

EMERGENCE OF COMPLEXITY IN BIOLOGICAL NETWORKS

from selection to tinkering

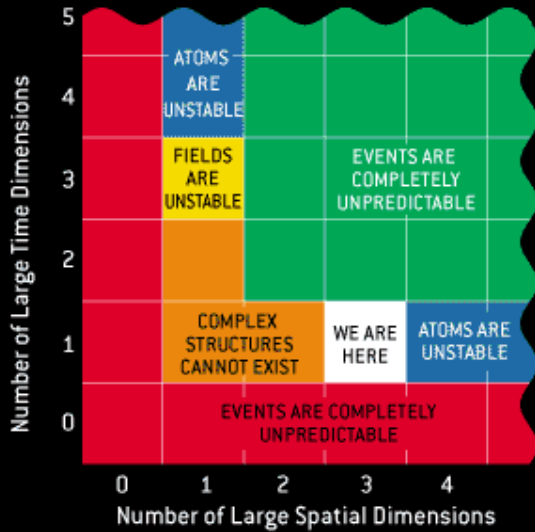


ECCS, Jerusalem, September 2008

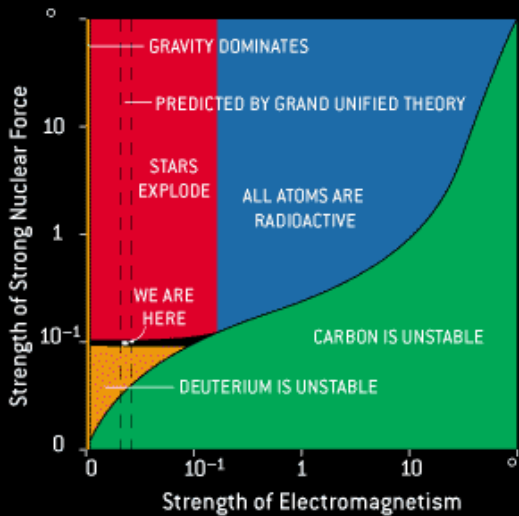
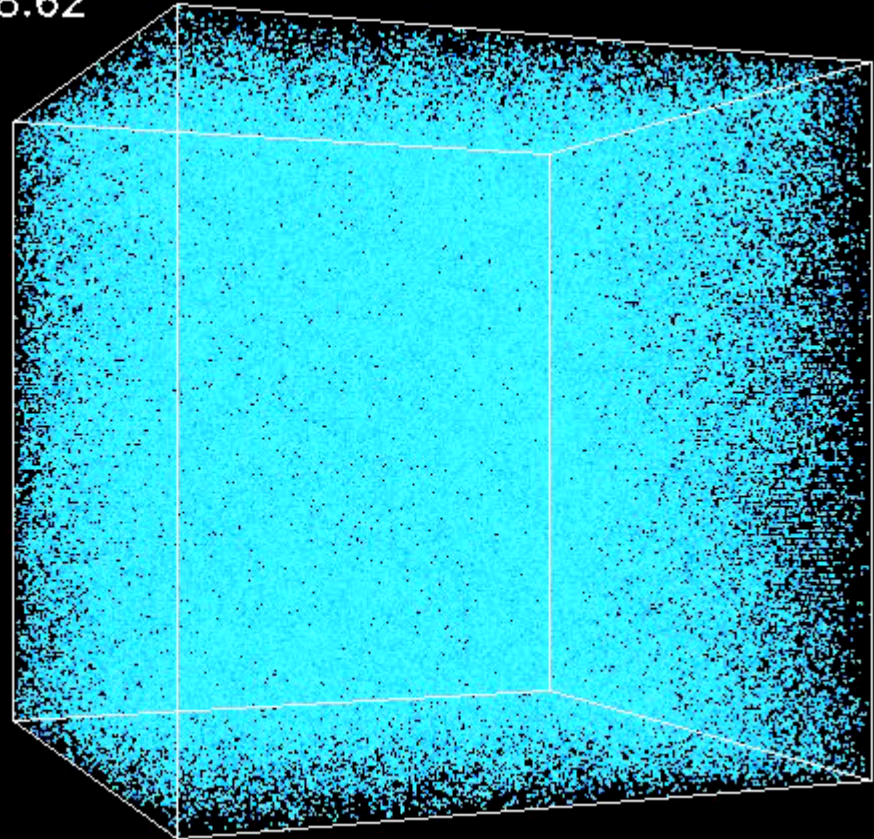
Ricard V. Solé

ICREA-Complex Systems Lab, UPF, Barcelona & Santa Fe Institute, USA

Evolution in physics?



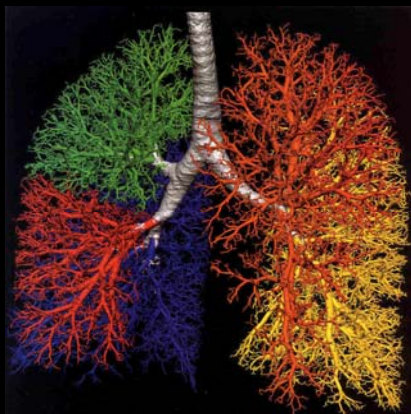
$Z=28.62$



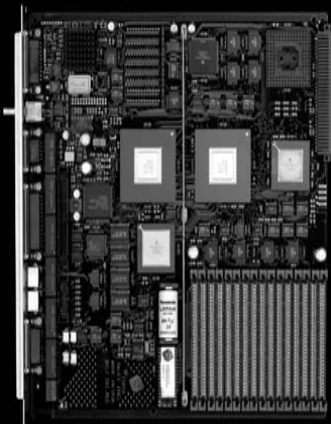
The Life of the Cosmos
 Lee Smolin, Oxford U. Press (1997)

Evolution: **how does** it take place?

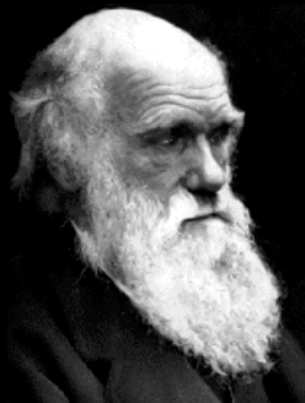
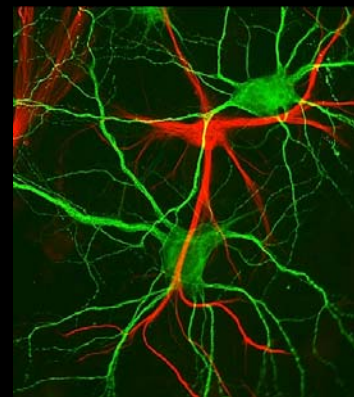
SELECTION



TINKERING



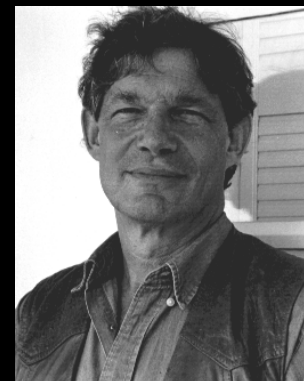
EMERGENCE



C. Darwin
 "The Origin of Species"
 (1859)



F. Jacob
 "Evolution and Tinkering"
 Science (1977)



S. Kauffman
 "Origins of Order"
 (1993)

Does evolution **occur at all?**



EVOLUTION IS A STUPID IDEA

THE BIG BANG IS A STUPID THEORY

TO BELIEVE IN LIFE ORIGINATED
FROM CHEMISTRY IS A STUPID IDEA

DON'T SUPPORT EVOLUTION IN SCHOOLS
PAYED WITH MY TAXES

Kent Hovind (creation “scientist”)

Can life be **created** in the lab?



