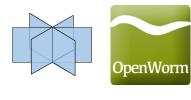
# Hard-to-Define Events in Biological Networks



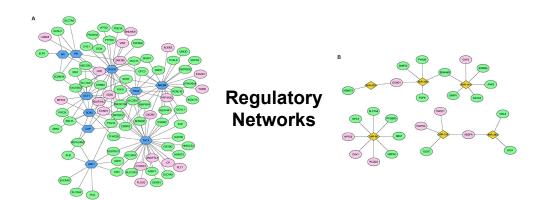
@ NetSci 2022: Shanghai and Virtual

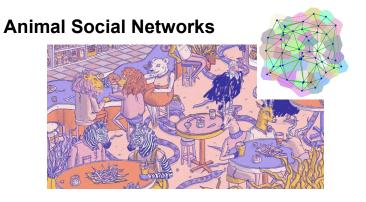


**Bradly Alicea** 

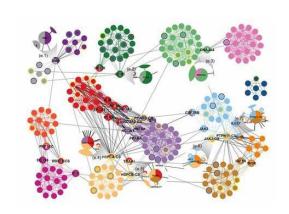
OpenWorm Foundation
Orthogonal Research and Education Lab

bradly-alicea.weebly.com

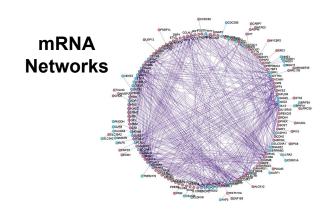




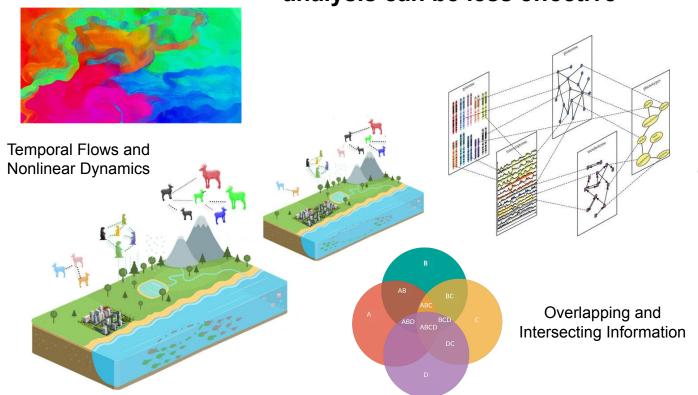
## Network modeling has been quite successful at the level of genes, proteins, cell population, and even individual organisms

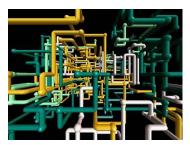


Protein-Protein Interaction Networks

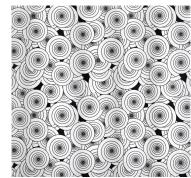


### In cases where the relationship between nodes, arcs, and temporal events are not clearly identifiable, network analysis can be less effective

