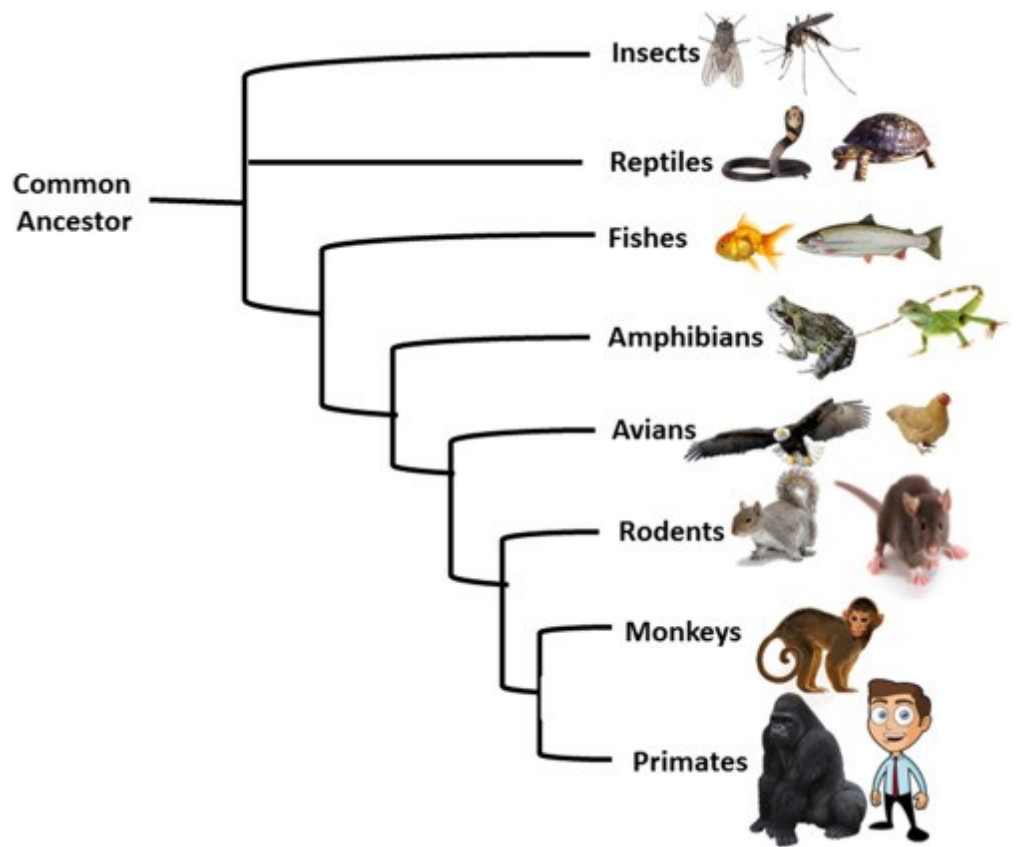


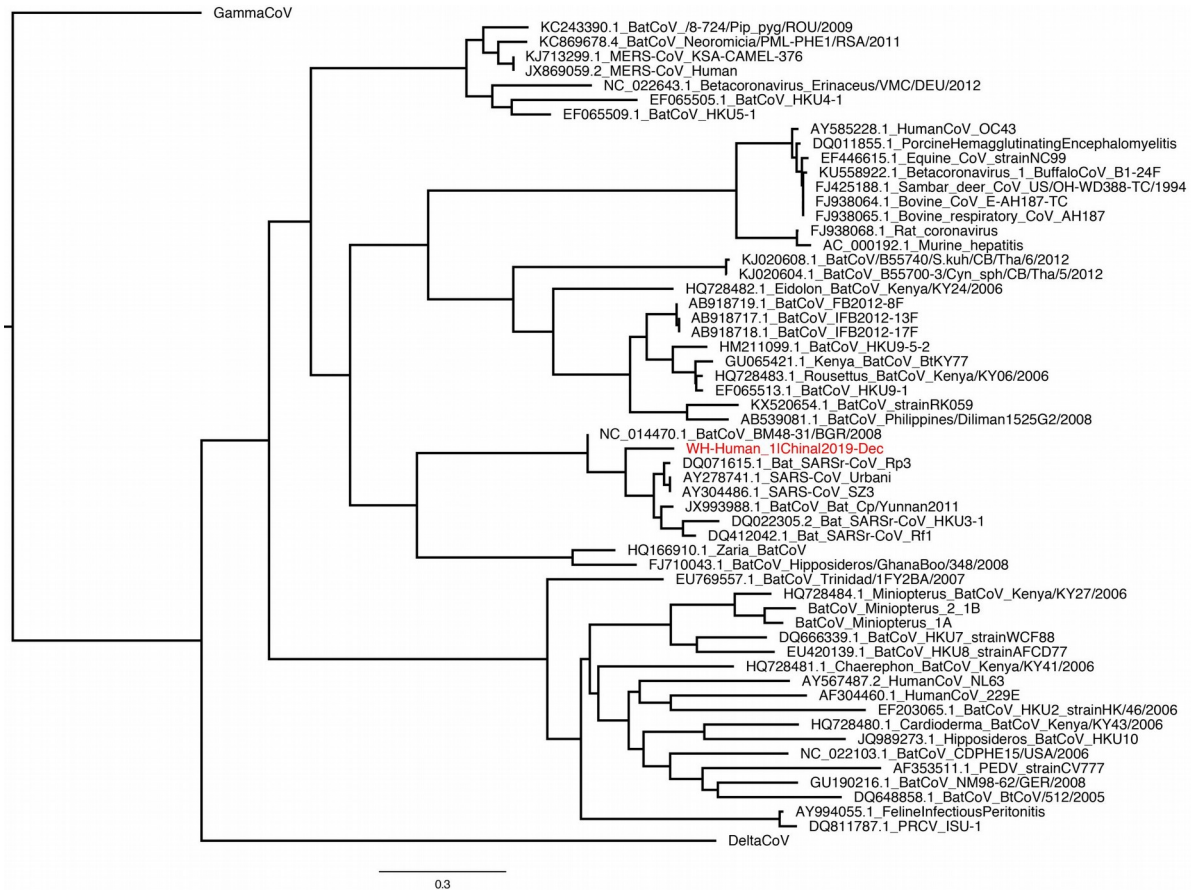


On the Elusiveness of Clusters

Steven M. Kelk Celine Scornavacca Leo van Iersel

Lukas Friedlos 22.04.21





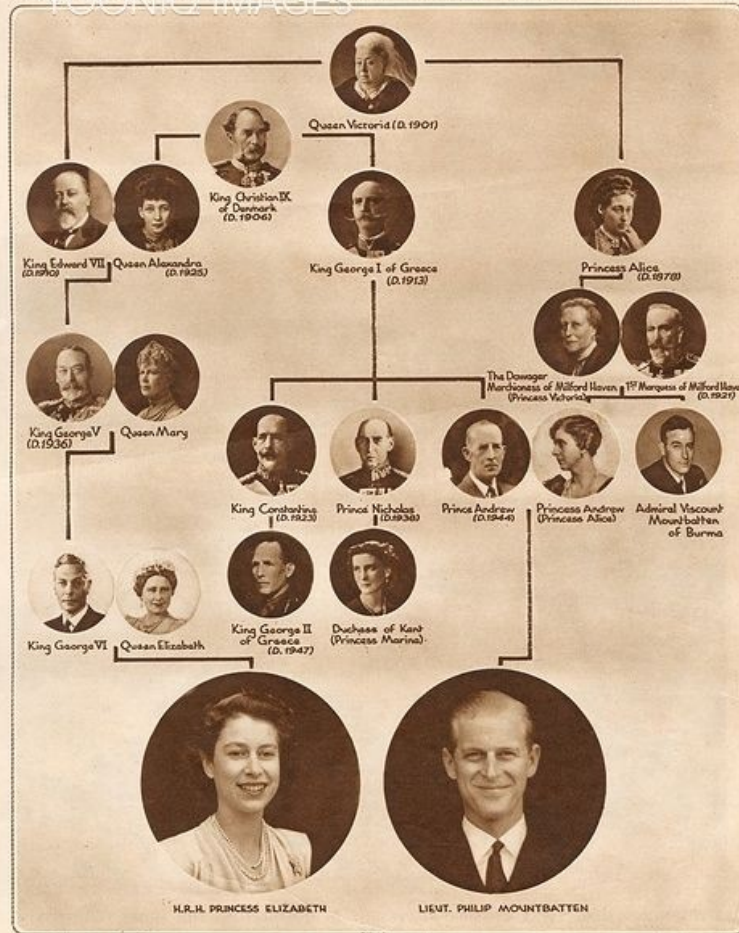
Preliminary maximum likelihood phylogenetic analysis of novel Wuhan, China human CoV in red, GenBank (accession MN908947)

Novel CoV seq data from: <http://virological.org/t/initial-genome-release-of-novel-coronavirus/319>. The Shanghai Public Health Clinical Center & School of Public Health, in collaboration with the Central Hospital of Wuhan, Huazhong University of Science and Technology, the Wuhan Center for Disease Control and Prevention, the National Institute for Communicable Disease Control and Prevention, Chinese Center for Disease Control, and the University of Sydney, Sydney, Australia.

