Dr. rer. nat. Matthias Elgeti

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ACADEMIC APPOINTMENTS

3/2023 Research Group Leader "Biomolecular EPR spectroscopy and signal transduction" - present Institute for Drug Discovery, University of Leipzig Medical School, Leipzig, Germany Interdisciplinary Center for Bioinformatics, University of Leipzig, Leipzig, Germany

- 7/2021 Assistant Research Professor & Principal Investigator
- 3/2023 Stein Eye Institute, University of California, Los Angeles

11/2018 Research Scientist

- 6/2021 Hubbell Laboratory, University of California, Los Angeles

EDUCATION & TRAINING

11/2013 - 11/2018	Postdoctoral Training , Hubbell Laboratory, University of California, Los Angeles Elucidation of common structure/function relationships of GPCR signaling based on the rhodopsin model system by means of EPR spectroscopy.	
11/2012 - 10/2013	Postdoctoral Training , Hofmann/Heck Laboratory, Charité – Universitätsmedizin Berlin Development of a titration assay to determine binding affinities of conformational states in equilibrium.	
11/2006 - 10/2012	Ph.D. in Biophysics , Humboldt Universität zu Berlin – <i>summa cum laude</i> Advisors: Drs. KP Hofmann/ FJ Bartl. Thesis title: " <i>Spectroscopic investigations of visual rhodopsin</i> "	
10/2005 - 9/2006	Diploma in Physics (minor in Mathematics), Freie Universität Berlin – <i>very good</i> Advisor: Dr. M Heyn. Thesis title: " <i>Investigation of the effect of UV light on the conformation of light-</i> <i>activated rhodopsin.</i> "	
2002	Pre-diploma in Physics (minor in Chemistry), Georg August Universität Göttingen	
GRANTS, FELLOWSHIPS & AWARDS		
1/2024 - 12/2027	SFB 1423, Collaborative Research Center Principal Investigator, Total €560.000 (€140.000/year) German Research Foundation <i>"The role of conformational dynamics for Y receptor activation"</i>	
10/2023 - 6/2027	Hyp*Mol, Transregional Collaborative Research Center Principal Investigator, Total €265.000 (€70.000/year) German Research Foundation <i>"Flavomaquettes with observer spin label for hyperpolarized EPR spectroscopy"</i>	

4/2021 **R01 GM137081**, Principal Investigator

- -3/2023 Total \$1,599,000 (\$319.800/year), accepted for 2 years National Institute of General Medical Sciences (NIGMS) *"Exploring the Conformational Landscape of G protein Coupled Receptors"*
- 2018 **Travel Award**, 18th International Conference on Retinal Proteins Toronto, Canada

2014	Postdoctoral Research Fellowship , German Research Foundation
- 2015	"Developing a common framework of structure/function relationships for G protein coupled receptors using site-directed spin labeling and EPR-spectroscopy".
2012	Poster Prize , 15 th International Conference on Retinal Proteins Monte Verita – Ascona, Switzerland
2006	Student Research Fellowship , Charité - Universitätsmedizin Berlin <i>"Time-resolved FTIR spectroscopy of rhodopsin deactivation"</i>

SELECTED PUBLICATIONS (*co-first author, [‡]corresponding author)

Belyaeva J, **Elgeti M**^{\ddagger}. (2024) Exploring protein structural ensembles: Integration of sparse experimental data from electron paramagnetic resonance spectroscopy with molecular modeling methods. *eLife*, accepted

Zhao J, **Elgeti M**^{*‡}, O'Brien ES, Sár CP, EI Daibani A, Heng J, Sun X, White E, Che T, Hubbell WL, Kobilka BK, Chen C. (2024) Ligand Efficacy Modulates Conformational Dynamics of the μ -Opioid Receptor. *Nature*, 1–7.

Khan F, **Elgeti M**^{*‡}; Grandfield S, Paz A, Marcoline FV, Wright EM, Grabe M, Hubbell WL, Abramson J (2023) Membrane Potential Increases Outward-facing Accessibility and Transport Rate of the Sodium-Glucose Transporter. *Nature Communications*, 14 (1), 7511.

Chen M, Kálai T, Cascio D, Bridges MD, Whitelegge JP, **Elgeti M**, Hubbell WL (2023) A Highly Ordered Nitroxide Side Chain for Distance Mapping and Monitoring Slow Structural Fluctuations in Proteins. *Applied Magnetic Resonance*.

Elgeti, M[‡] and Hubbell, WL. (2021) DEER Analysis of GPCR Conformational Heterogeneity. *Biomolecules*, 11:778.

