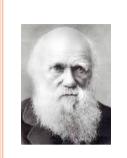


MOTIVATION

• Different systems "coevolve"

- hosts and their parasites or pathogens
- whole organisms and their genes
- geographical areas and the species which inhabit them
- cultural traditions and populations

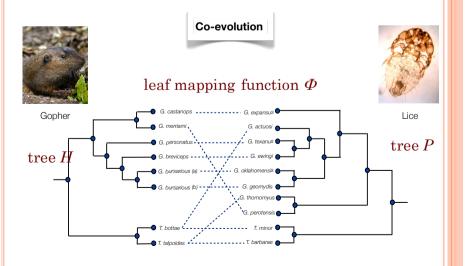
COEVOLUTION



"Thus I can understand how a flower and a bee might slowly become, either simultaneously or one after the other, modified and adapted to each other in the most perfect manner, by the continued preservation of all the individuals which presented slight deviations of structure mutually favourable to each other."

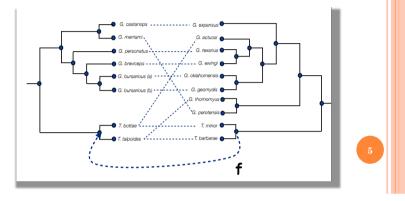
— Charles Darwin, The Origin of Species

THE INGREDIENTS



RECONCILIATIONS (1)

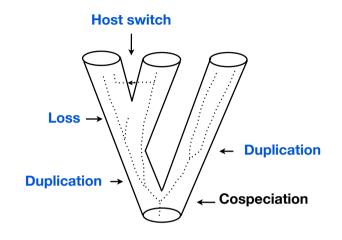
Very informally, a reconciliation is a mapping from the nodes of the parasite tree P to the nodes of the host tree H such that the leaf mapping function Φ is respected.



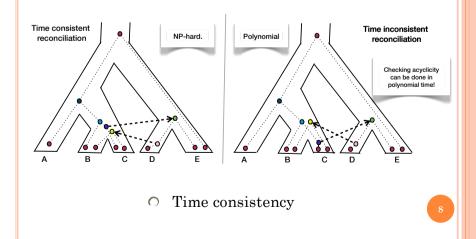
CO-PHYLOGENY RECONSTRUCTION PROBLEM (1)

- Determine reconciliations, given H, P and Φ
- Optimality of the solution: assigns a cost to each of the four types of events and then minimizes the total cost (Parsimony principle).
- Aim: generate all the optimal reconciliations

RECONCILIATIONS (2)



CO-PHYLOGENY RECONSTRUCTION PROBLEM (2)



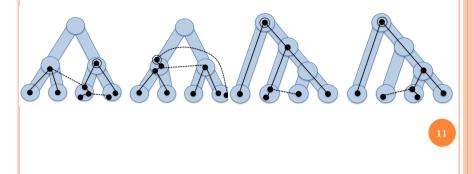
CO-PHYLOGENY RECONSTRUCTION PROBLEM (3)

- The number of optimal reconciliations increases rapidly even for small trees (exponential *in the size of the trees*).
- \bigcirc The size of the trees can be large.

• Many papers and tools trying to solve this issue 🧹

"SIMILAR" RECONCILIATIONS (1)

- Almost nothing in the literature:
 - similarity by event vectors
 - Similarity as the smallest number of operations needed to change one reconciliation into an- other

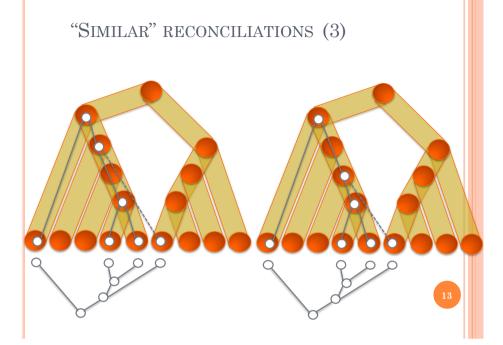


OUR PROBLEMS

- Biologists want to see all possible reconciliations, in order to understand which ones are biologically feasible and which ones are not.
- Either reduce the cardinality of or cluster the set of optimal reconciliations [with M. Gastaldello and M.-F. Sagot – CIBB '17]
- Visualize a given reconciliation in a "nice and clear" way [with M. Patrignani and V. Di Donato – GD '17]

"SIMILAR" RECONCILIATIONS (2)

- We introduced a new notion of metric to measure similarity between reconciliations:
 - take into account host-switches
 - *R*' and *R*" are the same reconciliation iff they have the same host-switches
 - Characteristic vector of each reconciliation (an item for each possible host-switch): relation with the Hamming distance of sufficiently long binary arrays (-> Hypercube)



CLUSTERING RECONCILIATIONS

• More open questions than solutions:

- We would like to exploit the similarity with the Hypercube, but the set of all the reconciliations represent only a subset of the nodes of the hypercube
- Do the connected components have any biological meaning? apparently no...
- How to choose the cluster head?
- Is the cluster head really representative?

