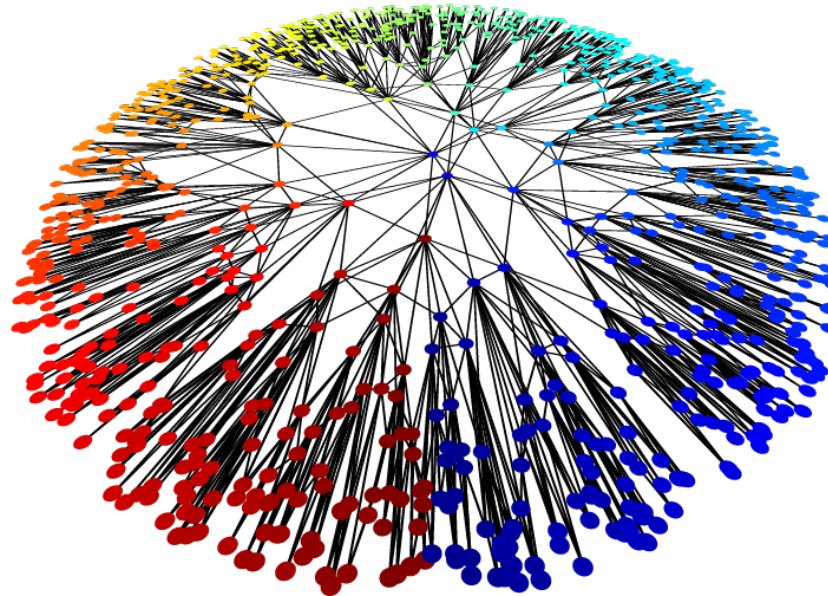


Complex systems and networks

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Thanks to Dr. Gregorio Alanis-Lobato for providing parts of the material for this course

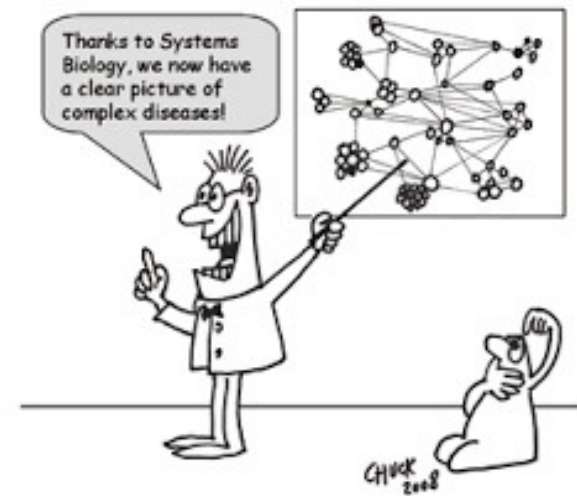
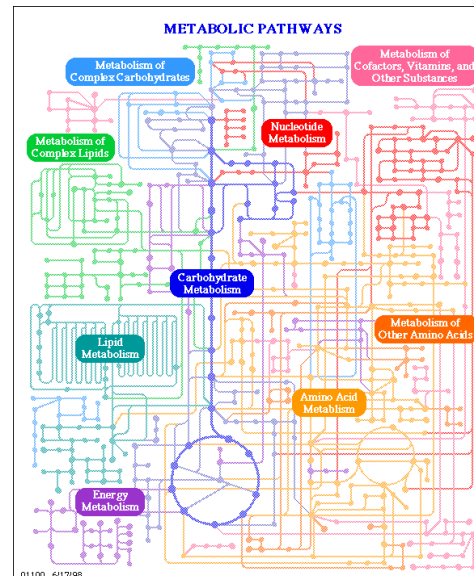
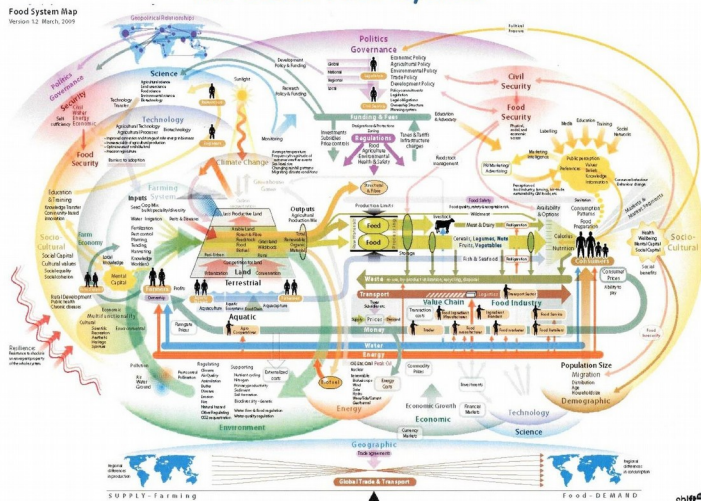
What is a complex system?

No generally accepted definition so far

Complex systems typically have several of the following properties:

- They are composed of many components
- Show emergent behavior
- Contain nonlinearities
- Are open systems
- Evolve far from equilibrium
- Show chaotic behavior

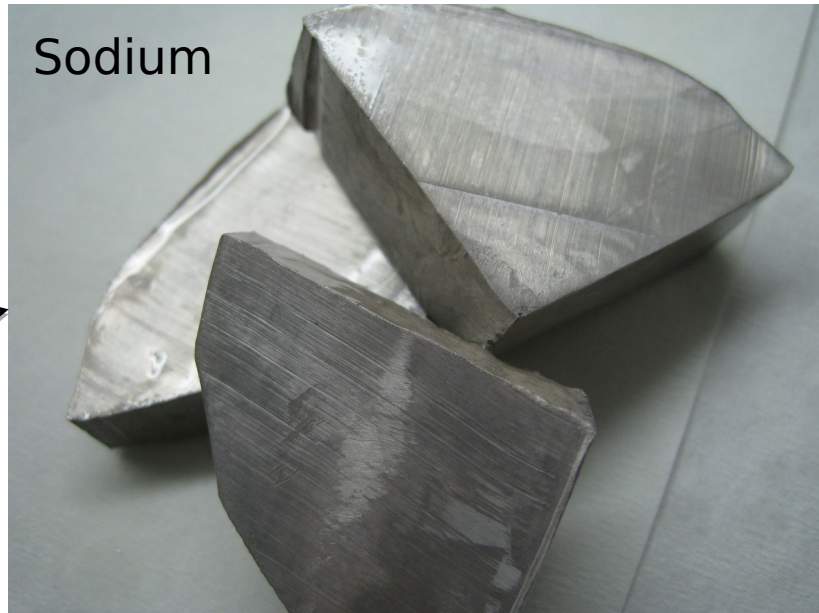
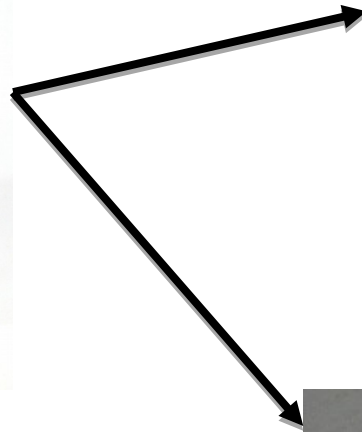
The Global Food System



