The derivatives  $D\hat{F}$  and  $D\hat{F}^{-1}$  are computed using numerical approximations for the first and last pieces, or as the gradient of the linear approximation for internal pieces. As its final step, the operator returns  $\ln |J|$ .

### Simple Distance

While ConstantDistance proposes internal node heights, SimpleDistance operates on the root. Let  $\mathcal{X}$  be the root node and let  $\mathcal{L}$  and  $\mathcal{R}$  be its two children.

<u>Step 1</u>. Propose a new height for  $t_{\mathcal{X}}$ :

$$t_{\mathcal{X}}' \leftarrow t_{\mathcal{X}} + s\Sigma. \tag{8}$$

Ensure that  $\max\{t_{\mathcal{L}}, t_{\mathcal{R}}\} < t_{\mathcal{X}}'$ , and if the constraint is broken then reject the proposal.

<u>Step 2</u>. Propose new rate quantiles for the two children  $\mathcal{C} \in \{\mathcal{L}, \mathcal{R}\}$ :

$$q_{\mathcal{C}'} \leftarrow \hat{F}\left(r_{\mathcal{C}'}\right)$$

$$\leftarrow \hat{F}\left(\frac{t_{\mathcal{X}} - t_{\mathcal{C}}}{t_{\mathcal{X}'} - t_{\mathcal{C}}} \times r_{\mathcal{C}}\right)$$

$$\leftarrow \hat{F}\left(\frac{t_{\mathcal{X}} - t_{\mathcal{C}}}{t_{\mathcal{X}'} - t_{\mathcal{C}}} \times \hat{F}^{-1}(q_{\mathcal{C}})\right).$$

$$(9)$$

These proposals ensure that the genetic distance between  $\mathcal{X}$  and its children  $\mathcal{C}$  remain constant after the operation by enforcing the constraint:

$$r_{\mathcal{C}}(t_{\mathcal{X}} - t_{\mathcal{C}}) = r_{\mathcal{C}}'(t_{\mathcal{X}}' - t_{\mathcal{C}}). \tag{10}$$

Ensure that  $0 < q_C' < 1$ , and if the constraint is broken then reject the proposal. <u>Step 3</u>. Finally, in order to calculate the Metropolis-Hastings-Green ratio, return the determinant of the Jacobian matrix:

$$J = \begin{bmatrix} \frac{\partial t_{\chi'}}{\partial t_{\chi}} & \frac{\partial t_{\chi'}}{\partial q_{\mathcal{L}}} & \frac{\partial t_{\chi'}}{\partial q_{\mathcal{L}}} \\ \frac{\partial q_{\mathcal{L}'}}{\partial t_{\chi}} & \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{L}}} & \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{R}}} \\ \frac{\partial q_{\mathcal{R}'}}{\partial t_{\chi}} & \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{L}}} & \frac{\partial q_{\mathcal{R}'}}{\partial q_{\mathcal{R}}} \end{bmatrix}$$

$$= \begin{bmatrix} \frac{\partial t_{\chi'}}{\partial t_{\chi}} & 0 & 0 \\ \frac{\partial q_{\mathcal{L}'}}{\partial t_{\chi}} & \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{L}}} & 0 \\ \frac{\partial q_{\mathcal{R}'}}{\partial t_{\chi}} & 0 & \frac{\partial q_{\mathcal{R}'}}{\partial q_{\mathcal{R}}} \end{bmatrix} . \tag{11}$$

As J is triangular, its determinant |J| is equal to the product of diagonal elements:

$$\ln |J| = \ln \left\{ \frac{\partial t_{\mathcal{X}'}}{\partial t_{\mathcal{X}}} \times \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{L}}} \times \frac{\partial q_{\mathcal{R}'}}{\partial q_{\mathcal{R}}} \right\} 
= \ln \frac{\partial t_{\mathcal{X}'}}{\partial t_{\mathcal{X}}} + \ln \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{L}}} + \ln \frac{\partial q_{\mathcal{R}'}}{\partial q_{\mathcal{R}}} 
= \ln 1 
+ \ln D \hat{F} \left( \frac{t_{\mathcal{X}} - t_{\mathcal{L}}}{t_{\mathcal{X}'} - t_{\mathcal{L}}} \times \hat{F}^{-1}(q_{\mathcal{L}}) \right) + \ln \frac{\partial}{\partial q_{\mathcal{L}}} \frac{t_{\mathcal{X}} - t_{\mathcal{L}}}{t_{\mathcal{X}'} - t_{\mathcal{L}}} \hat{F}^{-1}(q_{\mathcal{L}}) 
+ \ln D \hat{F} \left( \frac{t_{\mathcal{X}} - t_{\mathcal{R}}}{t_{\mathcal{X}'} - t_{\mathcal{R}}} \times \hat{F}^{-1}(q_{\mathcal{R}}) \right) + \ln \frac{\partial}{\partial q_{\mathcal{R}}} \frac{t_{\mathcal{X}} - t_{\mathcal{R}}}{t_{\mathcal{X}'} - t_{\mathcal{R}}} \hat{F}^{-1}(q_{\mathcal{R}}) 
= \ln D \hat{F} \left( \frac{t_{\mathcal{X}} - t_{\mathcal{L}}}{t_{\mathcal{X}'} - t_{\mathcal{L}}} \times \hat{F}^{-1}(q_{\mathcal{L}}) \right) + \ln D \hat{F}^{-1}(q_{\mathcal{L}}) + \ln \frac{t_{\mathcal{X}} - t_{\mathcal{L}}}{t_{\mathcal{X}'} - t_{\mathcal{L}}} 
+ \ln D \hat{F} \left( \frac{t_{\mathcal{X}} - t_{\mathcal{R}}}{t_{\mathcal{X}'} - t_{\mathcal{R}}} \times \hat{F}^{-1}(q_{\mathcal{R}}) \right) + \ln D \hat{F}^{-1}(q_{\mathcal{R}}) + \ln \frac{t_{\mathcal{X}} - t_{\mathcal{R}}}{t_{\mathcal{X}'} - t_{\mathcal{R}}}. \tag{12}$$

