## CINDEX: a software for protein ranking through network modeling based on graph theory

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## **Abstract**

Metabolic networks have increased in complexity throughout the evolution of species becoming strongly connected metabolic blocks, which seems to have given stability to the flux of information in these networks and, possibly, turning organisms into more relaxed ones to adapt. We present the first version of CINDEX, a software for protein ranking through the modeling of protein-protein interaction (PPI) networks based on graph theory; it can be downloaded at https://github.com/hugorody/cindex. The tool accepts as input an organism's subset of proteins provided by the user and uses PPI information from the KEGG Pathway database to model a specific metabolic network throughout a directed graph. The proteins are set as the nodes of the graph, whereas the connections among the nodes are given by arcs (directed edges) that are created when the product of a protein is a substrate to another. CINDEX then calculates the centrality degree of nodes using three different metrics - Degree Centrality, Closeness Centrality, and Betweenness Centrality -, which provides to the user different biological perspectives for protein (node) ranking. Additionally, our software searches for lethal bottleneck proteins – proteins represented by nodes that disconnect the network when removed, thus essential to keep the flux of information in a network and whose inactivation could be lethal to the organism. Finally, the clustering coefficient is calculated to indicate the presence of protein clusters within the networks; a subset of interacting proteins likely to control many cellular processes.

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