PhyML tree based on partial RdRp gene sequence (410bp), aligned with representative human and animal CoV sequences from Genbank compiled by Alice Latinne; tree by Kevin Olival.
Analysis by EcoHealth Alliance - 11 Jan 2020 (12:30pm EST)



HOW THE PRINCESS AND HER FIANCÉ DESCEND FROM QUEEN VICTORIA.

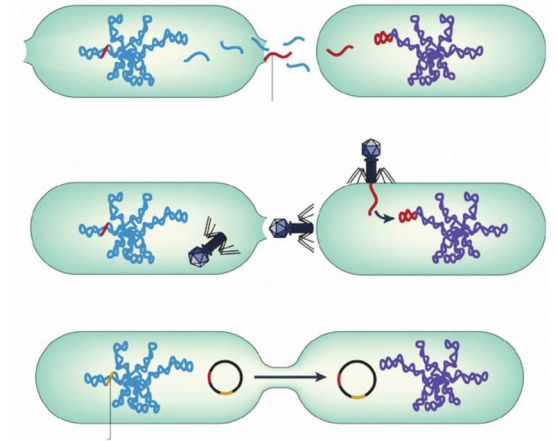


FIVE GENERATIONS OF THEIR ROYAL ANCESTRY: THE IMMEDIATE FORBEARS OF PRINCESS ELIZABETH AND LIEUT. MOUNTBATTEN.

The Helms-Princesses in the Throne is to marry a British Naval Officer of Royal ancestry, in whose veins flows the blood of one of the oldest dynasties in Europe; for though his own ancestry is not particularly noble recently on the throne of Greece, Lieut. Philip Mountbatten's is of old and illustrious origin in Denmark. From this great house those of a royal blood have chosen their consorts. Anne of Denmark was the Queen of James I.; Queen Anne's husband was Prince George of Denmark; and the lovely Queen Alexandra, consort of Edward VII., was the daughter of Christian IX., of Denmark. Princess Elizabeth and Lieut. Mountbatten are both great-grandchildren of Queen Victoria. Our tree illustrates how they descend from that great sovereign, and shows their distant kinship. Lieut. Mountbatten's father, the late Prince Andrew of Greece, was a nephew of Queen Alexandra, as she was a sister of George I., of the Hellenes, the Danish Prince William of Schleswig-Holstein-Sonderburg-Glücksburg, who in 1863 accepted the crown of Greece. His mother, Princess Andrew of Greece, was Princess Alice of Battenberg, sister of Admiral Viscount Mountbatten of Burma, and daughter of the first Marquis of Milford Haven (formerly Prince Louis of Battenberg), the great admiral to whom the British Navy owes so much. The Dowager Marchioness of Milford Haven, Lieut. Mountbatten's grandmother, is a granddaughter of Queen Victoria through her mother, Princess Alice, Grand Duchess of Hesse.

Cross-Hybridisation

- Hybrid speciation (through sexual reproduction)
 - Homo Sapiens ↔ Neanderthal
 - Polar Bear ↔ Brown Bear
 - Most common in plants
- Horizontal gene transfer



