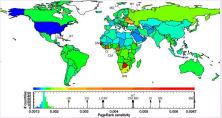
Maps of influence and interactions of infectious diseases from Wikipedia networks

Dima Shepelyansky (CNRS, Toulouse) www.quantware.ups-tlse.fr/dima

with J.Lages and G.Rollin (U Besancon)





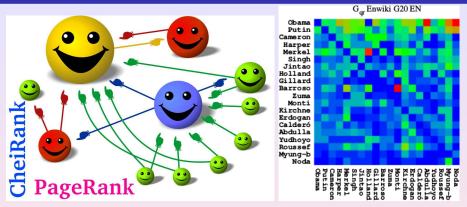
* Markov (1906) \rightarrow Brin and Page (1998)

* reduced Google matrix of directed networks (brief introduction)

* Applications: multiproduct world trade network (UN COMTRADE), Wikipedia Ranking of World Universitis (WRWU), protein-protein interactions, ...

* diseases and drugs influence from English Wikipedia 2017 (5.4 millions articles) Support: EC FET Open NADINE, APLIGOOGLE (CNRS) 2012-2017, LABEX NEXT THETRACOM (2017-2019) + Rev. Mod. Phys. 87, 1261 (2015)

(1906) Markov vs Wigner (1955)



1945: Nuclear physics \rightarrow Wigner (1955) \rightarrow Random Matrix Theory 1991: WWW, small world social networks \rightarrow Markov (1906) \rightarrow Google matrix

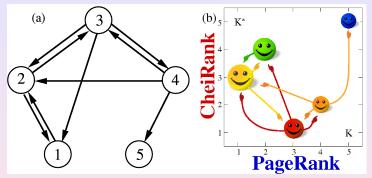
Despite the importance of large-scale search engines on the web, very little academic research has been done on them.

S.Brin and L.Page, Comp. Networks ISDN Systems 30, 107 (1998)

(Quantware group, CNRS, Toulouse)

Google matrix construction rules

Markov chains (1906) and Directed networks



For a directed network with *N* nodes the adjacency matrix **A** is defined as $A_{ij} = 1$ if there is a link from node *j* to node *i* and $A_{ij} = 0$ otherwise. The weighted adjacency matrix is

$$S_{ij} = A_{ij} / \sum_k A_{kj}$$

In addition the elements of columns with only zeros elements are replaced by 1/N.

Google matrix construction rules

Google Matrix and Computation of PageRank $P = SP \Rightarrow P$ = stationary vector of S; can be computed by iteration of S.To remove convergence problems:

• Replace columns of 0 (dangling nodes) by $\frac{1}{N}$:

	(0	1/2	1/3	0	1/5	$f \in \mathbb{N}$	(0	1/3	0	0	0 \
	1	0	1/3	1/3	$\left. \frac{1/5}{1/5} \right)$		1/2	0	1/2	0	0
S =	0	1/2	0	1/3	1/5	$S^* =$	1/2	1/3	0	1	0
	0	0	1/3	0	1/5		0	1/3	1/2	0	1
	(0	0	0	1/3	1/5	$S^* =$	0	0	0	0	0/

• To remove degeneracies of $\lambda = 1$, replace **S** by **Google matrix**

 $\mathbf{G} = \alpha \mathbf{S} + (\mathbf{1} - \alpha) \frac{\mathbf{E}}{N}$; $GP = \lambda P$ => Perron-Frobenius operator

- α models a random surfer with a random jump after approximately 6 clicks (usually α = 0.85); PageRank vector => P at λ = 1 (Σ_i P_j = 1).
- CheiRank vector P^* : $G^* = \alpha S^* + (1 \alpha) \frac{E}{N}$, $G^*P^* = P^*$ (S* with inverted link directions) Chepelianskii arXiv:1003.5455 (2010) ...

Reduced Google matrix

A selected network of interest with $N_r < N$ nodes called reduced network. Block structure of *G* matrix:

$$\mathbf{G} = \left(egin{array}{cc} \mathbf{G}_{rr} & \mathbf{G}_{rs} \ \mathbf{G}_{sr} & \mathbf{G}_{ss} \end{array}
ight)$$

with *s* index for scattering network $N_s = N - N_r$. Reduced G_R matrix

$$G_{\rm R}P_{\rm r}=P_{\rm r}~,~G_{\rm R}=G_{\rm rr}+G_{\rm rs}(1-G_{\rm ss})^{-1}G_{\rm sr}=G_{\rm pr}+G_{\rm rr}+G_{\rm qr}$$

Useful expansion

$$(\mathbf{1}-G_{ss})^{-1} = \mathcal{P}_c \frac{1}{1-\lambda_c} + \mathcal{Q}_c \sum_{l=0}^{\infty} \bar{G}_{ss}^l$$

with projector $\mathcal{P}_{c} = \psi_{B}\psi_{L}^{T}$ on eigenstate of maximal eigenvalue λ_{c} of G_{ss} , the complementary projector $\mathcal{Q}_{c} = \mathbf{1} - \mathcal{P}_{c}$ and $\bar{G}_{ss} = \mathcal{Q}_{c}G_{ss}\mathcal{Q}_{c}$. K.Frahm, DS arxiv:1602.02394 (2016)