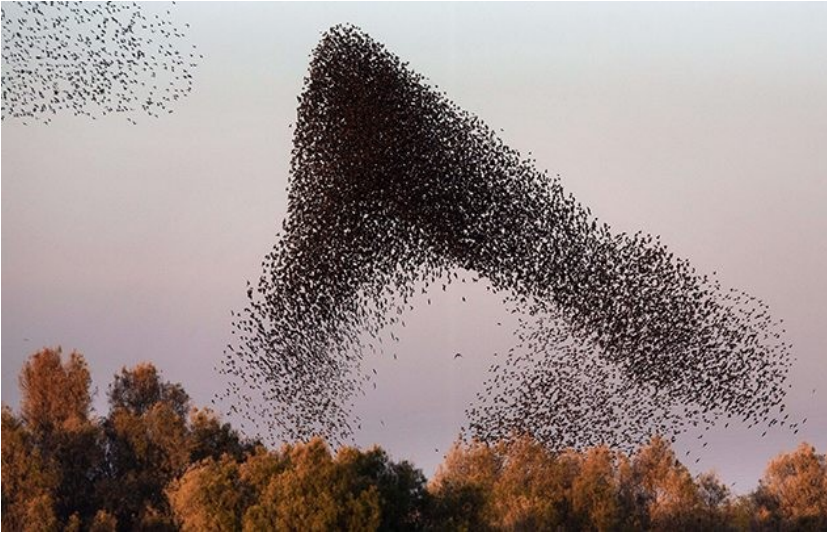
Emergent properties



The fallacy of division



Complex behaviours, simple rules



NatGeo

Flocking



Bored Panda

Complexity & Chaos

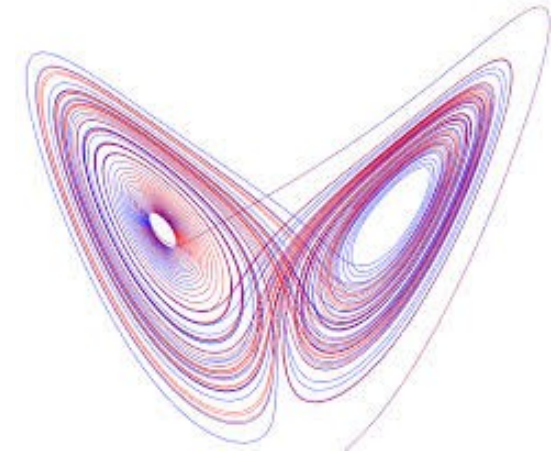
Chaotic systems:

- Difficult to predict, despite being deterministic
- Exact knowledge of equations and starting conditions allows exact simulation
- Slight differences in starting conditions or model errors can result in huge simulation errors

Chaotic vs. complex systems:

- Chaotic systems can be complex, but also “simple”
- Complex systems may show chaotic behavior, but don't have to

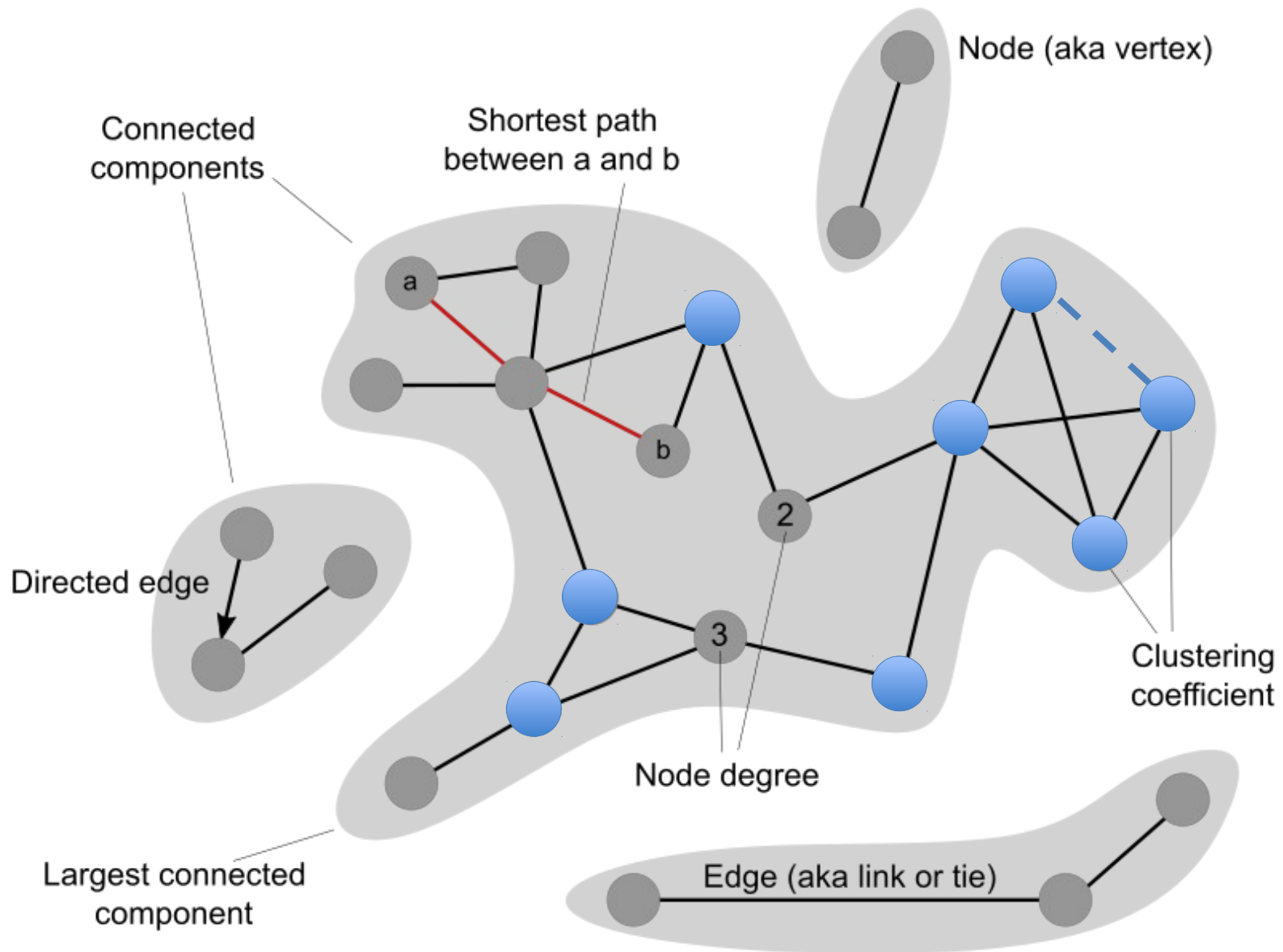
Example: Lorenz attractor



Simplified model for atmospheric convection

$$\begin{aligned}\frac{dx}{dt} &= \sigma(y - x), \\ \frac{dy}{dt} &= x(\rho - z) - y, \\ \frac{dz}{dt} &= xy - \beta z.\end{aligned}$$

