WILLIAM H. MILLER, PH.D.

132 Chinaberry Lane • Collegeville, PA 19426 • +1 610 724 1618
williamhmiller@verizon.net • whmiller@medchem-innovations.com
www.linkedin.com/in/william-h-miller-01b51014/

SUMMARY

Accomplished scientific leader with 29+ years of novel drug discovery contributions to Fortune 500 pharmaceutical companies. Expertise in medicinal and synthetic organic chemistry, preclinical drug discovery research, R&D leadership, building and managing successful, high-performing teams, 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. Significant US and overseas CRO outsourcing and management experience. Judicious, innovative, and effective.

- 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 (05/2019 to Present)
Principal Consultant

- 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 (formerly SMITHKLINE BEECHAM) | Collegeville, PA and King of Prussia, PA (12/1991 to 05/2019)
Senior Director, Medicinal Chemistry (04/2013 to 05/2019)

- 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 to 04/2013)

- 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 to 09/2002)

- 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) has completed 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 to 06/1998)

- Began as 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.
- 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 to 12/1991)
Research Scientist, then Senior Research Scientist, Medicinal Chemistry

- Lab 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 Director: 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 Director: 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 Director: 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

James L. Ellis, Ph.D. CSO Nocion Therapeutics, Inc. jim@nociontx.com (617) 230-4817 (mobile)	Zheng Li, Ph.D. VP, Head of Neuroexcitation DPU GlaxoSmithKline zheng.n.li@gsk.com (734) 730-2505 (mobile)
Karen A. Evans, Ph.D. Manager, Medicinal Chemistry GlaxoSmithKline karen.a.evans@gsk.com (610) 917-7764 (work)	Allen I. Oliff, MD SVP, Senior Discovery Consultant GlaxoSmithKline allen.i.oliff@gsk.com (484) 767-2784 (mobile)
Ryan M. Fox Operations Director GlaxoSmithKline ryan.m.fox@gsk.com (610) 220-6394 (mobile)	David Rubenstein, MD, Ph.D. CSO Dermavant Sciences, Inc. david.rubenstein@dermavant.com (919) 414-6127 (mobile)
Erding Hu, Ph.D. Exploratory Biology Director GlaxoSmithKline erding.hu@gsk.com (610) 324-2258 (mobile)	Peter Tummino, Ph.D. CSO Nimbus Therapeutics peter.tummino@nimbustx.com (610) 291-6238 (mobile)
Richard M. Keenan, Ph.D. Principal Consultant RMK Drug Discovery Consulting, LLC rickkeenan@gmail.com (610) 585-1183 (mobile)	

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PUBLICATIONS

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