I WORK FOR GOD, SO I DON'T HAVE TO
PAY TAXES

PAYING TAXES IS EVIL

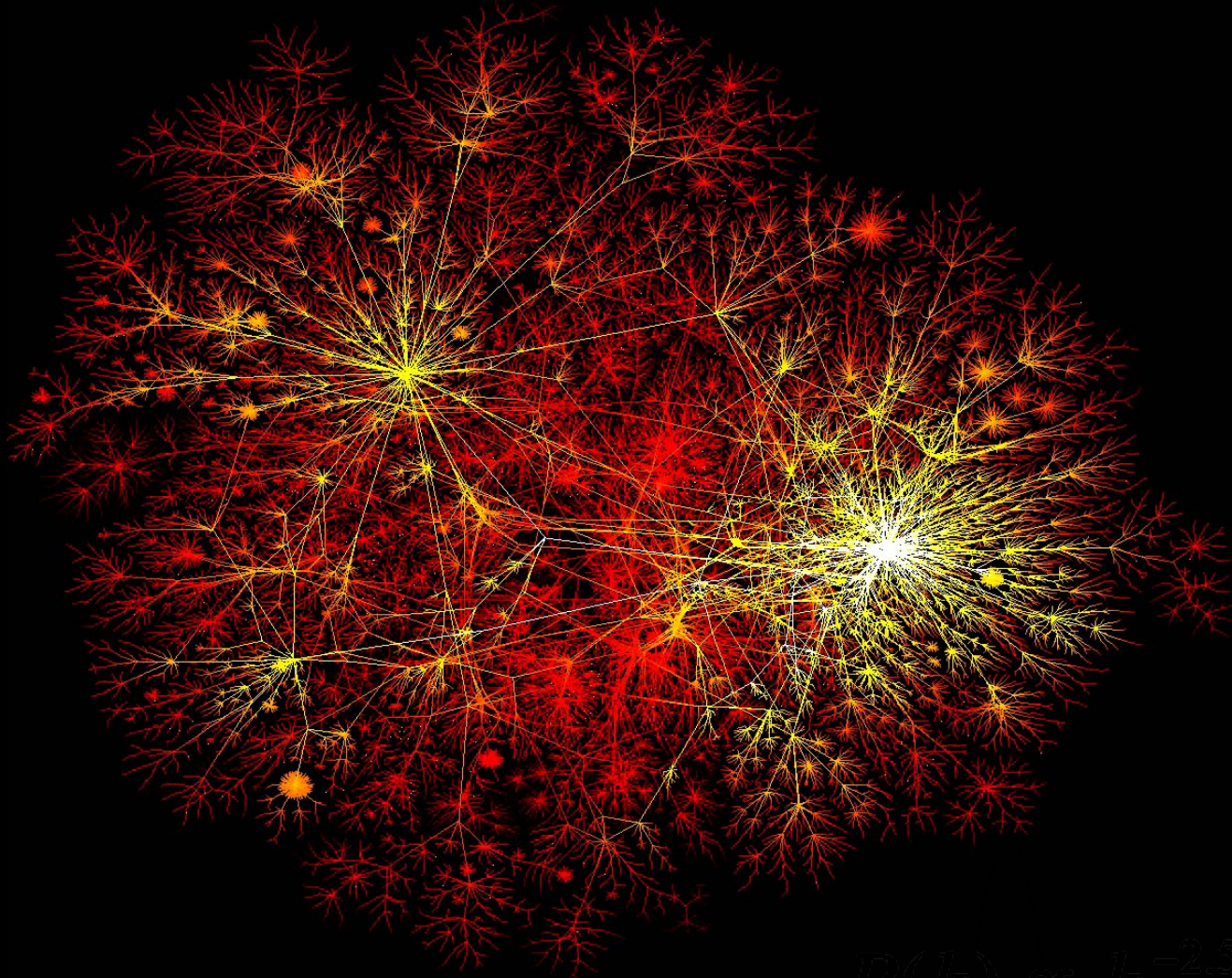
Kent Hovind (creation "scientist")

Evolution of complexity: **networks**

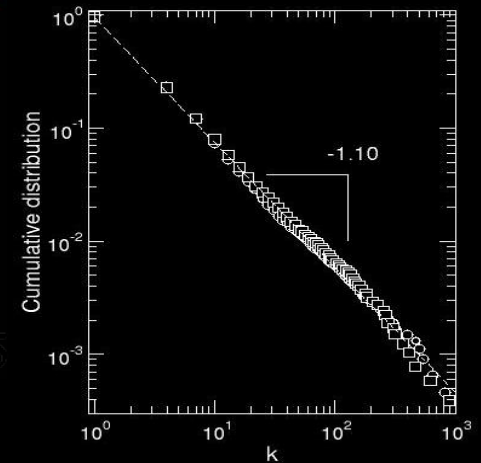


www.nrbsc.org/brainmovie/index.html

Complex networks are **scale-free**



$$P(k) \sim k^{-\gamma} \phi(k/\xi)$$



$$P(k) \propto k^{-2.5}$$

Cell Biology

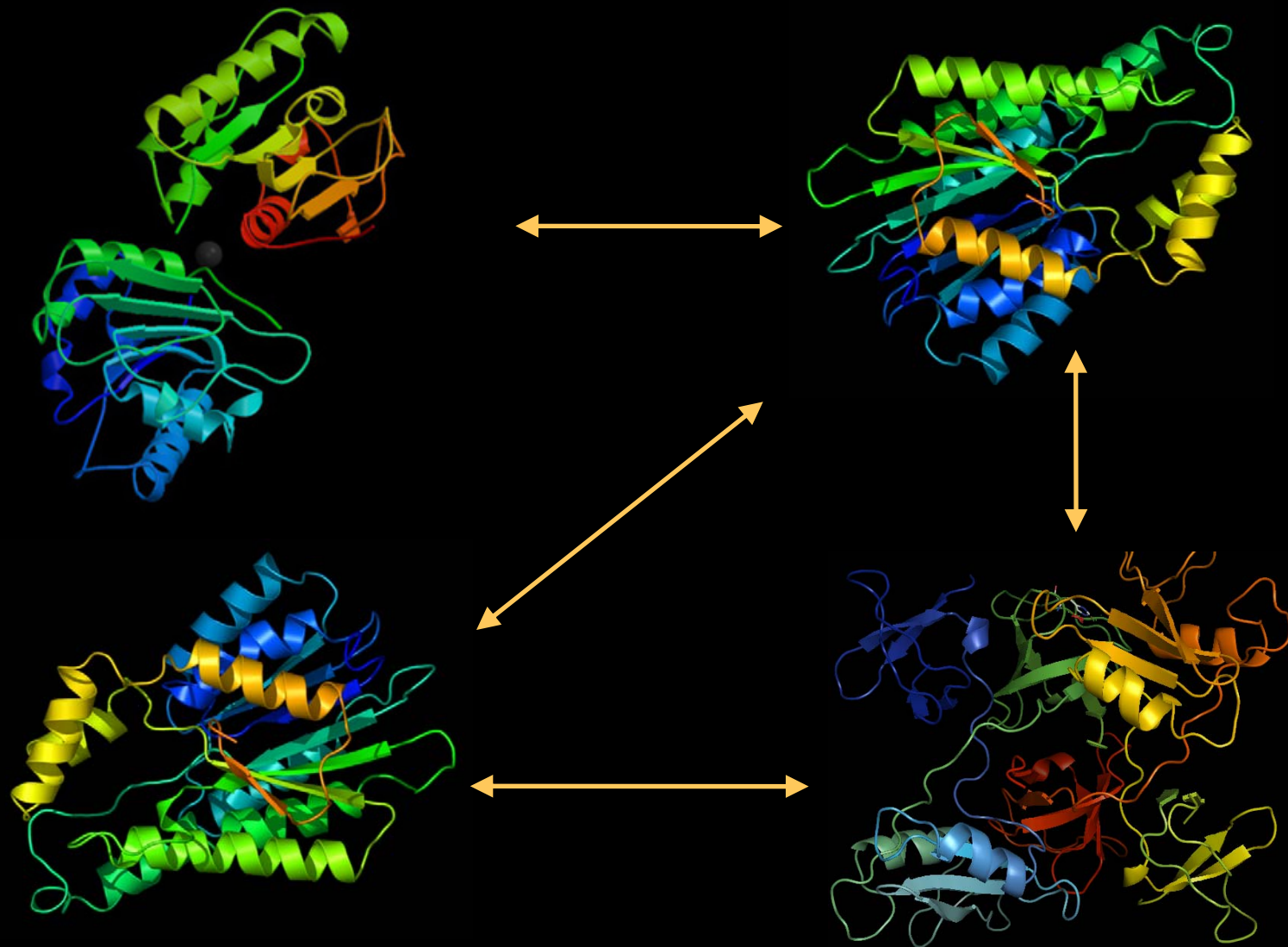
<http://multimedia.mcb.harvard.edu>

What is the architecture of cellular maps?

