Title (en)

MÉTHODS AND SYSTEMS FOR THE IDENTIFICATION OF COMPONENTS OF MAMMALIAN BIOCHEMICAL NETWORKS AS TARGETS FOR THERAPEUTIC AGENTS

Title (de)

VÉRFAHREN UND SYSTEME ZUM IDENTIFIZIEREN VON KOMPONENTEN VON BIOCHEMISCHEN NETZWERKEN FÜR SÄUGETIERE ALS TARGETS FÜR THERAPEUTISCHE MITTEL

Title (fr

PROCEDES ET SYSTEMES PERMETTANT D'IDENTIFIER DES COMPOSANTS DE RESEAUX BIOCHIMIQUES MAMMALIENS COMME ETANT DES CIBLES D'AGENTS THERAPEUTIQUES

Publication

EP 1454282 A4 20050406 (EN)

Application

EP 02773968 A 20021104

Priority

- US 0235301 W 20021104
- US 33599901 P 20011102
- US 40676402 P 20020829
- US 28637202 A 20021101

Abstract (en)

[origin: WO03040992A1] Systems and methods for modeling the interactions of the several genes, proteins and other components of a cell, employing mathematical techniques to represent the interrelationships between the cell components and the manipulation of the dynamics of the cell to determine which components of a cell may be targets for interaction with therapeutic agents. A first such method is based on a cell simulation approach in which a cellular biochemical network intrinsic to a phenotype of the cell is simulated by specifying its components and their interrelationships. The various interrelationships are represented with one or more mathematical equations which are solved to simulate a first state of the cell. The simulated network is then perturbed by deleting one or more components, changing the concentration of one or more components, or modifying one or more mathematical equations representing the interrelationships between one or more of the components. The equations representing the perturbed network are solved to simulate a second state of the cell which is compared to the first state to identify the effect of the perturbation on the state of the network, thereby identifying one or more components as targets. A second method for identifying components of a cell as targets for interaction with therapeutic agents is based upon an analytical approach, in which a stable phenotype of a cell is specified and correlated to the state of the cell and the role of that cellular state to its operation. A cellular biochemical network believed to be intrinsic to that phenotype is then specified by identifying its components and their interrelationships and representing those interrelationships in one or more mathematical equations. The network is then perturbed and the equations representing the perturbed network are solved to determine whether the perturbation is likely to cause the transition of the cell from one phenotype to another, thereby identifying one or more components as targets.

IPC 1-7

G06G 7/48; G06G 7/58; G01N 33/48; G01N 33/50

IPC 8 full level

G16B 5/30 (2019.01); G01N 33/50 (2006.01); G16B 5/10 (2019.01); G16B 5/20 (2019.01)

CPC (source: EP US)

G01N 33/5091 (2013.01 - EP); G16B 5/00 (2019.01 - EP); G16B 5/30 (2019.01 - EP US); G01N 2333/4739 (2013.01 - EP); G16B 5/20 (2019.01 - EP US)

Citation (search report)

- [PA] WO 02065119 A1 20020822 UNIV COLUMBIA [US]
- [XY] HUANG SUI: "Gene expression profiling, genetic networks, and cellular states: An integrating concept for tumorigenesis and drug discovery", JOURNAL OF MOLECULAR MEDICINE, SPRINGER VERLAG, DE, vol. 77, no. 6, June 1999 (1999-06-01), pages 469 480, XP002205218, ISSN: 0946-2716
- [XY] IDEKER T ET AL: "Integrated genomic and proteomic analyses of a systematically perturbed metabolic network", SCIENCE, AMERICAN ASSOCIATION FOR THE ADVANCEMENT OF SCIENCE,, US, vol. 292, 4 May 2001 (2001-05-04), pages 929 - 934, XP002963505, ISSN: 0036-8075
- [A] SCHUSTER STEFAN ET AL: "A general definition of metabolic pathways useful for systematic organization and analysis of complex metabolic networks", NATURE BIOTECHNOLOGY, vol. 18, no. 3, March 2000 (2000-03-01), pages 326 332, XP002316686, ISSN: 1087-0156
- [A] SCHERF U ET AL: "A gene expression database for the molecular pharmacology of cancer", NATURE GENETICS, NATURE AMERICA, NEW YORK, US, vol. 24, no. 3, March 2000 (2000-03-01), pages 236 244, XP002224798, ISSN: 1061-4036
- [A] HASTY JEFF ET AL: "Computational studies of gene regulatory networks: In numero molecular biology", NATURE REVIEWS GENETICS, vol. 2, no. 4, April 2001 (2001-04-01), pages 268 279, XP002316687, ISSN: 1471-0056
- [PX] WAGNER A: "How to reconstruct a large genetic network from n gene perturbations in fewer than n2 easy steps", BIOINFORMATICS, OXFORD UNIVERSITY PRESS, OXFORD, GB, vol. 17, no. 12, 2001, pages 1183 1197, XP002973480, ISSN: 1367-4803
- [PX] SOMOGYI ROLAND ET AL: "The dynamics of molecular networks: Applications to therapeutic discovery.", DRUG DISCOVERY TODAY, vol. 6, no. 24, 2001, pages 1267 1277, XP002316688, ISSN: 1359-6446
- [PX] SHMULEVICH ILYA ET AL: "Gene perturbation and intervention in Probabilistic Boolean Networks.", BIOINFORMATICS (OXFORD), vol. 18, no. 10, October 2002 (2002-10-01), pages 1319 1331, XP002316689, ISSN: 1367-4803
- See references of WO 03040992A1

Designated contracting state (EPC)

AT BE BG CH CY CZ DE DK EE ES FI FR GB GR IE IT LI LU MC NL PT SE SK TR

DOCDB simple family (publication)

WO 03040992 A1 20030515; CA 2501111 A1 20030515; EP 1454282 A1 20040908; EP 1454282 A4 20050406

DOCDB simple family (application)

**US 0235301 W 20021104**; CA 2501111 A 20021104; EP 02773968 A 20021104