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MEDICINAL CHEMISTRY AND DRUG DISCOVERY PROFESSIONAL

Accomplished scientific leader with extensive drug discovery experience at Fortune 500 pharmaceutical companies. Expertise in medicinal and synthetic organic chemistry, preclinical drug discovery research, and implementing medicinal chemistry and drug discovery strategies to accelerate the design, synthesis, evaluation and progression of high-quality drug candidates. Established track record for successfully advancing initial hits to viable development candidates, and for identifying creative solutions to complex drug discovery challenges. Proficient in R&D leadership, and in building and managing successful, high-performing teams. Significant US and global CRO outsourcing and management experience.

- Contributed to the identification of 18 development candidates, including the late-stage clinical assets Daprodustat, Gepotidacin, Afabicin and Lotrafiban.
- Inventor on 65+ issued US patents and authorship on 40+ journal publications.

EXPERTISE

- Medicinal and synthetic organic chemistry, and preclinical drug discovery research.
- Hit identification, advancing initial hits to sustainable leads, lead optimization, analog design, synthetic route design and optimization, structure-based drug design, fragment-based drug design, molecular modeling (MOE), DMPK, due diligence evaluations, and patent preparation.
- Scientific, managerial, and strategic leadership of multi-disciplinary teams, and teams of up to 26 internal chemists or 40+ external CRO scientists.
- Collaborating across multiple disciplines, including biology, biochemistry, medicinal chemistry, development chemistry, DMPK, safety assessment, and CMC, to deliver development candidates.
- Externalization / outsourcing of R&D activities, including medicinal chemistry, scale-up chemistry, biology and DMPK, at US and international (China and India) contract research organizations (CROs).
- Therapeutic areas: antibacterials, cardiovascular, dermatology, inflammation, neuroscience (pain), oncology, ophthalmology, osteoporosis, and supportive care.
- Target classes: enzymes (including bacterial enzymes, epigenetic enzymes, kinases, prolyl hydroxylase, sirtuins, thrombin), receptors (integrins), and ion channels.
- Oral, intravenous and topical routes of administration.

EXPERIENCE

MEDCHEM INNOVATIONS, LLC, Collegeville, PA
Principal Consultant

05/2019-Present

Partner with clients in need of medicinal chemistry support, to deliver cost-effective drug discovery solutions from hit identification through lead optimization to the identification and progression of high-quality development candidates.

GLAXOSMITHKLINE (SMITHKLINE BEECHAM), Collegeville, PA; King of Prussia, PA
Senior Director, Medicinal Chemistry (04/2013-05/2019)

12/1991-05/2019

Senior chemistry research director responsible for providing leadership and guidance for medicinal chemistry teams, the medicinal chemistry community, program teams, Discovery Performance Units (DPU), and the overall business.

- Member of the leadership team that planned, built, and led the Sirtuin Discovery Performance Unit (DPU), following the closure of Sirtris in Cambridge, MA.
- Contributed to the identification of developable SIRT1 activators, including an orally active development candidate, even though the target is a flat, lipophilic surface.
- Member of the leadership team that re-established the Neuroexcitation DPU, following the closure of this group at another GSK site.
- Contributed to the identification of novel inhibitors of voltage-gated sodium channels for the treatment of pain.
- Chemistry adviser for GSK Dermatology, assisting with development of a standard critical path for topical drug discovery, and identification of molecules with suitable properties for topical delivery.
- Early Development Leader for a GSK asset, with responsibility for leading a multidisciplinary development team through enabling, post-candidate-selection studies to support Phase I clinical investigation.

Director, Medicinal Chemistry (09/2002-04/2013)

Chemistry research director responsible for providing leadership and guidance for medicinal chemistry teams, the medicinal chemistry community, program teams, DPUs, and the overall business.

- Member of the leadership team that planned, built, and led the GSK Cancer Epigenetics DPU, an industry-leading oncology epigenetics drug discovery group.
- Contributed to the identification of multiple, epigenetics-targeted development candidates from internal efforts and external collaborations. Several of these agents have entered clinical trials.
- Contributed to the identification of the HIF prolyl hydroxylase inhibitor Daprodustat, currently in Phase III trials for the treatment of anemia.
- Led the US chemistry effort on the transnational (UK/US) BTI Program. The program identified the antibacterial agent Gepotidacin, currently entering Phase III trials.
- Contributed to the identification of an antibacterial peptide deformylase inhibitor that entered Phase II trials.
- Contributed to the identification of quality inhibitors/development candidates targeting various oncology kinases (AKT, Aurora-B, FAK, PAK1, PDK1).
- MedChem representative on teams that designed/built new chemistry lab space.
- Established a stringent review process for compounds advancing into toxicology studies.
- Developed a new, standardized/streamlined process for obtaining small molecule X-ray crystal structures from external sources.