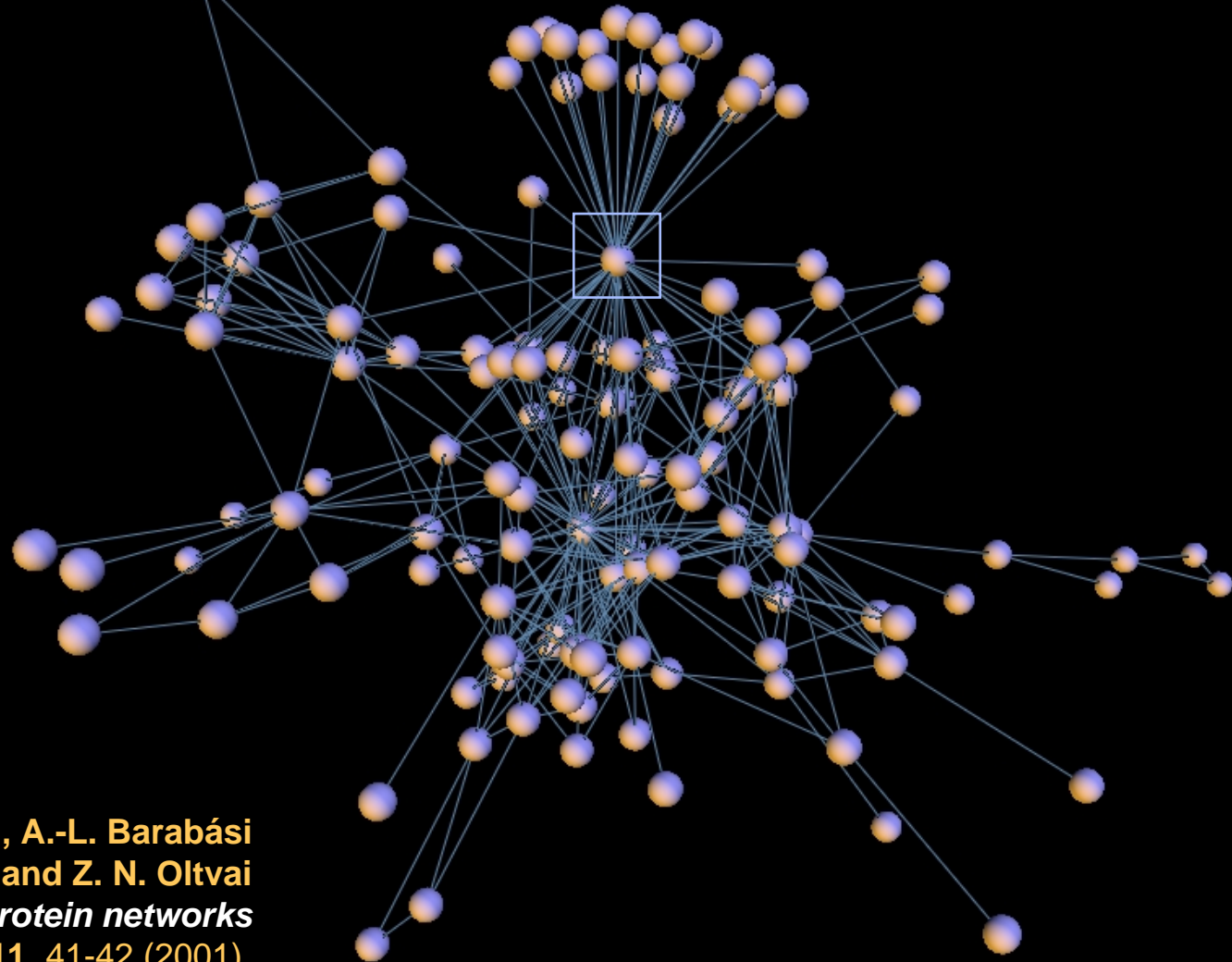
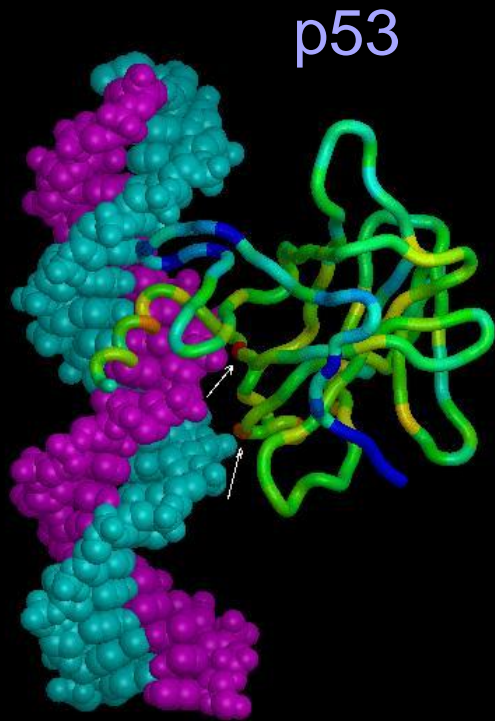
What are the consequences?

What are the origins?

The small world of **protein interactions**



Scale-free **genome** architecture



H. Jeong, S. Mason, A.-L. Barabási
and Z. N. Oltvai
Lethality and centrality in protein networks
Nature **411**, 41-42 (2001).

Proteome: small world, scale free

Topology, tinkering and evolution of the human transcription factor network

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1 ICREA-Complex Systems Laboratory, Universitat Pompeu Fabra, Barcelona, Spain

2 Department of Molecular Biology and Biochemistry, Faculty of Sciences, Universidad de Málaga, Spain

3 Santa Fe Institute, Santa Fe, New Mexico, USA

FEBS Journal 272 (2005) 6423-6434

Table 2. Description and functionality of transcriptions factor hubs. Transcription factor (TF), degree (k), betweenness centrality (b).

TF	Description	Associate disease	k	$b \times 10^3$
TBP	Basal transcription machinery initiator	Spinocerebellar ataxia [40]	27	17.3
p53	Tumor suppressor protein	Proliferative disease [68]	23	18.5
P300	Coactivator. Histone acetyltransferase	May play a role in epithelial cancer [69]	18	20.2
RXR- α	Retinoid X- α receptor	Hepatocellular carcinoma [70]	18	8
pRB	retinoblastoma suppressor protein. Tumour suppressor protein	Proliferative disease Bladder cancer. Osteosarcoma [71]	15	27.1
RelA	NF- κ B pathway	Hepatocyte apoptosis and foetal death [72]	14	6.6
c-jun	AP-1 complex (activator). Proto-oncogen	Proliferative disease [73]	14	4.1
c-myc	Activator. Proto-oncogen	Proliferative disease [74]	13	10.5
c-fos	AP-1 complex (activator). Proto-oncogen	Proliferative disease [75]	12	2

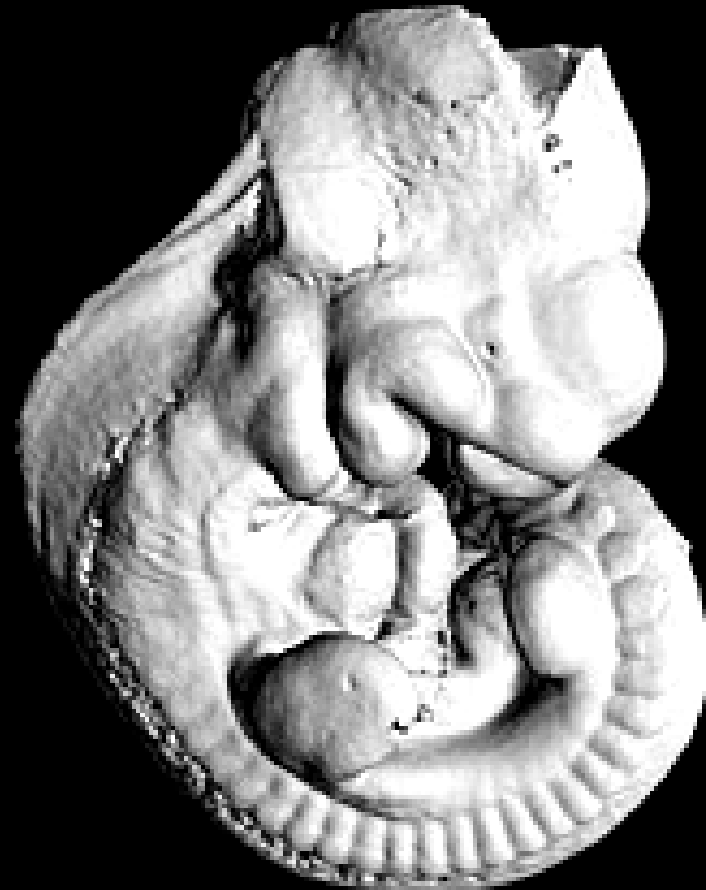
Evolution of genome : how to model?



