

Roman Manetsch, Ph.D.

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A. EDUCATION, POSITIONS AND HONORS

Education

- October 2002 Ph.D., Chemistry
Jointly at the Institute of Organic Chemistry at the University of Basel (Switzerland) and at the Department of Chemistry and Biochemistry at the University of Berne (Switzerland),
Advisor Professor Wolf-Dietrich Woggon and Co-Advisor Professor Jean-Louis Reymond
Thesis: *“Transition State Analogues for the Identification of the Enzyme Tocopherol Cyclase and for the Preparation of Catalytic Monoclonal Antibodies”*
- June 1998 Diploma in Chemistry
University of Basel (Switzerland), Studies in Chemistry (main subject) and Biology (minor subject)
Advisor: Professor Wolf-Dietrich Woggon
Thesis: *Synthesis of Potential Inhibitors of the Enzyme Carotene Oxygenase*

Positions and Employment

- 2002 – 2005 Postdoctoral Fellow, The Scripps Research Institute, La Jolla, CA
2005 – Present Assistant Professor, Department of Chemistry, University of South Florida, Tampa, FL

Honors

- 2002 Ph.D. Summa Cum Laude
2003 Swiss National Science Foundation Postdoctoral Fellowship
2003 Novartis Foundation (formerly the Ciba-Geigy Jubilee Foundation), Postdoctoral Fellowship
2004 Swiss National Science Foundation Postdoctoral Fellowship

Research Summary

- The research interests of the Manetsch laboratory represent a well-balanced blend of organic and biological chemistry, addressing in particular modern aspects of medicinal chemistry. The research focuses mainly on lead discovery and optimization using synthetic chemistry in close conjunction with liquid chromatography with mass spectrometry (LC-MS) and molecular modeling. The Manetsch laboratory develops straightforward strategies for fragment-based lead discovery and optimization using the protein of interest to assemble its own high-affinity bidentate ligand from a library of complimentary reacting fragments. The developed lead discovery methods are currently applied for the discovery of therapeutic agents targeting cancer, malaria and infectious diseases.
- The second research pillar of the Manetsch laboratory comprises the development of chemical tools to label and identify specific proteins in complex mixtures or entire proteomes. Currently, the Manetsch laboratory is developing various probes to investigate proteins related to cell-cell communication and energy metabolism.
- Current Research Group: two postdoctoral researchers (Dr. Niranjana Namelikonda, Dr. David Flanigan), eight graduate students (Richard Matthew Cross, Sameer Kulkarni, Arun Babu Kumar, Shikha Mahajan (co-advised by Professor David Merkler, Department of Chemistry, University of South Florida), Jordany Maignan Andrii Monastyrskyi, Katya Nacheva, Kurt Van Horn) and nine undergraduate students (Jordan Anderson, Renee Fleeman, Petoria Gayle, Lisa Luong, Anthony Melendez, David Z. Myers, Justin Sargent, Maylene Quiroga, Nicolas Zoumberos)

Editorial activities

Served as referee for *The Journal of Organic Chemistry*, *Combinatorial Chemistry & High Throughput Screening*, *Medicinal Research Reviews* and *Bioorganic and Medicinal Chemistry Letters*.

Teaching Summary (in chronological order)

- October 1998 to October 1999. Teaching assistant in the practical laboratory classes for students in Biochemistry, Biology and Pharmacy at the Institute of Organic Chemistry at the University of Basel (Switzerland)
- Spring 1999, Fall 2001 and Spring 2002. Supervision of three final year undergraduate students during their diploma research projects at the Institute of Organic Chemistry at the University of Basel (Switzerland)
- Fall 2005, Fall 2006, Fall 2007. Teaching of the graduate level course CHM6250/5225 Advanced Organic Chemistry I at the Department of Chemistry at the University of South Florida. Graduate level course with approximately 15 participants
- Spring 2007, Spring 2008, Fall 2008, Spring 2009. Teaching of the undergraduate level course CHM2210 Organic Chemistry I at the Department of Chemistry at the University of South Florida. Graduate level course with 210 participants
- Fall 2006, Spring 2007, Fall 2007, Spring 2008, Fall 2008, Spring 2009. Coordinating CHM6935 Graduate Seminar program of the Department of Chemistry. Course with 70 participants.

