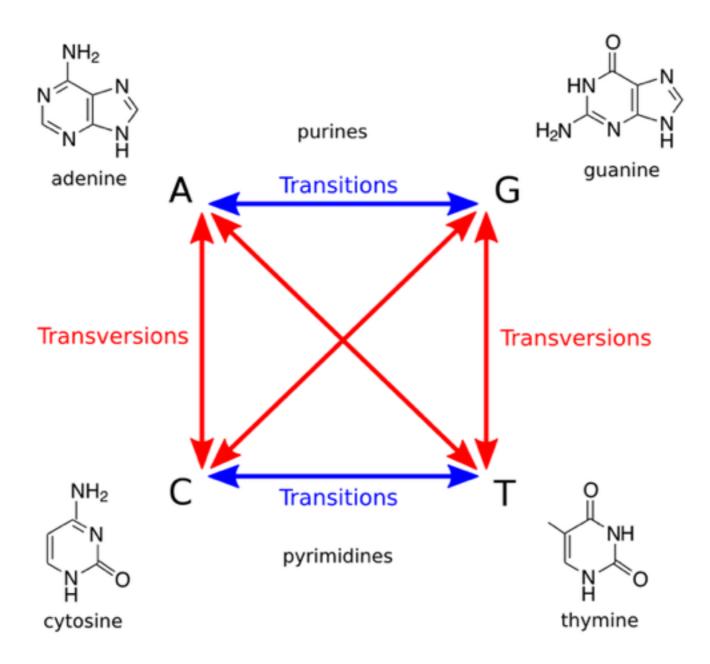
Y

| | | ϵ | G | C | T | Α | T | Α | C |
|-----|------------|------------|---|---|---|---|---|---|---|
| | ϵ | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 |
| | G | 1 | 0 | 1 | 2 | 3 | 4 | 5 | 6 |
| | C | 2 | 1 | 0 | 1 | 2 | 3 | 4 | 5 |
| X | G | 3 | 1 | 1 | 1 | 2 | 3 | 4 | 5 |
| 7 (| Т | 4 | 2 | 2 | 1 | 2 | 3 | 4 | 5 |
| | Α | 5 | 3 | 3 | 2 | 1 | 2 | 3 | 4 |
| | Т | 6 | 4 | 4 | 3 | 2 | 1 | 2 | 3 |
| | G | 7 | 5 | 5 | 4 | 3 | 2 | 2 | 3 |
| | C | 8 | 6 | 5 | 5 | 4 | 3 | 3 | 2 |



Human transition to transversion ratio (AKA ti/tv) is ~2.1



Human substitution rate ≈ 1 in 1,000



Small-gap rate is ≈ 1 in 3,000

Penalty matrix

| | Α | С | G | Т | _ |
|---|---|---|---|---|---|
| Α | 0 | 4 | 2 | 4 | 8 |
| С | 4 | 0 | 4 | 2 | 8 |
| G | 2 | 4 | 0 | 4 | 8 |
| T | 4 | 2 | 4 | 0 | 8 |
| _ | 8 | 8 | 8 | 8 | |

- 2 Transitions (A↔ G, C ↔ T)
- 4 Transversions
- 8 Gaps

$$\mathbf{edist}(\alpha x, \beta y) = \min \left\{ \begin{array}{l} \mathbf{edist}(\alpha, \beta) + \delta(x, y) \\ \mathbf{edist}(\alpha x, \beta) + 1 \\ \mathbf{edist}(\alpha, \beta y) + 1 \end{array} \right.$$

$$\mathbf{galign}(\alpha \mathbf{x}, \beta \mathbf{y}) = \min \left\{ \begin{array}{l} \mathbf{galign}(\alpha, \beta) + p(\mathbf{x}, \mathbf{y}) \\ \mathbf{galign}(\alpha \mathbf{x}, \beta) + p(\mathbf{x}, \mathbf{y}) \\ \mathbf{galign}(\alpha, \beta \mathbf{y}) + p(\mathbf{-}, \mathbf{y}) \end{array} \right.$$
Use penalty matrix