Network representation of complex systems

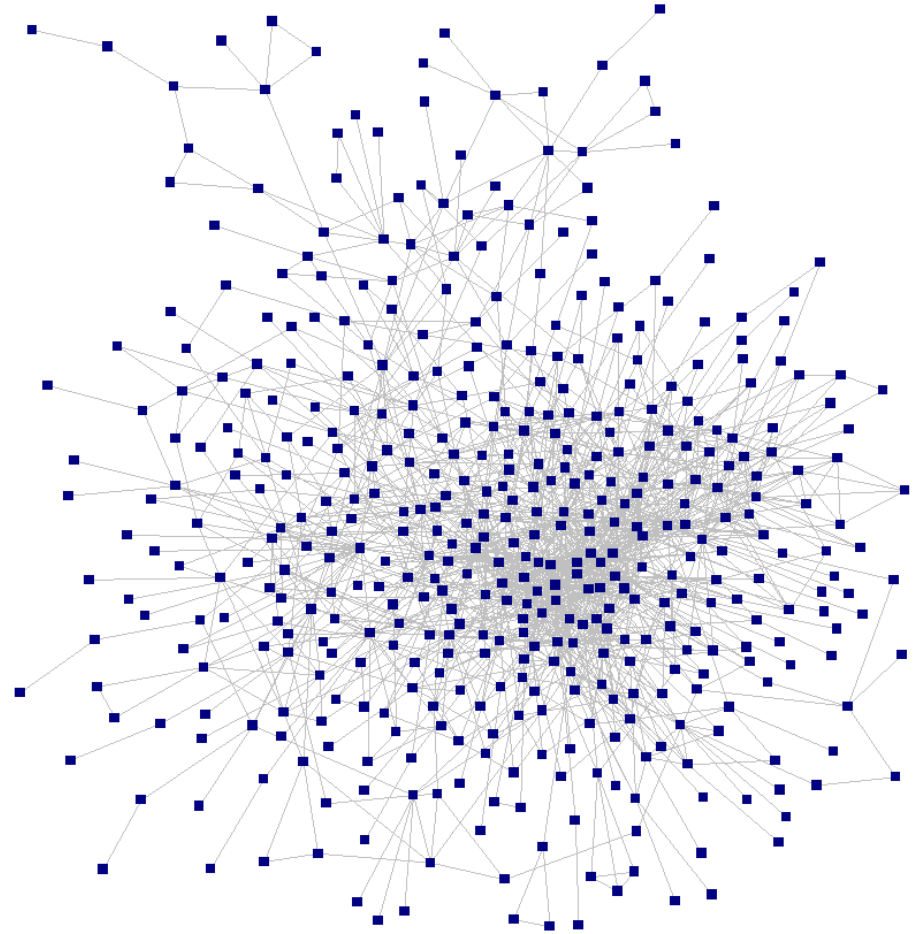


Graph:

$G=(V, E)$

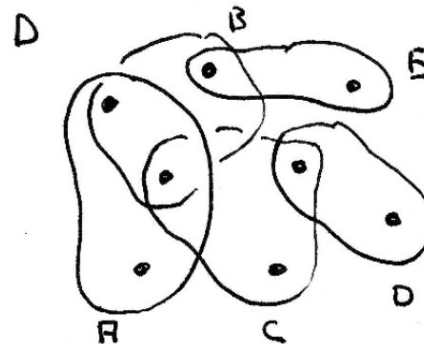
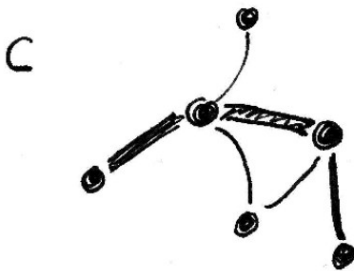
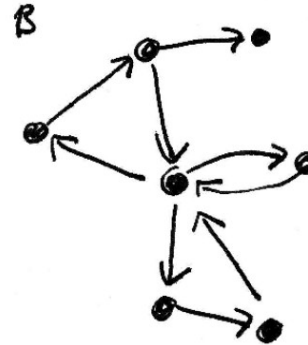
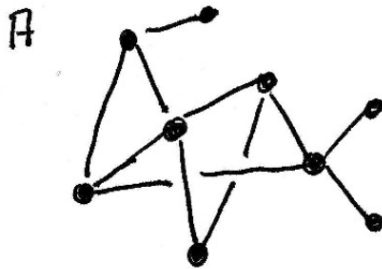
V: set of vertices

E: set of edges, each
associated with 2 vertices



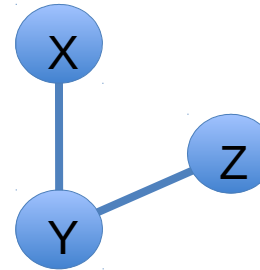
Network types

- A) Homogeneous undirected network → no direction, no weights
- B) Directed network
- C) Weighted network
- D) Hypergraphs and Bipartite network



Network representations

Drawing



Adjacency matrix

$$A = \begin{pmatrix} 0 & 1 & 0 \\ 1 & 0 & 1 \\ 0 & 1 & 0 \end{pmatrix}$$

Link table

X	Y
Y	Z

Linked list

X: {Y}
Y: {X,Z}
Z: {Y}

What about directed networks, weighted networks, and bipartite graphs?

Network properties

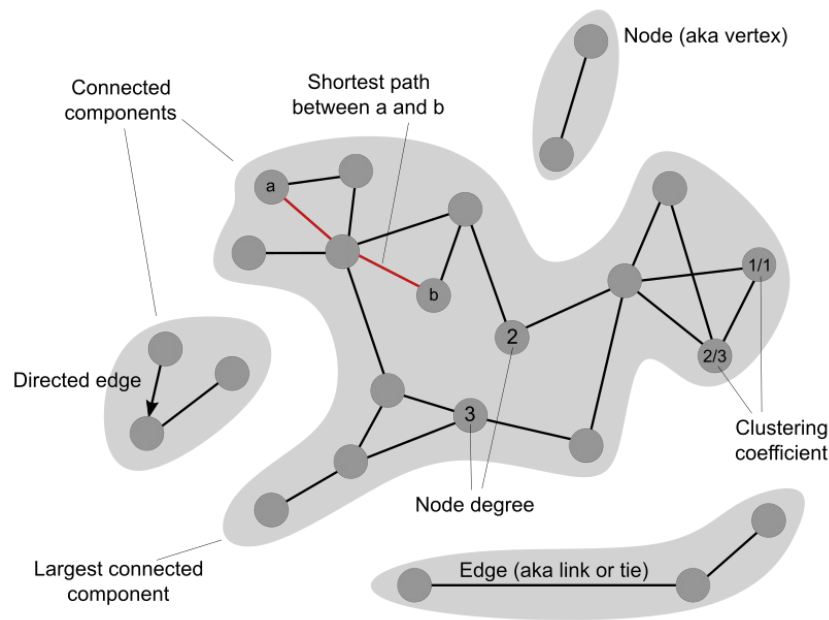
Node degree: Number of edges connected to a node
(in-degree and out-degree for directed networks)

Path: Set of edges

Shortest path: Path with minimal amount of edges connecting two nodes

(Connected) components: Set of nodes such that a path exists that connects each pair of nodes

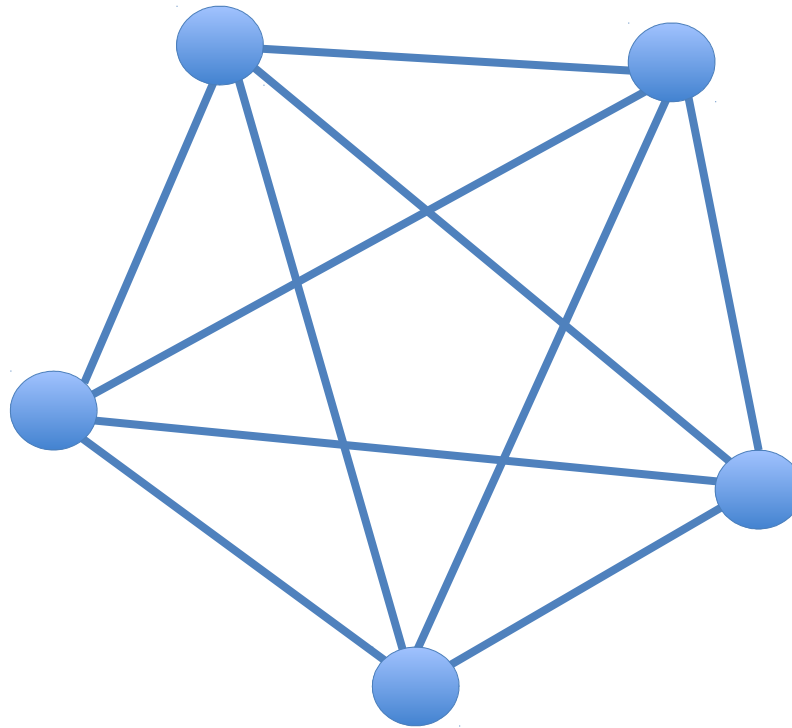
Clustering coefficient: Number of edges between neighboring nodes divided by maximum possible number of edges



