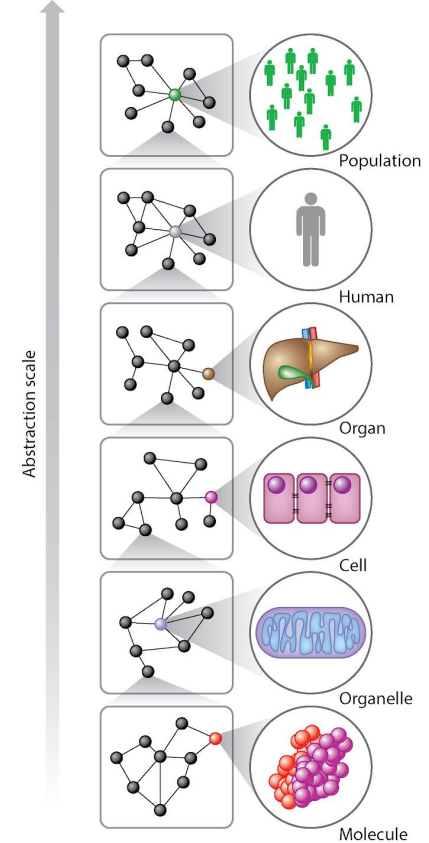
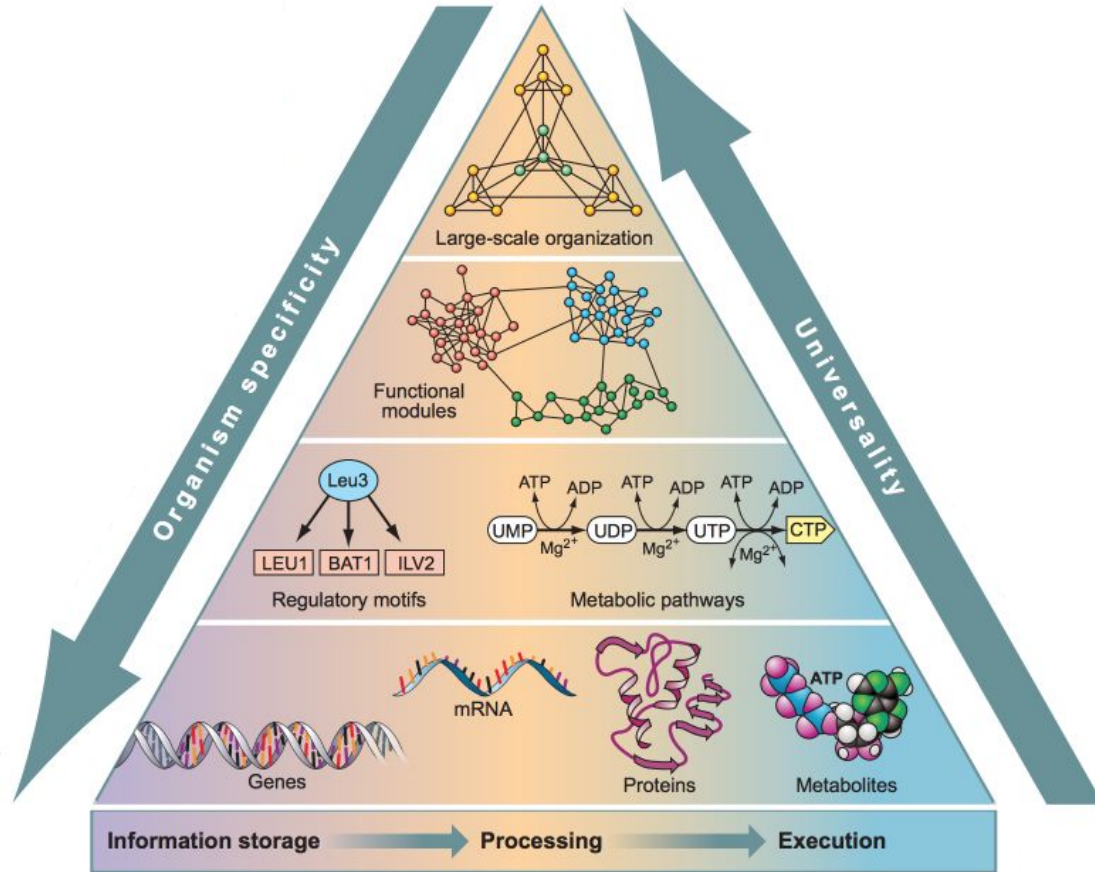