Global alignment

| | ϵ | T | Α | T | G | T | C | Α | T | G | C |
|------------|------------|----|----|----|----|----|----|----|----|----|----|
| ϵ | 0 | 8 | 16 | 24 | 32 | 40 | 48 | 56 | 64 | 72 | 80 |
| Т | 8 | 0 | 8 | 16 | 24 | 32 | 40 | 48 | 56 | 64 | 72 |
| A | 16 | 8 | 0 | 8 | 16 | 24 | 32 | 40 | 48 | 56 | 64 |
| C | 24 | 16 | 8 | 2 | 10 | 18 | 24 | 32 | 40 | 48 | 56 |
| G | 32 | 24 | 16 | 10 | 2 | 10 | 18 | 26 | 34 | 40 | 48 |
| T | 40 | 32 | 24 | 16 | 10 | 2 | 10 | 18 | 26 | 34 | 42 |
| C | 48 | 40 | 32 | 24 | 18 | 10 | 2 | 10 | 18 | 26 | 34 |
| A | 56 | 48 | 40 | 32 | 26 | 18 | 10 | 2 | 10 | 18 | 26 |
| G | 64 | 56 | 48 | 40 | 32 | 26 | 18 | 10 | 6 | 10 | 18 |
| C | 72 | 64 | 56 | 48 | 40 | 34 | 26 | 18 | 12 | 10 | 10 |

| | Α | С | G | Т | _ |
|---|---|---|---|---|---|
| Α | 0 | 4 | 2 | 4 | 8 |
| C | 4 | 0 | 4 | 2 | 8 |
| G | 2 | 4 | 0 | 4 | 8 |
| Т | 4 | 2 | 4 | 0 | 8 |
| _ | 8 | 8 | 8 | 8 | |

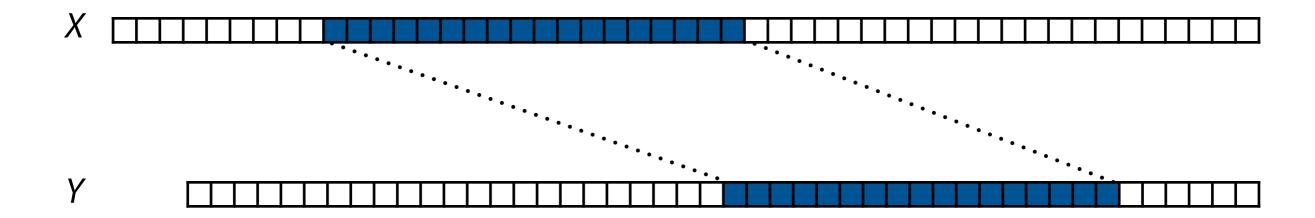
Global alignment

| | ϵ | T | Α | Т | G | Т | C | Α | Т | G | C |
|------------|------------|----|----|----|----|----|----|----|----|----|----|
| ϵ | 9 | 8 | 16 | 24 | 32 | 40 | 48 | 56 | 64 | 72 | 80 |
| Τ | 8 | 8 | 8 | 16 | 24 | 32 | 40 | 48 | 56 | 64 | 72 |
| A | 16 | 8 | 8 | 8 | 16 | 24 | 32 | 40 | 48 | 56 | 64 |
| C | 24 | 16 | 8 | 5 | 10 | 18 | 24 | 32 | 40 | 48 | 56 |
| G | 32 | 24 | 16 | 10 | 3 | 10 | 18 | 26 | 34 | 40 | 48 |
| Т | 40 | 32 | 24 | 16 | 10 | 3 | 10 | 18 | 26 | 34 | 42 |
| C | 48 | 40 | 32 | 24 | 18 | 10 | 3 | 10 | 18 | 26 | 34 |
| A | 56 | 48 | 40 | 32 | 26 | 18 | 10 | 15 | 10 | 18 | 26 |
| G | 64 | 56 | 48 | 40 | 32 | 26 | 18 | 10 | 6 | 10 | 18 |
| C | 72 | 64 | 56 | 48 | 40 | 34 | 26 | 18 | 12 | 10 | |

| | Α | С | G | Т | _ |
|---|---|---|---|---|---|
| Α | 0 | 4 | 2 | 4 | 8 |
| C | 4 | 0 | 4 | 2 | 8 |
| G | 2 | 4 | 0 | 4 | 8 |
| Т | 4 | 2 | 4 | 0 | 8 |
| _ | 8 | 8 | 8 | 8 | |