Complex networks, simple rules

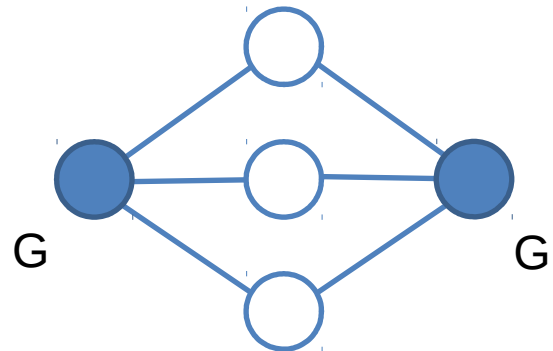
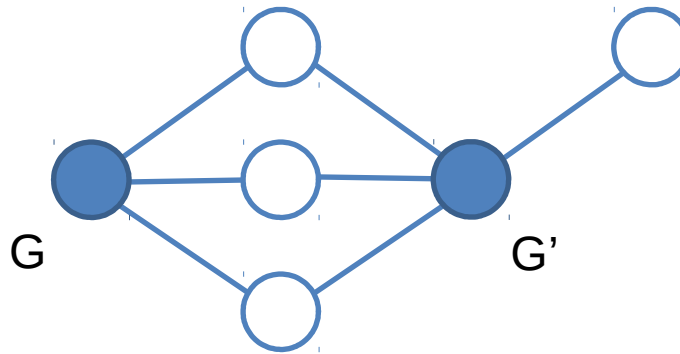
Random network: Form links between nodes with probability p

$$p = 1/6$$



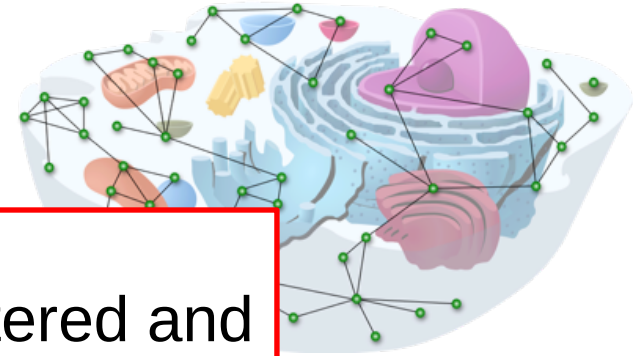
Complex networks, simple rules

Duplication-divergence



Retain links with probability p

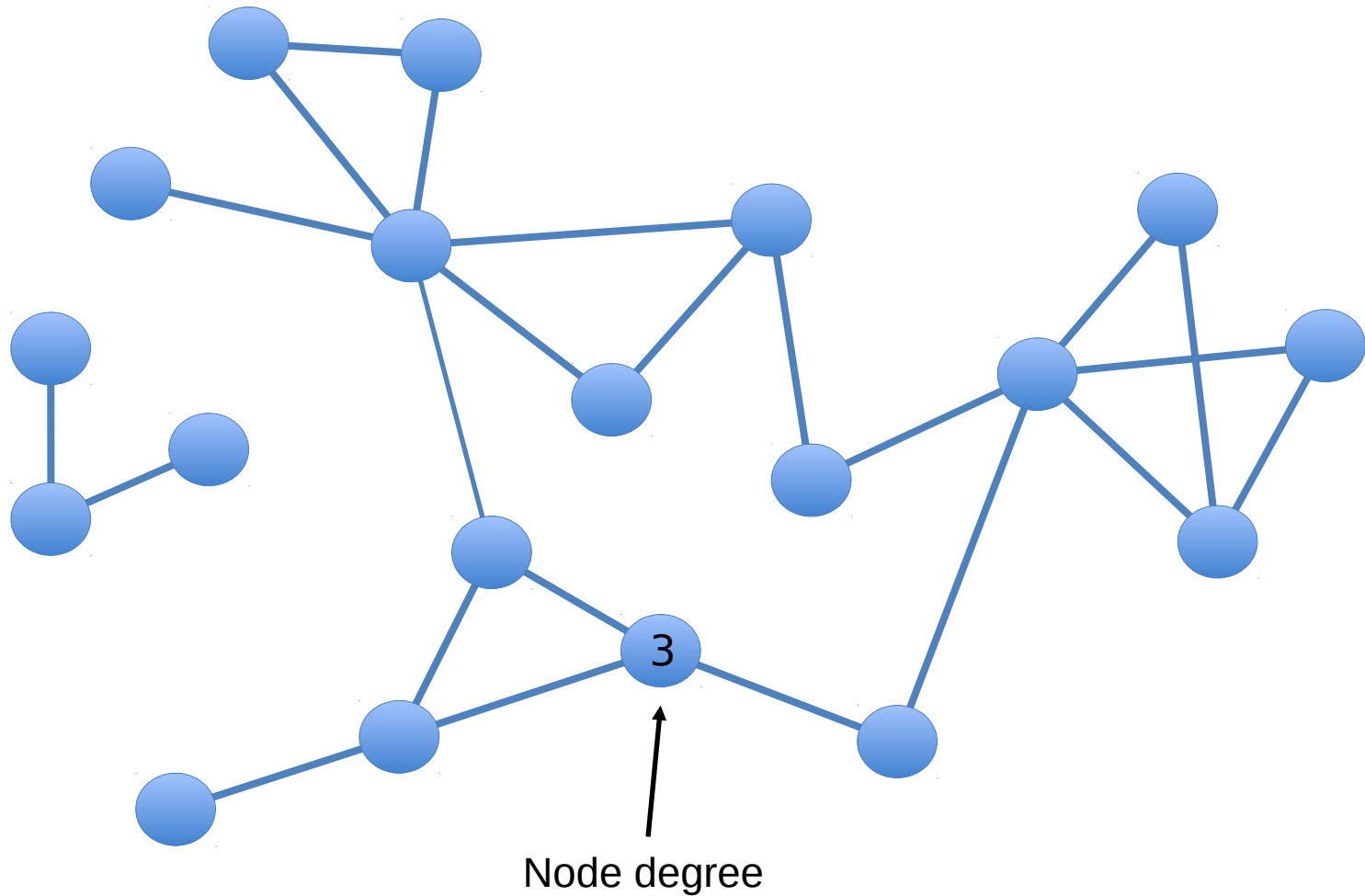
Different systems, same network structural properties



Scale-free, strongly clustered and small-world.