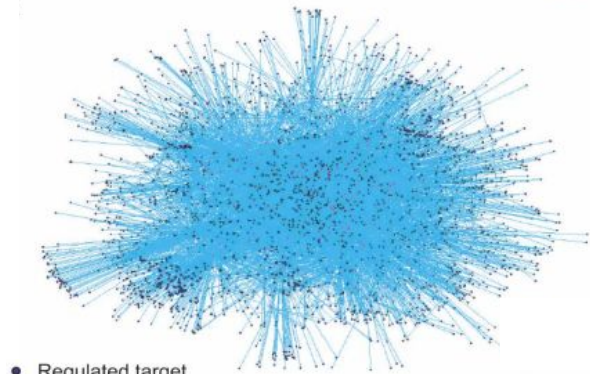
Week 15: Large-scale biological networks

- Network topology
- Network motifs
- Condition-specific networks
- Network reconstruction
- Network propagation

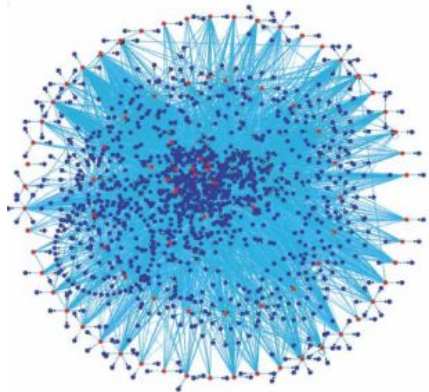
Layers of complexity and Network representations



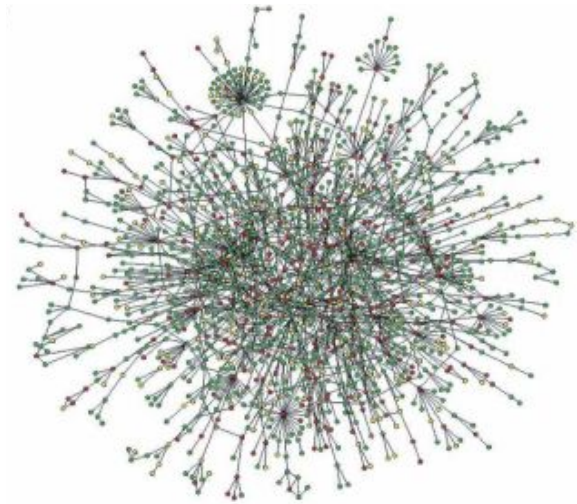
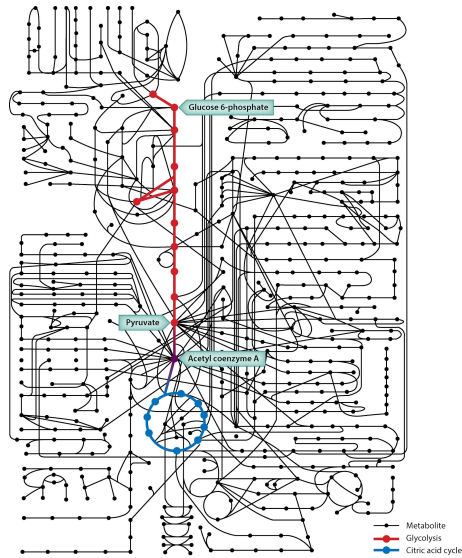
Molecular networks