Horizontal gene transfer
in bacteria

Heliconius heurippa

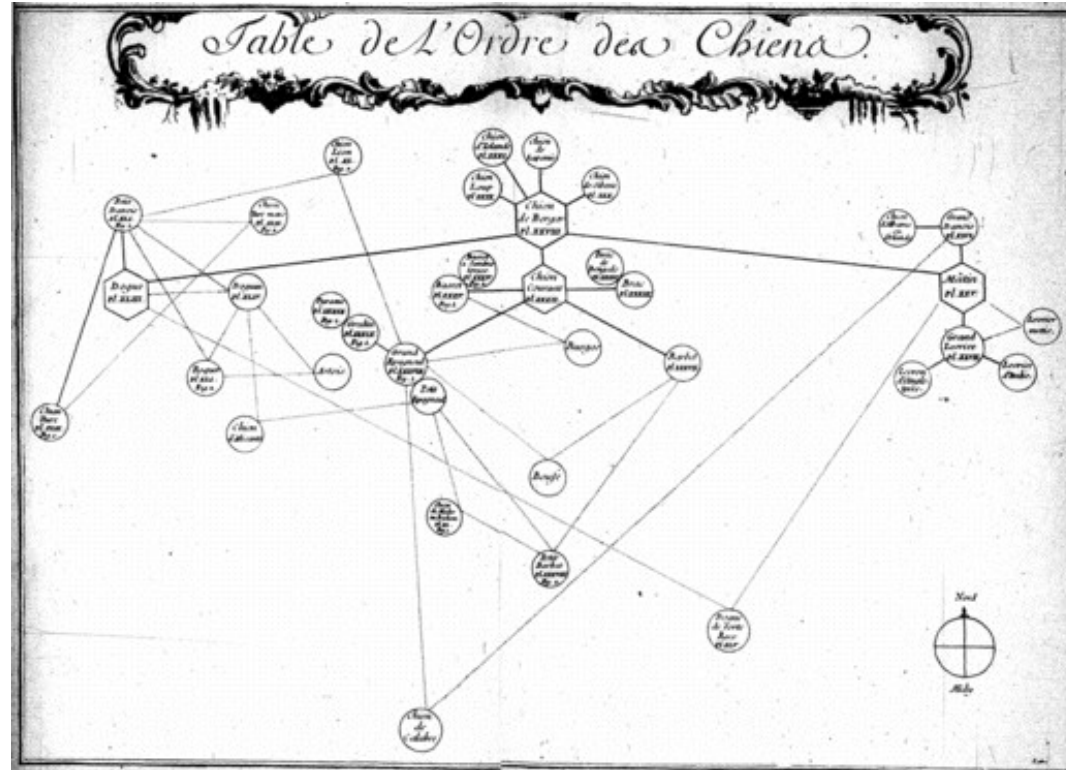


Jaguma

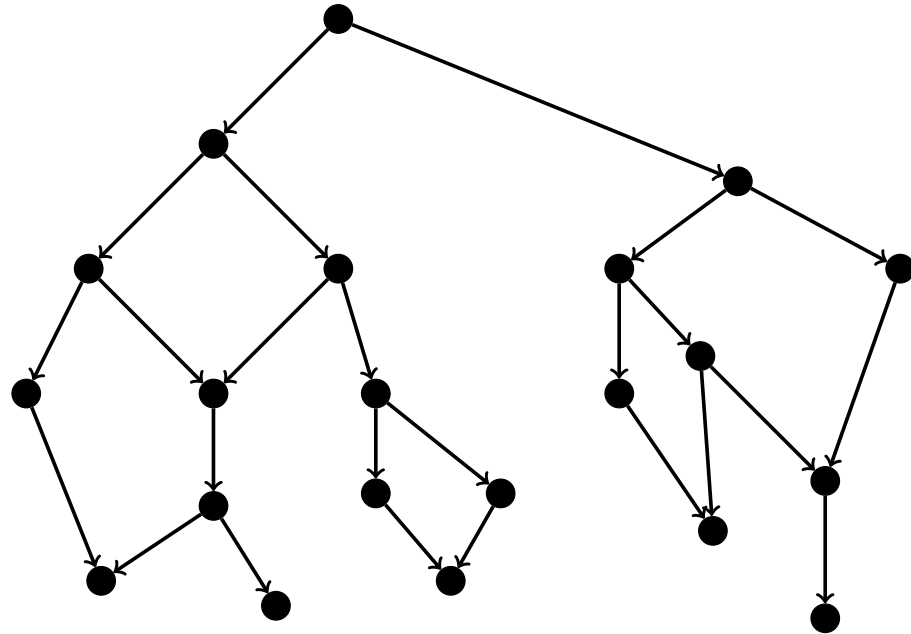


Prizzly

Phylogenetic Networks



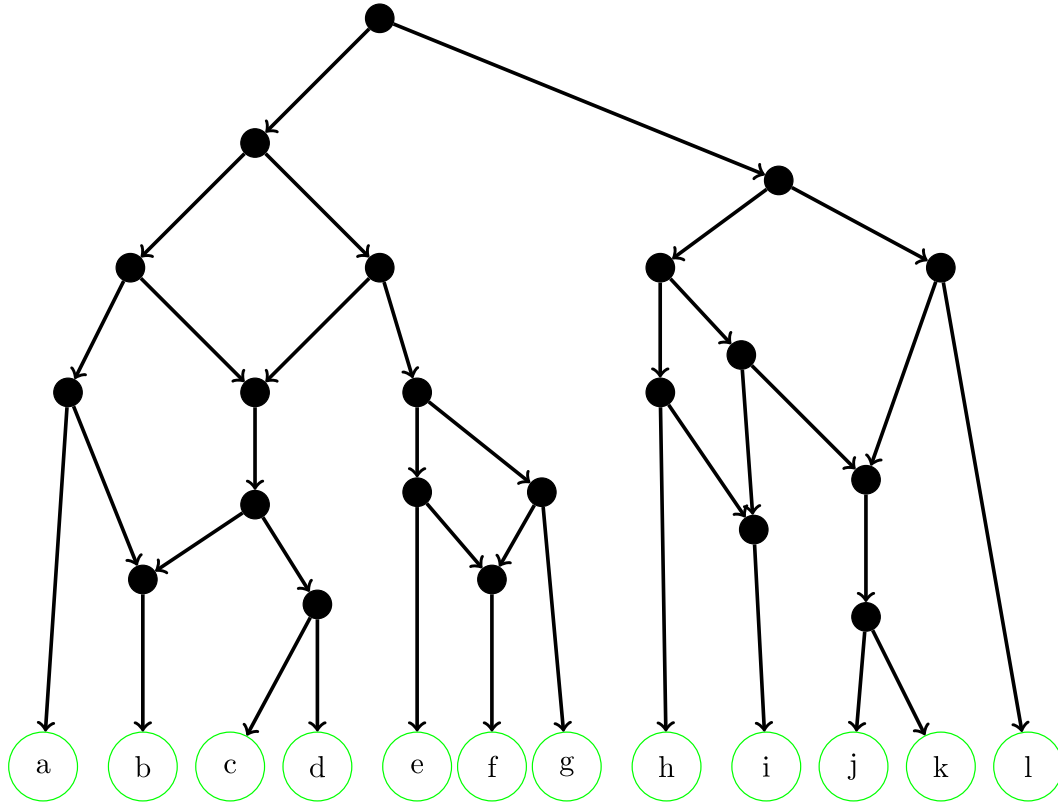
Phylogenetic Networks



Definition

- rooted directed acyclic graph
Henceforth network \mathcal{N}

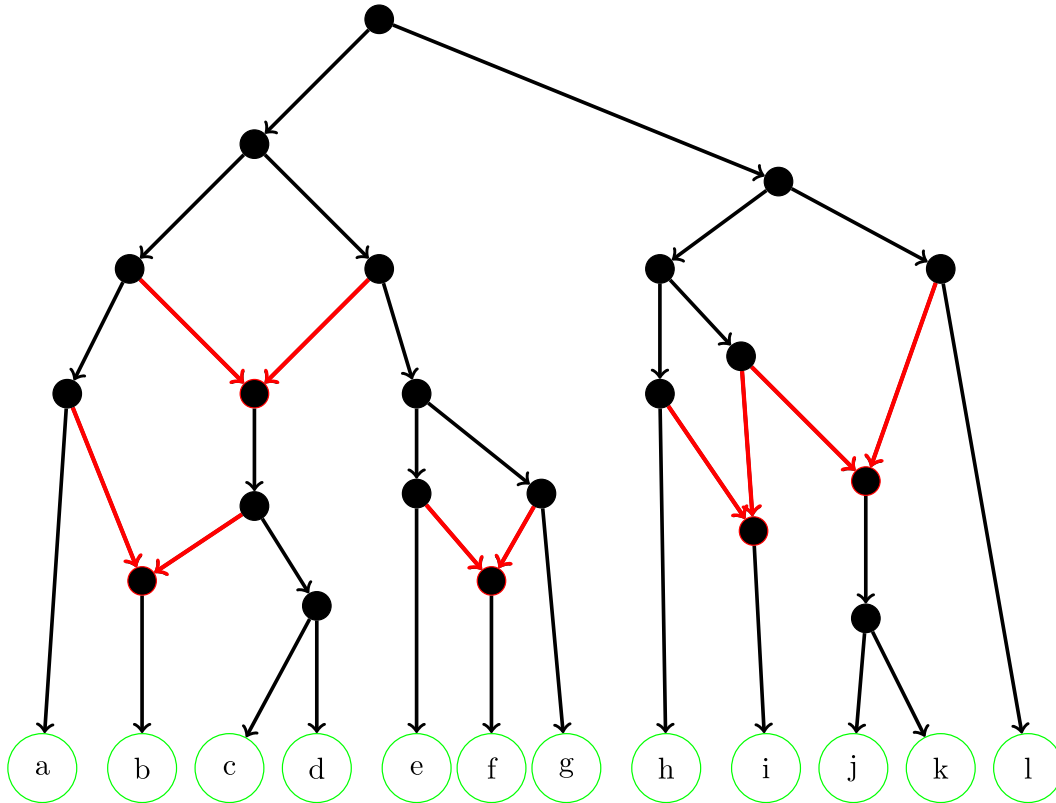
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 $\mathcal{X} = \{a, b, c, d, e, f, g, h, i, j, k, l\}$

Phylogenetic Networks



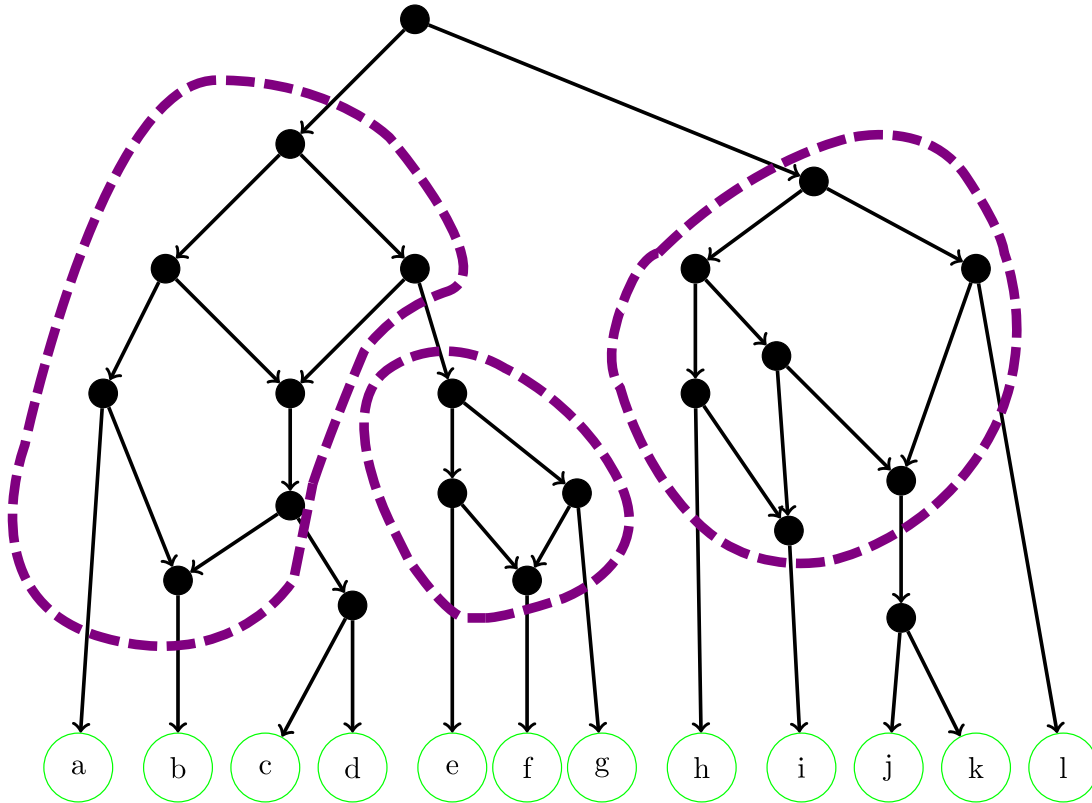
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Terminology

