

DEPARTMENT OF BIOCHEMISTRY, MOLECULAR BIOLOGY, & BIOPHYSICS, THE UNIVERSITY OF MINNESOTA  
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## MICHAEL F. FREEMAN

### EDUCATION

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*Johns Hopkins University*, Baltimore, Maryland, USA  
Ph.D. Biology, 2008

*University of Massachusetts*, Amherst, Massachusetts, USA  
B.S. Biochemistry and Molecular Biology, 2001  
GPA: 3.9/4.0

### RESEARCH EXPERIENCE

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| 2016-present | <b>Assistant Professor at the University of Minnesota</b><br>Department of Biochemistry, Molecular Biology, and Biophysics & The Biotechnology Institute   |
| 2010-2015    | <b>Post-doctoral Research with Professor Jörn Piel</b><br>Eidgenössische Technische Hochschule (ETH) Zurich, Zurich, Switzerland<br>University of Bonn, Bonn, Germany<br><i>Discovery and Elucidation of Marine Natural Products</i> |
| 2008-2010    | <b>Post-doctoral Research with Professor Craig A. Townsend</b><br>Johns Hopkins University, Baltimore, Maryland, USA<br><i>Regulation of <math>\beta</math>-Lactam Antibiotics in <i>Streptomyces</i></i>                            |
| 2001-2008    | <b>Graduate Research with Professor Craig A. Townsend</b><br>Johns Hopkins University, Baltimore, Maryland, USA<br><i>Characterization of Enzymes Involved in Thienamycin Biosynthesis</i>   |
| 1999-2001    | <b>Undergraduate Research with Professor Craig T. Martin</b><br>University of Massachusetts, Amherst, Massachusetts, USA<br><i>Directed Evolution of T7 RNA Polymerase Promoters</i>   |

### RESEARCH FUNDING

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| 2018-2019 | <i>Co-PI</i> , University of Minnesota Biocatalysis Initiative for Advancing Biomanufacturing for the Environment, Health and Industry |
| 2017-2019 | University of Minnesota Biocatalysis Initiative for Advancing Biomanufacturing for the Environment, Health and Industry                |
| 2011-2014 | Human Frontier Science Program (HFSP) Long-Term Fellowship<br><i>Harnessing the Bacterial Biodiversity of Marine Invertebrates</i>     |

## PUBLICATIONS, PATENTS, AND PRESENTATIONS

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*Publications*                      \* *Equal contribution of authors*, § *Co-corresponding authors*, † *Corresponding author*