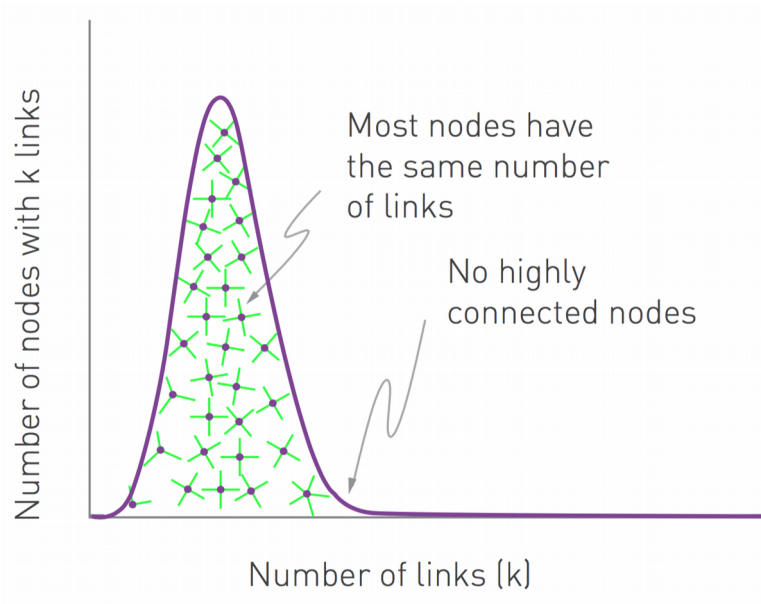


Complex networks are scale-free

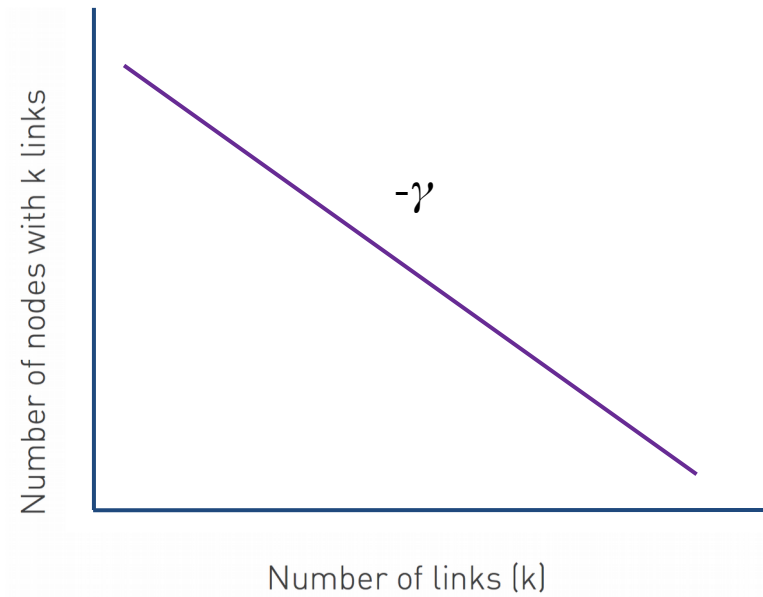


Complex networks are scale-free

Random network

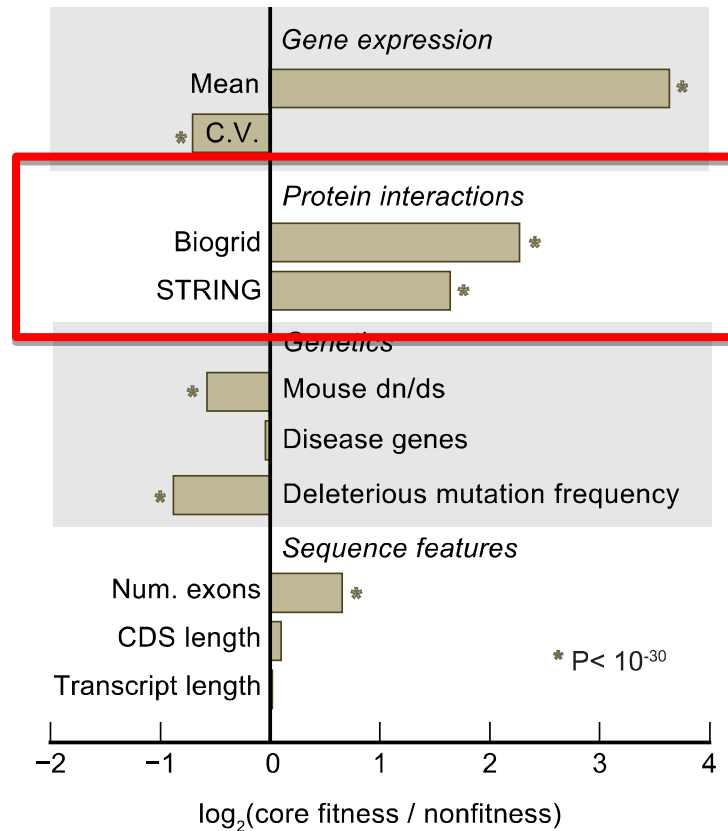


Complex network
Log-log scale



High-degree proteins: core fitness genes

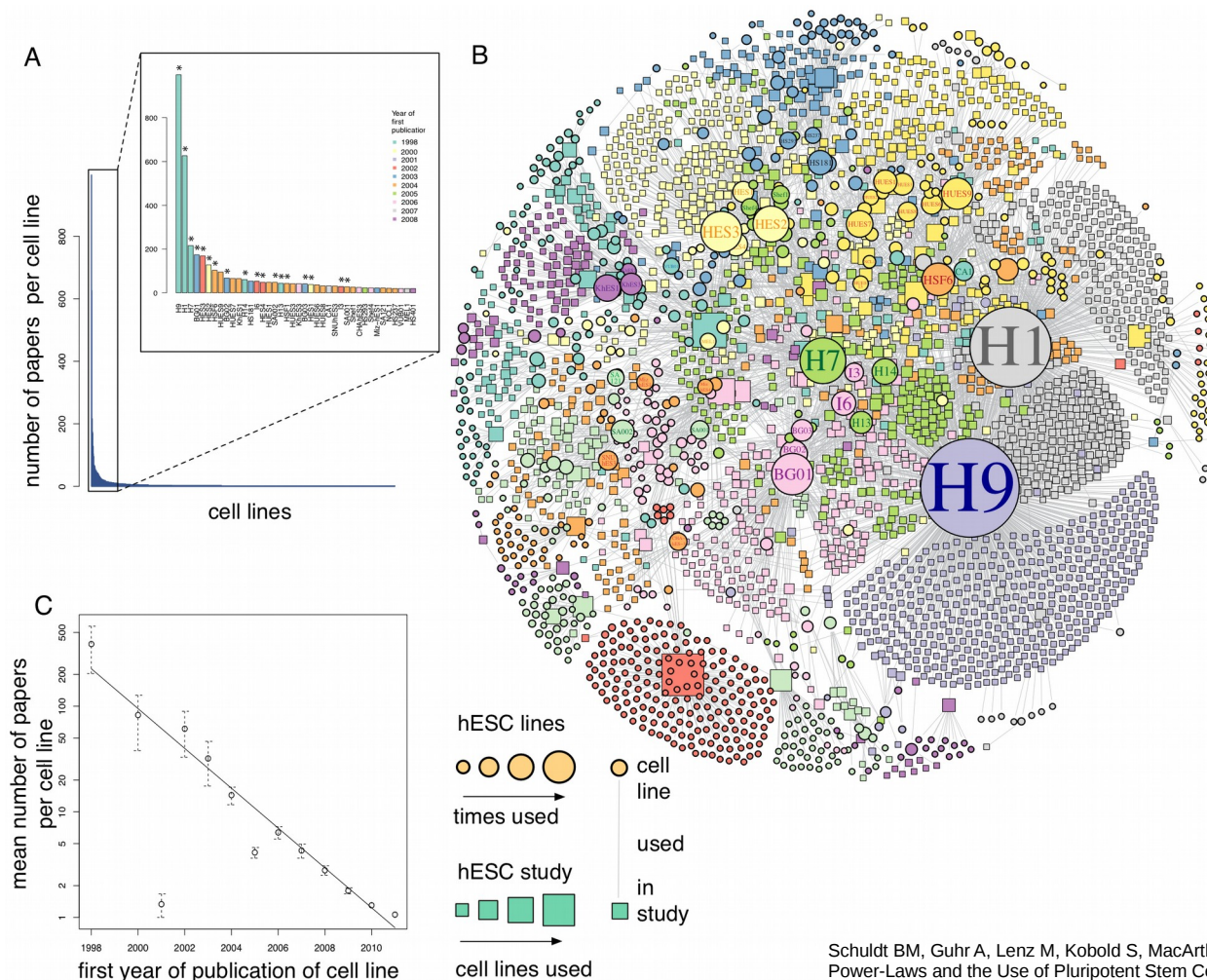
Fitness gene: Gene whose perturbation decreases cell growth and proliferation.



