Lawrence Berkeley National Laboratory Joint Genome Institute

Title

Network analysis of genes and their association with diseases

Permalink

https://escholarship.org/uc/item/7mp855jw

Journal

Gene, 590(1)

ISSN

0378-1119

Authors

Kontou, Panagiota I Pavlopoulou, Athanasia Dimou, Niki L <u>et al.</u>

Publication Date

2016-09-01

DOI

10.1016/j.gene.2016.05.044

Peer reviewed



Contents lists available at ScienceDirect

Data in Brief





Data Article

Data and programs in support of network analysis of genes and their association with diseases



Panagiota I. Kontou^{a,1}, Athanasia Pavlopoulou^{a,1}, Niki L. Dimou^a, Georgios A. Pavlopoulos^b, Pantelis G. Bagos^{a,*}

ARTICLE INFO

Article history: Received 2 June 2016 Received in revised form 6 July 2016 Accepted 13 July 2016 Available online 19 July 2016

Keywords: Gene-disease associations Gene-gene networks Disease-disease networks

ABSTRACT

The network-based approaches that were employed in order to depict the relationships between human genetic diseases and their associated genes are described. Towards this direction, monopartite disease-disease and gene-gene networks were constructed from bipartite gene-disease association networks. The latter were created by collecting and integrating data from three diverse resources, each one with different content, covering from rare monogenic disorders to common complex diseases. Moreover, topological and clustering graph analyses were performed. The methodology and the programs presented in this article are related to the research article entitled "Network analysis of genes and their association with diseases" [1].

© 2016 Published by Elsevier Inc. This is an open access article under the CC BY license

(http://creativecommons.org/licenses/by/4.0/).

^a Department of Computer Science and Biomedical Informatics, University of Thessaly, Papasiopoulou 2-4, Lamia 35100, Greece

^b Lawrence Berkeley Lab, Joint Genome Institute, United States Department of Energy, 2800 Mitchell Drive, Walnut Creek, CA 94598, USA

DOI of original article: http://dx.doi.org/10.1016/j.gene.2016.05.044

^{*} Cooresponding author. Fax: +30 223 106 6915. E-mail address: pbagos@compgen.org (P.G. Bagos).

¹ These authors contributed equally to this work.

Specifications Table

Subject area	Systems biology
More specific sub- ject area	Gene-disease networks
Type of data	Figure, text files, Cytoscape Network file
How data were	Data were acquired from the publicly available databases: OMIM, GAD,
acquired	GWAS, UniProtKB, ICD, HGNC
Data format	Processed, analyzed
Experimental	Gene-disease association data were analyzed using Perl and R scripts and
factors	Cytoscape.
Experimental	Gene-gene and disease-disease networks were constructed.
features	
Data source	Department of Computer Science and Biomedical Informatics, University of
location	Thessaly, Lamia, Greece
Data accessibility	Data are provided with this article.

Value of the data

- The need for integrating complementary data from different sources to biological networks is further highlighted in this study.
- Important, previously unknown, associations between genes and diseases were revealed.
- Based on the constructed disease-disease networks, diseases with apparently distinct phenotypic
 manifestations were found to share a common genetic background. This finding could be utilized
 in network pharmacology.

1. Data

The overall procedure of the data analysis is shown illustratively in Fig. 1. The Perl (Supplementary Files 1-5) and R (Supplementary File 6) programs used for data analysis are indicated. A complete description of the data and methodology is presented in [1].

2. Experimental design, materials and methods

2.1. Data collection

Disease-gene association data were collected and integrated from three diverse publicly available, comprehensive resources (NCBI's OMIM [2], NIH's GAD [3] and NHRI GWAS Catalog [4]). As a given disease can be associated with more than one gene, a script was written in Perl to separate the multiple entries (Supplementary File 1; separate.pl).