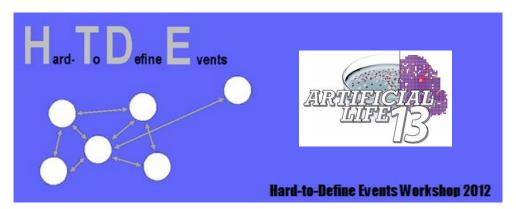


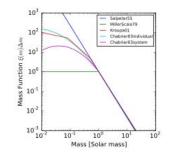


Cryptic Temporal and Emergent Processes



#### Hard-to-Define Events (HTDE): 10 years out

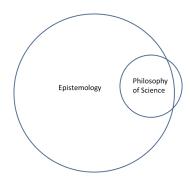




Modeling power laws and long-tailed phenomena



Rare and out-of-distribution empirical observation



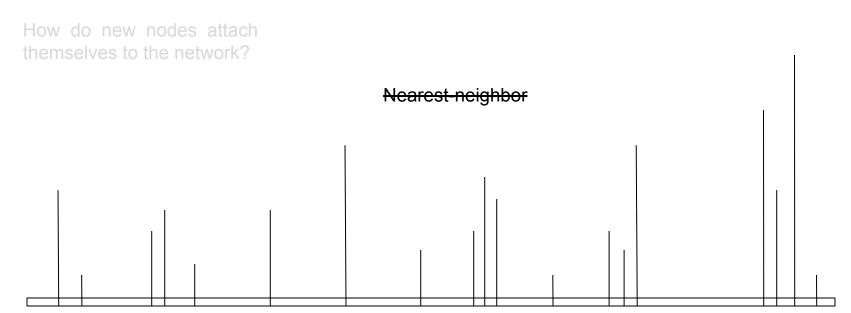
Epistemology of Science

### Systems with rare events, or out-of-distribution events (occur extremely rarely in a given spatiotemporal context)

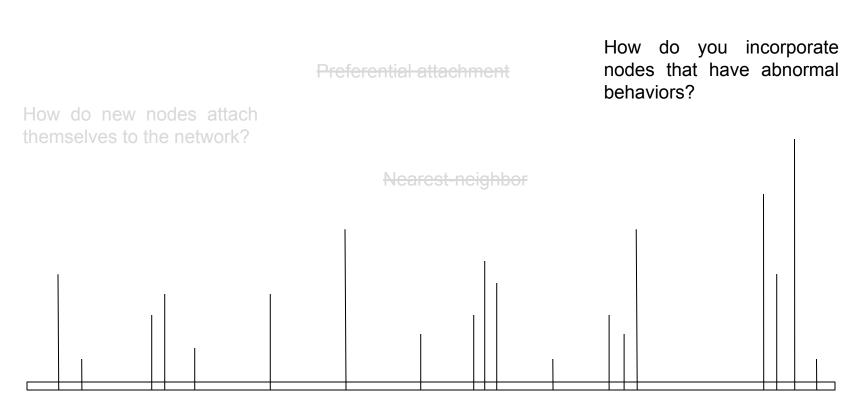
How do new nodes attach themselves to the network?

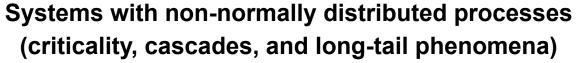
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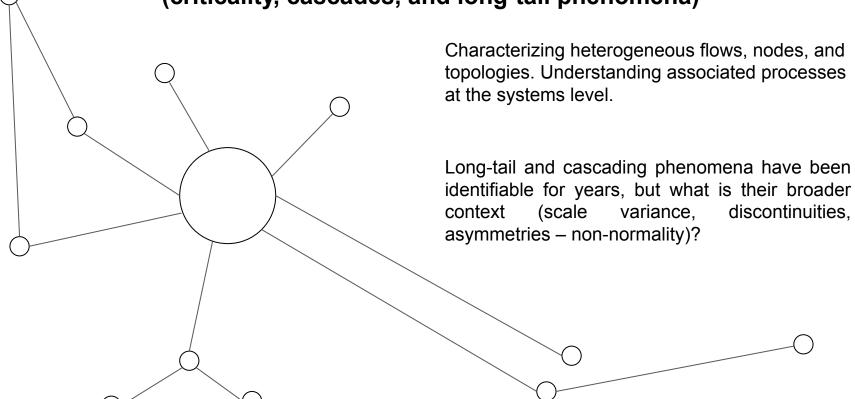
#### Preferential attachment



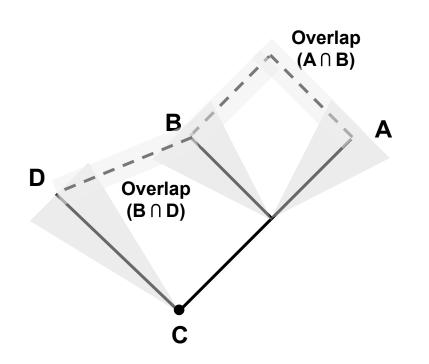
## Systems with rare events, or out-of-distribution events (occur extremely rarely in a given spatiotemporal context)







### Cases in which nodes are poorly defined in terms of their compartmentalization and/or epistemology



Edges can belong to different categories simultaneously.

