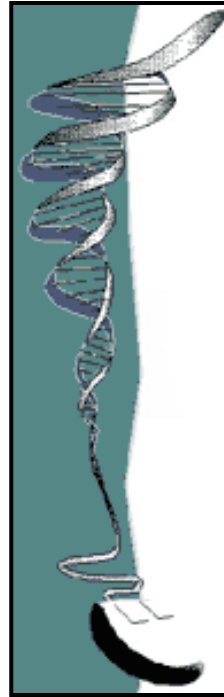
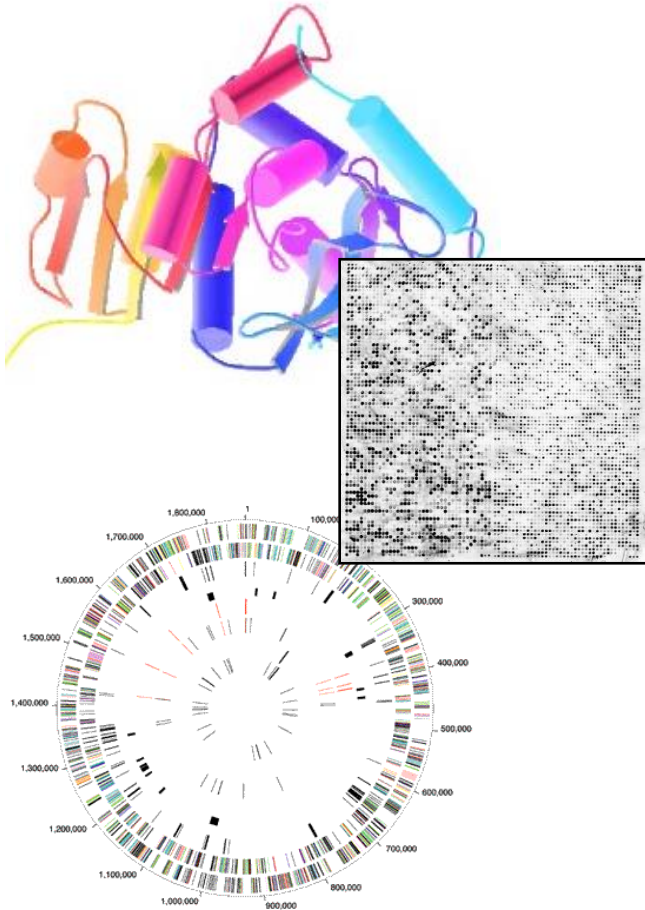


# Biomedical Data Science: Analysis of Network Topology -- Network Generation Models



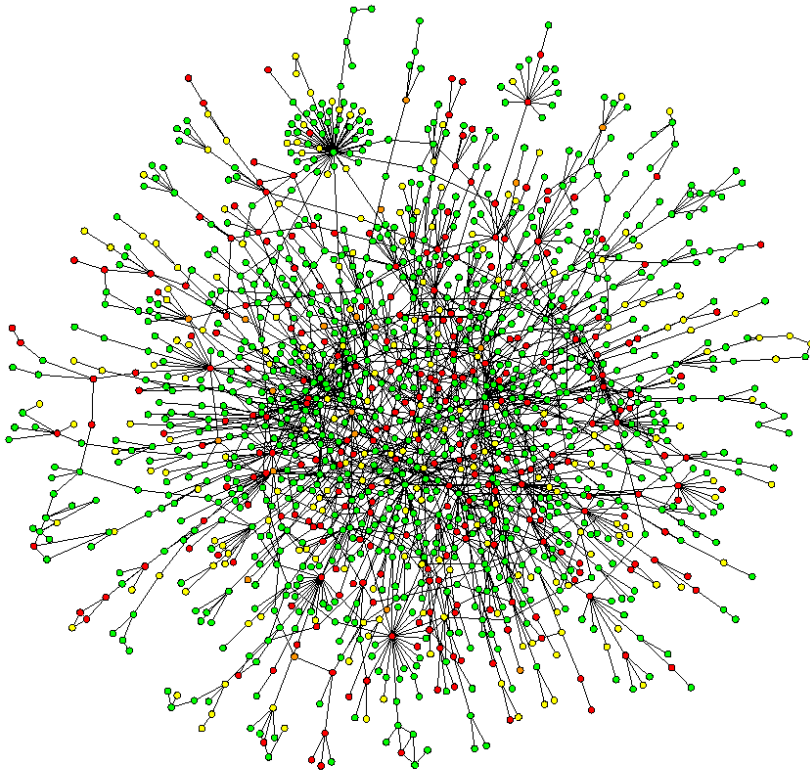
Mark Gerstein, Yale University  
[gersteinlab.org/courses/452](http://gersteinlab.org/courses/452)

