

The Pharmacophore Concept and Its Applications in Computer-Aided Drug Design: Pharmaceutical design concepts based on drug-like properties.

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Abstract (Limit 600words)

Many computer-aided drug design workflows now include pharmacophore-based methodologies, which have been effectively and widely used for tasks including virtual screening, de novo design, and lead optimization. Pharmacophore models are abstract descriptions of fundamental non-bonded interactions that often occur between small-molecule ligands and macromolecular targets. They can be obtained both from receptors and from ligands. Pharmacophores are well-suited for efficient computer processing and easy to comprehend for life and physical scientists due to their simplistic and abstract nature. As a result, they've proven to be a useful tool for communicating between computational and medicinal chemists. The purpose of this chapter is to give a quick review of the pharmacophore idea and its uses in modern computer-aided drug design. The chapter is broken down into three sections. The first section provides an overview of the pharmacophore idea. The second section describes the most prevalent types of nonbonded interactions and how they are represented as pharmacophoric characteristics. It also provides an overview of the various methodologies for pharmacophore production as well as major pharmacophore-based drug design tools. This section finishes with some contemporary pharmacophore concept research and development examples. The final section is devoted to a review of natural product chemistry research that has been conducted using pharmacophore-based drug discovery methodologies. Antibody drug conjugates (ADCs) have evolved as a significant pharmaceutical class of medications that combine antibody specificity with the potency of small molecule therapies. The antibody, linker, and payload are the three basic components of ADCs, and the majority of early research focused on increasing their functionality. With the goal of manufacturing more homogenous ADCs, researchers have recently concentrated their efforts on developing techniques to manage the site and quantity of linker/drug attached to the antibody. In this article, we look at some of the most common conjugation methods and highlight some of the more modern ones, such as "click" conjugation and enzymatic ligation. We review current linker technology, contrasting the qualities of cleavable and non-cleavable linkers, and outline the key aspects of ADC payloads, with a focus on chemotherapeutics. Furthermore, we discuss developments in characterizing to determine physicochemical features and purifying to achieve homogeneous products. The development of a set of selection and analytical criteria will aid in the translation of innovative ADCs and ensure the manufacturing of high-quality biosimilars.

The pharmaceutical sector is under increasing pressure to produce safer and more effective drugs. Drug development efforts have traditionally been driven entirely by potency, regardless of the characteristics. As a result, developing non-drug-like compounds was expensive, risky, and had a low success rate. To tackle the obstacles, the threshold for drug candidates has been raised. To advance to clinical development, they must not only be active, but also drug-like. Drug-like qualities like solubility, permeability, metabolic stability, and transporter activities are crucial for drug candidates' success. Oral bioavailability, metabolism, clearance, toxicity, and in vitro pharmacology are all affected. In enzyme and cell-based experiments, insoluble and impermeable substances can lead to erroneous biological data and incorrect SAR. Fast clearance, short half-life, low systemic exposure, and insufficient effectiveness might result from rapid enzyme metabolism and high efflux by transporters. Early property information aids teams in making well-informed decisions and prevents the waste of valuable resources. Relationships between structure and property are critical for guiding structural alteration to improve qualities. In parallel with

activity screening, high throughput ADME/TOX assays have been established and are now frequently employed to drive drug discovery programmes. The contemporary drug discovery paradigm has made property design an integral and inseparable aspect of it. The strategy has been demonstrated to be successful.

Biography (Limit 200words)

Edward Yersal is a Senior Nurse Therapist and a researcher who has developed a technique called Rebinding of the Body which helps people recover from hormones; growth factors learn self-help techniques and lead more productive lives. Her intersubjective ethnographic study has been published in a text called, "pharmaceutical chemistry and drug design, Connection and disconnections in pharmaceutical chemistry and drug design treatment". He has published several articles in child and family psychiatry including an extensive literature review called "The Health Impact". Presently, she has a small private practice and she works as a consultant for Cogenz and Thought Leadership and Innovation Foundation. She graduated from the University of Western Ontario with Doctor of Philosophy in Nursing in 2009. Her dissertation was "Seeking and Obtaining Help for pharmaceutical chemistry and drug design.