- Regulated target
- Transcription factor

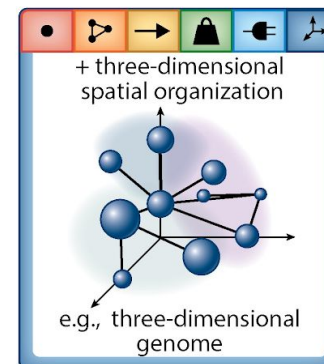
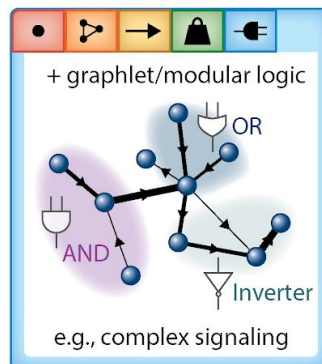
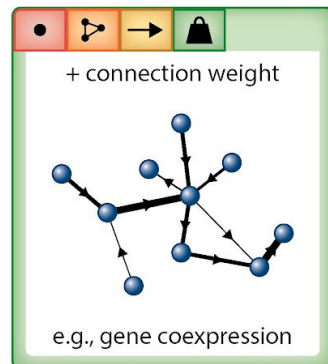
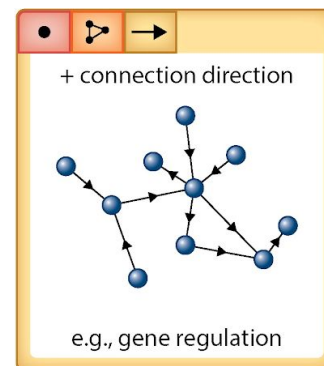
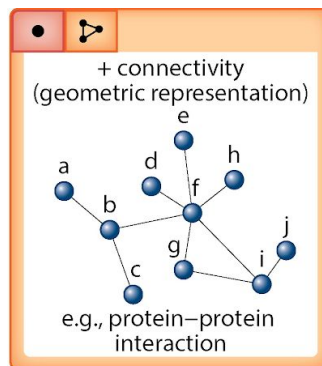
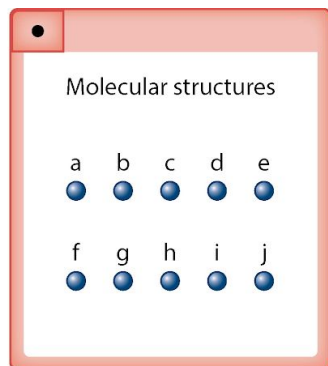


- Kinase
- Regulated target



- Lethal
- Slow growth
- Unknown
- Non-lethal

Network description



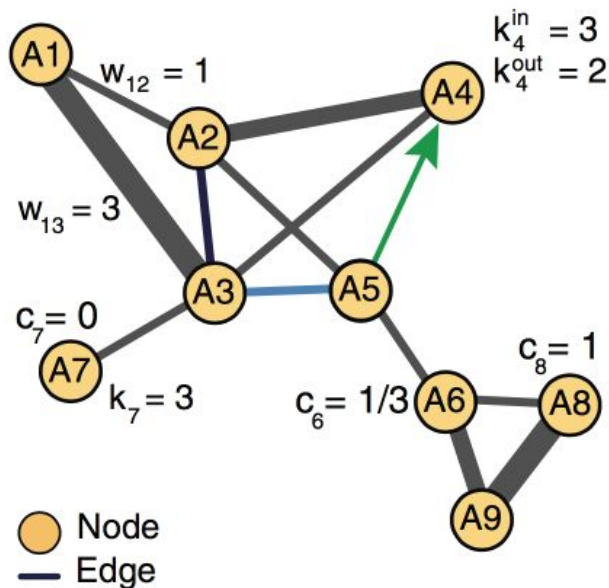
Network representations

Relationships

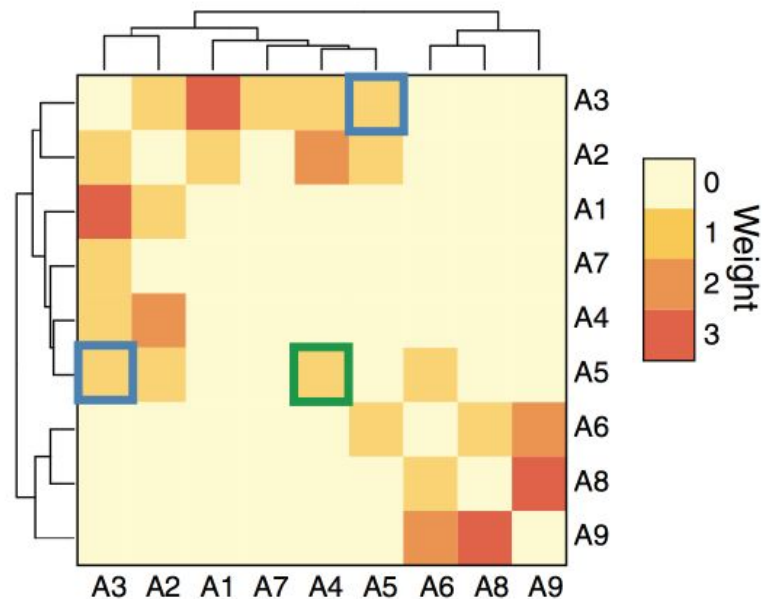
Optional weight

A1 ↔ A2	1
A1 ↔ A3	3
A2 ↔ A3	1
A2 ↔ A4	2
A2 ↔ A5	1
A3 ↔ A4	1
A3 ↔ A5	1
A3 ↔ A7	1
A5 → A4	1
A5 ↔ A6	1
A6 ↔ A8	1
A6 ↔ A9	2
A8 ↔ A9	3