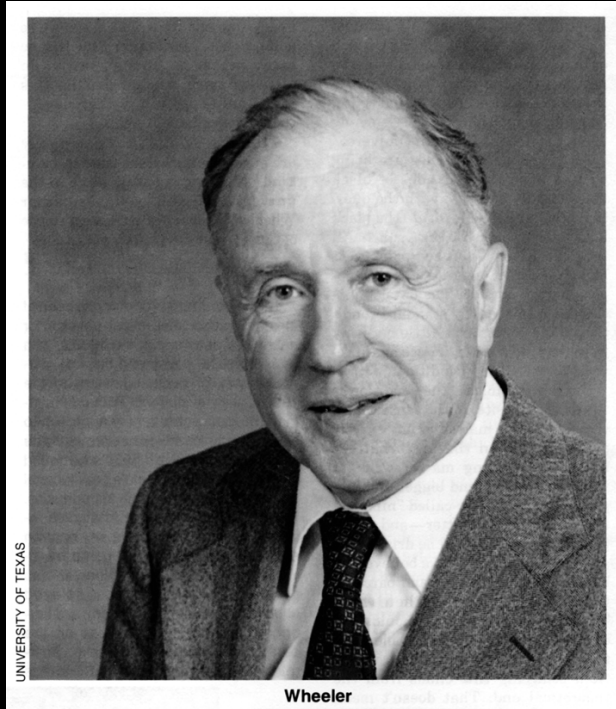
François Jacob (1976)

“Natural selection does not work as an engineer but as a tinkerer, limited by the constraints present at all levels of biological organization”

Science 196: 1161-1166



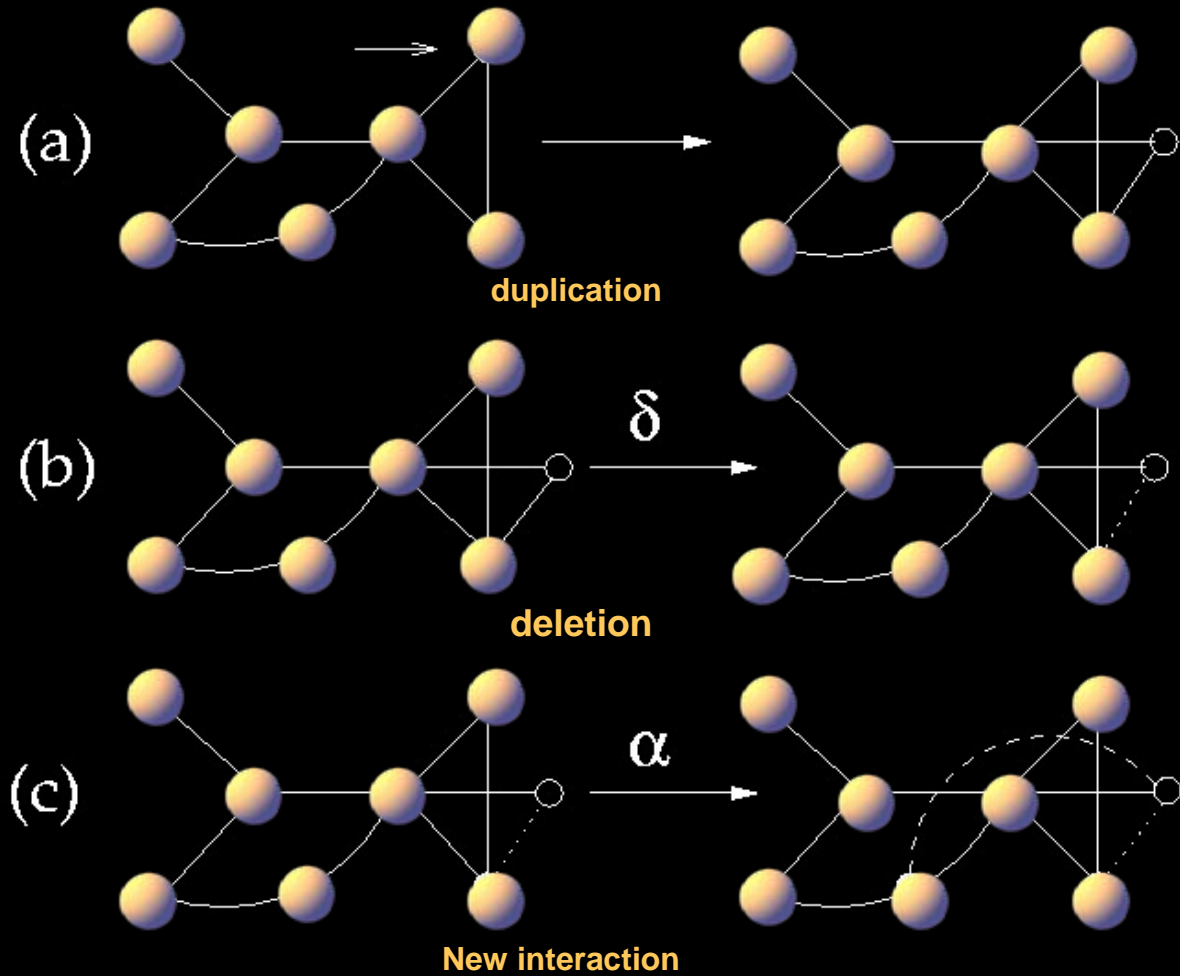
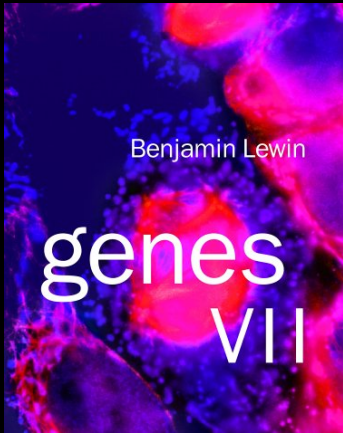
Complexity “for free”?



“If you haven’t found something strange during the day,
it hasn’t been much of a day”

John Wheeler

Origins of scale-free regulatory maps



Origins of scale-free regulatory maps

$$\begin{aligned}\frac{dn_k(t)}{dt} = & \frac{n_k}{N} + \frac{\delta}{N} [(k+1)n_{k+1} - kn_k] \\ & + \frac{1-\delta}{N} [(k-1)n_{k-1} - kn_k] \\ & + \frac{2\beta}{N} [n_{k-1} - n_k].\end{aligned}$$

$$p_k \sim (k_0 + k)^{-\gamma} e^{-k/k_c}$$

with

$$\gamma = -k_0 = 1 - \frac{2\beta}{1-\delta}, \quad k_c = \frac{1}{\ln\left(\frac{\delta}{\delta-1}\right)}$$