B. PUBLICATIONS (in chronological order)

- 1) Manetsch R, Zheng L, Reymond MT, Woggon WD, Reymond JL*. A Catalytic Antibody against a Tocopherol Cyclase Inhibitor. *Chem Eur J* 2004; 10: 2487-2506.
- 2) Manetsch R, Krasinski A, Radic Z, Raushel J, Taylor P, Sharpless KB, Kolb HC*. In Situ Click Chemistry: Enzyme Inhibitors Made to Their Own Specifications. *J Am Chem Soc* 2004; 126: 12809-12818.
- 3) Zheng L, Manetsch R, Woggon WD, Baumann U, Reymond JL*. Mechanistic Study of Proton Transfer in Catalytic Antibody 16E7 by Site-directed Mutagenesis and Homology Modeling. *Bioorg Med Chem* 2005; 13: 1021-1029.
- 4) Krasinski A, Radic Z, Manetsch R, Raushel J, Taylor P, Sharpless KB, Kolb HC*. Click Chemistry Screening In Situ: Target-guided Optimization of Acetylcholinesterase Inhibitors. *J Am Chem Soc* 2005; 127: 6686-6692.
- 5) Radic Z, Manetsch R, Krasinski A, Raushel J, Yamauchi J, Garcia C, Kolb HC, Sharpless KB, Taylor P*. Molecular basis of interactions of cholinesterases with tight binding inhibitors. *Chem-Biol Interact* 2005; 157: 133-141.
- 6) Sharpless KB, Manetsch R*. In Situ Click Chemistry: A Powerful Means for Lead Discovery (Review). *Expert Opinion on Drug Discovery* 2006; 1: 525-538.
- 7) Radic Z, Manetsch R, Fournier D, Sharpless KB, Taylor P*. Probing Gorge Dimensions of Cholinesterases by Freeze-Frame Click Chemistry. *Chem-Biol Interact* 2008; 175: 161-165.
- 8) Hu X, Sun J, Wang H-G, Manetsch R*. Bcl-X_L-Templated Assembly of its Own Protein-Protein Interaction Modulator from Fragments Decorated with Thio Acids and Sulfonyl Azides. *J Am Chem Soc* 2008; 130: 13820-13821.

C. PATENTS, BOOK CHAPTERS and OTHERS

- Two International Patent Applications on two compound series targeting proteins of the Bcl-2 family as anti-cancer agents were filed February 23, 2009 through the Patent Cooperation Treaty.
- In Situ Click Chemistry. Book chapter in a book on click chemistry. Editor Valery V. Fokin, Wiley, Fall 2009.

D. INVITED TALKS

- 1) Mass Spectrometry-Guided Chemistry Approaches Targeting Cancer and Malaria
The Skaggs School of Pharmacy and Pharmaceutical Sciences at the University of California of San Diego
San Diego, California, April 8, 2009
- 2) Conventional Lead Discovery Approaches and Kinetic Target-Guided Synthesis Targeting Cancer and Malaria
Department of Chemistry and Biochemistry, Florida State University
Tallahassee, Florida, November 20, 2008
- 3) Quinolones and 1,2,3,4-Tetrahydroacridinols Chemotypes for Malaria Drug Discovery.
American Chemical Society, Florida Section
Orlando, May 8 - 10, 2008
- 4) Target-Guided Synthesis: A New Approach for Drug Discovery
BioStat International / Molecular Medicine Seminar Series, College of Medicine, University of South Florida
Tampa, March 31, 2006