Manager (GSK)/Assistant Director (SB), Medicinal Chemistry (06/1998-09/2002)

Chemistry research manager responsible for providing leadership and guidance for multiple medicinal chemistry teams and multiple program teams.

- Co-leader/chemistry team leader of the Vitronectin Program, which advanced three compounds into development (two of which entered Phase I trials).
- Co-leader/chemistry team leader of the FabI Program, which identified novel, potent FabI-directed antibacterial agents. A molecule from this effort (Afabicin) is in Phase II trials.
- Built and led the Upper Providence MedChem Safety Program, which received company-wide recognitions for best safety practices and achieved 50/50 on safety audits.
- Performance review: "...substantial intellectual contributions to the efforts...extraordinary creativity, efficiency and productivity".

Associate Senior Investigator, then Senior Investigator, Medicinal Chemistry (12/1991-06/1998)

Initially I was a lab chemist designing/synthesizing small molecule integrin antagonists, and advanced to program co-leader/chemistry team leader. Led chemistry efforts and collaborated with biology co-leaders to oversee the overall drug discovery efforts of program teams.

- Instrumental in the identification of potent, selective integrin $\alpha v\beta 3/\alpha v\beta 5$ antagonists with high oral bioavailability, solving a long-standing challenge.
- Contributed significantly to the investigation of benzodiazepine-based screening leads and the identification/optimization of three series of highly potent, orally active $\alpha v\beta 3/\alpha v\beta 5$ antagonists (benzazepines, dibenzocycloheptenes, and phenylbutyrates).
- Investigated small molecule antagonists of integrin $\alpha IIb\beta 3$ as a lab chemist in the Fibrinogen Program, which identified Lotrafiban (terminated in Phase III trials).
- Coordinated and contributed significantly to the successful development of the first enantioselective synthesis of Lotrafiban.
- Collaborated with attorneys and other chemists to design and implement a new priority patent filing process.

DUPONT MERCK PHARMACEUTICAL COMPANY, Wilmington, DE

09/1989-12/1991

Research Scientist, then Senior Research Scientist, Medicinal Chemistry

Laboratory chemist involved in the design and synthesis of target molecules.

- Investigated inhibitors of IL-1 biosynthesis, and cartilage degradation, for treatment of arthritis, and boropeptide-based thrombin inhibitors for treatment of clotting disorders.

EDUCATION**HARVARD UNIVERSITY | NIH Postdoctoral Fellow – Synthetic Organic Chemistry**

Research Adviser: Professor Yoshito Kishi

Research: Conformational analysis of C- and O-disaccharides
Synthetic studies towards the taxane diterpenes

YALE UNIVERSITY | Ph.D. – Synthetic Organic Chemistry

Research Adviser: Professor Samuel J. Danishefsky

Dissertation: An Approach to the Total Synthesis of Anguidine

UNIVERSITY OF CONNECTICUT | B.S. in Chemistry with Honors – Graduated *Magna Cum Laude*

Research Adviser: Professor Gary A. Epling

Honors Thesis: Photochemistry from Upper Excited States

AFFILIATIONS

American Chemical Society, Organic and Medicinal Chemistry Divisions of the American Chemical Society,
Philadelphia Organic Chemist's Club

TECHNOLOGY

MOE, ScFinder, Reaxys, Microsoft Office Suite (Word, Excel, PowerPoint), ChemDraw.