Nodes can be intersections  $\{A, B, D\}$  or defined elements of a set  $\{C\}$ .

Nodes can also overlap {A and B, B and D}.

Nodes and edges that do not conform to transitivity  $\{A < B \rightarrow B < C \rightarrow A < C\}$ .

## Sampling of Approaches in the Network Science Literature (since 2005)

Lawyer, G. (2015). Understanding the influence of all nodes in a network. Scientific Reports, 5, 8665.

Asllani, M., Lambiotte, R., and Carletti, T. (2018). Structure and dynamical behavior of non-normal networks. *Scientific Advances*. 4. eaau9403.

Allen, R.J., Warren, P.B., and ten Wolde, P.R. (2005). Sampling rare switching events in biochemical networks. *Physical Review Letters*, 94, 018104.

Hindes, J. and Schwartz, I.B. (2017). Rare events in networks with internal and external noise. *Europhysics Letters*, 120, 56004.

Sheridan, P. and Onodera, T. (2018). A Preferential Attachment Paradox: how preferential attachment combines with growth to produce networks with log-normal in-degree distributions. *Scientific Reports*, 8, 2811.

Guimera, R., Sales-Pardo, M., and Amaral, L.A.N. (2007). Module identification in bipartite and directed networks. *Physical Review E*, 76, 036102.

Lawyer, G. (2015). Understanding the influence of all nodes in a network. Scientific Reports, 5, 8665.

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Allen, R.J., Wonetworks. Phy

State of the Art: identification of HTDE phenomena within problem domains

Hindes, J. and Letters, 120, 5

Future Challenge: refining network science techniques for the HTDE problem class.

Europhysics

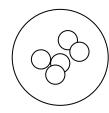
Sheridan, P. and Onogera, T. (2018). A Preferential Attachment Paradox: now preferential attachment

Guimera, R., Sales-Pardo, M., and Amaral, L.A.N. (2007). Module identification in bipartite and directed networks. *Physical Review E*, 76, 036102.

combines with growth to produce networks with log-normal in-degree distributions. Scientific Reports, 8,

2811.

#### A future role for theory-building and model epistemology

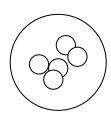


Nodes that represent quantitative constructs, overlapping states (experiments with developmental hypergraphs).



Edges that represent multiple properties simultaneously (experiments with Contextual Geometric Structures).

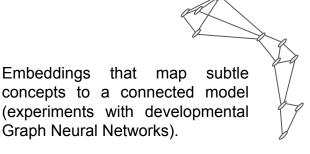
#### A future role for theory-building and model epistemology

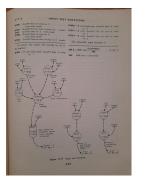


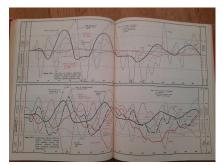
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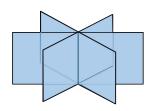




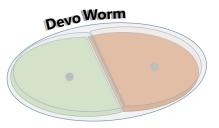
Networked dynamics of systems models.

Forrester. Industrial Dynamics (1958). Images: Trace and Traceability (trace-and- traceability.org)

#### Thank You for Your Attention!



Cybernetics and Systems Dynamics, Contextual Geometric Structures (CGS)



Developmental Graph Neural Networks (D-GNNs), Developmental Hypergraphs July 27, 1:15 UTC

Hypergraphs Demonstrate

Anastomoses During Divergent

Integration

