Joseph E. Wedekind, Ph.D.

Professor

Department of Biochemistry & Biophysics,
Center for AIDS Research and Center for RNA Biology
601 Elmwood Avenue, MC 712
University of Rochester School of Medicine & Dentistry
Rochester, New York 14642
e-mail: joseph.wedekind@rochester.edu

https://www.urmc.rochester.edu/labs/wedekind.aspx



EDUCATION

Ph.D., **Biochemistry** 1995 (Structural Enzymology) University of Wisconsin, Madison

B.S. Biochemistry 1990

University of California, Riverside

EXPERIENCE

2002 – present

Department of Biochemistry & Biophysics, University of Rochester, New York Professor (2014 – present), Associate Professor (2006 – 2014, tenured 2010), Assistant Professor (2002-2006)

1999 - 2002

Department of Biochemistry & Molecular Biology, Rosalind Franklin University, Illinois Assistant Professor

1995-1999

Department of Structural Biology, Stanford University, California Post-Doctoral Research Fellow (Prof. David B. McKay)

1990-1995

Institute for Enzyme Research, University of Wisconsin at Madison Graduate Research Assistant (Advisor: Prof. Ivan Rayment; co-advisor: Prof. Perry A Frey)

1988-1990

Department of Biochemistry, University of California at Riverside *Undergraduate Research Assistant* (Prof. Alexander McPherson Jr.)

1989

Department of Pathology, University of Southern California, Los Angeles *Edmondson Summer Research Fellow* (Prof. Alan Epstein)

1986-1989

United States Department of Agriculture and Department of Entomology, Riverside, California *Federal Work Study Program* (Prof. Dan Gonzalez)

HONORS

05,'10,'15, '19	Teaching Award in Biophysics, University of Rochester
2012-13	Excellence in Research Award, University of Rochester Medical Center Faculty
1996-99	Burroughs Wellcome Fund Fellow of the Life Sciences Research Foundation
1995	Dean's Fellow, Stanford University School of Medicine
1992-95	NIH Biophysics Training Grant T32, U.W. Madison
1990	B.S., Magna Cum Laude, 3.80 GPA, U.C. Riverside (top 2% of class)
1990	Elected to Phi Beta Kappa, lota Chapter of California
1986-90	Dean's List, Pell Grant, Cal Grant A

SERVICE

2019-2022 *Member*, Strategic Advisory Committee, University of Rochester

2018-22 Seated Member, MSFB Study Section, NIH Center for Scientific Review 2017-19 Elected Secretary, Medical Faculty Council, University of Rochester

2013-present Scientific Advisory Board, Functional Proteomics Core, University of Rochester

2011-16 Editorial Board Member, Journal of Biological Chemistry

2010-present Founder & Co-Director, Struct. Biol. & Biophys. Facility, University of Rochester

2008-10 Elected Secretary, Board of Directors, The Pittsburgh Diffraction Society

ONGOING RESEARCH SUPPORT

R01 Al150463 Wedekind (PI) 08/01/17 – 07/31/21

NIH/NIAID

Cyclic Peptide Inhibitors of HIV-1 Proliferation

Goals: The goal of this project is to develop novel constrained peptides that target HIV-1 TAR based on our library of lab-evolved $\underline{T}AR$ $\underline{B}inding$ $\underline{P}roteins$ (TBPs). The novel co-crystal structure of TAR in complex with variant TBP6.7 suggests a minimal lab-evolved β 2- β 3 peptide is sufficient for binding. We will optimize this peptide, followed by testing for anti-viral activity. We will also work to identify other novel peptides that bind TAR based on our TBP library. The results should provide new compounds that block viral infectivity, leading to a functional cure. Role: PI

R01 GM63162 Wedekind (PI) 07/01/17 – 06/30/21

NIH/NIGMS

Mechanisms of Action of Non-Coding RNA Molecules

Goals: The major goal of this project is to elucidate riboswitch signal-transduction pathways for the family of translational preQ₁ riboswitches. We will use: (*i*) crystallography and calorimetry to characterize on-path mutants, (*ii*) an innovative live-cell chemical modification analysis coupled to a GFP reporter assay, and (*iii*) single-molecule kinetic analysis of RNA transient structure (SiM-KARTS) to probe Shine-Dalgarno sequence accessibility. The results will reveal unprecedented structure-function relationships that integrate atomic-resolution structural and biophysical information with effector-mediated RNA conformational changes in live cells.

Role: PI

3R01GM063162-15S1 Wedekind (PI) 09/01/19 – 08/31/20

NIH/NIGMS

Mechanisms of Action of Non-Coding RNA Molecules

Goals: This equipment supplement was awarded to support the acquisition of a Sorvall LYNX 6000 superspeed centrifuge with six accompanying carbon fiber and titanium rotors.