"SIMILAR" RECONCILIATIONS (4)

Dataset	# solutions	$\# \sim_1$
GL	2	2
RH	2208	368
FD	408	180
COG2085	37568	3200
COG3715	9	7
COG4964	36	4
COG4965	640	576
PP	72	72
SFC	40	16
EC	18	18
PMP	2	2
PML	2	2
Wolbachia	$\sim 1.01\cdot 10^{47}$	$\sim 3.77\cdot 10^{44}$

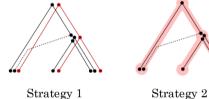
With a cost vector promoting co-speciation and discouraging host-switch: (0,1,2,1)

FURTHER PROBLEMS

- Reconciliation comparison:
 - new metrics
 - exact and approximate algorithms
- More realistic models:
 - deal with errors in phylogenetic trees (the phylogenetic trees are assumed to be correct, which may be not the case...)
 - ϕ is not a function: multiple hosts multiple parasites (a single parasite can infect more than one host...)
 - handle unrooted trees (many phylogenetic tree reconstruction algorithms produce unrooted trees; the outgroup method has the problem of the availability of a proper outgroup)

VISUALIZING RECONCILIATIONS (1)

- Given *H*, *P*, φ and a reconciliation *R*, we have to: draw *H* and *P* (on *H*) to highlight φ and *R* in a nice and clear way
- Three main strategies 0
 - 1. representing two paired trees
 - parasites are drawn inside their hosts 2.
 - 3. host tree is made of pipes and parasites are drawn into the pipes



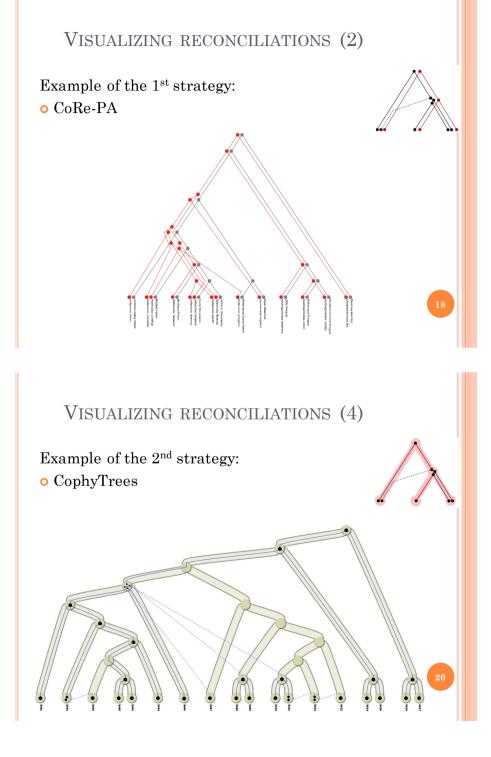




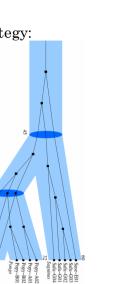
Strategy 1

Strategy 3

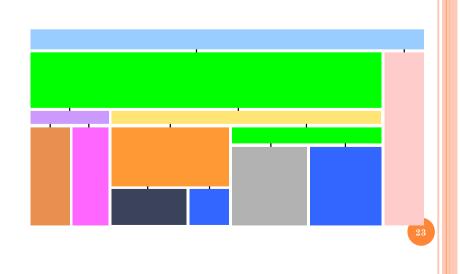
VISUALIZING RECONCILIATIONS (3) Example of the 1st strategy: • Jane 4