- reticulation
 $\text{in-deg}(v) \geq 2$

Phylogenetic Networks



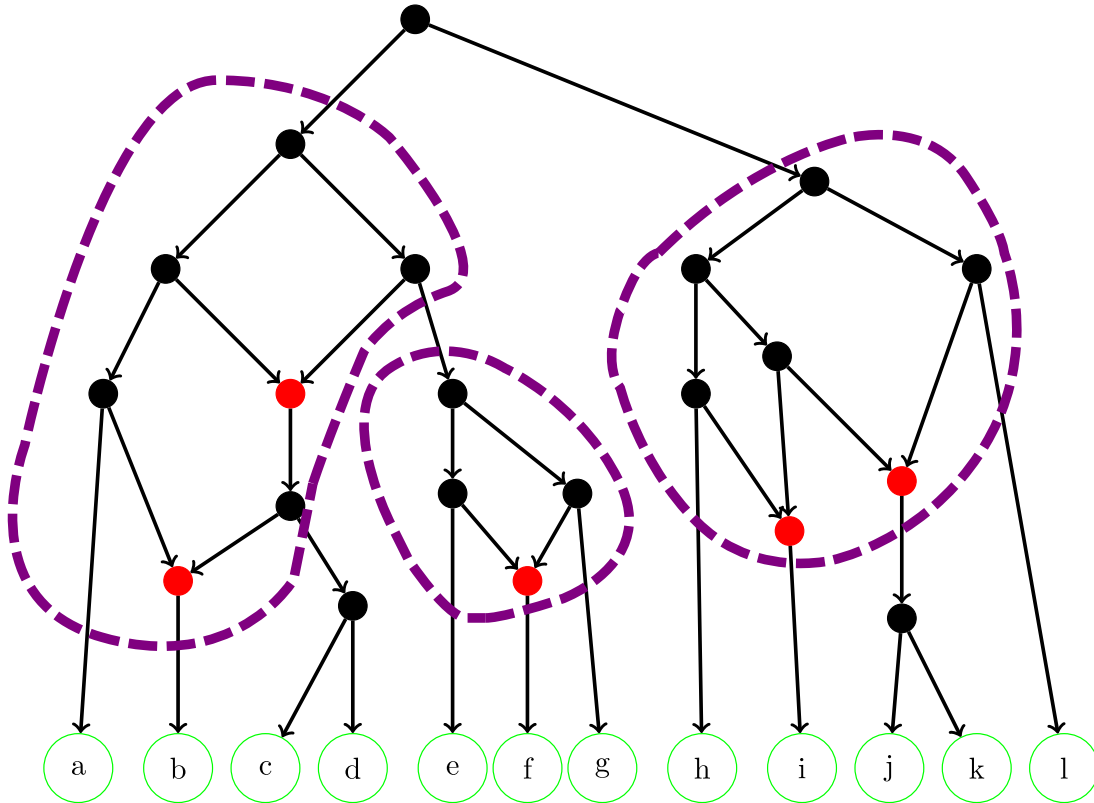
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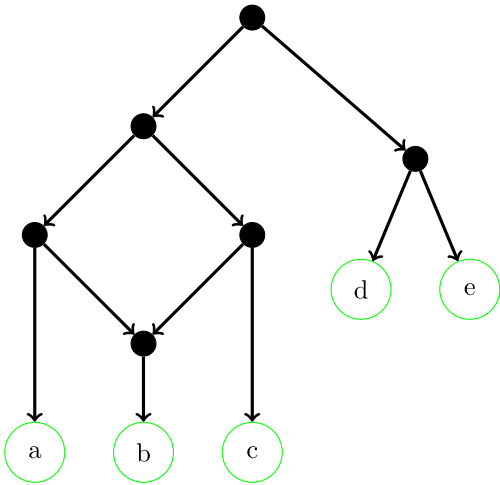
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Terminology