E. RESEARCH SUPPORT

Current

- 1) Agency FCoE-BITT Seed Grant
Funding Period 06/01/2009-05/31/2010
Role in Project Co-PI Manetsch (42.5 %)
Other Contributors PI John Adams (57.5 %) USF College of Public Health
Title Evaluation of a phosphotyrosine phosphatase as an antimalarial drug target
Total Amount \$75,000 (direct \$75,000; no indirect)
Manetsch Lab (direct) \$31,874
- 2) Agency FCoE-BITT Seed Grant
Funding Period 06/01/2009-05/31/2010
Role in Project Co-PI Manetsch (42.5 %)
Other Contributors PI Andreas Seyfang (57.5 %) USF College of Medicine, Molecular Medicine
Title Characterization of Candida Cytochrome b5 Reductase as Pharmacological Target
Total Amount \$75,000 (direct \$75,000; no indirect)
Manetsch Lab (direct) \$31,874
- 3) Agency Medicines For Malaria Venture
Funding Period 11/01/08-12/31/2009
Role in Project PI Roman Manetsch (51 %)
Other Contributors Co-PI Dennis Kyle (49 %) USF Department of Global Health
Title 4(1*H*)-Quinolone and 1,2,3,4-Tetrahydroacridone Chemotypes for Malaria Drug Discovery
Total Amount \$399,025 (direct \$362,750; indirect 10% \$36,275)
Manetsch Lab (direct) \$187,899
- 4) Agency Bankhead-Coley Biomedical Research Program, Florida Department of Health
Funding Period 07/01/2008-06/30/2011
Role in Project PI Manetsch (50 %)
Other Contributors Co-PI David Merkler (50 %) USF Chemistry
Title Chemical Tools for Proteomic Profiling
Total Amount \$375,000 (direct \$347,222; indirect 8% \$27,778)
Manetsch Lab (direct) \$173,610

- 5) Agency Johnnie B. Byrd, Sr. Alzheimer's Center and Research Institute
 Funding Period 7/01/2008-12/31/2009
 Role in Project PI Manetsch (50 %)
 Other Contributors Co-PI David Merkler (50 %) USF Chemistry
 Title Adenyloomics and Caffeyloomics
 Total Amount \$40,793 (direct \$31,869; indirect 28% \$8,924)
 Manetsch Lab (direct) \$15,934
- 6) Agency James and Esther King Biomedical Research Program, Florida Department of Health
 Funding Period 07/01/2007-06/30/2010
 Role in Project PI Manetsch (95 %)
 Other Contributors Mentor Wayne Guida (5 %) USF Chemistry
 Title Bcl-X_L-templated Assembly of Compounds Modulating Bcl- X_L-Protein Interactions
 Total Amount \$375,000 (direct \$347,222; indirect 8% \$27,778)
 Manetsch Lab (direct) \$329,859

Expired

- 1) Agency FCoE-BITT Seed Grant, GALS007
 Funding Period 05/01/2008-04/30/2009
 Role in Project Co-PI Manetsch (50 %)
 Other Contributors PI David Merkler (50 %) USF Chemistry
 Title Adenyloomics
 Total Amount \$75,000 (direct \$75,000; no indirect)
 Manetsch Lab (direct) \$37,500
- 2) Agency FCoE-BITT Seed Grant, GALS008
 Funding Period 05/01/2008-04/30/2009
 Role in Project PI Manetsch (51 %)
 Other Contributors Co-PI Dennis Kyle (49 %) USF Department of Global Health
 Title SAR Study of Quinolones and 1,2,3,4-Tetrahydroacridinols for the Development of Novel Chemotypes Targeting Atovaquone Resistant Malaria Parasites
 Total Amount \$75,000 (direct \$38,000; no indirect)
 Manetsch Lab (direct) \$38,000
- 3) Agency Interdisciplinary Research Development Grant, University of South Florida
 Current Status expired
 Funding Period 03/01/2006-02/29/2008
 Role in Project PI Manetsch (33.3 %)
 Other Contributors Co-PIs Edwin Rivera (33.3 %) USF Chemistry and Van Olphen (33.3 %) USF Center for Biological Defense
 Title Development of Novel Antiviral Compounds Targeting Non-structural Protein 1
 Total Amount \$50,000 (direct \$50,000; no indirect)
 Manetsch Lab (direct) \$16,667
- 4) Agency American Cancer Society Institutional Grant Program, Cycle 20, Fall 2005
 Current Status expired
 Funding Period 04/01/2006-03/31/2007
 Role in Project PI Manetsch (100 %)
 Other Contributors None
 Title Bcl-X_L-templated Assembly of Compounds Modulating Bcl-X_L
 Total Amount \$20,000 (direct \$20,000; no indirect)
 Manetsch Lab (direct) \$20,000

5) Agency USF Health Office of Research
Current Status expired
Funding Period 02/06-01/07
Role in Project Co-PI Manetsch (33.3 %)
Co-PIs Edwin Rivera (33.3 %) USF Chemistry and PI Van Olphen (33.3 %) USF Center
Other Contributors for Biological Defense
Title Development of Novel Antiviral Compounds against Influenza
Total Amount \$19,994 (direct \$19,994; no indirect)
Manetsch Lab (direct) \$6,665