As its final step, the operator returns  $\ln |J|$ .

#### Small Pulley

Just like the previous operator, SmallPulley operates on the root. Let  $\mathcal{X}$  be the root node and let  $\mathcal{L}$  and  $\mathcal{R}$  be its two children. However, unlike SimpleDistance, this operator alters the two genetic distances  $d_{\mathcal{L}} = r_{\mathcal{L}}(t_{\mathcal{X}} - t_{\mathcal{L}}) = \hat{F}^{-1}(q_{\mathcal{L}})(t_{\mathcal{X}} - t_{\mathcal{L}})$  and  $d_{\mathcal{R}} = r_{\mathcal{R}}(t_{\mathcal{X}} - t_{\mathcal{R}}) = \hat{F}^{-1}(q_{\mathcal{R}})(t_{\mathcal{X}} - t_{\mathcal{R}})$ , while conserving their sum  $d_{\mathcal{L}} + d_{\mathcal{R}}$ .

Step 1. Propose new genetic distances for  $d_{\mathcal{L}}$  and  $d_{\mathcal{R}}$ :

$$d_{\mathcal{L}}' \leftarrow d_{\mathcal{L}} + s\Sigma \tag{13}$$

$$d_{\mathcal{R}}' \leftarrow d_{\mathcal{R}} - s\Sigma \tag{14}$$

Ensure that  $0 < d_{\mathcal{L}}' < d_{\mathcal{L}} + d_{\mathcal{R}}$ , and if the constraint is broken then reject the proposal.

Step 2. Propose new rate quantiles for the two children  $\mathcal{L}$  and  $\mathcal{R}$ :

$$q_{\mathcal{L}'} \leftarrow \hat{F}\left(\frac{d_{\mathcal{L}'}}{t_{\mathcal{X}} - t_{\mathcal{L}}}\right)$$

$$\leftarrow \hat{F}\left(\frac{\hat{F}^{-1}(q_{\mathcal{L}})(t_{\mathcal{X}} - t_{\mathcal{L}}) + \Sigma}{t_{\mathcal{X}} - t_{\mathcal{L}}}\right)$$

$$q_{\mathcal{R}'} \leftarrow \hat{F}\left(\frac{d_{\mathcal{R}'}}{t_{\mathcal{X}} - t_{\mathcal{R}}}\right)$$

$$\leftarrow \hat{F}\left(\frac{\hat{F}^{-1}(q_{\mathcal{R}})(t_{\mathcal{X}} - t_{\mathcal{R}}) - \Sigma}{t_{\mathcal{X}} - t_{\mathcal{R}}}\right).$$
(15)

Step 3. Return the determinant of the Jacobian matrix:

$$J = \begin{bmatrix} \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{L}}} & \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{R}}} \\ \frac{\partial q_{\mathcal{R}'}}{\partial q_{\mathcal{L}}} & \frac{\partial q_{\mathcal{R}'}}{\partial q_{\mathcal{R}}} \end{bmatrix}$$

$$= \begin{bmatrix} \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{L}}} & 0 \\ 0 & \frac{\partial q_{\mathcal{R}'}}{\partial q_{\mathcal{R}}} \end{bmatrix}. \tag{17}$$