Network

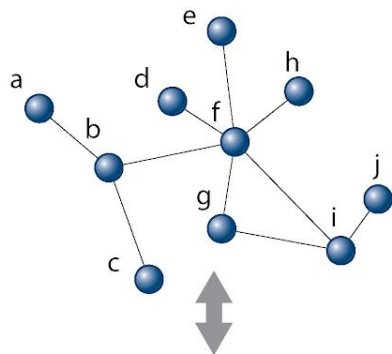


Heat map



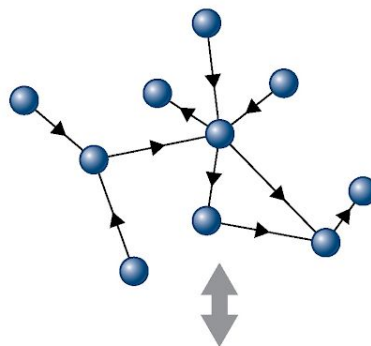
Network representations

Connectivity
e.g., protein–protein interaction



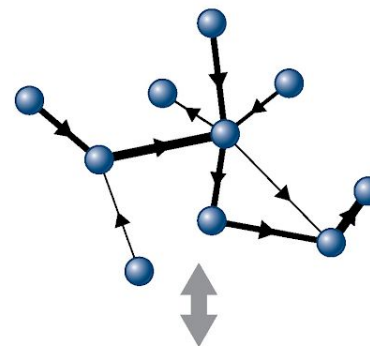
a		1								
b	1		1			1				
c		1								
d					1					
e						1				
f		1		1	1		1	1	1	
g						1			1	
h							1			
i						1	1			1
j									1	
a	b	c	d	e	f	g	h	i	j	

Connection direction
e.g., gene regulation



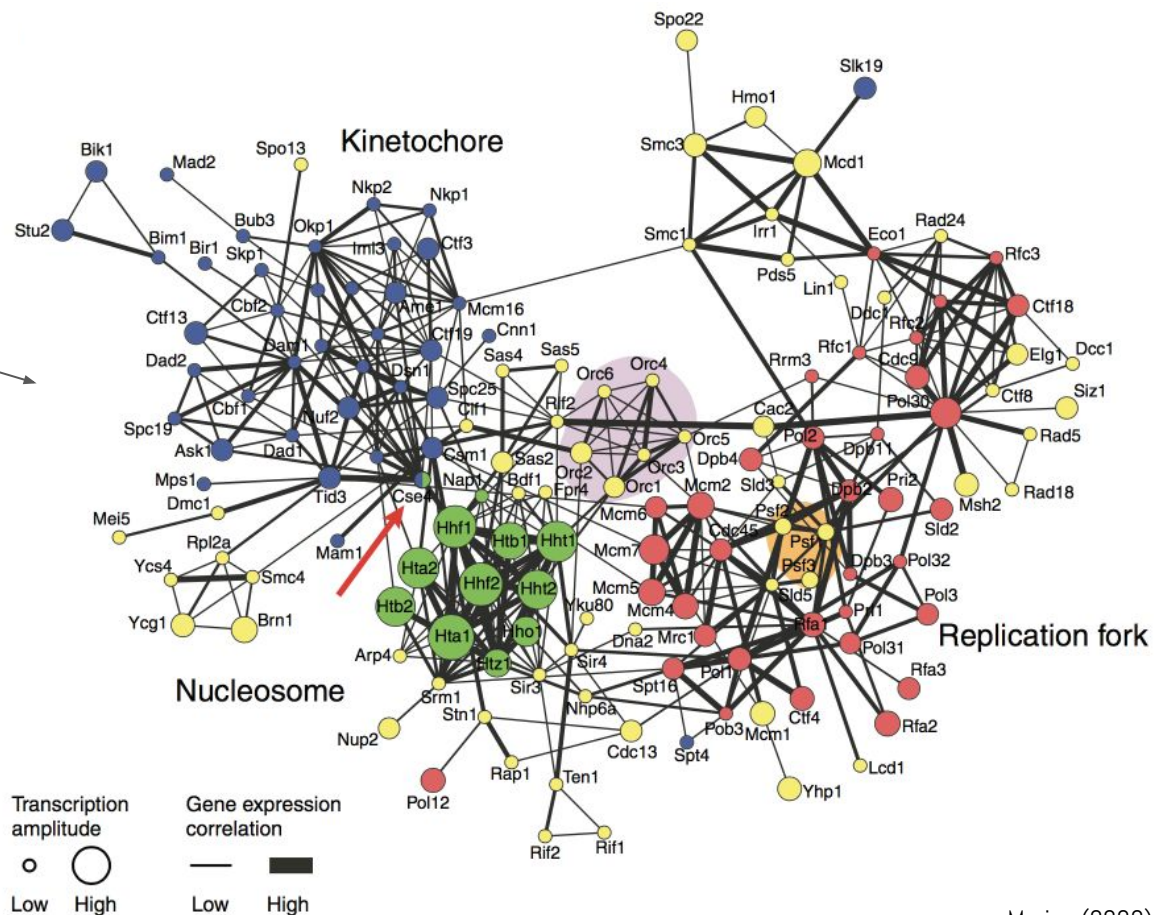
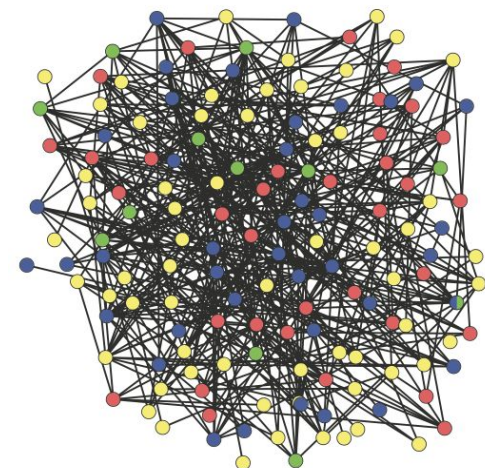
a		1								
b					1					
c		1								
d										
e					1					
f			1			1	1	1		
g								1		
h					1					
i									1	
j										1
a	b	c	d	e	f	g	h	i	j	