(Last edit in spring '22; pack 22m10c, very similar to M10c from '21)

# Network Topology

**Simple Mathematical Models  
for Interpreting Complex  
Topology: ER Model & Small  
World Networks**

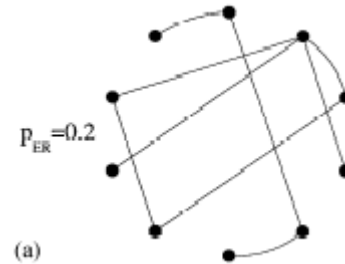
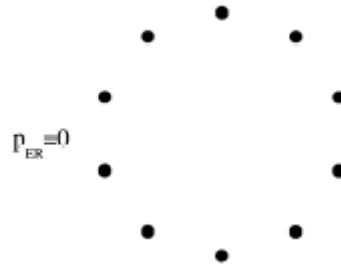
## Models for networks of complex topology



- Erdos-Renyi (1960)
- Watts-Strogatz (1998)
- Barabasi-Albert (1999)

**A Barabási & R Albert  
"Emergence of scaling in  
random networks,"  
*Science* 286, 509-512 (1999).**

# The Erdős-Rényi [ER] model (1960)

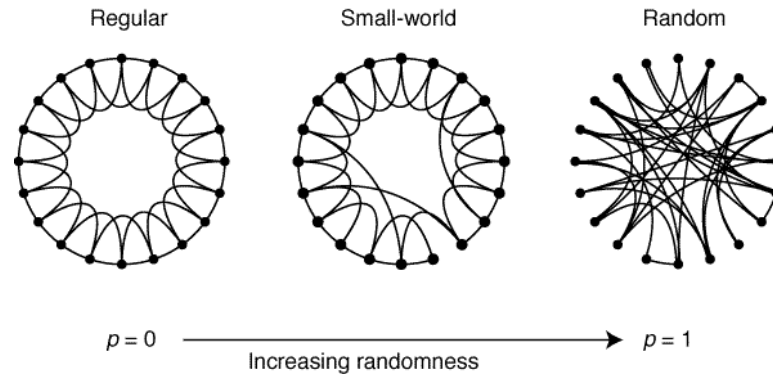


- Start with  $N$  vertices and no edges
- Connect each pair of vertices with probability  $P_{ER}$

**Important result:** many properties in these graphs appear quite suddenly, at a threshold value of  $P_{ER}(N)$

- If  $P_{ER} \sim c/N$  with  $c < 1$ , then almost all vertices belong to isolated trees
- Cycles of all orders appear at  $P_{ER} \sim 1/N$

# The Watts-Strogatz [WS] model (1998)



- Start with a regular network with  $N$  vertices
- Rewire each edge with probability  $p$

For  $p=0$  (Regular Networks):

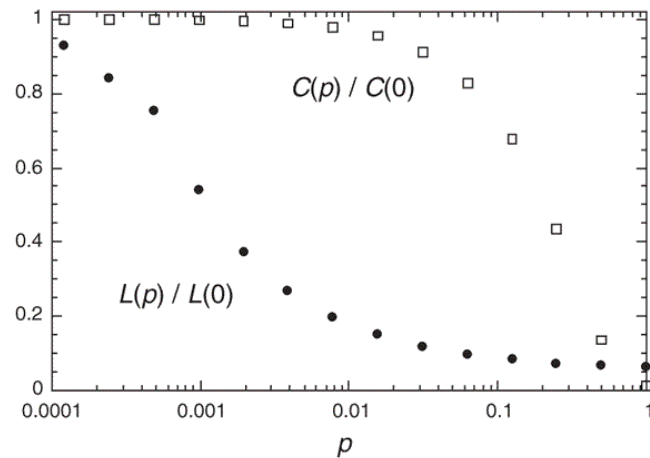
- high clustering coefficient
- high characteristic path length

For  $p=1$  (Random Networks):

- low clustering coefficient
- low characteristic path length

QUESTION: What happens for intermediate values of  $p$ ?