REFERENCES

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ISSUED US PATENTS

1. Miller, W. H. H-[1,2]Benzisothiazolo[2,3a]quinoline-5-ones and analogs as anti-inflammatory agents. US 5,225,418, issued 06 July 1993.
2. Miller, W. H. 5H-[1,2]Benzisothiazolo[2,3a]quinoline-5-ones and analogs as anti-inflammatory agents. US 5,393,756, issued 28 February 1995.
3. Amparo, E. C.; Miller, W. H.; Pacofsky, G. J.; Wityak, J.; Weber, P. C.; Duncia, J. J. V.; Santella III, J. B. Boronic acid and ester inhibitors of thrombin. US 5,563,127, issued 08 October 1996.
4. Amparo, E. C.; Miller, W. H.; Pacofsky, G. J.; Wityak, J.; Weber, P. C.; Duncia, J. J. V.; Santella III, J. B. Boronic acid and ester inhibitors of thrombin. US 5,698,538, issued 16 December 1997.
5. Keenan, R. M.; Miller, W. H. Substituted benzimidazoles which inhibit platelet aggregation. US 5,741,804, issued 21 April 1998.
6. Ali, F. E.; Bondinell, W. E.; Huffman, W. F.; Lago, M. A.; Keenan, R. M.; Kwon, C.; Miller, W. H.; Nguyen, T. T.; Takata, D. T. Benzimidazoles/imidazoles linked to a fibrinogen receptor antagonist template having vitronectin receptor antagonist activity. US 5,977,101, issued 02 November 1999.
7. Bondinell, W. E.; Miller, W. H. Integrin receptor antagonists. US 6,008,213, issued 28 December 1999.
8. Kwon, C.; Miller, W. H. Bicyclic compounds. US 6,008,214, issued 28 December 1999.
9. Galemmo, R. A., Jr.; Abelman, M. M.; Amparo, E. C.; Fevig, J. M.; Knabb, R. M.; Miller, W. H.; Pacofsky, G. J.; Weber, P. C. Cacciola, J. Electrophilic peptide analogs as inhibitors of trypsin-like enzymes. US 6,060,462, issued 09 May 2000.
10. Ali, F. E.; Bondinell, W. E.; Calvo, R. R.; Ku, T. W.; Miller, W. H.; Samanen, J.; Venslavsky, J. W.; Yellin, T. O. Fibrinogen receptor antagonists. US 6,069,143, issued 30 May 2000.
11. Miller, W. H.; Bondinell, W. E.; Ku, T. W. Vitronectin receptor antagonists. US 6,069,158, issued 30 May 2000.
12. Ali, F. E.; Bondinell, W. E.; Keenan, R. M.; Ku, T. W.; Miller, W. H.; Samanen, J. M. Vitronectin receptor antagonists. US 6,159,964, issued 12 December 2000.
13. Miller, W. H.; Bondinell, W. E.; Ku, T. W. Vitronectin receptor antagonists. US 6,191,304, issued 20 February 2001.
14. Cousins, R. D.; Keenan, R. M.; Kwon, C.; Miller, W. H.; Uzinskas, I. N. Vitronectin receptor antagonists. US 6,458,784, issued 01 October 2002.
15. Manley, P. J.; Miller, W. H. Vitronectin receptor antagonists. US 6,458,814, issued 01 October 2002.
16. Miller, W. H.; Manley, P. J. Vitronectin receptor antagonist. US 6,495,560, issued 17 December 2002.
17. Miller, W. H.; Newlander, K. A.; Seefeld, M. A. FabI inhibitors. US 6,503,903, issued 07 January 2003.
18. Miller, W. H.; Newlander, K. A.; Seefeld, M. A. Antibacterial compounds. US 6,573,272, issued 03 June 2003.
19. Bondinell, W. E.; Miller, W. H.; Heerding, D.; Samanen, J. M. Vitronectin receptor antagonists. US 6,576,643, issued 10 June 2003.
20. Miller, W. H.; Newlander, K. A.; Seefeld, M. A.; Uzinskas, I. N. FabI inhibitors. US 6,730,684, issued 04 May 2004.
21. Miller, W. H.; Newlander, K. A.; Seefeld, M. A. FabI inhibitors. US 6,762,201, issued 13 July 2004.
22. Miller, W. H.; Newlander, K. A.; Seefeld, M. A. FabI inhibitors. US 6,765,005, issued 20 July 2004.

23. Callahan, J. F.; Cousins, R. D.; Keenan, R. M.; Kwon, C.; Miller, W. H.; Uzinskas, I. N. Vitronectin receptor antagonists. US 6,825,188, issued 30 November 2004.
24. Miller, W. H.; Newlander, K. A.; Seefeld, M. A.; Uzinskas, I. N.; DeWolf, W. E., Jr.; Jakas, D. R. FabI inhibitors. US 6,846,819, issued 25 January 2005.
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26. Miller, W. H.; Newlander, K. A.; Seefeld, M. A. Antibacterial compounds. US 6,964,970, issued 15 November 2005.
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36. Miller, W. H.; Newlander, K. A.; Seefeld, M. A.; Uzinskas, I. N.; DeWolf, W. E., Jr.; Jakas, D. R. FabI inhibitors. US 7,557,125, issued 07 July 2009.
37. Miller, W. H.; Rouse, M. R.; Seefeld, M. A. Antibacterial agents. US 7,592,334, issued 22 September 2009.
38. Miller, W. H.; Seefeld, M. A. Antibacterial agents. US 7,605,169, issued 20 October 2009.
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50. Miller, W. H.; Newlander, K. A.; Seefeld, M. A.; Uzinskas, I. N.; DeWolf, W. E., Jr.; Jakas, D. R. FabI inhibitors. US 7,790,716, issued 07 September 2010.
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52. Miller, W. H.; Newlander, K. A.; Seefeld, M. A.; Uzinskas, I. N.; DeWolf, W. E., Jr.; Jakas, D. R. FabI inhibitors. US 8,173,646, issued 08 May 2012.
53. Miller, W. H.; Tian, X.; Verma, S. K. Indoles. US 8,536,179, issued 17 September 2013.
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PUBLICATIONS

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