Connection weight
e.g., gene coexpression



a		3								
b						4				
c		1								
d										
e						3				
f			1				3	1		
g								3		
h						2				
i										4
j										
a	b	c	d	e	f	g	h	i	j	

Network layout

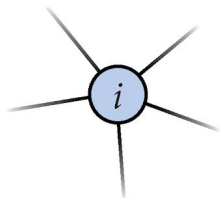


Network topology

Degree

$$d_i$$

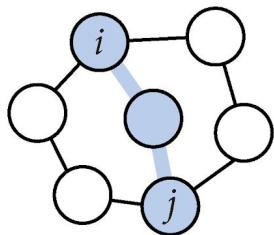
Number of nodes bound to node i



Shortest path distance

$$d_{ij} = \min\{|e_p| \mid e_p \in E_{ij}\}$$

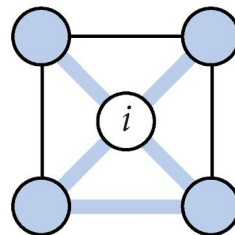
E_{ij} : all edge sets connecting nodes i and j



Clustering coefficient

$$c_i / \binom{n_i}{2}$$

c_i : edges connecting all n_i nodes bound to i

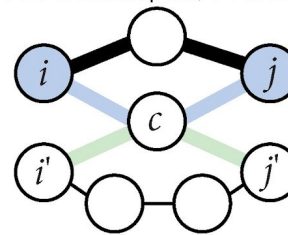


Betweenness centrality

$$b_c = \sum_i \sum_j I_{ij} / s_{ij}$$

s_{ij} : total number of shortest paths between i and j

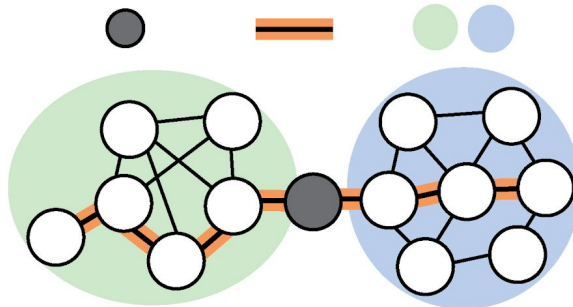
I_{ij} : 1 if c is within path; 0 otherwise