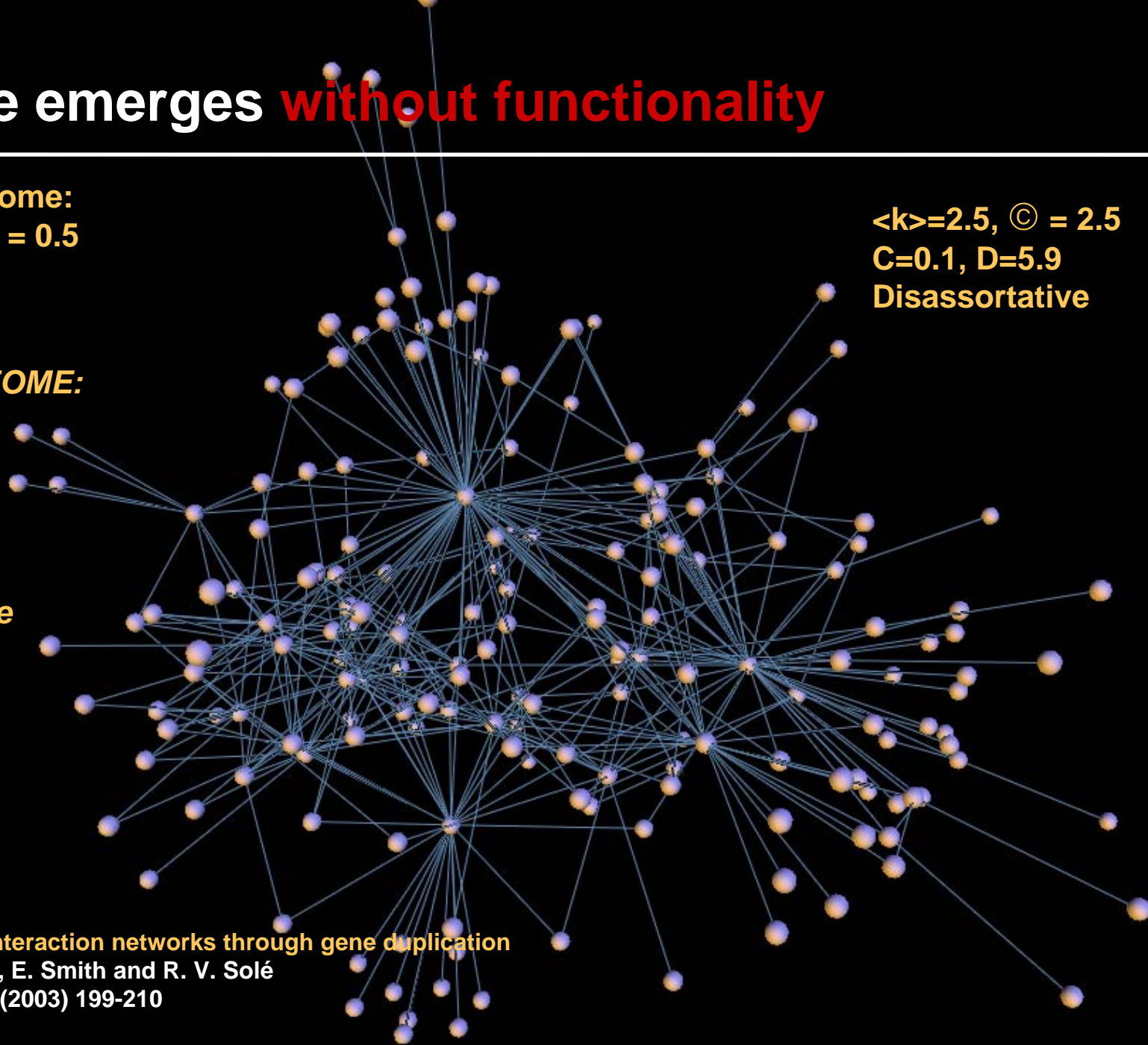
Structure emerges **without functionality**

Model proteome:
assuming $\tau^M = 0.5$

$\langle k \rangle = 2.5$, $\langle C \rangle = 2.5$
 $C = 0.1$, $D = 5.9$
Disassortative

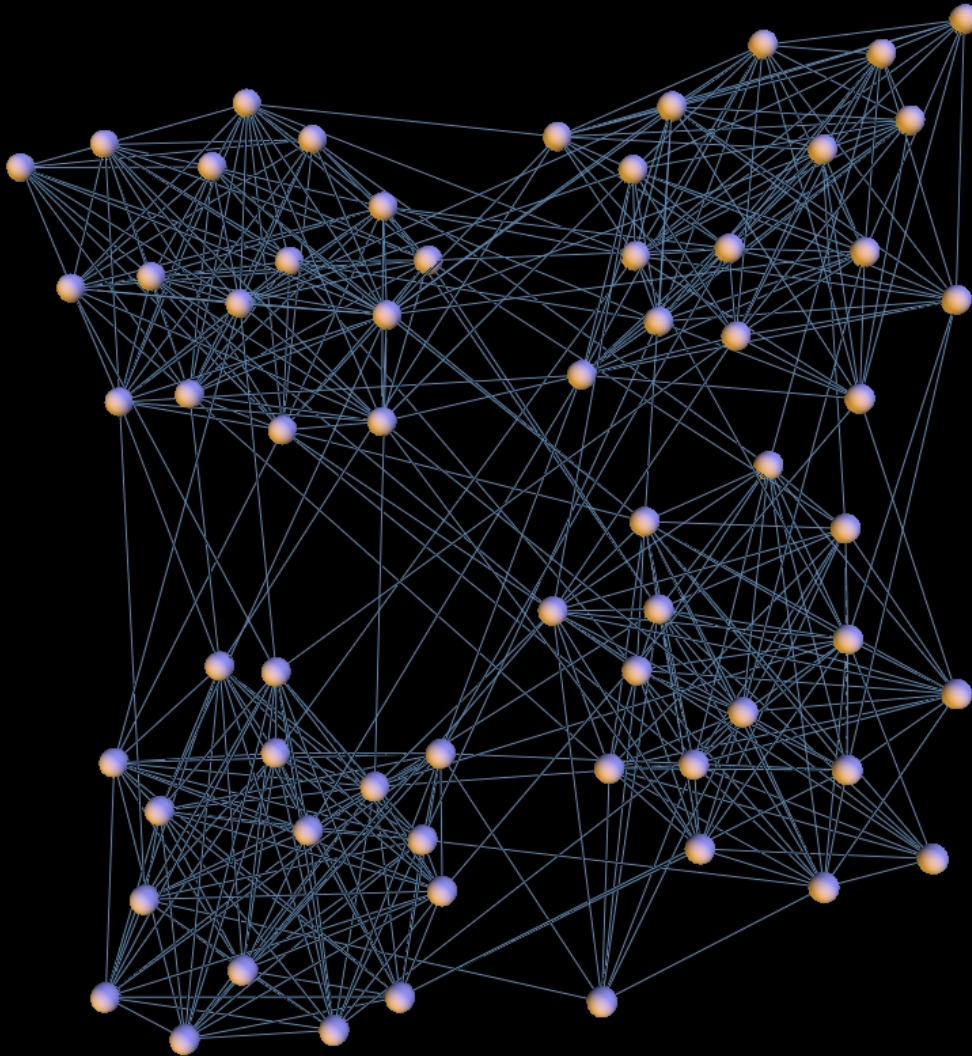
REAL PROTEOME:

$\langle k \rangle = 2.4$
 $\langle C \rangle = 2.5$
 $C = 0.7$
 $D = 6.81$
Disassortative
Modular
Hierarchical



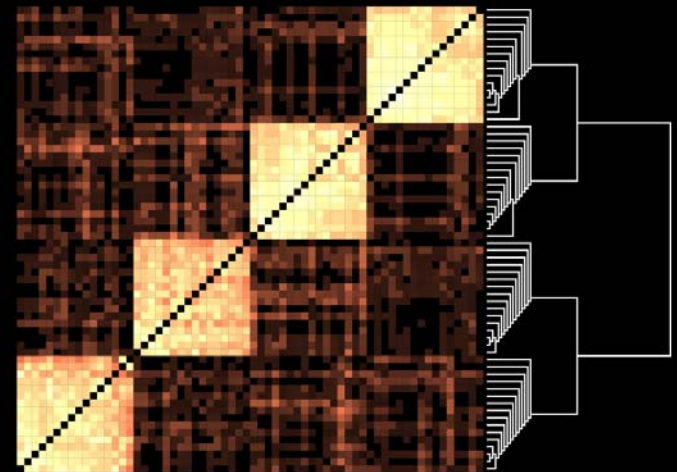
Evolving protein interaction networks through gene duplication
R. Pastor-Satorras, E. Smith and R. V. Solé
J. Theor. Biol. 222 (2003) 199-210

