ulm university universität **UUU**





On Reversal and Transposition Medians

Martin Bader | June 25, 2009

Genome Rearrangements

- During evolution, the gene order in a chromosome can change
- Gene order of two land snail mitochondrial DNAs

 $\begin{array}{c} \hline Cepaea \ nemoralis\\ \hline Coption Cepaea \ nemoralis\\ \hline Copti$

Genome Rearrangements

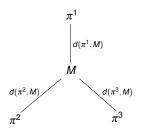
- During evolution, the gene order in a chromosome can change
- Gene order of two land snail mitochondrial DNAs

 $\begin{array}{c} \hline Cepaea \ nemoralis\\ \hline Coption Cepaea \ nemoralis\\ \hline Copti$

- Reconstruct evolutionary events
- Use as distance measure
- Use for phylogenetic reconstruction

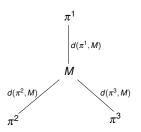
The Median Problem

- Given gene orders π^1, π^2, π^3
- Find *M* where $\sum_{i=1}^{3} d(\pi^{i}, M)$ is minimized



The Median Problem

- Given gene orders π^1, π^2, π^3
- Find *M* where $\sum_{i=1}^{3} d(\pi^{i}, M)$ is minimized



NP-hard even for the most simple distance measures

Our contribution

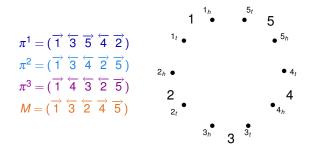
Exact algorithms for the Transposition Median Problem
Exact algorithm for the weighted Reversal and Transposition
Median Problem

(Extension of Reversal Median solver, Caprara 2003)

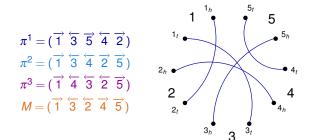
 Improved exact algorithm for pairwise distances (Improvement of Christie 1998)

- Edge-colored graph
- Contains neighborhood relations for each gene order

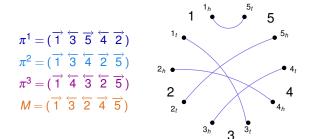
- Edge-colored graph
- Contains neighborhood relations for each gene order
- Example:



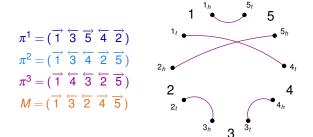
- Edge-colored graph
- Contains neighborhood relations for each gene order
- Example:



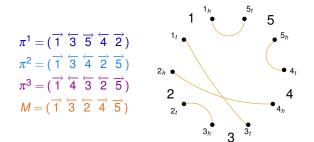
- Edge-colored graph
- Contains neighborhood relations for each gene order
- Example:



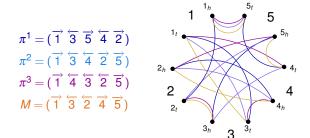
- Edge-colored graph
- Contains neighborhood relations for each gene order
- Example:



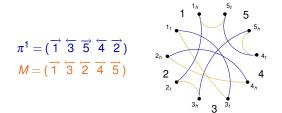
- Edge-colored graph
- Contains neighborhood relations for each gene order
- Example:



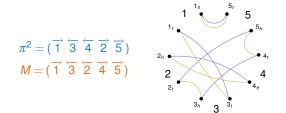
- Edge-colored graph
- Contains neighborhood relations for each gene order
- Example:



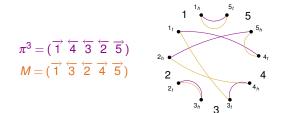
Edges of two colors form cycles



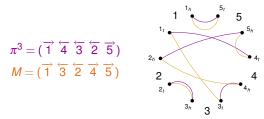
Edges of two colors form cycles



Edges of two colors form cycles



Edges of two colors form cycles



Distances closely related to number of cycles

$$egin{aligned} & d_r = n-c+h+f \ & d_t \geq rac{n-c_{odd}}{2} \ & d_w \geq rac{w_t}{2}(n-c_{odd}-(2-rac{2w_r}{w_t})c_{even}) \end{aligned}$$

Sketch of the algorithm

- Solve Cycle Median Problem
- Verify solution

Sketch of the algorithm

- Solve Cycle Median Problem
 - Start with empty M
 - Subsequently add edges
 - Estimate lower bound for partial solution
 - Continue with partial solution with least lower bound (branch and bound)
 - NEW: Consider cycle lengths
- Verify solution

Sketch of the algorithm

- Solve Cycle Median Problem
 - Start with empty M
 - Subsequently add edges
 - Estimate lower bound for partial solution
 - Continue with partial solution with least lower bound (branch and bound)
 - NEW: Consider cycle lengths
- Verify solution
 - Calculate edge weights
 - ... either by an exact algorithm for pairwise distances
 - ... or by an approximation algorithm (faster)

Experiments

- Create random input
 - Start with *id* of size n (n = 37 and n = 100)
 - Create 3 sequences of operations of length r (2 \leq $r \leq$ 15)
 - Use these sequences to obtain π^1 , π^2 , and π^3

Experiments

- Create random input
 - Start with *id* of size n (n = 37 and n = 100)
 - ► Create 3 sequences of operations of length r (2 ≤ r ≤ 15)
 - Use these sequences to obtain π^1 , π^2 , and π^3
- Testing
 - Most inputs could be solved within a few seconds
 - Verifying solutions with approximation algorithm is very accurate
 - Much faster than previous algorithm for the Transposition Median Problem (Yue et al. 2008)

Conclusion

We presented an algorithm that ...

- can solve the TMP and wRTMP exactly
- is fast enough for practical use
- ▶ is FREE SOFTWARE (GPL v3.0)

 \Rightarrow download it from

http://www.uni-ulm.de/fileadmin/website_uni_ulm/iui.

inst.190/Mitarbeiter/bader/phylo-1.0.1.tar.gz

Conclusion

We presented an algorithm that ...

- can solve the TMP and wRTMP exactly
- is fast enough for practical use
- ▶ is FREE SOFTWARE (GPL v3.0)

 \Rightarrow download it from

http://www.uni-ulm.de/fileadmin/website_uni_ulm/iui.

inst.190/Mitarbeiter/bader/phylo-1.0.1.tar.gz

Thanks for your attention!