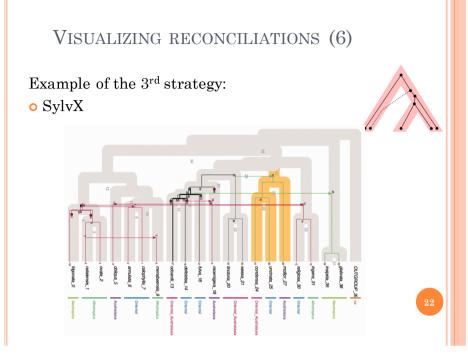


Example of the 2nd/3rd strategy: • Primetv

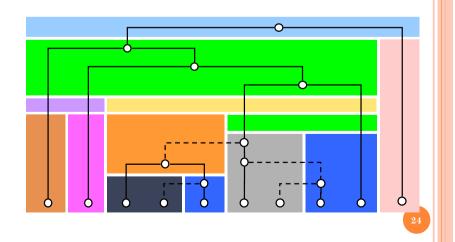


NEW METAPHOR: HP-DRAWINGS

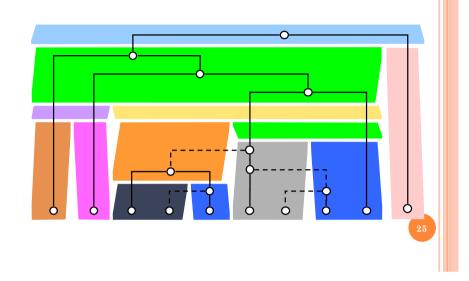




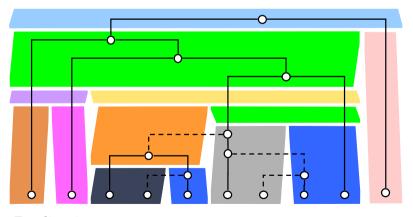
NEW METAPHOR: HP-DRAWINGS



NEW METAPHOR: HP-DRAWINGS



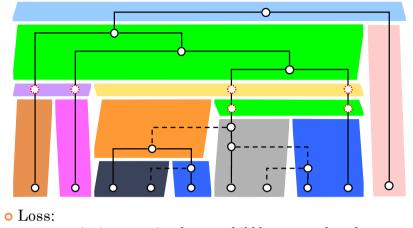
EVOLUTIONARY PHENOMENA



• Duplication:

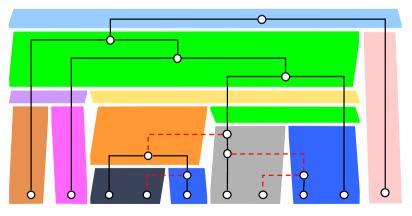
• both the children of a node go down in the same "direction"

EVOLUTIONARY PHENOMENA



• a parasite is transmitted to one child but not to the other child

EVOLUTIONARY PHENOMENA

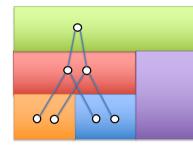


• Host switch:

• a parasite is transmitted to a host that is not a descendant of the current one

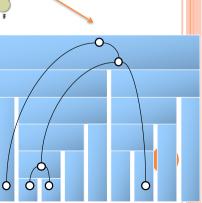
OPTIMIZATIONS (1)

- Given H, P, φ and a reconciliation R, we would like to:
 - 2. minimize the **crossing number** (not always possible to avoid crossings)



OPTIMIZATIONS (3)

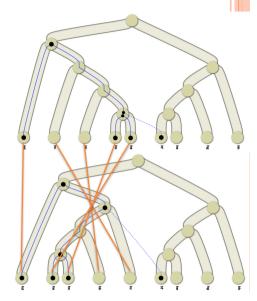
Note: Our model makes easier to understand what happens and keep the mental map, while trying to minimize the crossing number.

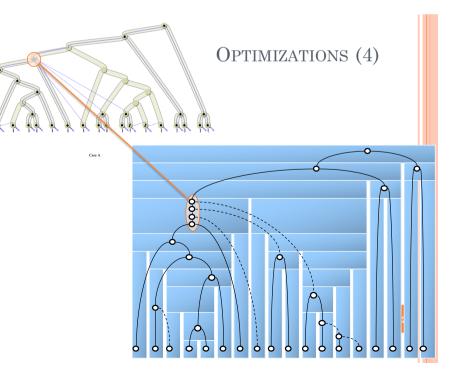


OPTIMIZATION (2)

• Given H, P, ϕ and a reconciliation R, we would like to:

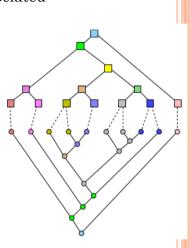
3. keep the **mental map** passing from a reconciliation to another one (not so in CophyTrees)





PLANAR INSTANCES

- Given H, P, φ , construct the associated tanglegram:
- Theorem: Every reconciliation on H, P, φ admits a planar representation iff the associated tanglegram is planar.
- So, we speak about planar and not planar instances.
- Theorem: deciding whether a time-consistent reconciliation γ admits an drawing with at most *k* crossings is NP-complete.

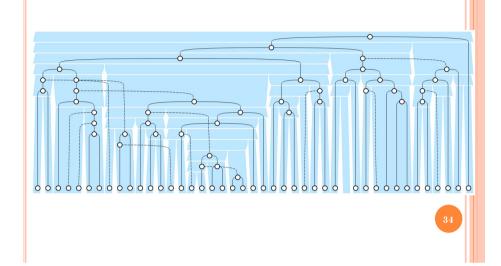


FURTHER PROBLEMS

• Test the tool:

- Do biologists like this metaphor?
- Models with more information:
 - handle additional information (e.g. geography) colors?

AN EXAMPLE



DISTANCES BETWEEN PHYLOGENETIC TREES

- Different phylogenetic trees for the same group of species are often produced either by procedures that use diverse optimality criteria (parsimony, compatibility, max likelihood, ...) or from different genes.
- Comparing these trees to find similarities (consensus) and dissimilarities (distance) is an important issue.
- There are a lot of known distances between rooted trees
 - RF Robinson-Foul
 - NNI Nearest neighbor interchange NP-hard
 - SPR Subtree Prune & Regraft NP-hard
 - DL based on reconciliations where host switches are not allowed (LCA)
 - ...
- Not very well studied their relations

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FURTHER PROBLEMS

- Compare different metrics
- Generalize to the unrooted case
 - (In the case of species-gene trees) Plateau property