Modular networks



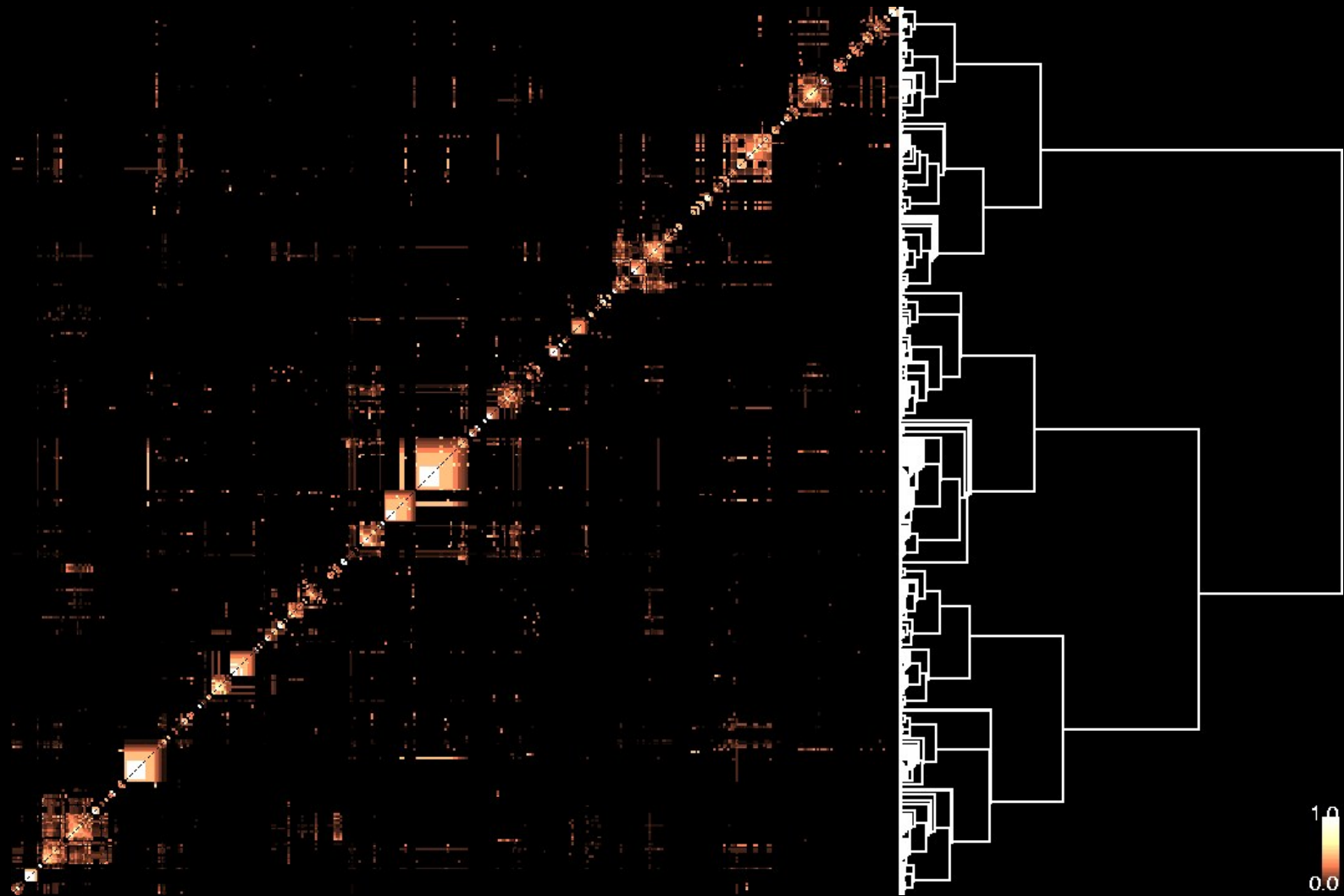
E. Ravasz, A. L. Somera, D. A. Mongru, Z. N. Oltvai and A.-L. Barabási
Hierarchical Organization of Modularity in Metabolic Networks
Science 297, 1551-1555 (2002).

$$O(i,j) = \frac{J(i,j)}{\text{Min}\{k(i),k(j)\}}$$



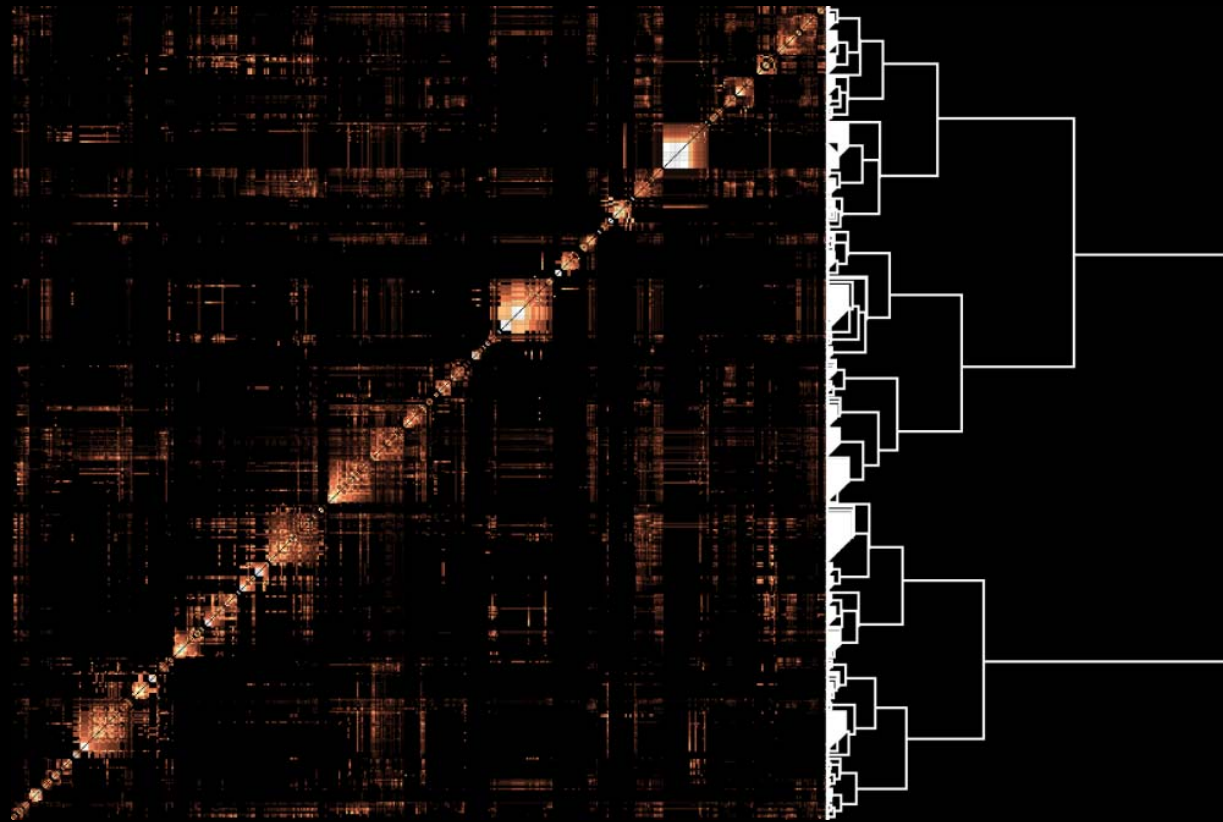
Overlap map

Cellular networks are modular



Modular architecture of the yeast interaction map

Modularity *for free*?



Spontaneous emergence of modularity in cellular networks

R. V. Solé and S. Valverde, *J. Roy. Soc. Interface* 2007

Are network motifs the spandrels of cellular complexity?