1) There is a broad interval of  $p$  for which  $L$  is small but  $C$  remains large



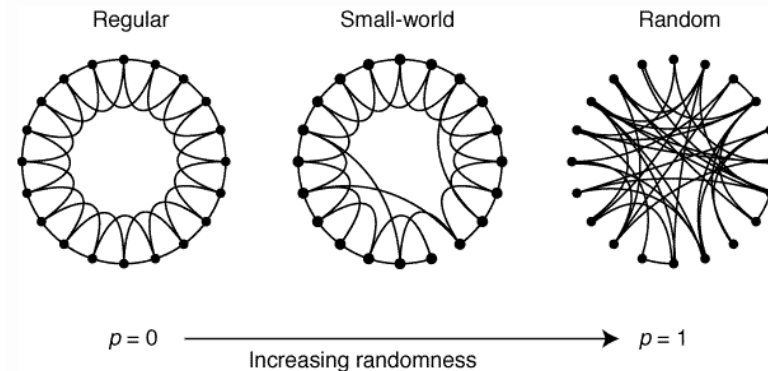
2) Small world networks are common :

**Table 1 Empirical examples of small-world networks**

	$L_{\text{actual}}$	$L_{\text{random}}$	$C_{\text{actual}}$	$C_{\text{random}}$
Film actors	3.65	2.99	0.79	0.00027
Power grid	18.7	12.4	0.080	0.005
<i>C. elegans</i>	2.65	2.25	0.28	0.05

# Small world network

- A simple connected graph  $G$  exhibiting two properties:
  - **Large Clustering Coefficient:** Each vertex of  $G$  is linked to a relatively well-connected set of neighboring vertices, resulting in a large value for the clustering coefficient  $C(G)$ ;
  - **Small Characteristic Path Length:** The presence of short-cut connections between some vertices results in a small characteristic path length  $L(G)$ .



- local connectivity and global reach

# Network Topology

**Simple Mathematical Models  
for Interpreting Complex  
Topology: BA Model & Scale  
Free Networks**



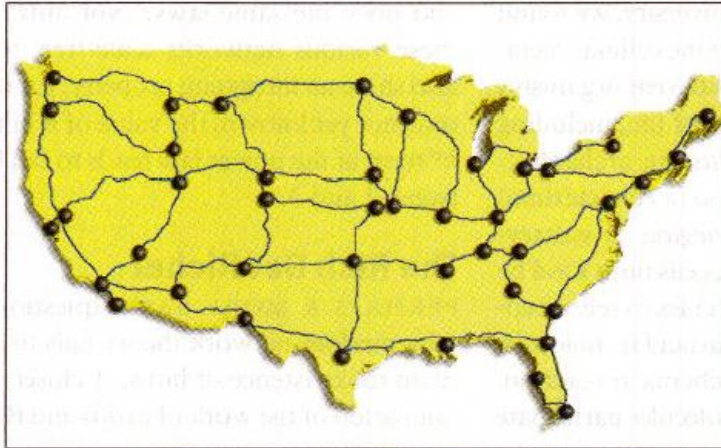
# Random v Scale-free Networks

RANDOM NETWORKS, which resemble the U.S. highway system (*simplified in left map*), consist of nodes with randomly placed connections. In such systems, a plot of the distribution of node linkages will follow a bell-shaped curve (*left graph*), with most nodes having approximately the same number of links.