Lerch, MT, Matt RA, Masureel M, **Elgeti M**, Kumar KK, Hilger D, Foys B, Kobilka BK, Hubbell WL (2020) Viewing Rare Conformations of the β_2 -Adrenergic Receptor with Pressure-Resolved DEER Spectroscopy. *Proceedings of the National Academy of Sciences of the U.S.A.* 117:31824–31.

McMahon C, Staus DP, Wingler LM, Wang J, Skiba MA, **Elgeti M**, Hubbell WL, Rockman HA, Kruse AC, Lefkowitz RJ (2020). Synthetic Nanobodies as Angiotensin Receptor Blockers. *Proceedings of the National Academy of Sciences of the U.S.A.* 117:20284–91.

Wingler LM, **Elgeti**, M*, Hilger, D, Latorraca, NR, Lerch, MT, Staus, DP, Dror, RO, Kobilka, BK, Hubbell, WL, Lefkowitz, RJ (2019). Angiotensin Analogs with Divergent Bias Stabilize Distinct Receptor Conformations. *Cell* 176:468-78.

Elgeti, M[‡], Kazmin, R, Rose, AS, Szczepek, M, Hildebrand, PW, Bartl, FJ, Scheerer, P, Hofmann, KP (2018). The Arrestin-1 Finger Loop Interacts with Two Distinct Conformations of Active Rhodopsin. *The Journal of Biological Chemistry* 293:4403–10.

Kazmin R, Rose, AS, Szczepek, M, **Elgeti, M**, Ritter, E, Piechnick, R, Hofmann, KP, Scheerer, P, Hildebrand, PW, Bartl, FJ (2015). The Activation Pathway of Human Rhodopsin in Comparison to Bovine Rhodopsin. *The Journal of Biological Chemistry* 290:20117–27.

Sommer ME, **Elgeti**, **M**, Hildebrand, PW, Szczepek, M, Hofmann, KP, Scheerer, P (2015). Structure-Based Biophysical Analysis of the Interaction of Rhodopsin with G protein and Arrestin. *Methods in Enzymology* 556:563–608.

Szczepek, M. Beyrière, F, Hofmann, KP, **Elgeti, M**, Kazmin, R, Rose, AS, Bartl, FJ, von Stetten, D, Heck, M, Sommer, ME, Hildebrand, PW, Scheerer, P (2014). Crystal Structure of a Common GPCR-Binding Interface for G protein and Arrestin. *Nature Communications* 5:4801.

Rose AS, **Elgeti, M**, Zachariae, U, Grubmüller, H, Hofmann, KP, Scheerer, P, Hildebrand, PW (2014). Position of Transmembrane Helix 6 Determines Receptor G protein Coupling Specificity. *Journal of the American Chemical Society* 136:11244–7.

Elgeti, **M**[‡], Rose, AS, Bartl, FJ, Hildebrand, PW, Hofmann, KP, Heck, M (2013). Precision vs Flexibility in GPCR Signaling. *Journal of American Chemical Society* 135:12305–12.

Elgeti, M[‡], Kazmin, R, Heck, M, Morizumi, T, Ritter, E, Scheerer, P, Ernst, OP, Siebert, F, Hofmann, KP, Bartl, FJ (2011). Conserved Tyr223(5.58) Plays Different Roles in the Activation and G-protein Interaction of Rhodopsin. *Journal of the American Chemical Society* 133:7159–65.

Ritter E, Elgeti, M, Bartl FJ (2008). Activity Switches of Rhodopsin. Photochemistry Photobiology 84:911–20.

Elgeti, M, Ritter E, Bartl FJ (2008). New Insights into Light-Induced Deactivation of Active Rhodopsin by SVD and Global Analysis of Time-Resolved UV/Vis- and FTIR-Data. *Zeitschrift für Physikalische Chemie* 222:1117–29.

Ritter E, **Elgeti**, **M***, Hofmann KP, Bartl FJ (2007). Deactivation and Proton Transfer in Light-Induced Metarhodopsin II/Metarhodopsin III Conversion: A Time-Resolved Fourier Transform Infrared Spectroscopic Study. *The Journal of Biological Chemistry* 282:10720–30.