Role: PI

T32AI049815 Wedekind (PI) 07/01/11 – 06/30/21

NIH/NIAID

Training in HIV Replication and Pathogenesis

Major Goal: This program seeks to train predoctoral students for careers as outstanding research scientists in the field of HIV/AIDS.

Role: PI

PUBLICATIONS

78 Total (71 manuscripts + 7 book chapters https://www.urmc.rochester.edu/people/23504899-joseph-e-wedekind/publications)

h-index = 34; i10 = 61 (https://scholar.google.com/citations?user=-FWrFpoAAAAJ&hl=en)

Selected Publications

SS Chavali, R Bonn-Breach, JE <u>Wedekind</u>* (2019) Face-time with TAR: Portraits of an HIV-1 RNA with diverse modes of effector recognition relevant for drug discovery. *J. Biol. Chem.* **294**, 9326-9341. [featured in JBC Reviews roundup].

- IA Belashov, DW Crawford, CE Cavender, P Dai, PC Beardslee, DH Mathews, BL Pentelute, BR McNaughton, JE <u>Wedekind</u>* (2018) Structure of HIV TAR in complex with a Lab-Evolved RRM provides insight into duplex RNA recognition and synthesis of a constrained peptide that impairs transcription. *Nucl. Acids Res.* **46**, 6401-6415. [*Breakthrough article & cover*].
- JA Liberman, KC Suddala, A Aytenfisu, D Chan, IA Belashov, M Salim, DH Mathews, RC Spitale, NG Walter, JE <u>Wedekind</u>* (2015) Structural analysis of a class III preQ₁ riboswitch reveals an aptamer distant from a ribosome-binding site regulated by fast dynamics. *Proc. Nat'l Acad. Sci.* **112**, E3485-E3494. [*PNAS Plus manuscript*]
- JA Liberman, M Salim, J Krucinska, JE <u>Wedekind</u>* (2013) Structure of a class II preQ₁ riboswitch reveals ligand recognition by a new fold. *Nat. Chem. Biol.* **9**, 353-355. [cover art]
- M Guo, RC Spitale, R Volpini, J Krucinska, G Cristalli, PR Carey, JE <u>Wedekind</u>* (2009) Direct Raman Measurement of an Elevated Base pK_a in the Active Site of a Small Ribozyme in a Precatalytic Conformation. *J. Am. Chem. Soc.* **131**, 12908-12909 [C&E News Highlight].
- JE <u>Wedekind</u>*, R Gillilan, A Janda, J Krucinska, JD Salter, RP Bennett, J Raina, HC Smith (2006) Nanostructures of APOBEC3G support a hierarchical assembly model of high molecular mass ribonucleoprotein particles from dimeric subunits. *J. Biol. Chem.* **281**, 38122-38126. [accelerated publication].
- AK Ghosh, PR Sridhar, S Leshchenko, AK Hussain, J Li, AY Kovalevsky, DE Walters, JE <u>Wedekind</u>, V Grum-Tokars, D Das, Y Koh, K Maeda, H Gatanaga, IT Weber, H Mitsuya (2006) Structure-based design of novel HIV-1 protease inhibitors to combat drug resistance. *J. Med. Chem.* **49**, 5252-5261.
- K Xie, MP Sowden, GSC Dance, AT Torelli, HC Smith, JE <u>Wedekind</u>* (2004) The structure of a yeast RNA-editing deaminase provides insight into the fold and function of activation-induced deaminase and APOBEC-1. *Proc. Nat'l Acad. Sci.* **101**, 8114-8119.
- JE <u>Wedekind</u>, DB McKay (1999) Crystal structure of a lead-dependent ribozyme revealing metal binding sites relevant to catalysis. *Nat. Struct. Molec Biol.* **6**, 261-268. [commentary by E. Westhof].
- PC Babbitt, MS Hasson, JE <u>Wedekind</u>, DRJ Palmer, WC Barrett, GH Reed, I Rayment, G D Ringe, GL Kenyon, JA Gerlt (1996) The enolase superfamily: a general strategy for enzyme-catalyzed abstraction of the α-protons of carboxylic acids. *Biochemistry* **35**, 16489-16501.
- JE <u>Wedekind</u>, PA Frey, I Rayment (1996) The Structure of Nucleotidylated Histidine-166 of Galactose-1-phosphate Uridylyltransferase Provides Insight into Phosphoryl Group Transfer, *Biochemistry* **35**, 11560-11569.