- reticulation
 $\text{in-deg}(v) \geq 2$
- biconnected components
- k -level

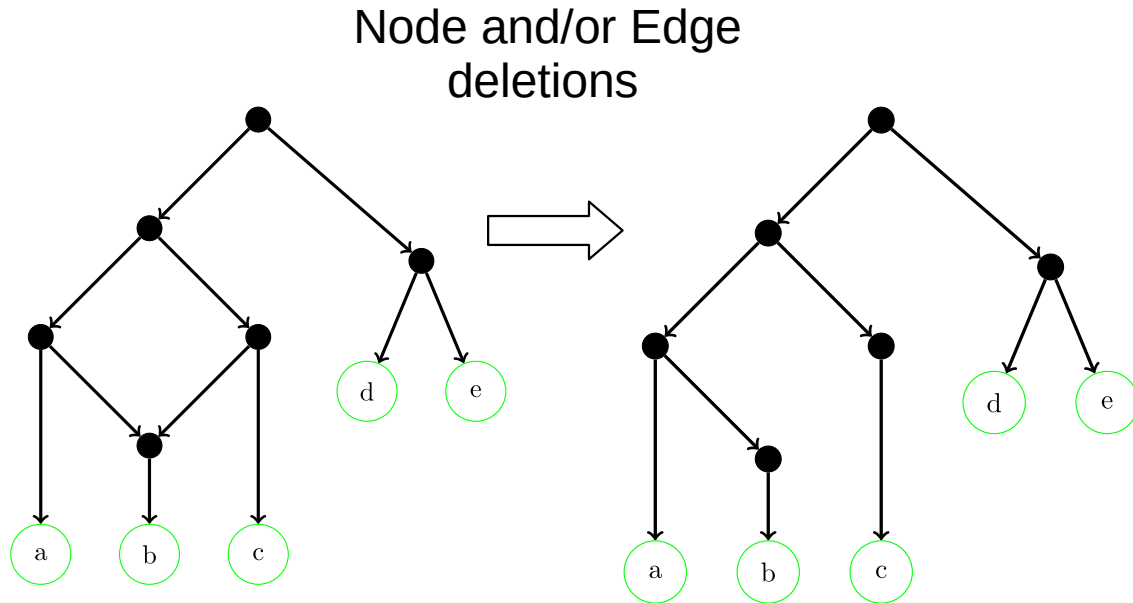
Network 'Displaying' a Tree

A Network N displays a tree T , if it can be obtained from N via



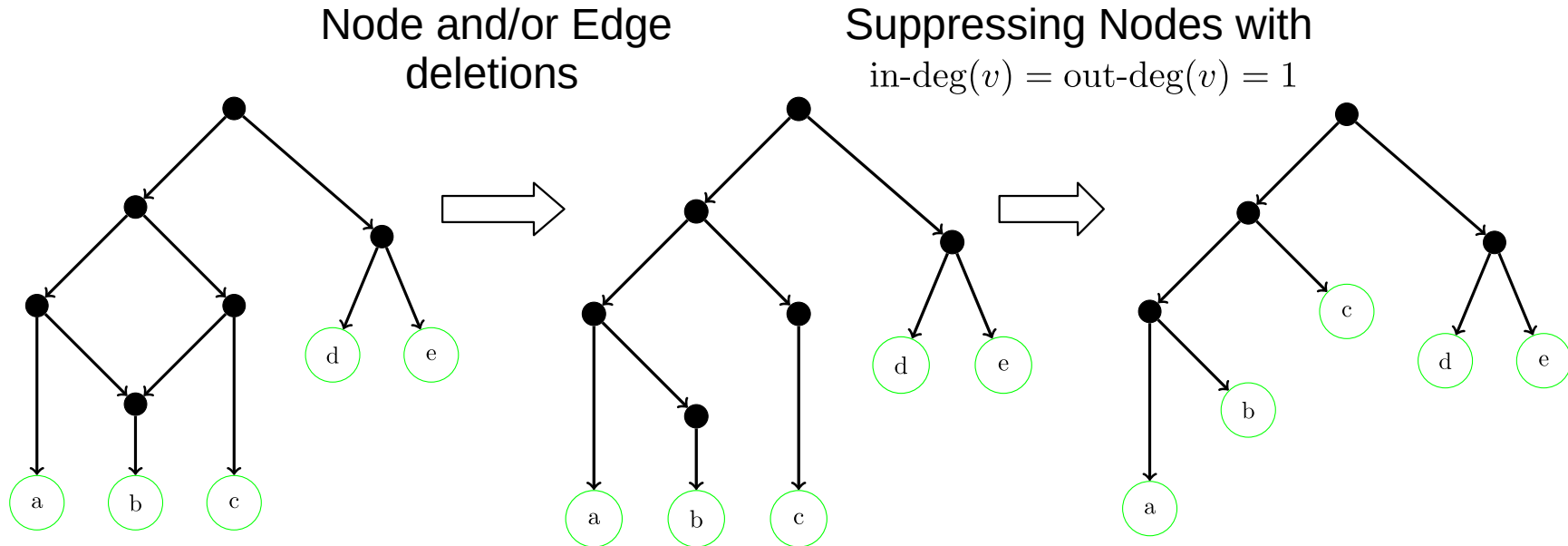
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Clusters

Given a set of taxa $\mathcal{X} = \{a, b, c, d, e\}$

A cluster $C \subset \mathcal{X}$ is a proper subset of all taxa, e.g. $\{a, b, c\}$

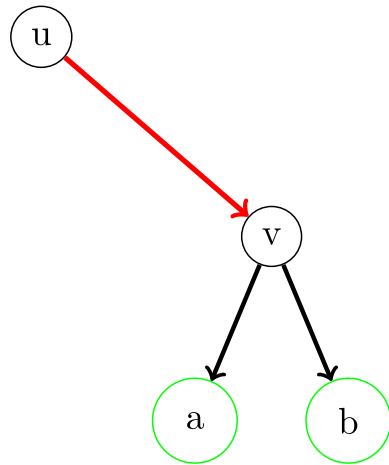
Taxa are in a cluster \implies Taxa have a common ancestor

A set of clusters \mathcal{C} on a set of taxa, e.g. $\{\{a, b, c\}, \{a, b\}, \{d, e\}\}$

Networks represent Clusters

Representing Clusters

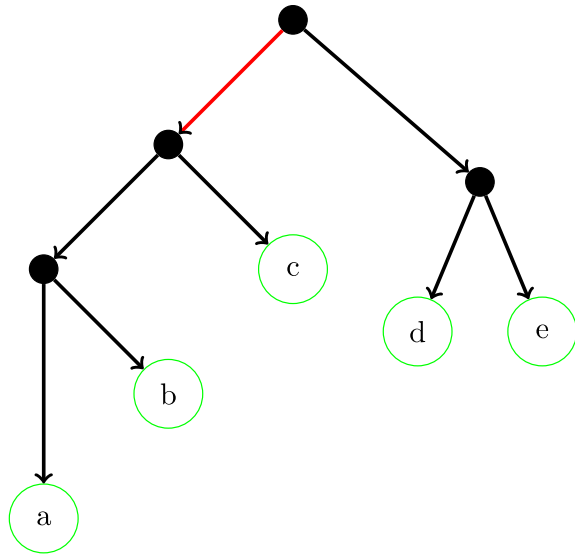
We say that an **edge** (u, v) represents a cluster C if C is the set of leaf descendants of v .



$$C = \{a, b\}$$

Representing Clusters

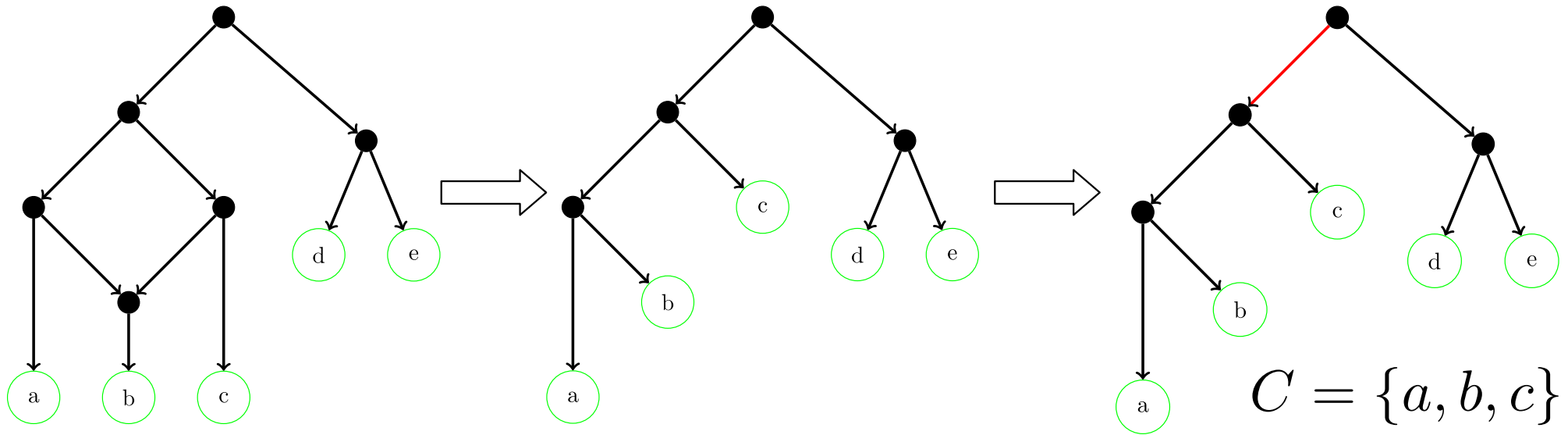
We say that a tree T represents a cluster C if T has an edge that represents C .



$$C = \{a, b, c\}$$

Representing Clusters

We say that a network N represents a cluster C if N displays a tree that represents C .





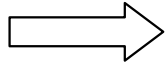
A Theoretical Polynomial-Time Algorithm for Constructing Level-k Networks

Goal

- Given:
- set of clusters \mathcal{C}
 - fixed $k \geq 0$

Goal

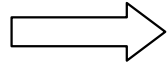
Given: • set of clusters \mathcal{C}
• fixed $k \geq 0$



Construct level- k network N representing \mathcal{C} ,
if such a network exists.

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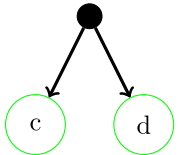
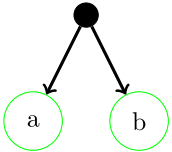
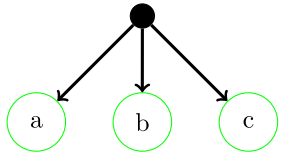
Possible in Polynomial-Time!