As J is triangular/diagonal, its determinant |J| is equal to the product of diagonal elements:

$$\ln |J| = \ln \left\{ \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{L}}} \times \frac{\partial q_{\mathcal{R}'}}{\partial q_{\mathcal{R}}} \right\} 
= \ln \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{L}}} + \ln \frac{\partial q_{\mathcal{R}'}}{\partial q_{\mathcal{R}}} 
= \ln D \hat{F} \left( \frac{\hat{F}^{-1}(q_{\mathcal{L}})(t_{\mathcal{X}} - t_{\mathcal{L}}) + \Sigma}{t_{\mathcal{X}} - t_{\mathcal{L}}} \right) + \ln \frac{\partial q_{\mathcal{L}'}}{\partial q_{\mathcal{L}}} \frac{\hat{F}^{-1}(q_{\mathcal{L}})(t_{\mathcal{X}} - t_{\mathcal{L}}) + \Sigma}{t_{\mathcal{X}} - t_{\mathcal{L}}} 
+ \ln D \hat{F} \left( \frac{\hat{F}^{-1}(q_{\mathcal{R}})(t_{\mathcal{X}} - t_{\mathcal{R}}) - \Sigma}{t_{\mathcal{X}} - t_{\mathcal{R}}} \right) + \ln \frac{\partial q_{\mathcal{R}'}}{\partial q_{\mathcal{R}}} \frac{\hat{F}^{-1}(q_{\mathcal{R}})(t_{\mathcal{X}} - t_{\mathcal{R}}) - \Sigma}{t_{\mathcal{X}} - t_{\mathcal{R}}} 
= \ln D \hat{F} \left( \frac{\hat{F}^{-1}(q_{\mathcal{L}})(t_{\mathcal{X}} - t_{\mathcal{L}}) + \Sigma}{t_{\mathcal{X}} - t_{\mathcal{L}}} \right) + \ln D \hat{F}^{-1}(q_{\mathcal{L}}) 
+ \ln D \hat{F} \left( \frac{\hat{F}^{-1}(q_{\mathcal{R}})(t_{\mathcal{X}} - t_{\mathcal{R}}) - \Sigma}{t_{\mathcal{X}} - t_{\mathcal{R}}} \right) + \ln D \hat{F}^{-1}(q_{\mathcal{R}}). \tag{18}$$

Thus, as its final step, the operator returns  $\ln |J|$ .

# 3 CisScale operator

CisScale was originally introduced by Zhang and Drummond 2020 for the *real* parameterisation (therein named ucldstdevScaleOperator). Under the *quant* configuration, the CisScale operator works as follows.

Step 1. Propose a new value for the relaxed clock standard deviation  $\sigma$ 

$$\sigma' \leftarrow \sigma \times e^{s\Sigma}.\tag{19}$$

<u>Step 2</u>. Recalculate all branch substitution rate quantiles q such that their rates r remain constant

$$let r = \hat{F}^{-1}(q|\sigma) \tag{20}$$

$$let r' = r (21)$$

$$q' \leftarrow \hat{F}(r'|\sigma') = \hat{F}(\hat{F}^{-1}(q|\sigma)|\sigma'). \tag{22}$$

<u>Step 3</u>. Return the log Hastings-Green ratio of this proposal. If  $\Sigma$  was drawn from a symmetric proposal kernel (such as the Bactrian distribution), this is equal to:

$$|J| = \log(e^{s\Sigma}) + \log\left(\frac{\delta}{\delta q}\hat{F}(\hat{F}^{-1}(q|\sigma)|\sigma')\right)$$
(23)

$$= s\Sigma + \log D\hat{F}(\hat{F}^{-1}(q|\sigma)|\sigma') + \log D\hat{F}^{-1}(q|\sigma), \tag{24}$$

where derivatives  $D\hat{F}$  and  $D\hat{F}^{-1}$  can be approximated using either the piecewise linear model or standard numerical libraries.

# 4 Narrow exchange rates

The NarrowExchangeRate operator is also compatible with rate quantiles. This operator behaves the same as presented in the main article however the Hastings-Green ratio requires further augmentation due to changes in dimension throughout the proposal.

- <u>Step 1</u>. Apply NarrowExchange to the current tree topology as described in the main article. This will return a Hastings ratio H due to the asymmetry of this proposal.
- <u>Step 2</u>. Compute the relevant branch rates  $r_i$  for  $r \in \{A, B, C, D\}$  of the current state from their respective quantile parameters.

$$r_i = \hat{F}^{-1}(q_i). {25}$$

<u>Step 3</u>. Propose new rates and node heights and compute the Hastings-Green ratio of the real-space component of the proposal (e.g. Algorithms 1-2 of the main article).

$$(r'_A, r'_B, r'_C, r'_D, t'_D, |J_r|) \leftarrow \text{PROPOSAL}(r_A, r_B, r_C, r_D, t_D).$$
 (26)

Step 4. Transform the rates back into quantiles.

$$q_i' = \hat{F}(r_i'). \tag{27}$$

 $\underline{Step\ 5}$ . Compute the log Hastings-Green ratio of the interconversion between rates and quantiles.

$$\log |J_q| = \log \hat{F}(q) + \log \hat{F}^{-1}(r'). \tag{28}$$

<u>Step 6</u>. Return the total log Hastings-Green ratio of this proposal:  $\log H + \log |J_r| + \log |J_q|$ .

## References

[1] Zhang R, Drummond A. Improving the performance of Bayesian phylogenetic inference under relaxed clock models. BMC Evolutionary Biology. 2020;20:1–28.