R. V. Solé and S. Valverde, *Trends Ecol. Evol.* 2006

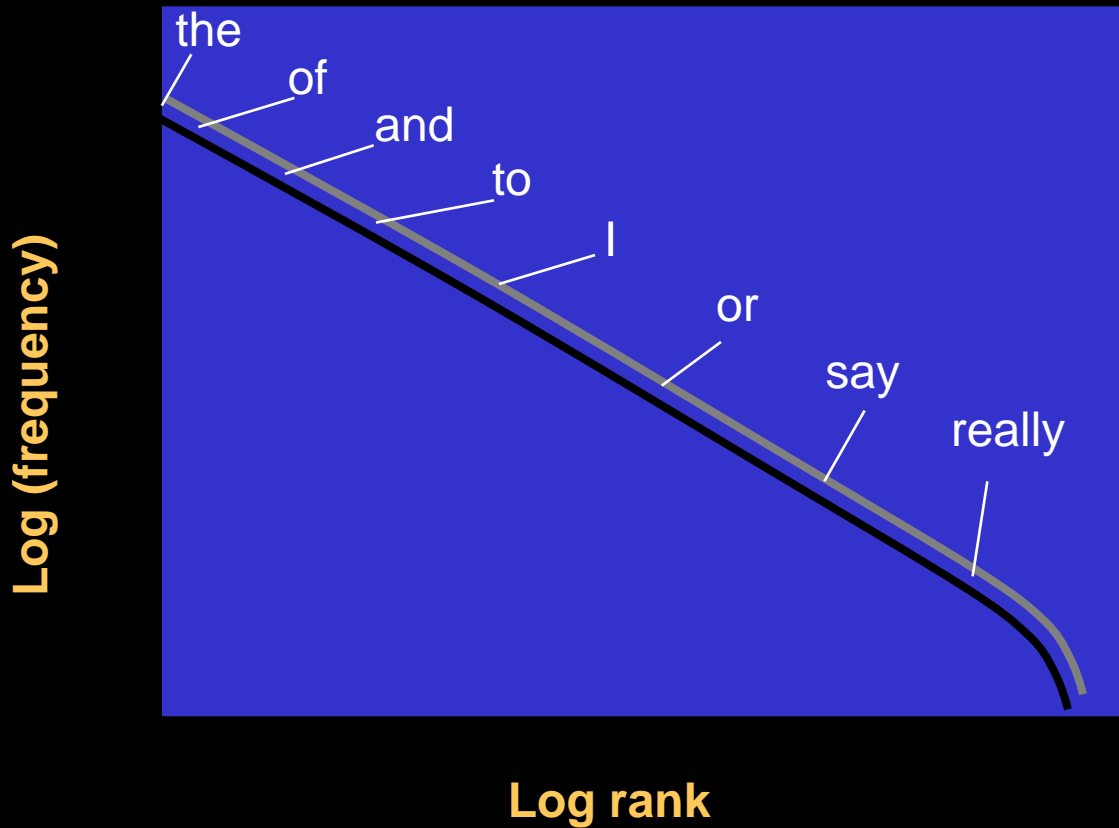
Language

What is the topology of language networks?

What are the consequences?

What are the origins?

Language **universals**



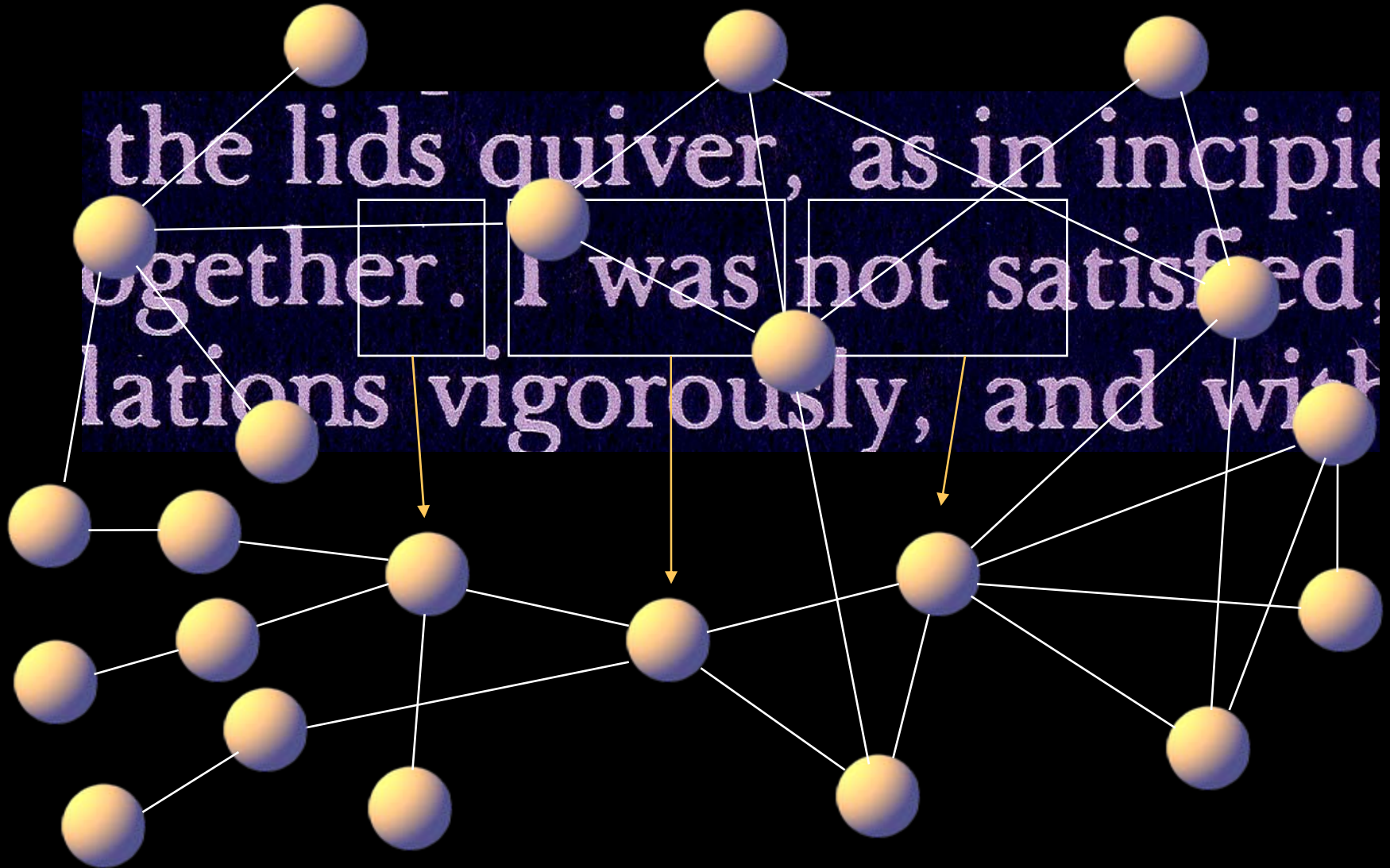
Zipf's law: all languages display it, why?
R. Ferrer and R. Solé, PNAS (2003)

Network of word **co-occurrences**

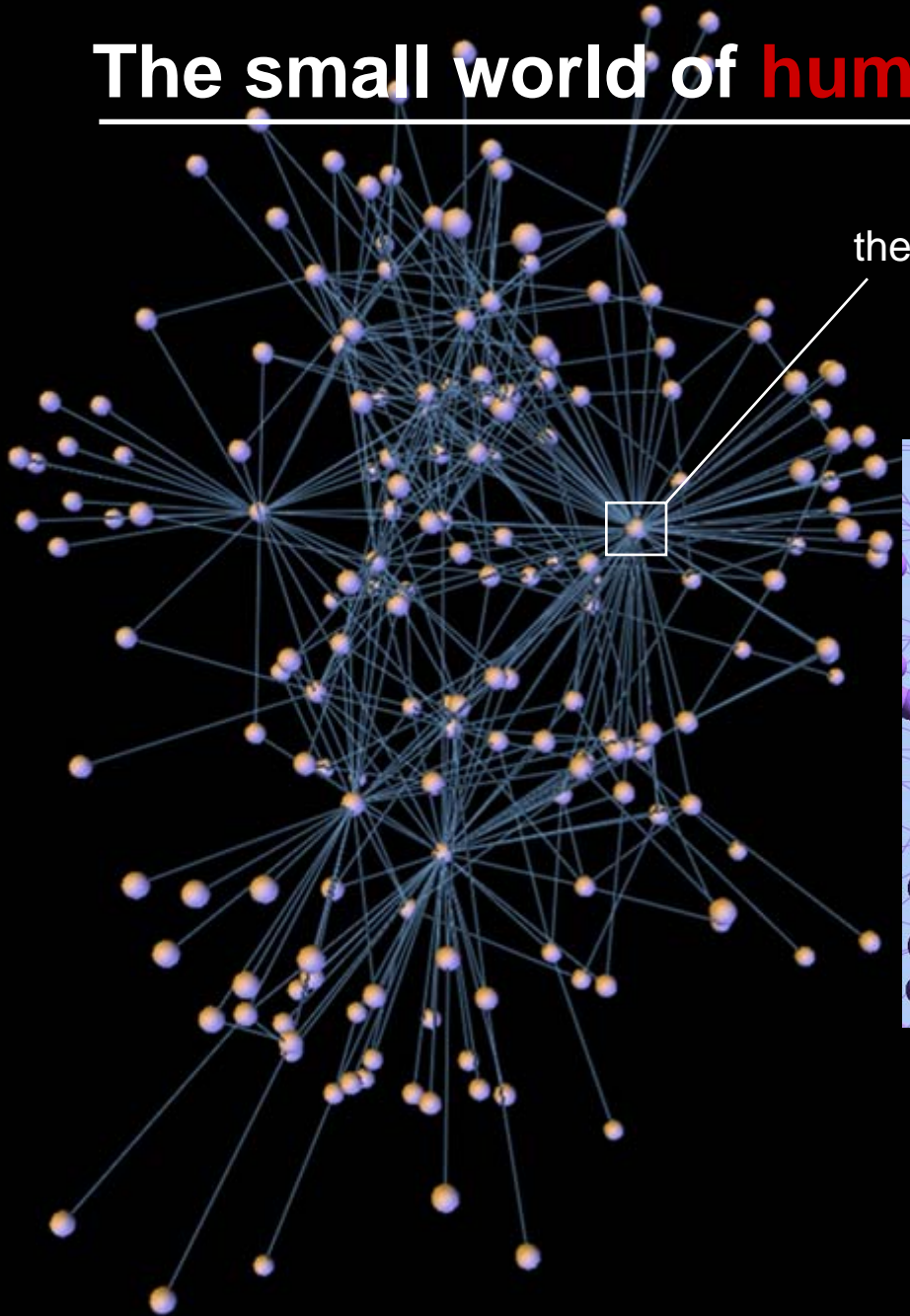
At five minutes before eleven I perceived unequivocal signs of the mesmeric influence. The glassy roll of the eye was changed for that expression of uneasy *inward* examination which is never seen except in cases of sleep-waking, and which it is quite impossible to mistake. With a few rapid lateral passes I made the lids quiver, as in incipient sleep, and with a few more I closed them altogether. I was not satisfied, however, with this, but continued the manipulations vigorously, and with the fullest exertion of the will, until I had completely stiffened the limbs of the slumberer, after placing them in a seemingly easy position. The legs were at full length; the arms were nearly so, and reposed on the bed at a moderate distance from the loins. The head was very slightly elevated.