MANUSCRIPTS SUBMITTED OR IN PREPARATION

Helabad MB, Belyaeva J, Elgeti M. Targeting Hidden Conformational States of the Type 1 Angiotensin II Receptor. In preparation

Elgeti M, Wingler LM, Lefkowitz RJ, Hubbell WL. Segmental Coupling in G protein Coupled Receptors Revealed by Pressure-Resolved DEER. *In preparation*

Elgeti M. The Affinity Framework of Molecular Efficacy. In preparation

INVITED TALKS (selection)

9/2024	The affinity framework of molecular efficacy. Annual Meeting of the German Biophysical Society Leipzig, Germany
9/2024	Studying membrane protein conformational dynamics using EPR spectroscopy and integrative modeling. 45th FGMR Annual Meeting and Joint Conference of the German, Danish, Polish, and Swedish Magnetic Resonance Communities University of Rostock, Germany
11/2022	Exploring the conformational landscapes of G protein coupled receptors with EPR spectroscopy. 50 th Middle German Magnetic Resonance Meeting Leipzig University, Germany
11/2020	GPCR conformational dynamics investigated by double electron-electron resonance (DEER). Institute for Drug Discovery (virtual) Leipzig University
10/2019	Structural underpinnings of biased agonism in G protein coupled receptors. Symposium of the Collaborative Research Center 1078 Free University Berlin, Germany
9/2018	A structure/function framework of GPCR activation based on the rhodopsin model. 18 th International Conference on Retinal Proteins Hockley Valley Resort, Toronto, Canada
1/2018	Protein flexibility and its important role in signal transduction. Basic Sciences Seminars University of California, Los Angeles, United States

7/2015	Precision vs. flexibility in GPCR signaling: A case study of visual rhodopsin. "Unifying Concepts in Catalysis" Colloquium Technical University Berlin, Germany
10/2012	GPCR signaling investigated by FTIR spectroscopy. 15 th International Conference on Retinal Proteins Monte Verita – Ascona, Switzerland
10/2010	Signal transfer from rhodopsin to its G protein: Insights from spectroscopic and structural studies. 14 th International Conference on Retinal Proteins University of California, Santa Cruz, United States
6/2010	<i>The different roles of Tyr223^{5.58} for the activation and G protein interaction of bovine rhodopsin.</i> Symposium of the Collaborative Research Center 498 Free University Berlin, Germany

TEACHING & MENTORING

Summer 2024	Coordinator and lecturer "From the active substance to the medicinal product" (11-BCH-0815) M.Sc. Biochemistry, Leipzig University Medical School
Winter 2023/24	Coordinator and lecturer for laboratory course "Inorganic chemistry BIO-103", B.Sc. Biology, University of Leipzig
Summer 2023	Coordinator and lecturer for laboratory course "Fluoreszenz-Spektroskopie" M.Sc. Pharmacy, Leipzig University Medical School
Summer 2023	Lecturer for Scientific Module "Integration of Experimental Data with Artificial Intelligence for the Investigation of Membrane Proteins" <u>https://research.uni-leipzig.de/vanderbilt/</u>
Summer 2023	Coordinator and lecturer "From the active substance to the medicinal product" (11-BCH-0815) M.Sc. Biochemistry, Leipzig University Medical School
Winter 2022/23	Lecture and laboratory course "EPR Spectroscopy" Modul Strukturanalytik, B.Sc. Biochemistry
Summer 2017	Supervision of Undergraduate Student Research Project (Chem99) "Conformational equilibria of a ligand-free GPCR investigated by EPR spectroscopy" University of California, Los Angeles
Summer 2017	"Entering Mentoring" Training Program University of California, Los Angeles https://bioscience.ucla.edu/ucla-entering-mentoring-training-program/
Spring 2017	Course Development and Educational Leadership Training Program University of California, Los Angeles https://www.biomedpostdoc.ucla.edu/bioscience-postdoc-educational-leadership-program/
11/2006 - 11/2012	Teaching Assistant, Biophysics (M.Sc. program) Humboldt University Berlin <i>Biophysics of signal transduction (Laboratory course)</i>
11/2006 - 05/2012	Teaching Assistant, Medical School Charité – Universitätsmedizin Berlin <i>Introduction to experimental physics (Lecture and Laboratory course)</i> Maintenance of the online learning platform Blackboard

2011 **Author/Instructor,** www.chemgapedia.de "Analysis of a protein microswitch using FTIR difference spectroscopy" Course available online at <u>www.chemgapedia.de</u> (in German)

ACADEMIC SERVICE

9/2023 Organizing Committee <u>Hyp23</u> International Conference on Hyperpolarization.

Ad hoc reviewer:

Applied Magnetic Resonance Biomolecules Biology Helvetica Chimica Acta Journal of Biological Chemistry Journal of Physical Chemistry B Marine Drugs Molecules Pharmaceuticals Proceedings in the National Academy of Sciences of the U.S.A Nature - Signal Transduction and Targeted Therapy

eLife - Early Career Reviewer in Structural Biology and Molecular Biophysics