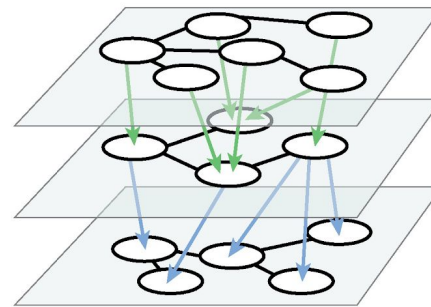
Bottleneck

Diameter

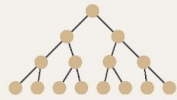
Modules



Network hierarchy



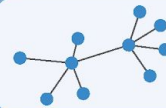
Large-scale network topology



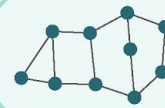
Hierarchical



Small world

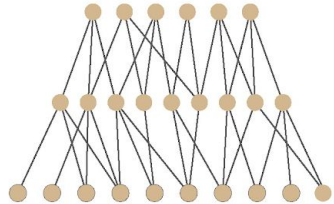


Scale-free



Geometric random

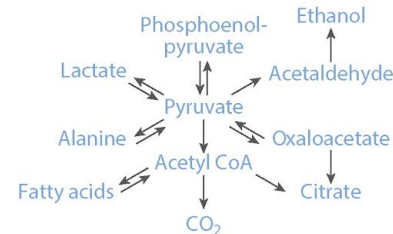
Transcription factor regulation



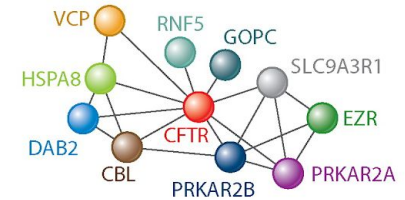
Immune regulation



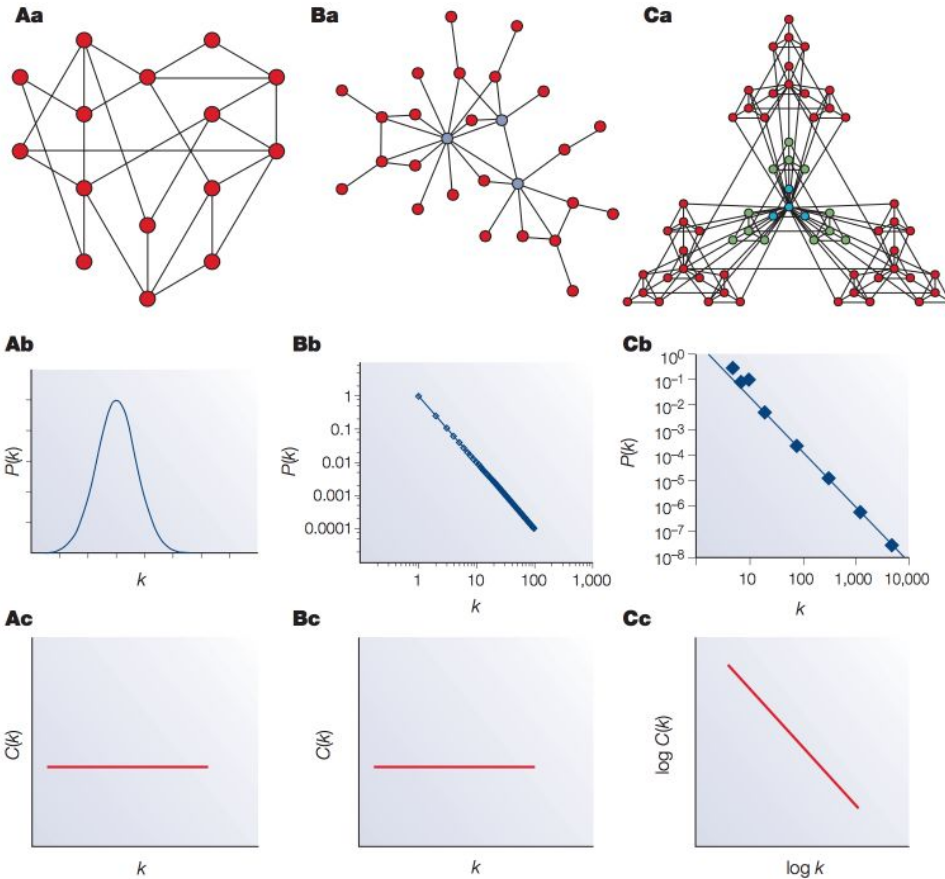
Metabolic network



Protein–protein interaction



Network topology



Network topology

Yeast TF-target network