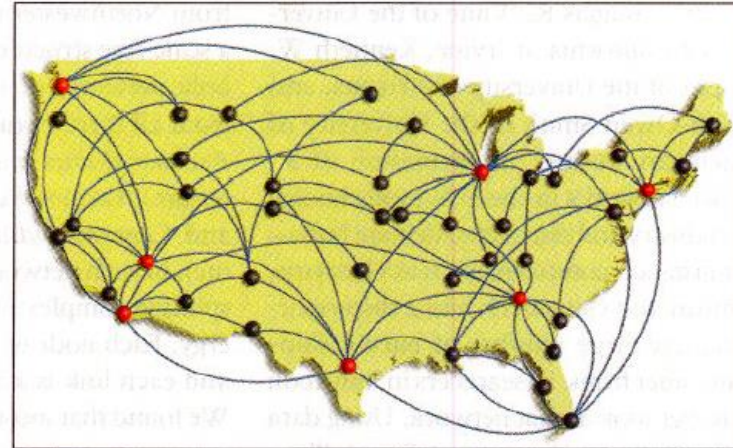
In contrast, scale-free networks, which resemble the U.S. airline system (*simplified in right map*), contain hubs (red)—

nodes with a very high number of links. In such networks, the distribution of node linkages follows a power law (*center graph*) in that most nodes have just a few connections and some have a tremendous number of links. In that sense, the system has no “scale.” The defining characteristic of such networks is that the distribution of links, if plotted on a double-logarithmic scale (*right graph*), results in a straight line.

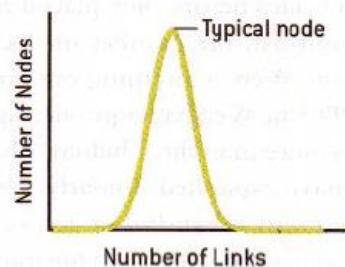
Random Network



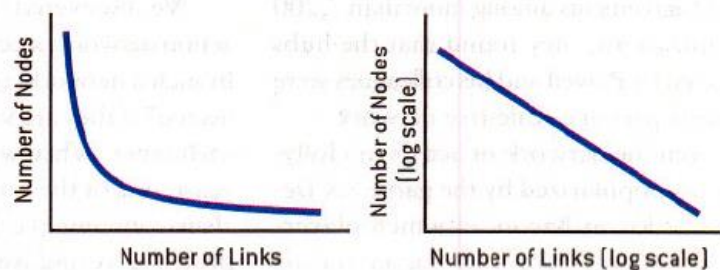
Scale-Free Network



Bell Curve Distribution of Node Linkages

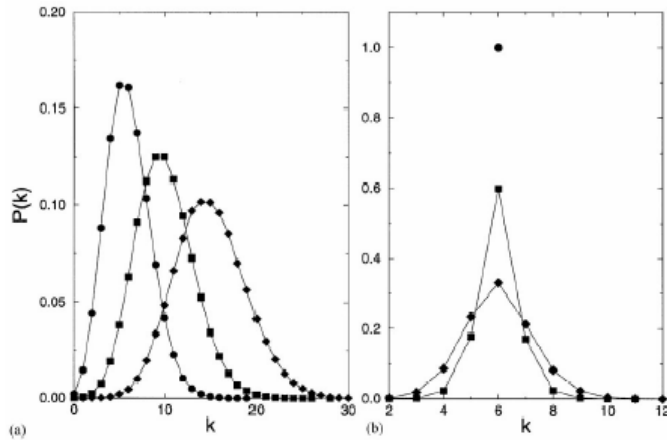


Power Law Distribution of Node Linkages



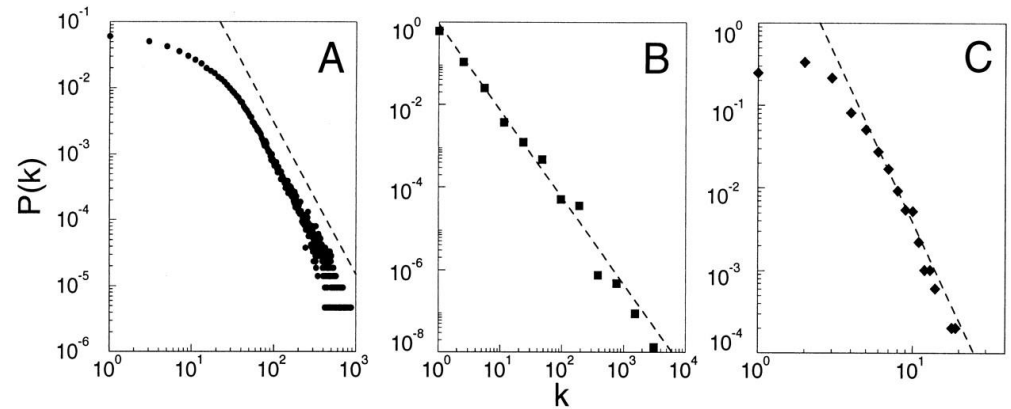
# The Barabási-Albert [BA] model (1999)