20. Song, H, van der Velden, NS, Shiran, SL, Bleiziffer, P, Zach, C, Sieber, R, Imani, AS, Krausbeck, F, Aebi, M, **Freeman, MF**, Riniker, S§, Kuenzler, M§, Naismith, JH§ (2018) A molecular mechanism for the enzymatic methylation of nitrogen atoms within peptide bonds, *Sci. Adv.* 4(8): eaat2720. DOI: 10.1126/sciadv.aat2720.
19. Miller, FS, **Freeman, MF**† (2018) Impact of synthetic biology on secondary metabolite biosynthesis, *Modern biocatalysis: Advances towards synthetic biological systems*. RSC Publishing Cambridge, UK. Chapter 11: 287-320. DOI: 10.1039/9781788010450.
18. **Freeman, MF**† (2018) Cobalamin-dependent C-methyltransferases from marine microbes: accessibility via rhizobia expression, *Methods Enzymol.* 604:259-286. DOI: 10.1016/bs.mie.2018.02.013.
17. Imani, AS, **Freeman, MF**† (2018) RiPPing apart the rules for peptide natural products, *Syst. Synth. Biotechnol.* 3(2): 81-82. DOI: 10.1016/j.synbio.2018.03.002.
16. van der Velden, NS, Kaelin, N, Helf, MJ, Piel, J, **Freeman, MF**§, Kuenzler, M§ (2017) Autocatalytic backbone N-methylation in a family of ribosomal peptide natural products, *Nat. Chem. Biol.* 13(8): 833-835. DOI: 10.1038/nchembio.2393.  
[News and Views: *Nat. Chem. Biol.* 2017 13: 821-822.]  
[Highlight: *Angew. Chem. Int. Ed. Engl.* 2017 56: 2-5.]
15. Morinaka, BI, Verest, M, **Freeman, MF**, Gugger, M, Piel, J (2017) An orthogonal D<sub>2</sub>O-based induction system provides insights into D-amino acid pattern formation by radical S-adenosylmethionine peptide epimerases, *Angew. Chem. Int. Ed. Engl.* 56: 762-766. DOI: 10.1002/anie.201609469.
14. **Freeman, MF**§, Helf, MJ, Bhushan, A, Morinaka, BI, Piel, J§ (2017) Seven enzymes create extraordinary molecular complexity in an uncultivated bacterium, *Nat. Chem.* 9: 387-395. DOI: 10.1038/nchem.2666.  
[Research Highlight: *Nat. Chem. Biol.* 2017 13: 129.]  
[Leading Edge: *Cell* 2017 169: 373.]
13. **Freeman, MF**, Vagstad, AL, Piel, J (2016) Polytheonamide biosynthesis showcasing the metabolic potential of sponge-associated uncultivated 'Entotheonella', *Curr. Opin. Chem. Biol.* 31: 8-14. DOI: 10.1016/j.cbpa.2015.11.002.
12. Buller, AR, **Freeman, MF**, Schilbach, JF, Townsend, CA (2014) Exploring the role of conformational heterogeneity in *cis*-autoproteolytic activation of ThnT, *Biochemistry* 53(26): 4273-4281. DOI: 10.1021/bi500385d.
11. Morinaka, BI, Vagstad, AL, Helf, MJ, Gugger, M, Kegler, C, **Freeman, MF**, Bode, HB, Piel, J (2014) Radical S-adenosyl methionine epimerases: regioselective introduction of diverse D-amino acid patterns into peptide natural products, *Angew. Chem. Int. Ed.* 53(32): 8503-8507. DOI: 10.1002/anie.201400478.
10. Cai, X, Teta, R, Kohlhas, C, Cruesemann, M, Ueoka, R, Mangoni, A, **Freeman, MF**§, Piel, J§ (2013) Manipulation of regulatory genes reveals complexity and fidelity in hormaomycin biosynthesis, *Chem. Biol.* 20(6): 839-846. DOI: 10.1016/j.chembiol.2013.04.018.