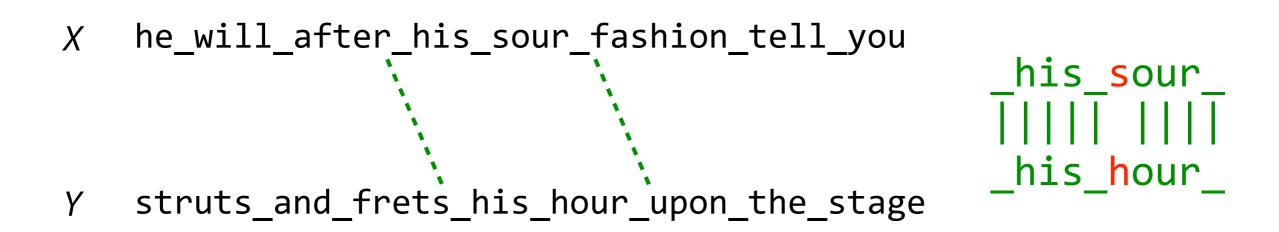
Local alignment

Find the most similar pair of substrings from X and Y



Local alignment

Find the most similar *pair of substrings* from X and Y



$$\label{eq:lalign} \begin{aligned} \textbf{lalign}(\alpha, \beta) + s(x, y) \\ \textbf{lalign}(\alpha x, \beta y) &= \max \left\{ \begin{array}{l} \textbf{lalign}(\alpha, \beta) + s(x, y) \\ \textbf{lalign}(\alpha x, \beta) + s(x, -) \\ \textbf{lalign}(\alpha, \beta y) + s(-, y) \\ 0 \end{array} \right. \end{aligned}$$

Scoring matrix: matches are positive, differences negative

| | А | С | G | Т | ı |
|---|----|----|----|----|----|
| Α | 2 | -4 | -4 | -4 | -6 |
| С | -4 | 2 | -4 | -4 | -6 |
| G | -4 | -4 | 2 | -4 | -6 |
| Т | -4 | -4 | -4 | 2 | -6 |
| _ | -6 | -6 | -6 | -6 | |

Y

| | | ϵ | Т | Α | Т | Α | Т | G | C | G | G | C | G | Т | Т | Т |
|---|------------|------------|---|---|---|---|---|---|----|---|---|----|----|---|----|---|
| | ϵ | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | G | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 2 | 2 | 0 | 2 | 0 | 0 | 0 |
| | G | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 2 | 4 | 0 | 2 | 0 | 0 | 0 |
| | Τ | 0 | 2 | 0 | 2 | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 4 | 2 | 2 |
| | Α | 0 | 0 | 4 | 0 | 4 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | Т | 0 | 2 | 0 | 6 | 0 | 6 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 2 | 2 |
| | G | 0 | 0 | 0 | 0 | 2 | 0 | 8 | 2 | 2 | 2 | 0 | 2 | 0 | 0 | 0 |
| | C | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 10 | 4 | 0 | 4 | 0 | 0 | 0 | 0 |
| V | Τ | 0 | 2 | 0 | 2 | 0 | 2 | 0 | 4 | 6 | 0 | 0 | 0 | 2 | 2 | 2 |
| Λ | G | 0 | 0 | 0 | 0 | 0 | 0 | 4 | 0 | 6 | 8 | 2 | 2 | 0 | 0 | 0 |
| | G | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 2 | 8 | 4 | 4 | 0 | 0 | 0 |
| | C | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 4 | 0 | 2 | 10 | 4 | 0 | 0 | 0 |
| | G | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 0 | 6 | 2 | 4 | 12 | 6 | 0 | 0 |
| | C | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 4 | 0 | 2 | 4 | 6 | 8 | 2 | 0 |
| | Т | 0 | 2 | 0 | 2 | 0 | 2 | 0 | 0 | 0 | 0 | 0 | 0 | 8 | 10 | 4 |
| | Α | 0 | 0 | 4 | 0 | 4 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 2 | 4 | 6 |
| | | · | | | | | · | | | | | | | | | |

Y

| E 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 | 0 (| 0 |
|---|-------|---|
| G Q Q Q Q Q Q Q Q Q Q Q Q Q Q Q Q Q Q Q | | 9 |
| | | U |
| G 0 0 0 0 0 0 2 0 2 4 0 2 0 | 0 6 | 0 |
| T 0 2 0 2 0 2 0 0 0 0 0 4 | 2 2 | 2 |
| A 0 0 4 0 4 0 0 0 0 0 0 0 0 | 0 0 | 0 |
| T 0 2 0 6 0 0 0 0 0 0 2 | 2 2 | 2 |
| G 0 0 0 0 2 0 2 2 2 0 2 0 | 0 0 | 0 |
| C 0 0 0 0 0 0 2 10 4 0 4 0 0 | 0 0 | 0 |
| T 0 2 0 2 0 2 0 4 6 0 0 2 | 2 2 | 2 |
| G 0 0 0 0 0 4 0 6 8 2 2 0 | 0 (| 0 |
| G 0 0 0 0 0 0 2 0 2 3 4 4 0 | 0 0 | 0 |
| C 0 0 0 0 0 0 0 4 0 2 10 4 0 | 0 0 | 0 |
| G 0 0 0 0 0 0 2 0 6 2 4 2 6 | 0 0 | 0 |
| C 0 0 0 0 0 0 0 4 0 2 4 6 8 | 2 (| 0 |
| T 0 2 0 2 0 2 0 0 0 0 0 8 | 10 4 | 4 |
| A 0 0 4 0 4 0 0 0 0 0 0 0 2 | 4 (| 6 |