Look at the distribution of degrees



ER Model

WS Model



actors

power grid

www

The probability of finding a highly connected node decreases exponentially with  $k$

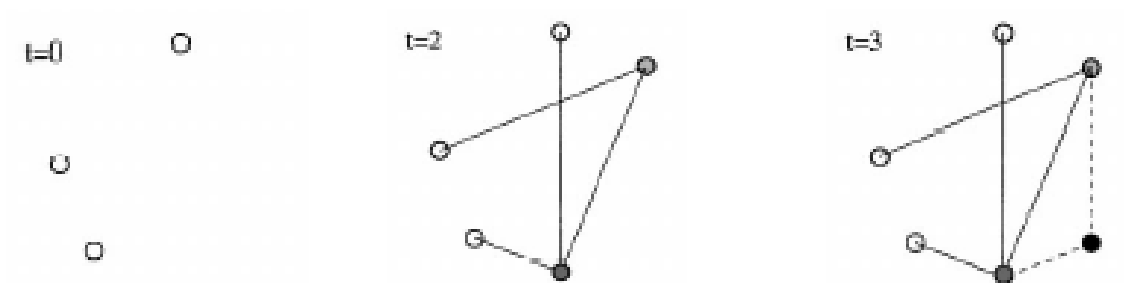
$$P(K) \sim K^{-\gamma}$$

- two problems with the previous models:
  1. N does not vary
  2. the probability that two vertices are connected is uniform

- **GROWTH:** starting with a small number of vertices  $m_0$  at every timestep add a new vertex with  $m \leq m_0$

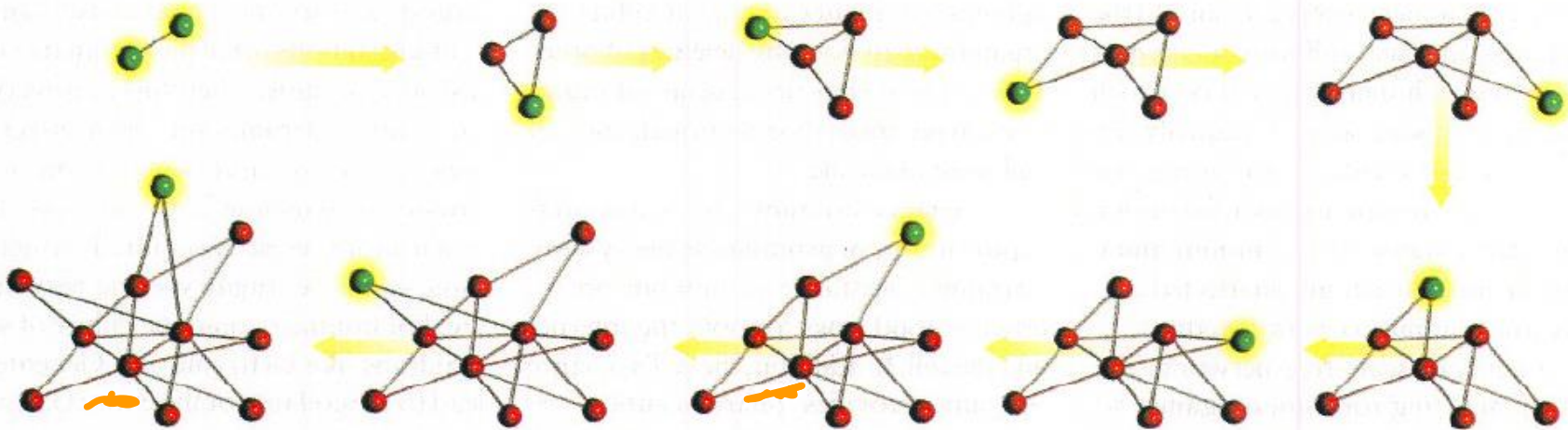
- **PREFERENTIAL ATTACHMENT:** the probability  $\Pi$  that a new vertex will be connected to vertex  $i$  depends on the connectivity of that vertex:

$$\Pi(k_i) = \frac{k_i}{\sum_j k_j}$$



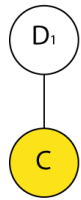
# Birth of Scale-Free Network

A SCALE-FREE NETWORK grows incrementally from two to 11 nodes in this example. When deciding where to establish a link, a new node [*green*] prefers to attach to an existing node [*red*] that already has many other connections. These two basic mechanisms—growth and preferential attachment—will eventually lead to the system's being dominated by hubs, nodes having an enormous number of links.

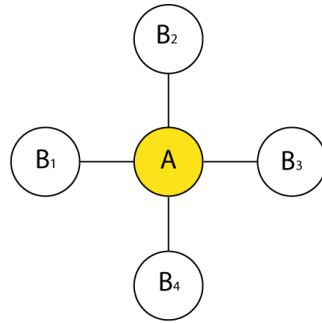


# SCALE FREENESS GENERALLY EVOLVES THROUGH PREFERENTIAL ATTACHMENT (THE RICH GET RICHER)

## The Duplication Mutation Model



Gene duplication



The interaction partners of A are more likely to be duplicated

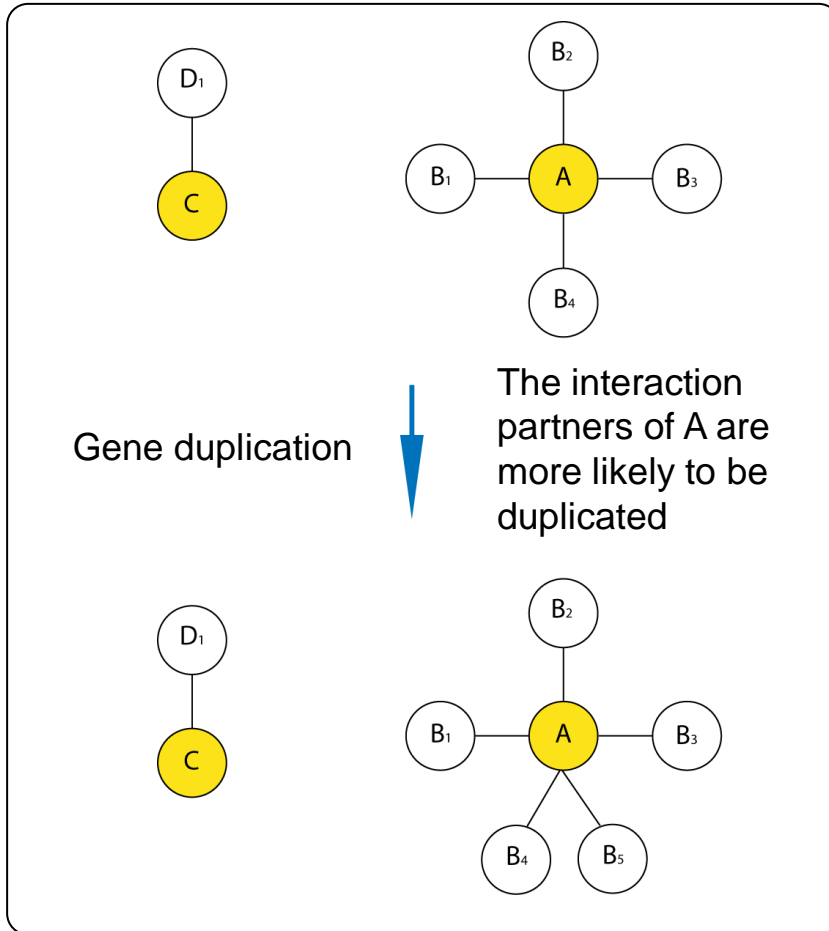
## Description

- Theoretical work shows that a mechanism of preferential attachment leads to a scale-free topology (“The rich get richer”)
- In interaction network, gene duplication followed by mutation of the duplicated gene is generally thought to lead to preferential attachment
- Simple reasoning: The partners of a hub are more likely to be duplicated than the partners of a non-hub



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