From Clusters to Networks (Trivial)

E.g. $\mathcal{X} = \{a, b, c, d\}$, $\mathcal{C} = \{\{a, b, c\}, \{a, b\}, \{c, d\}\}$

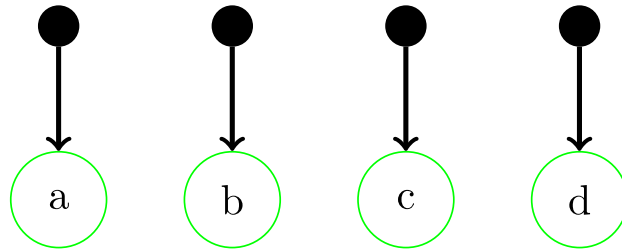
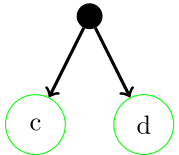
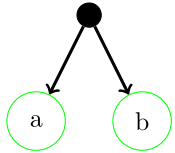
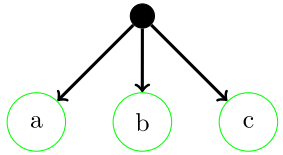
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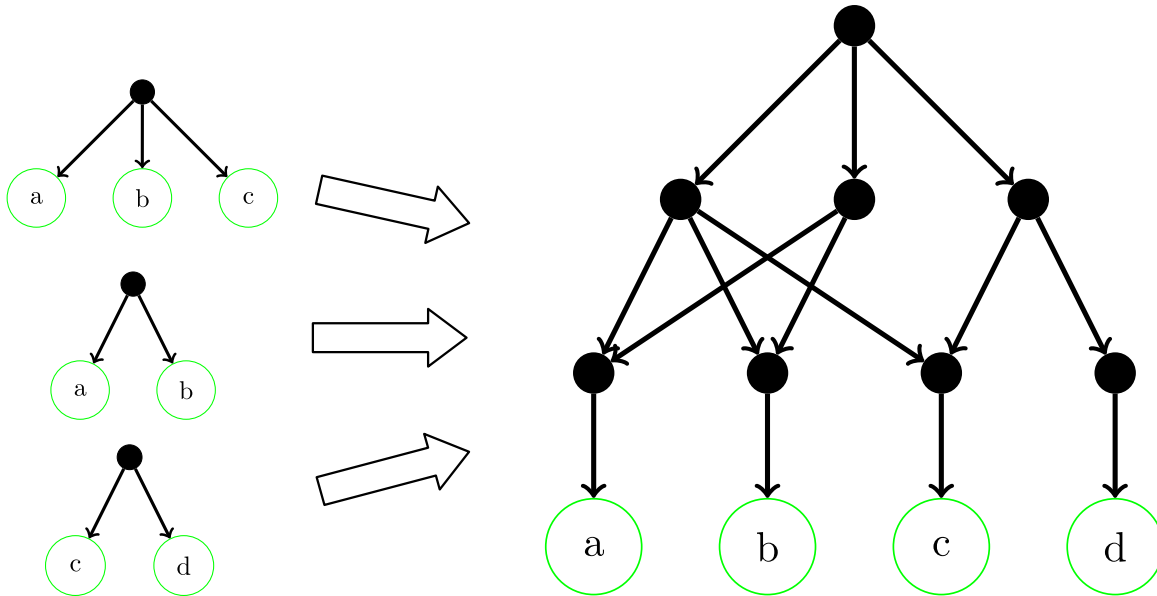
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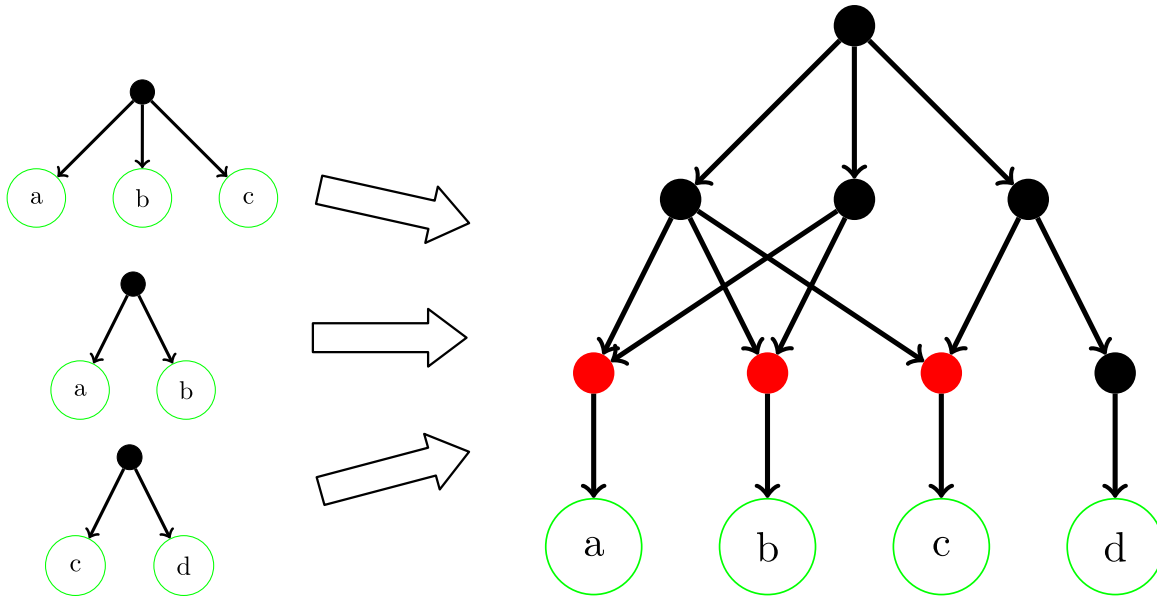
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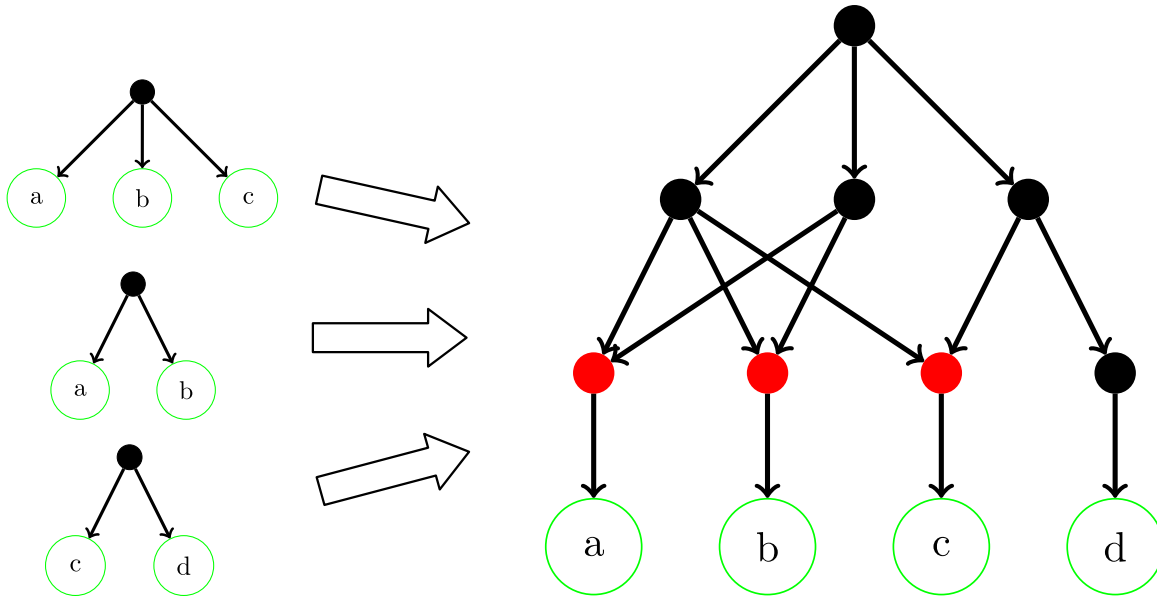
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$k = 3$

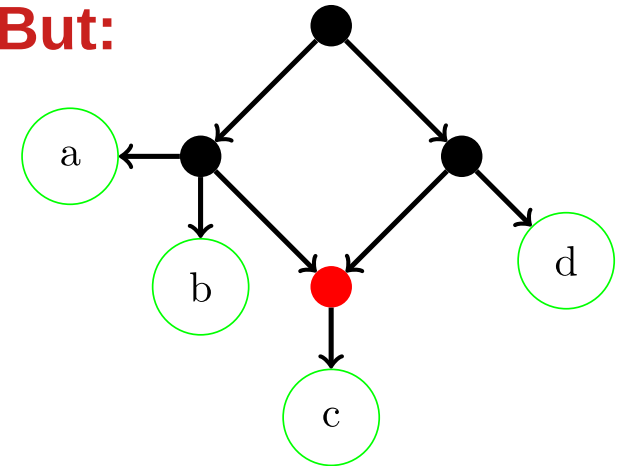
From Clusters to Networks (Trivial)