Importance of Research (Limit 200words)

When it comes to precision medicine, medicinal chemistry is confronting new problems. Medicinal chemists today have access to a number of sophisticated new tools, as well as upgrades to existing techniques, to aid in the process of drug development, from a hit molecule to a clinically utilised medicine. The idea of evaluating folding intermediates or a protein's catalytic mechanism as a target for discovering new hits has emerged as one of the new tools. Furthermore, machine learning is a new and useful method for medicinal chemists to uncover new hits. Other talents, ranging from a better grasp of the time development of biochemical processes to a better understanding of the biological meaning of data derived from genetic analysis, are on their way to improving patient care in the drug discovery area. In this regard, advances in understanding the metabolic pathways for a rising number of medications and linking them to the genetic traits of patients, as well as innovative techniques to the administration of drugs targeted to the central nervous system, represent significant developments in the field.

About institute (Limit 200words)

Santa Rosa Junior College (SRJC) is a public community college located in Santa Rosa, California, with a second campus in Petaluma and regional centres across Sonoma County. Santa Rosa Junior College was created with the intention of serving as a feeder institution for the University of California system (a "junior" version of nearby University of California, Berkeley, with the Bear Cub mascot modelled after Oski). The Sonoma County Community College District manages SRJC. The main campus of SRJC is 52 miles (84 kilometres) north of San Francisco, on a 100-acre (0.40 km²) campus with ivy-covered brick buildings in the heart of Santa Rosa, California. The site also features a Planetarium, the Robert F. Agrella Art Gallery, the Summer Repertory Theatre, and the Santa Rosa Junior College Museum, in addition to administration buildings, classroom facilities, and laboratories.



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References (Limit 15 – 20)

1. Takahashi K, Tanabe K, Ohnuki M, et al. Induction of pluripotent stem cells from adult human fibroblasts by defined factors. *Cell* 2007;131:861-72. [10.1016/j.cell.2007.11.019](https://doi.org/10.1016/j.cell.2007.11.019)
2. Jie Yang, Yi Yuan, Jing Gu, Ao Li, Zhao Wei, Oin Ouvang. Drug Synthesis and Analysis of an Acetylcholinesterase Inhibitor: A Comprehensive Medicinal Chemistry Experience for Undergraduates. *Journal of Chemical Education* 2021, 98 (3) , 991-995.
3. Valeria V. Acuna, Rachel M. Hopper, Ryan J. Yoder. Computer-Aided Drug Design for the Organic Chemistry Laboratory Using Accessible Molecular Modeling Tools. *Journal of Chemical Education* 2020, 97 (3) , 760-763.
4. Laura Rowe . Green Fluorescent Protein-Focused Bioinformatics Laboratory Experiment Suitable for Undergraduates in Biochemistry Courses. *Journal of Chemical Education* 2017, 94 (5) , 650-655.
5. Travis R. Helgren and Timothy J. Hagen . Demonstration of AutoDock as an Educational Tool for Drug Discovery. *Journal of Chemical Education* 2017, 94 (3) , 345-349.
6. Travis R. Helgren and Timothy J. Hagen . Demonstration of AutoDock as an Educational Tool for Drug Discovery. *Journal of Chemical Education* 2017, 94 (3) , 345-349.
7. Katherine Locock, Hue Tran, Rachel Codd, and Robin Allan . Hands-On Approach to Structure Activity Relationships: The Synthesis, Testing, and Hansch Analysis of a Series of Acetylcholinesterase Inhibitors. *Journal of Chemical Education* 2015, 92 (10) , 1745-1750.
8. Julie B. Ealy. Assessment of students' external representations of mmCIF entries and their biochemical knowledge. *Biochemistry and Molecular Biology Education* 2018, 46 (6) , 634-643.
9. Marco Eissen, Donata Backhaus. Pharmaceuticals in the environment: an educational perspective. *Environmental Science and Pollution Research* 2011, 18 (9) , 1555-1566.
10. Angela L. Mahaffey. A Complementary Laboratory Exercise: Introducing Molecular Structure–Function Topics to Undergraduate Nursing Health Professions Students. *Journal of Chemical Education* 2019, 96 (10) , 2188-2193.

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