Hart *et al.* (2015) Cell

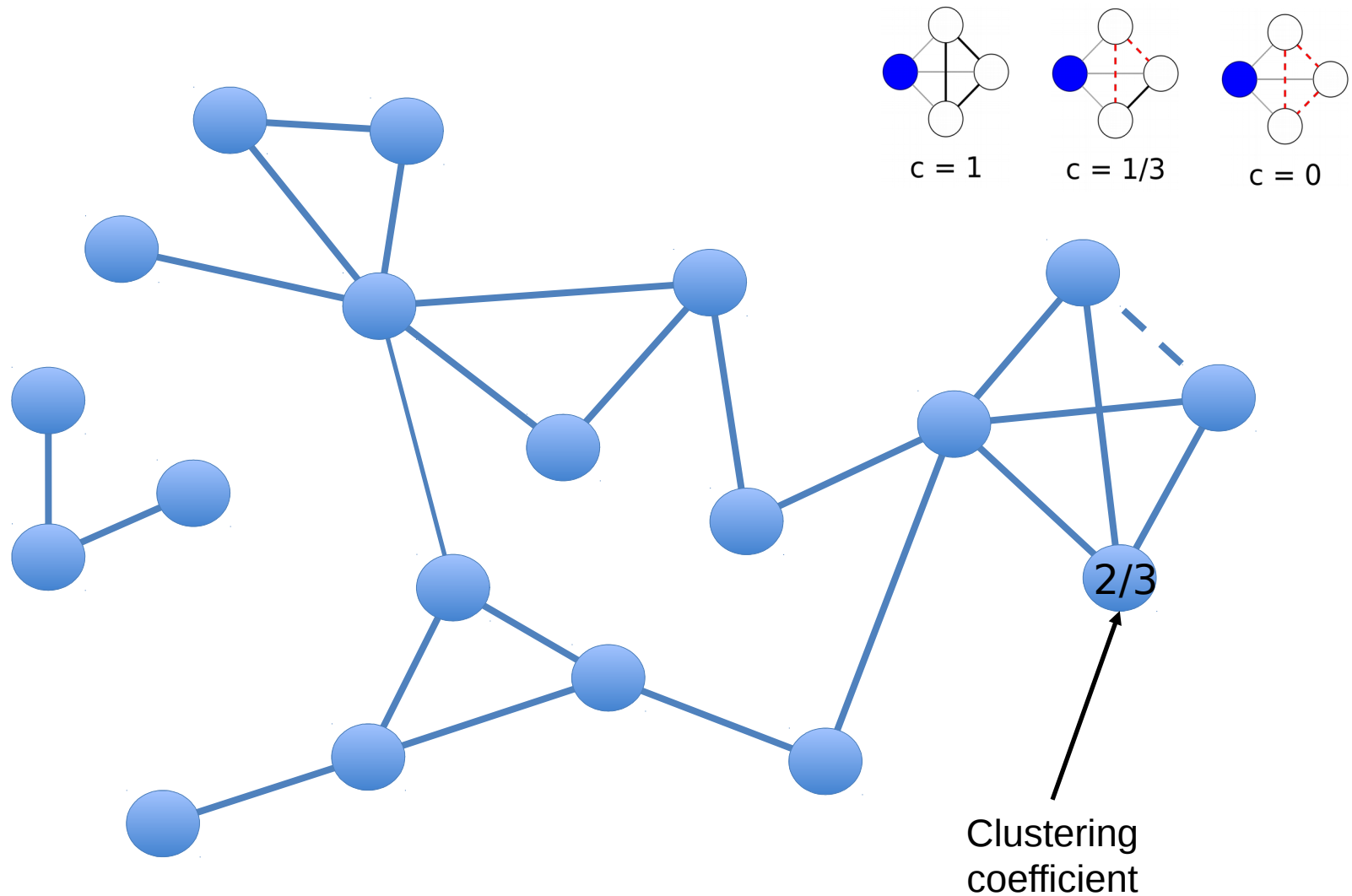
Scale free networks and preferential attachment

Example: Citation networks
Use of pluripotent stem cell lines

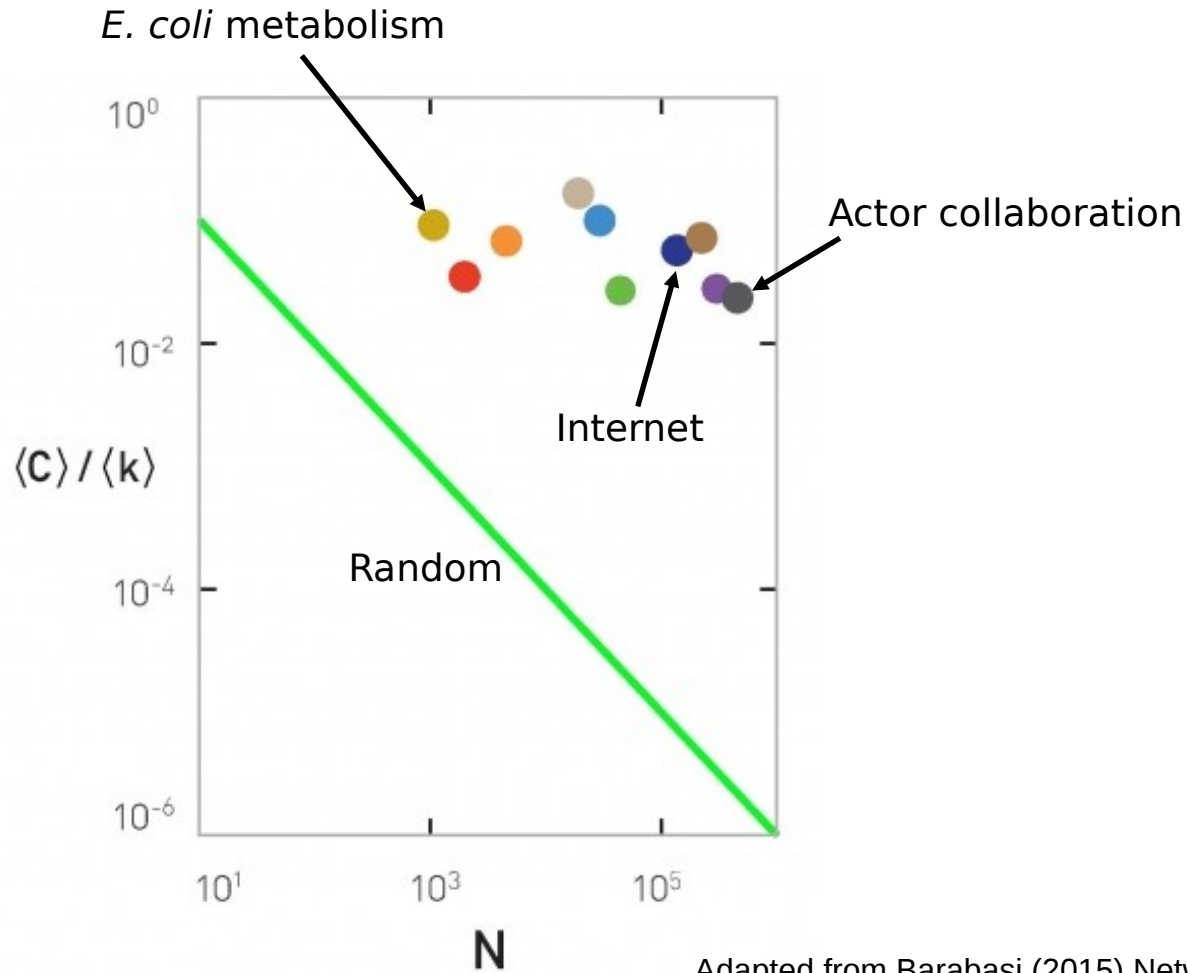


Schuldt BM, Guhr A, Lenz M, Kobold S, MacArthur BD, Schuppert A, et al. (2013) Power-Laws and the Use of Pluripotent Stem Cell Lines. PLoS ONE 8(1): e52068.