E.g. $\mathcal{X} = \{a, b, c, d\}$, $\mathcal{C} = \{\{a, b, c\}, \{a, b\}, \{c, d\}\}$



$k = 3$

But:



$k = 1$



From Clusters to Networks

- Constructing a network representing clusters is trivial

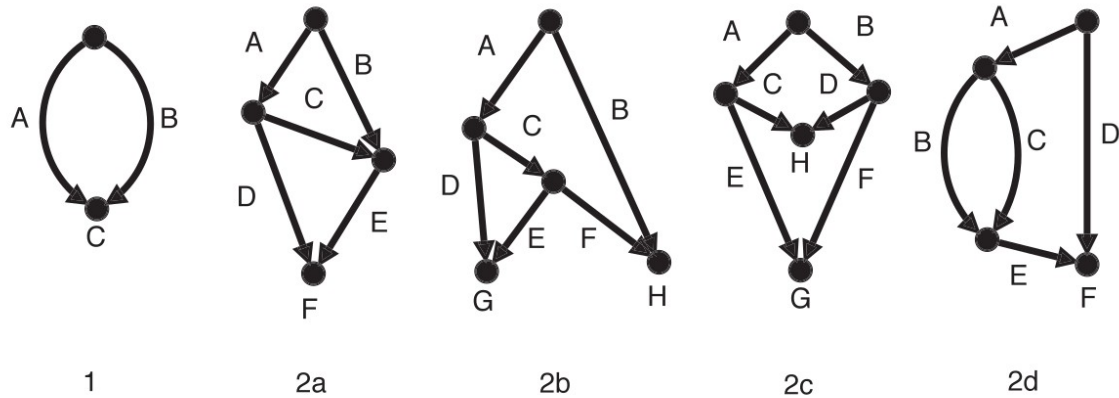


From Clusters to Networks

- Constructing a network representing clusters is trivial
- Constructing a network representing clusters with minimal level is NP-hard

Generators

Any level- k network can be reduced to a level- k generator



Edges and nodes with out-deg = 0 are called *sides*



Algorithm Outline

- Guess a generator for the network
- Build the network from the generator with guesses

Disclaimer: This algorithm is purely theoretical and doesn't lend itself for practical implementation

Algorithm

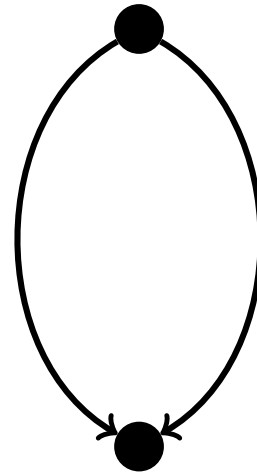
$$\mathcal{X} = \{a, b, c, d\}, \mathcal{C} = \{\{a, b, c\}, \{a, b\}, \{c, d\}\}, k = 1$$

1. Guess a generator

Algorithm

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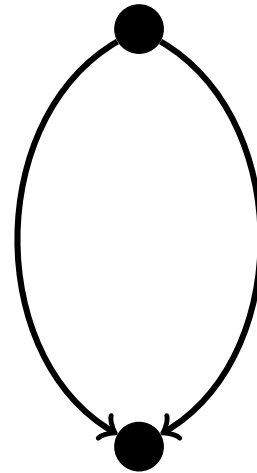
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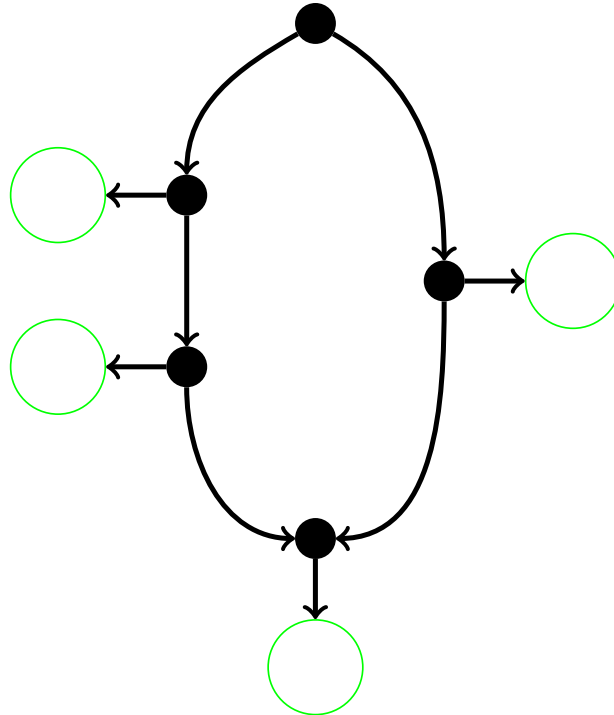
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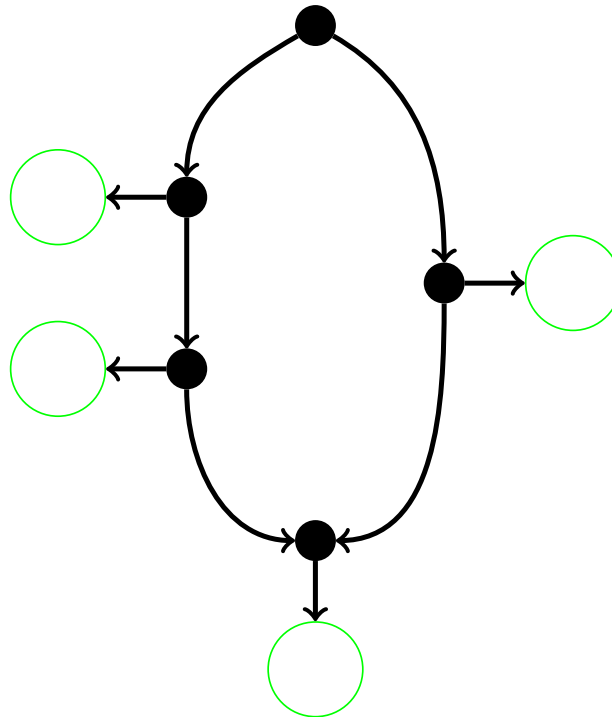
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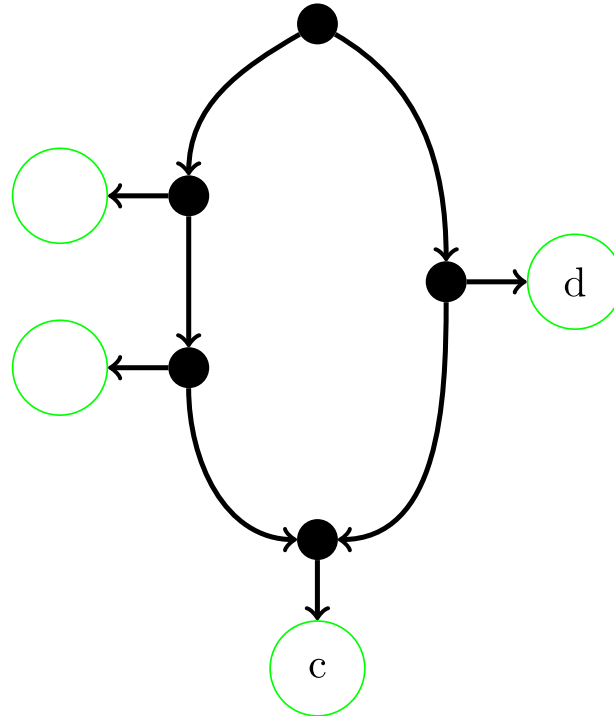
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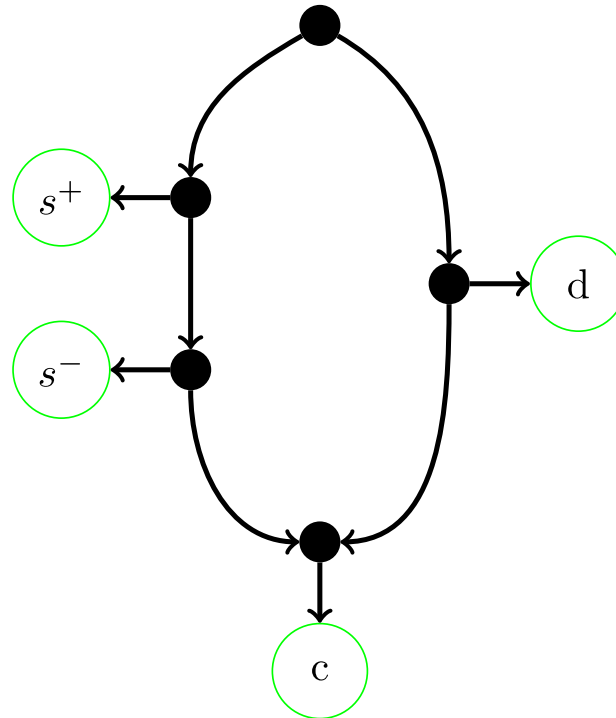
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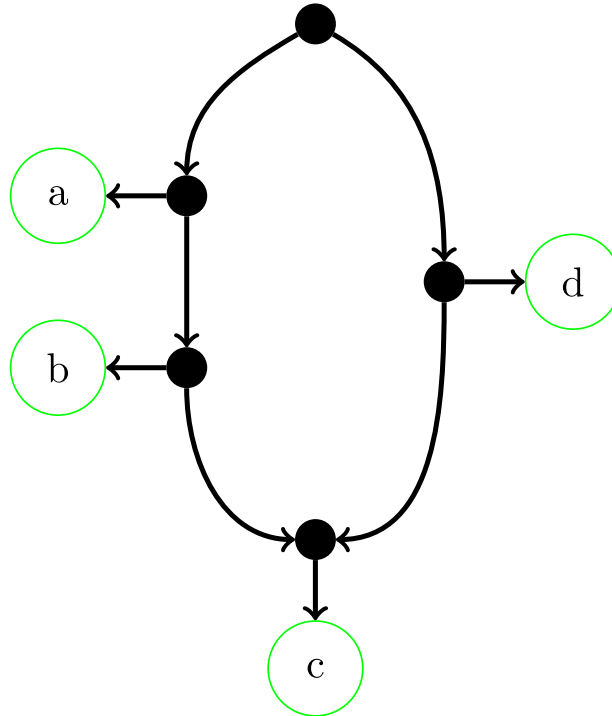
1. Guess a generator
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4. Guess top s^+ and bottom s^- of the sides with ≥ 2 leaves