When I had accomplished this, it was fully midnight, and I requested the gentlemen present to examine M. Valdemar's condition. After a few experiments, they admitted him to be in an unusually perfect state of mesmeric trance. The curiosity of both the physicians was greatly excited.

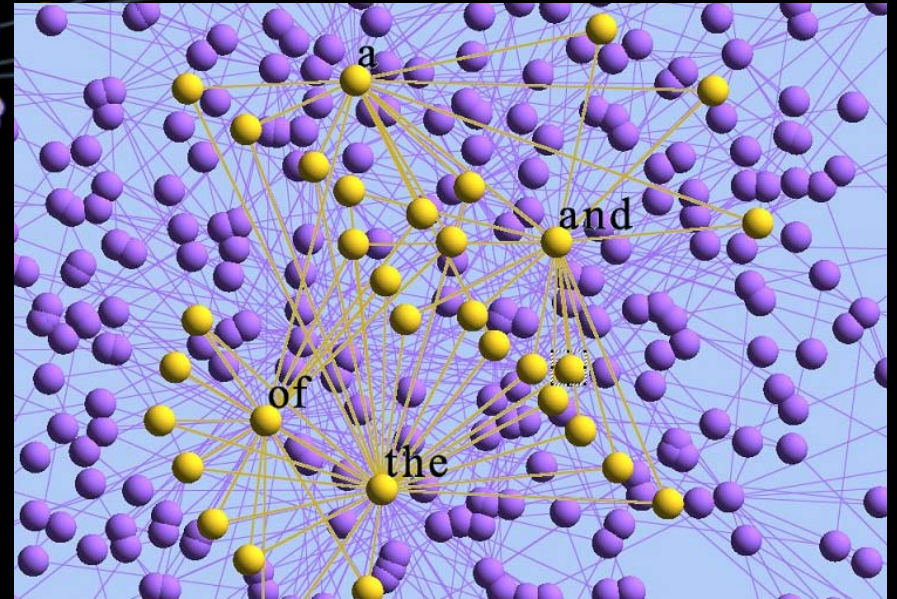
Network of word **co-occurrences**



The small world of **human language**



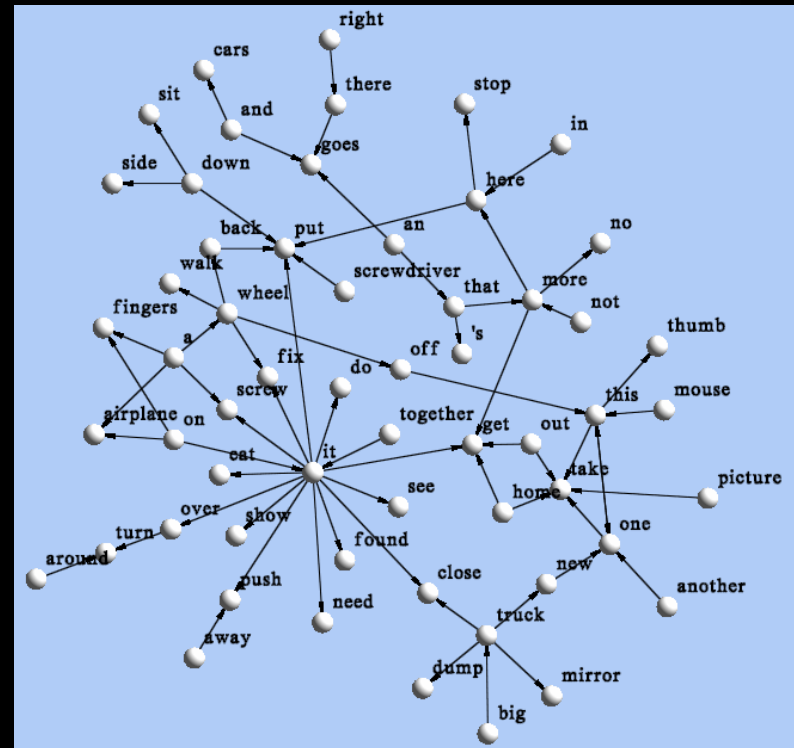
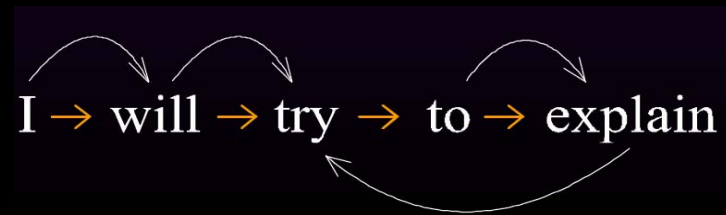
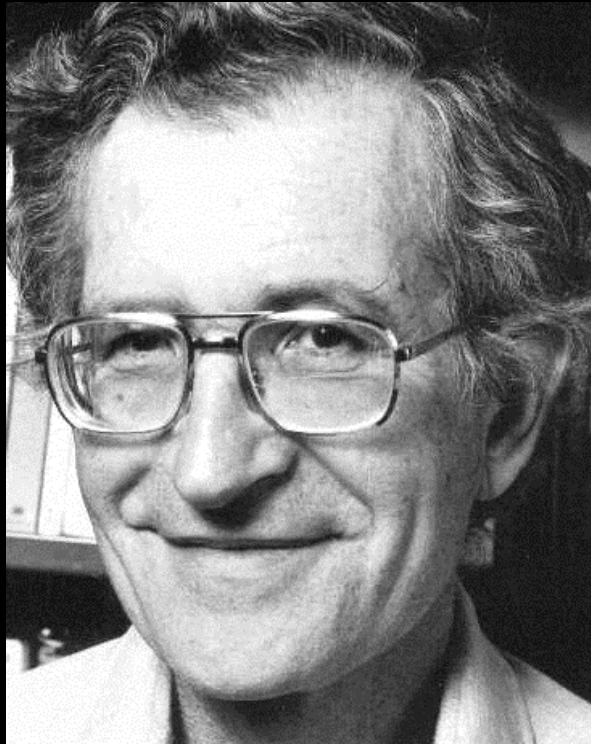
Less than **3 words** in a universe of
50000-100000 words: $L=2.5$



“The small world of human language”

R. Ferrer and R.V.Solé
Proc.Roy.Soc.London B (2002)

The **ontogeny** of human language



The SN provides a global, coarse picture of syntactic organization.
How do syntax networks change through language development?



Complex Systems Lab

ICREA-Universitat Pompeu Fabra



<http://complex.upf.es>



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