9. **Freeman, MF\***, Gurgui, C\*, Helf, MJ, Uria, AR, Oldham, NJ, Sahl, HG, Matsunaga, S, Piel, J (2012) Metagenome mining reveals polytheonamides as posttranslationally modified ribosomal peptides, *Science* 338: 387-390. DOI: 10.1126/science.1226121.  
[Concentrates: *Chem. Eng. News* 2012 90(3): 26.]
8. Buller, AR, Labonte, JW, **Freeman, MF**, Wright, NT, Schildbach, JF, Townsend, CA (2012) Autoproteolytic activation of ThnT results in structural reorganization necessary for substrate binding and catalysis, *J. Mol. Biol.* 422(4): 508-518. DOI: 10.1016/j.jmb.2012.06.012.
7. Labonte, JW, Kudo, F, **Freeman, MF**, Raber, ML, Townsend, CA (2012) Engineering the synthetic potential of  $\beta$ -lactam synthetase and the importance of catalytic group dynamics, *MedChemComm.* 3: 960-966. DOI: 10.1039/C2MD00305H.
6. Buller, AR, **Freeman, MF**, Wright, NT, Schildbach, JF, Townsend, CA (2011) Insights into cis-autoproteolysis reveal a reactive state formed through conformation rearrangement, *Proc. Natl. Acad. Sci. U.S.A.* 109(7): 2308-2313. DOI: 10.1073/pnas.1113633109.
5. Bodner, MJ, Li, R, Phelan, RM, **Freeman, MF**, Moshos, KA, Lloyd, E, Townsend, CA (2011) Definition of the common and divergent steps in carbapenem  $\beta$ -lactam antibiotic biosynthesis, *ChemBioChem.* 12(14): 2159-2165. DOI: 10.1002/cbic.201100366.
4. Gulder, TAM, **Freeman, MF**, Piel, J (2011) The catalytic diversity of multimodular polyketide synthases: natural product biosynthesis beyond textbook assembly rules, *Top. Curr. Chem.* (pp. 1-53) Springer-Verlag Berlin Heidelberg. DOI: 10.1007/128\_2010\_113. DOI: 10.1007/128\_2010\_113.
3. Bodner, MJ, Phelan, R, **Freeman, MF**, Li, R, Townsend, CA (2010) Non-heme iron oxygenases generate natural structural diversity in carbapenem antibiotics, *J. Am. Chem. Soc.* 132(1): 12-13. DOI: 10.1021/ja907320n.
2. Raber, ML, **Freeman, MF**, Townsend, CA (2009) Dissection of the stepwise mechanism to  $\beta$ -lactam formation and elucidation of a rate-determining conformational change in  $\beta$ -lactam synthetase, *J. Biol. Chem.* 284(1): 207-217. DOI: 10.1074/jbc.M805390200.
1. **Freeman, MF**, Moshos, KA, Bodner, MJ, Li, R, Townsend, CA (2008) Four enzymes define the role of coenzyme A in thienamycin biosynthesis, *Proc. Natl. Acad. Sci. U.S.A.* 105(32): 11128-11133. DOI: 10.1073/pnas.0804500105.  
[Research Highlight: *ACS Chem. Biol.* 2008 3(9): 522.]

## *Patents*

3. Kuenzler, J., van der Velden, N, **Freeman, MF**, Piel, J, Aebi, N, Kaelin, N, Novel Multiply Backbone N-Methyl Transferases and Uses Thereof, WO2017EP58327, October 12, **2017**.
2. Piel, J, Gurgui, C, **Freeman, MF**, Uria, AR, Helf, MJ, Biosynthetic Gene Cluster for the Production of Peptide/Protein Analogues, WO2013034579 A1, March 14, **2013**.
1. Townsend, CA, Bodner, MJ, Phelan, RM, **Freeman, MF**, Method for Late Introduction of the (8R)-Hydroxyl Group in Carbapenem Beta-lactam Antibioitc Synthesis, EP2513112 A2, October 24, **2012**.

## *Oral Presentations*

**Freeman, MF** Invited speaker, SIMB Annual Meeting, Chicago, IL, August 16, **2018**.

**Freeman, MF** Invited speaker, ASBMB / Experimental Biology 2018, San Diego, CA, April 23, **2018**.

**Freeman, MF** MinnCrest training seminar “How to start a lab?”, University of Minnesota, April 18, **2018**.

**Freeman, MF** Invited speaker, MycoNet seminar, University of Minnesota, MN April 11, **2018**.

**Freeman, MF** BTI lunch training seminar, University of Minnesota, MN, March 26, **2018**.

**Freeman, MF** Invited speaker, ACS National Conference, New Orleans, LA, March 22, **2018**.

**Freeman, MF** Invited speaker, Developmental Biology Center Seminar, University of Minnesota, December 7, **2017**.

**Freeman, MF** Invited speaker, Bug Club, University of Minnesota, October 13, **2017**.

**Freeman, MF** BMBB annual retreat, Itasca State Park, University of Minnesota, October 1, **2017**.

**Freeman, MF** Joint Symposium on Microbial Biotechnology, University of Minnesota, August 8, **2017**.

**Freeman, MF** Invited speaker, Biofilm club, University of Minnesota, April 4, **2017**.

**Freeman, MF** Directing Biosynthesis V, Norwich England, March 22, **2017**. (*Lightning talk*)

**Freeman, MF** Co-organizer, 2017 Microbial Factories Symposium, University of Minnesota, February 6, **2017**.

**Freeman, MF** Invited speaker, Microbial Communication Colloquium, Friedrich Schiller University, Jena, Germany, November 30, **2016**.