Algorithm

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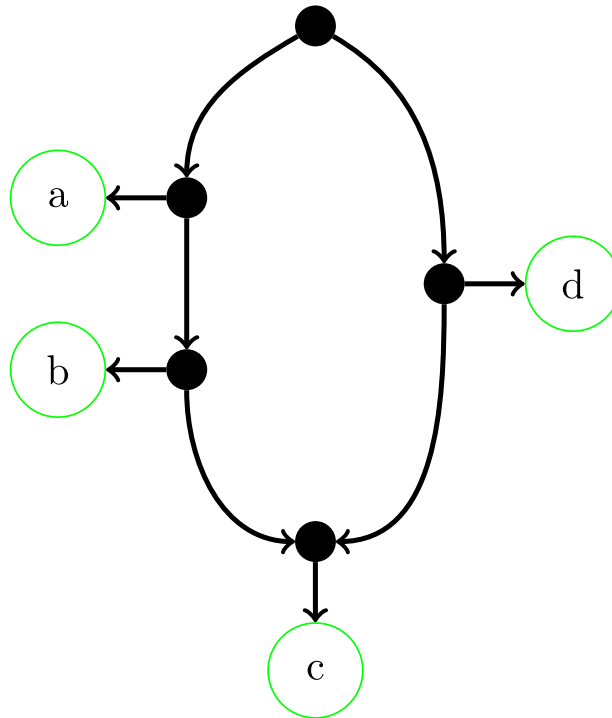
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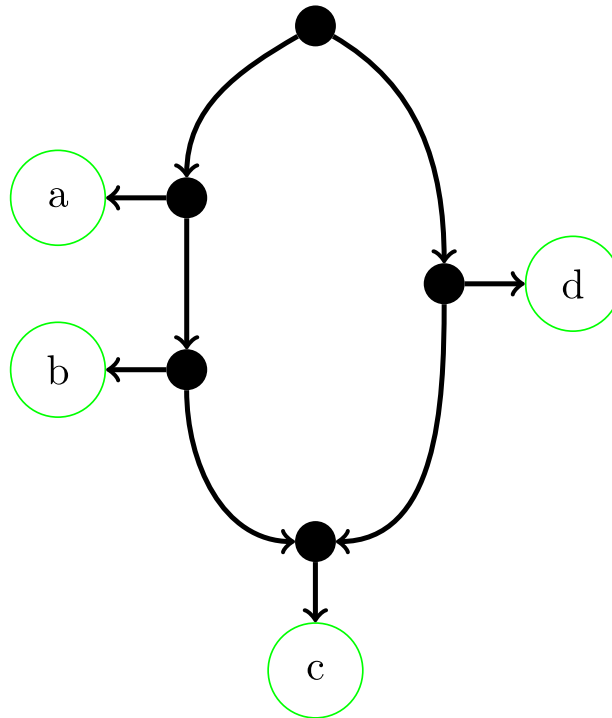
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5. Add remaining leaves
6. Verify that the network actually represents \mathcal{C}





Adding Remaining Leaves

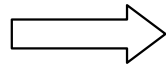
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Adding Remaining Leaves

$$\mathcal{C} = \{\{a, b, c\}, \{a, b\}, \{c, d\}\} \implies \begin{array}{l} a \rightarrow b \\ b \rightarrow a \\ c \rightarrow d \\ d \rightarrow c \end{array} \quad \begin{array}{l} a \rightarrow b \\ = \\ \text{"Every cluster containing } a, \\ \text{also contains } b\text{"} \end{array}$$

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$$\begin{aligned} a &\rightarrow b \\ b &\rightarrow a \\ c &\rightarrow d \\ d &\rightarrow c \end{aligned}$$

$$a \rightarrow b$$

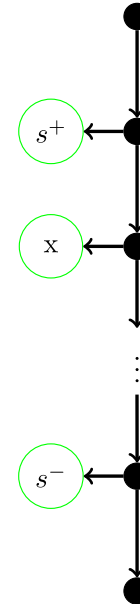
=

“Every cluster containing a ,
also contains b ”

Start with the lowest side

Based on s^+ , s^- and the relationships from
above, you can determine the leaf to be inserted
in polynomial time

$$s^+ \rightarrow x \rightarrow s^- \text{ must hold}$$



Adding Remaining Leaves

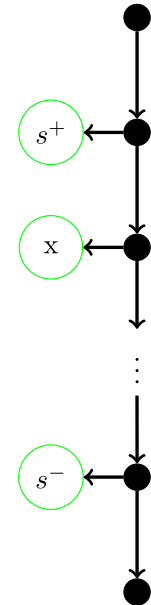
$$C = \{\{a, b, c\}, \{a, b\}, \{c, d\}\} \implies \begin{matrix} a \rightarrow b \\ b \rightarrow a \\ c \rightarrow d \\ d \rightarrow c \end{matrix} \qquad \begin{matrix} a \rightarrow b \\ = \\ \text{"Every cluster containing } a, \\ \text{also contains } b\text{"} \end{matrix}$$

Start with the lowest side

Based on s^+ , s^- and the relationships from above, you can determine the leaf to be inserted in polynomial time

$$s^+ \rightarrow x \rightarrow s^- \text{ must hold}$$

If no leaf can be added \implies Start new



Complexity Recap

- Guess from a **constant** number of generators
- For a **constant** number of sides, guess how many leaves from 4 options
- $\mathcal{O}(n^2)$ guesses for top and bottom
- Polynomial time for the remaining leaves
- Polynomial time to verify

 **Very polynomial-time algorithm!**



Issues



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- # generators explodes

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$$[2^{k-1}, k!^2 50^k]$$

k	generators
1	1
2	4
3	65
4	1993
5	91454

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In sum this makes the algorithm practically unfeasible :(



The End

Thank you for your attention!