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Aleš Prokop · Seth Michelson

# Systems Biology in Biotech & Pharma A Changing Paradigm



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# Systems Biology in Biotech & Pharma

A Changing Paradigm

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*Dedicated to my “teachers” and friends, the late Zdeněk Fencl (Prague), Arthur E. Humphrey, Elmer L. Gaden and Godfred E. Tong (USA) who influenced and shaped my professional career (AP)*

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# Abbreviations

|            |   |
|------------|---|
| 3D         | Three-dimensional   |
| 3D/4D QSAR | Three dimensional/four dimensional QSAR                         |
| 3D-QSAR    | Three dimensional QSAR  |
| ABC        | Bayesian computation  |
| ABM        | Agent-based methods   |
| ADMET      | Absorption, distribution, metabolism, excretion, and toxicity   |
| AMS        | Accelerator mass spectrometry                                   |
| ANN        | Artificial Neural Networks                                      |
| anti-CD40L | Antibody raised against CD40L region                            |
| ARACNE     | Algorithm for the reconstruction of accurate cellular networks  |
| BI         | Bioinformatics  |
| BLA        | Biologic application  |
| BOSS       | Biological objective solution search                            |
| BSR        | Biochemical system (network) reconstruction                     |
| CCA        | Canonical correlation analysis                                  |
| CD14-/-    | Type of mice  |
| CDD        | Controlled drug delivery  |
| CellML     | Mark-up language  |
| CG         | Coarse-graining or CG computing                                 |
| CNI        | Correlation network inference                                   |
| COAST      | Complex automata for modeling and simulation of complex systems |
| COMBINE    | Comparative binding energy                                      |
| CSB        | Computational systems biology                                   |
| CSDD       | Center for the study of drug development (Tufts)                |
| DD         | Drug discovery  |
| DDD        | Drug discovery and development                                  |

|              |  |
|--------------|--|
| DDS          | Drug delivery systems                                  |
| DDv          | Drug development                                       |
| DOS          | Diversity-oriented synthesis                           |
| dsRNA        | Double strand RNA                                      |
| EBI          | European bioinformatics institute                      |
| ED           | Enrichment designs                                     |
| EGFR         | Endothelial growth factor receptor                     |
| EPR          | Passive uptake   |
| ERK          | Extracellular regulated kinase                         |
| FBA          | Flux balance analysis                                  |
| FBDD         | Fragment based DD                                      |
| FCR          | Fluorochromatic reaction fluorescence                  |
| FDA          | Food and drug association                              |
| FRET         | Fluorescence resonance energy transfer                 |
| GNR          | Gene regulatory network                                |
| GO           | Gene ontology  |
| hPXR         | Humanized transgenic mice                              |
| HQSAR        | Hologram quantitative structure-activity relationships |
| HT           | High throughput  |
| HTS          | High throughput screening                              |
| iFBA         | Integrated dynamic FBA                                 |
| IFN- $\beta$ | Interferon beta  |
| IL-12        | Interleukin 12   |
| IL-15        | Interleukin 15   |
| JWS          | Journal of web semantics                               |
| KEGG         | Kyoto encyclopedia of genes and genomes                |
| KNN          | K-nearest neighbors                                    |
| LDA          | Linear discriminant analysis                           |
| MARS         | Splines  |
| MBDD         | Model based drug design                                |
| MCA          | Metabolic control analysis                             |
| MD           | Molecular dynamics                                     |
| MINDy        | Modulator inference by network dynamics                |
| miRNA        | MicroRNA   |
| MM           | Molecular mechanics                                    |
| MMR          | DNA mismatch repair                                    |
| MoA          | Mode of action   |
| MS           | Multiple sclerosis                                     |
| MVDA         | Multivariate data analysis                             |
| NB           | Naïve Bayes  |
| NDA          | New drug application                                   |
| NME          | New medical entity                                     |
| NMR          | Nuclear magnetic resonance                             |

|            |  |
|------------|--|
| NOD        | Nude mouse   |
| OBRC       | Online bioinformatics resources collection   |
| ODE        | Ordinary differential equation   |
| OiCR       | Ontario institute for cancer research  |
| OMICS      | Discipline of science and engineering for analyzing the interactions of biological information objects |
| PAT        | Process analytical technology  |
| PCA        | Principal components analysis  |
| PD         | Parkinson disease  |
| PD         | Pharmacodynamics   |
| PEGylation | Attachment of polyethylene glycol (PEG)  |
| PEM        | Protein epitope mimetic  |
| PET        | Positron emission tomography   |
| PGN        | Pharmacogenomics   |
| PhRMA      | Pharmaceutical research and manufacturers of America   |
| PI3K       | Phosphoinositide-3-kinase  |
| PK         | Pharmacokinetics   |
| PKPD       | Combined PK and PD   |
| PLS        | Partial least-squares  |
| PM         | Pharmacometrics  |
| PromoLign  | Simulation tool  |
| PTEN       | Phosphatase and tensin homolog   |
| PupaSNP    | Simulation tool  |
| QbD        | Quality by design  |
| QSAR       | Quantitative structure-activity relationship   |
| R&D        | Research and development   |
| R03        | Rule of three  |
| RA         | Entelos rheumatoid arthritis   |
| RAW        | Mouse leukaemic monocyte macrophage cell line  |
| RDD        | Re-randomization design  |
| ReguLign   | Simulation tool  |
| R-L        | Receptor-ligand  |
| RNAi       | RNA interference   |
| RNI        | Reaction network inference   |
| RNIDD      | Reaction network inference for drug discovery  |
| Ro5        | Rule of five   |
| ROT        | Rule-of-thumb  |
| RPART      | Recursive partitioning and regression trees  |
| SAR        | QSAR   |
| SB         | Systems biology  |
| SBML       | Systems biology markup language  |
| siRNA      | Small interfering RNA  |
| SG         | Systems genetics   |

|               |                                   |
|---------------|-----------------------------------|
| SNP           | Single nucleotide polymorphism    |
| SSM           | Scale separation map              |
| SVM           | Support vector machines           |
| TGF- $\beta$  | Transformation growth factor beta |
| TNF- $\alpha$ | Tumor necrosis factor alfa        |
| uHTS          | Ultra high throughput screening   |
| WW2           | Second world war                  |

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