Protein-protein interactions for cancer networks

PLOS ONE

Reduced Google matrix for directed biological networks

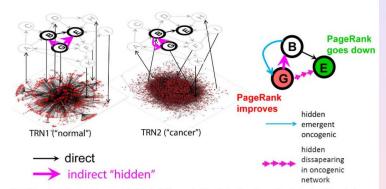


Fig 1. Using reduced Google matrix approach for inferring hidden causal relations in signaling pathways. Here the structure of the contextdependent global regulatory network is symbolically shown as consisting of two layers: the upper (nodes A-K) is the global signaling network whose structure does not depend on the context and the lower is a symbolic view of the contextual transcriptional regulatory network (TRN) whose structure can change between a "normal" and a "cancer" cell. Thick node borders denote a pathway embedded into the global signaling network. Black arrows denote direct physical interactions. Pink arrows denote inferred hidden directed regulations through the global regulatory network (bath kayers). In the final representation of the pathway (on the right), one can show those hidden regulations through the global regulatory network (bath kayers). In the structure. Also, the color of the pathway nodes can show those hidden regulations through the effect of the pageRank decreased in the cancer network while red corresponds to the opposite.

https://doi.org/10.1371/journal.pone.0190812.g001

Lages, DS, Zinovyev (PLOS ONE 2018)

PageRank of Infectious diseases - EnWiki2017

Type of infectious diseases : bacterial, viral, parasitic, fongic, prionic, multiple origin, other

Rank	Disease	
1	Tuberculosis	
2	HIV/AIDS	
3	Malaria	
4	Pneumonia	
5	Smallpox	
6 Cholera		
7	Influenza	
8	Measles	
9	Typhoid fever	
10	Syphilis	
11	Yellow fever	
12	Bubonic plague	
13	Poliomyelitis	
14	Leprosy	
15	Sepsis	
16	Meningitis	
17	Plague	
18	Typhus	
19	Diphtheria	
20	Severe acute respiratory syndrome	

230 diseases, 195 countries (5.4/122 million articles/links)

(Quantware group, CNRS, Toulouse)

Network structure, PageRank-CheiRank plane

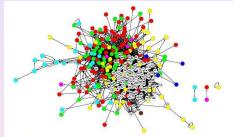
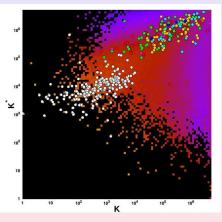
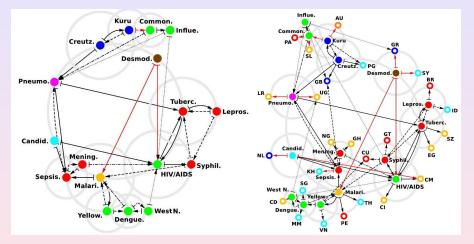


FIGURE 1. Subnetwork of the 425 articles devoted to countries and infectious diseases in 2017 English Wikipedia. The bulk of Wikipedia articles is not shown. Articles devoted to countries are presented by empty nodes with country codes (see Tab. 3). Articles devoted to infectious diseases are presented by colored nodes with the following color code: bacterial diseases, viral diseases, parasitic diseases, fonglic diseases, prionic diseases, diseases with multiple origins, and other kind of diseases.



Network of close "friends"

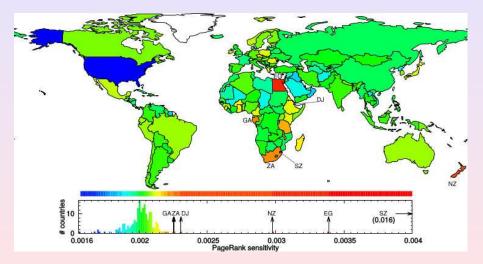


Left: top category diseases and their 2 best "friend" (full circles); Right: same but also 2 friend countries (open circles)

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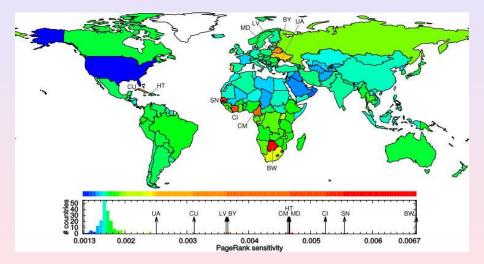
PageRank sensitivity of countries to tuberculosis

$S = d \ln(P_c)/d\delta_{tub}$ (weight variation disease-country)



PageRank sensitivity of countries to HIV/AIDS

$S = d \ln(P_c)/d\delta_{hiv}$ (weight variation disease-country)



Cancers, drugs and countries

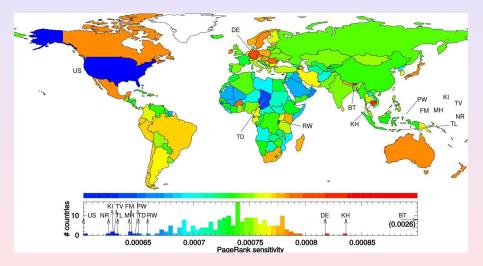
37 cancers, 203 drugs, 195 countries; PageRank order

Cancers and Drugs

Rank	Cancer	Drug				
1	Lung cancer	Methotrexate				
2	Breast cancer	Thalidomide				
3	Leukemia	Paclitaxel				
4	Prostate cancer	Prednisone				
5	Colorectal cancer	Cisplatin				
6	Brain tumor	Dexamethasone				
7	Pancreatic cancer	Doxorubicin				
8	Melanoma	Propranolol				
9	Stomach cancer	Interleukin 2				
10	Ovarian cancer	Cyclophosphamide				

PageRank sensitivity of countries to lung cancer

$S = d \ln(P_c)/d\delta_{lung}$ (weight variation disease-country)



Further applications of Markov chains and Google matrix ? → Artificial intelligence for bionetworks



Google matrix: fundamentals, applications and beyond, IHES workshop 15-18 Oct 2018 (see www.ihes.fr)