Complex networks are strongly clustered

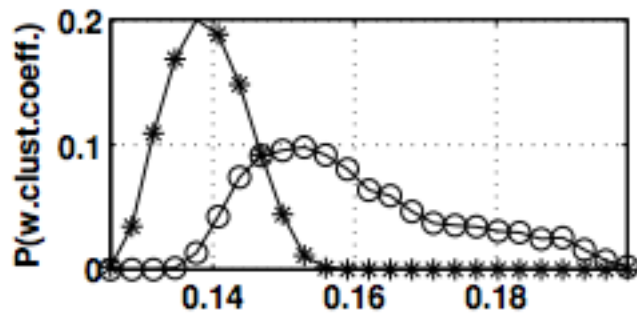


Real networks: strongly clustered

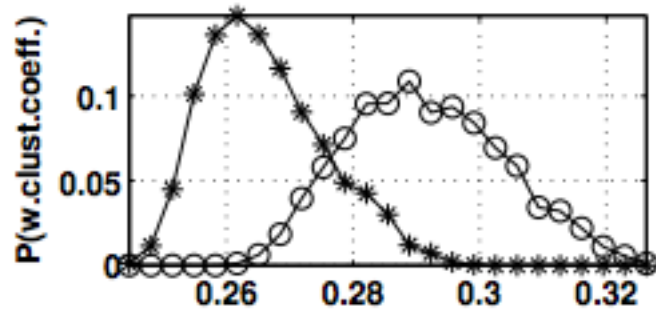


Adapted from Barabasi (2015) Network Science

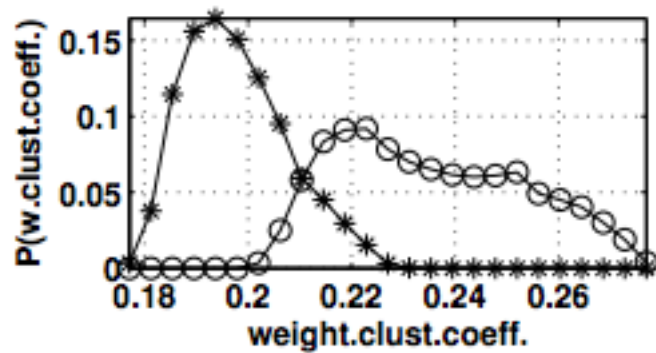
Cancer cells: reduced clustering



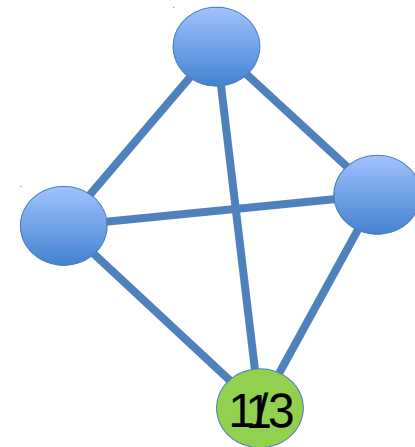
Liver



Breast



Lymphoma



- Genes in normal cells
- * Genes in cancer cells

Strong clustering motifs: protein complexes

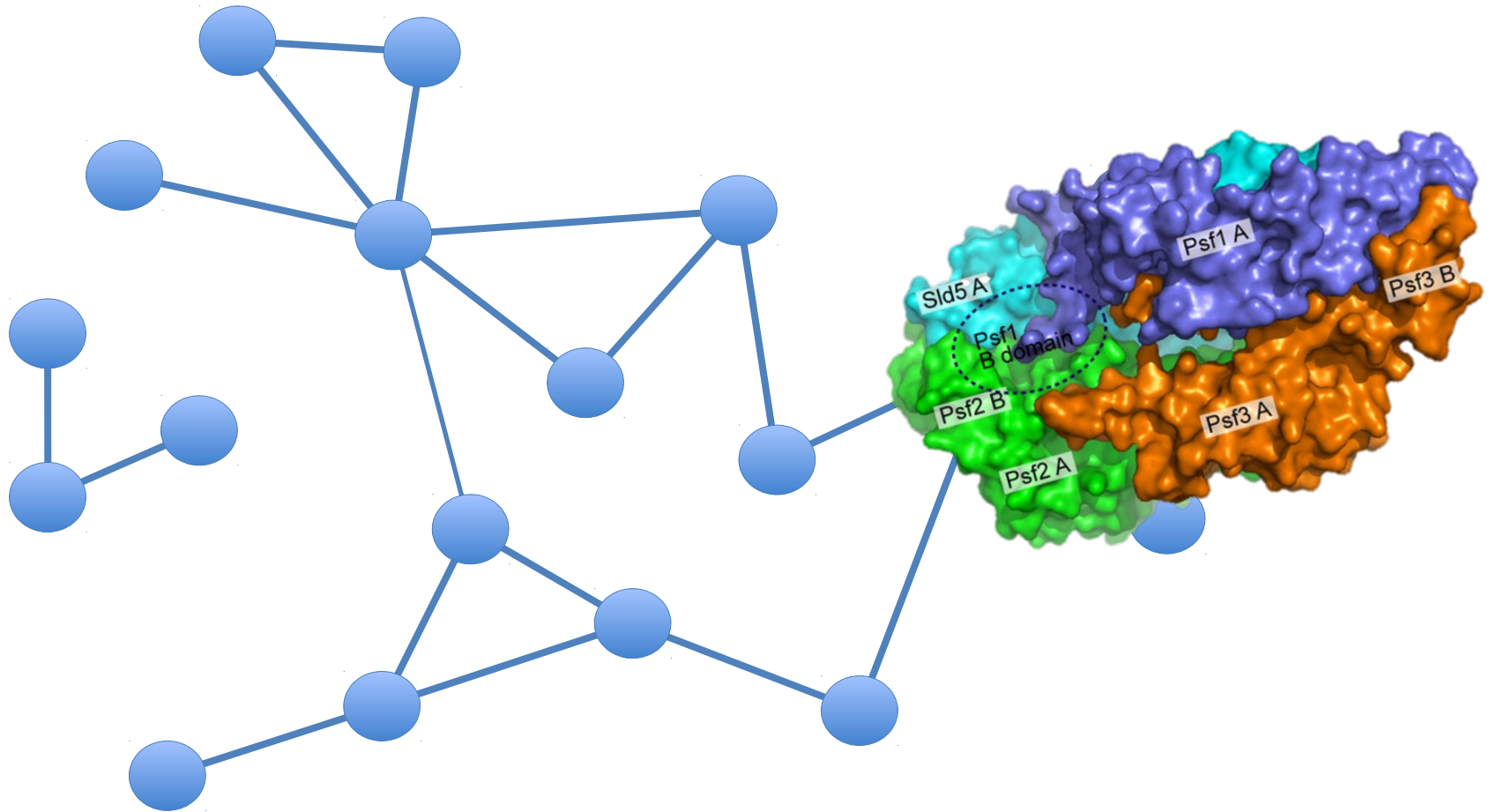
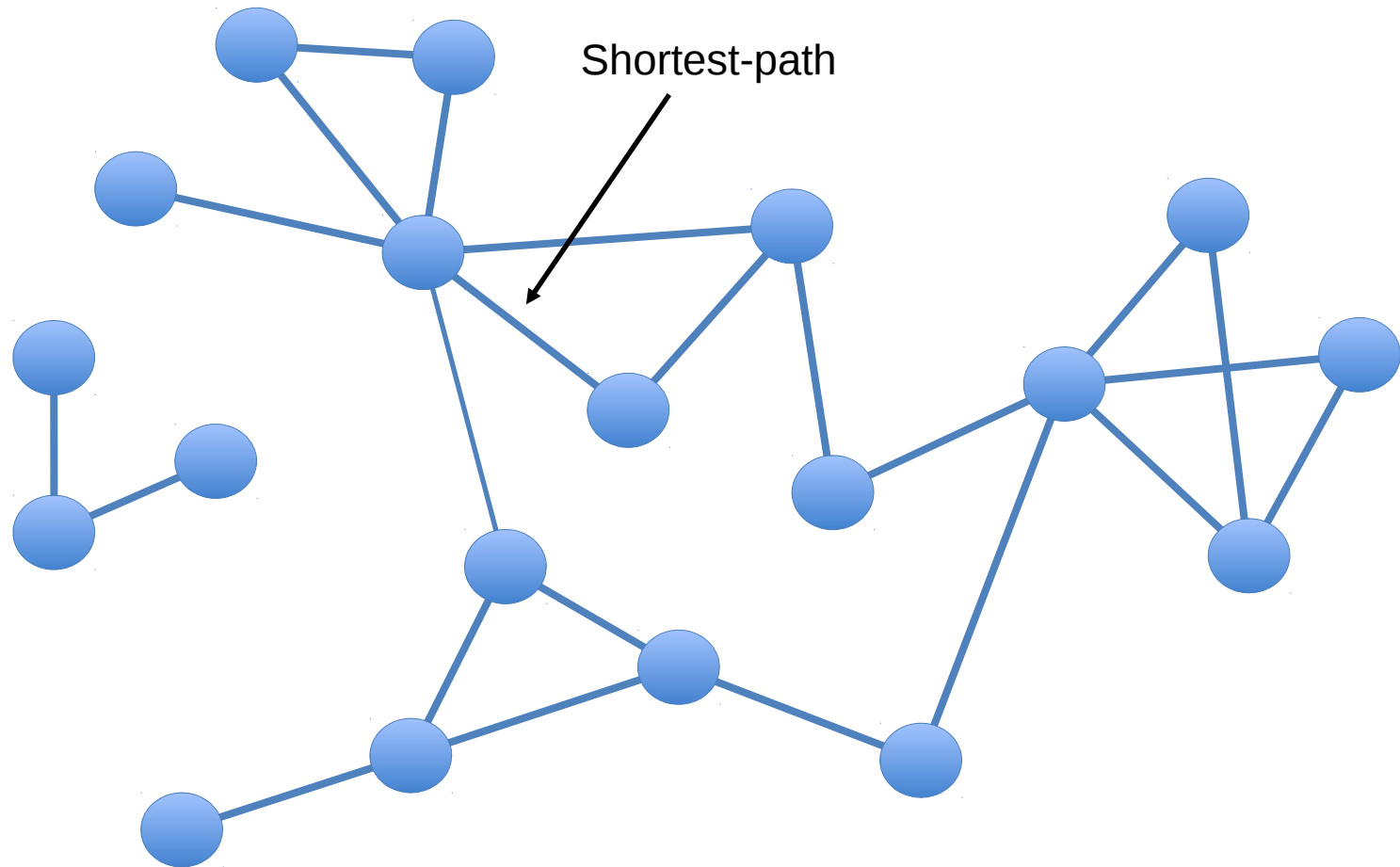
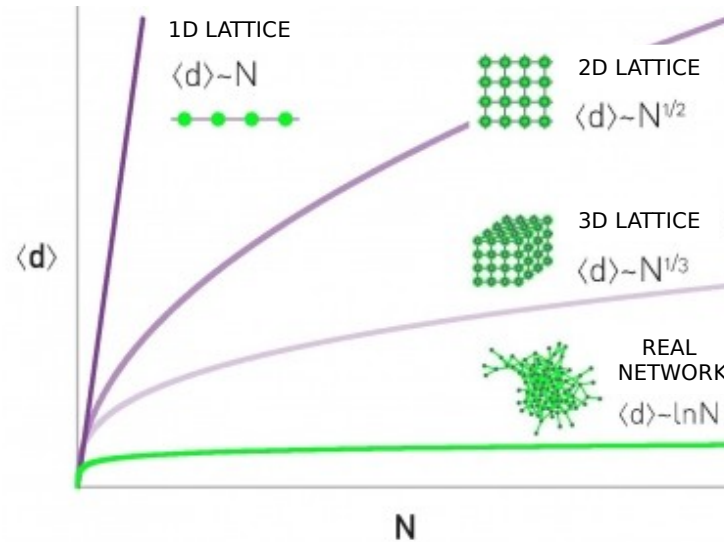


Figure from RIKEN

Complex networks are small-world

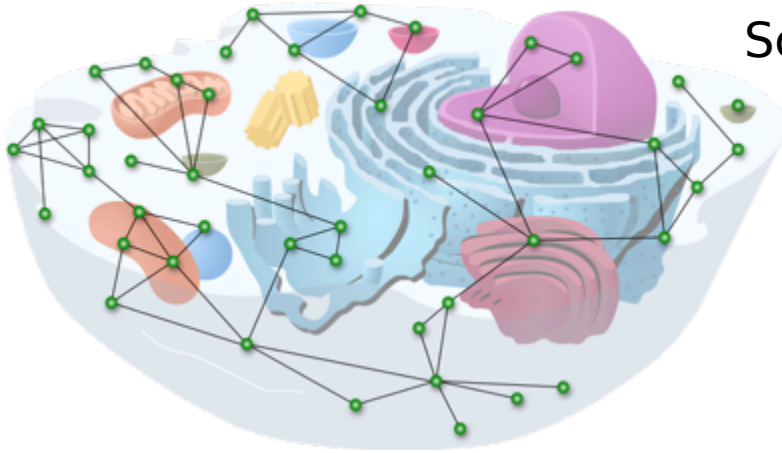


Shortest-paths and small worlds

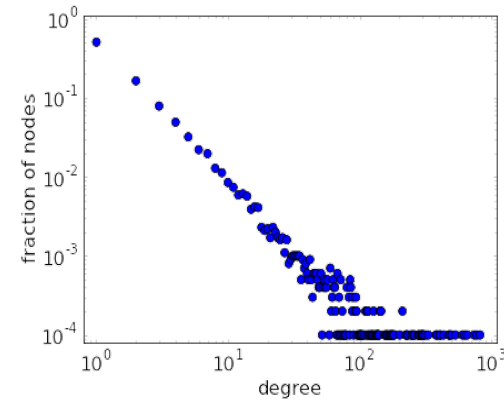


Adapted from Barabasi (2015) Network Science

Protein interaction network in the living cell



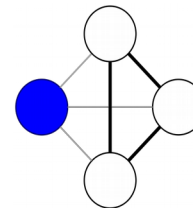
Scale-free/power law node degree distribution



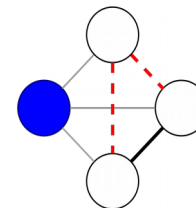
Small-world property

$$L \propto \log N$$

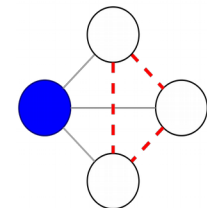
Strong average clustering coefficient



$$c = 1$$



$$c = 1/3$$



$$c = 0$$

Take-home message

A wide range of network-based approaches have been and are being developed to address problems with relevance to biology and human health. Some of the scenarios where network analysis is playing an important role are:

- Gene function prediction
- Detection of protein complexes and other modular structures
- Prediction of new interactions
- Analysis of disease modules

THANK YOU

GRACIAS

DANKE

شكراً

KÖSZÖNÖM

ви благодарам

MERCI

धन्यवाद

GRAZIE

谢谢