2.2. Disease and gene nomenclature

In order to maintain a consistent nomenclature and classification for diseases in our analysis, the naming conventions described in the International Classification of Diseases (ICD) were used. The disease terms from the three databases were converted to ICD terms with the use of a Perl script (Supplementary File 2; ICD.pl). Moreover, in order to maintain a uniform nomenclature across all datasets, all genes from our three databases along with the ones from UniProtKB [5] were converted to the official HGNC (HUGO Gene Nomenclature Committee) [6] gene symbols using a Perl script (Supplementary File 3; Hugo.pl).

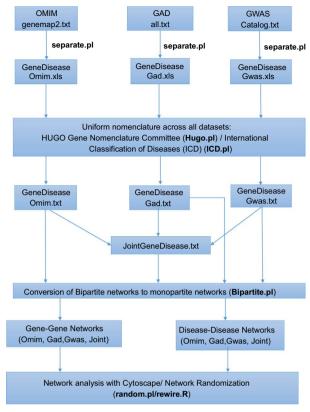


Fig.1. Flow Diagram of the data analysis.

2.3. Network processing and analysis

The bipartite networks of gene-disease associations were converted to monopartite networks of gene-gene and disease-disease interactions, by using a Perl script (Supplementary File 4; Bipartite.pl). This functionality is not available in other network analysis packages and we incorporated it in a publicly available web-server, PowerClust, which is available at: http://www.compgen.org/tools/powerclust. PowerClust, is an easy-to-use web application for clustering analysis, network processing and visualization. Moreover, randomization procedures were performed in order to determine whether the highly connected nodes in the original networks have a degree that cannot occur simply by chance given the other properties of the networks (Supplementary File 5; Random.pl). Finally, the robustness of the topological features of the projected gene-gene and disease-disease networks was assessed by employing a bipartite-specific rewiring algorithm [7] to test whether the degree distributions of the projected monopartite networks are kept stable in the randomized gene-gene/disease-disease networks compared to the initial ones (Supplementary File 6; Rewire.R). The JOINT gene-disease network (generated by combing data from the individual databases) is provided as a cytoscape network file.

Acknowledgments

The present work was funded by the SYNERGASIA 2009 PROGRAMME. This Programme is co-funded by the European Regional Development Fund and National resources (Project Code 09SYN-13-999),

General Secretariat for Research and Technology of the Greek Ministry of Education and Religious Affairs. Culture and Sports.

Transparency document. Supporting information

Transparency data associated with this article can be found in the online version at http://dx.doi. org/10.1016/j.dib.2016.07.022.

Appendix A. Supporting information

Supplementary data associated with this article can be found in the online version at http://dx.doi.org/10.1016/j.dib.2016.07.022.

References

- [1] P.I. Kontou, A. Pavlopoulou, N.L. Dimou, G.A. Pavlopoulos, P.G. Bagos, Network analysis of genes and their association with diseases, Gene 590 (2016) 68–78.
- [2] J.S. Amberger, C.A. Bocchini, F. Schiettecatte, A.F. Scott, A. Hamosh, OMIM.org: online Mendelian Inheritance in Man (OMIM (R)), an online catalog of human genes and genetic disorders. Nucleic Acids Res. 43 (2015) D789–D798.
- [3] H.J. Cordell, D.G. Clayton, Genetic association studies, Lancet 366 (2005) 1121-1131.
- [4] D. Welter, J. MacArthur, J. Morales, T. Burdett, P. Hall, H. Junkins, et al., The NHGRI GWAS Catalog, a curated resource of SNP-trait associations, Nucleic Acids Res. 42 (2014) D1001–D1006.
- [5] S. Poux, M. Magrane, C.N. Arighi, A. Bridge, C. O'Donovan, K. Laiho, et al., Expert curation in UniProtKB: a case study on dealing with conflicting and erroneous data, Database (Oxf.) 2014 (2014).
- [6] K.A. Gray, B. Yates, R.L. Seal, M.W. Wright, E.A. Bruford, Genenames.org: the HGNC resources in 2015, Nucleic Acids Res. 43 (2015) D1079–D1085.
- [7] A. Gobbi, F. Iorio, K.J. Dawson, D.C. Wedge, D. Tamborero, L.B. Alexandrov, et al., Fast randomization of large genomic datasets while preserving alteration counts, Bioinformatics 30 (2014) i617–23.