**Freeman, MF** Invited speaker, Mini-symposium: Structural aspects of synthetic biology systems, University of Minnesota, St. Paul, Minnesota, USA, August 3, **2016**.

**Freeman, MF** Invited Speaker, Biofilm Club Symposium, University of Minnesota, St. Paul, Minnesota, USA, May 20, **2016**.

**Freeman, MF** Invited Speaker, Science on the Spot, University of Minnesota, St. Paul, Minnesota, USA, April 14, **2016**.

**Freeman, MF** Invited Speaker, Host: Prof. Dr. Yaniv Brandvain. PBS Colloquium, University of Minnesota, St. Paul, Minnesota, USA, April 5, **2016**.

**Freeman, MF** Invited Speaker, Host: Prof. Dr. Michael H. Walter. University of Northern Iowa, Cedar Falls, Iowa, February 29, USA, **2016**.

**Freeman, MF** BMBB annual retreat, Itasca State Park, University of Minnesota, September 25, **2015**.

## TEACHING

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2017, Fall	BioC3960 Research Topics in Biochemistry, Guest lecturer, University of Minnesota
2017, Summer	Co-director of MCSB graduate research course, Itasca Biological Station, University of Minnesota.
2017, Spring	BioC5309 Biocatalysis and Biodegradation, Guest lecturer, University of Minnesota
2016-present	Research mentor to post-docs (2), Ph.D. students (3), Master's students (1), undergraduates (5), and High School students (4).

## AWARDS AND HONORS

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2016-2017	Residence Workplace Agreement, The Institute of Microbiology, Eidgenössische Technische Hochschule (ETH) Zurich, Zurich, Switzerland
2015	ETH Institute of Microbiology Performance Award
2011-2014	Human Frontier Science Program (HFSP) Long-Term Fellowship recipient <i>Harnessing the Bacterial Biodiversity of Marine Invertebrates</i>

## SERVICE

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2018-2019	HHMI Faculty Fellows for Inclusive Excellence Program
2018	LSSURP mock graduate student interviewee
2018	Session leader in round-table discussion for MinnCRest event: 'How to start your new lab?'
2016-present	Mentor for Minnetonka Research – Minnetonka High School
2016-present	Manuscript reviewer for <i>Nat. Chem. Biol.</i> (2), <i>Angew. Chem. Int. Ed.</i> , <i>Metab. Eng.</i> , <i>Nucleic Acids Res.</i> , <i>ACS Chem. Biol.</i> , <i>Biochemistry</i> (2), <i>Chem. Eur. J.</i>
2016-present	Ph.D. thesis committee member (6)
2016-present	BMBB graduate student recruitment committee
2016-present	Microbial Engineering graduate student recruitment committee
2016-2017	Co-organizer: Biocatalysis Initiative – Microbial Factories Symposium
2016	Outside reviewer for NSF Career Award
2015	AAAS Webinar: Careers for US Scientists in Europe and China

## MEMBERSHIPS

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2018-present	Society for Industrial Microbiology and Biotechnology
2018-present	University of Minnesota Biotechnology Training Grant mentor
2018-present	American Chemical Society (ACS)
2017-present	American Society for Biochemistry and Molecular Biology (ASBMB)
2017-present	Royal Society of Chemistry (RSC)
2017-present	American Society of Pharmacognosy
2016-present	Microbial and Plant Genomics Institute faculty member, University of Minnesota
2016-present	Microbial Engineering graduate program faculty member, University of Minnesota
2016-present	Minnesota Craniofacial Research Training Program (MinnCRest) mentor, University of Minnesota
2015-2016	American Association for the Advancement of Science (AAAS)