BIOGRAPHICAL SKETCH

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NAME: Juchem, Christoph

eRA COMMONS USER NAME (credential, e.g., agency login): CJUCHEM

POSITION TITLE: Associate Professor, Departments of Biomedical Engineering and Radiology

EDUCATION/TRAINING (Begin with baccalaureate or other initial professional education, such as nursing, include postdoctoral training and residency training if applicable. Add/delete rows as necessary.)

INSTITUTION AND LOCATION	DEGREE (if applicable)	Completion Date MM/YYYY	FIELD OF STUDY
University of Bonn, Germany		1995-1997	Physics
University of Madrid, Spain		1997-1998	Physics
University of Bonn, Germany	M.Sc.	1998-2001	Physics
Max-Planck Institute for Biological Cybernetics, Tübingen, Germany	Ph.D.	2001-2006	MR Physics / Neuroscience
Yale University, New Haven, CT, USA	PostDoc	2007-2008	MR Physics / MR Spectroscopy

A. Personal Statement

I am Associate Professor in the Departments of Biomedical Engineering and Radiology at Columbia University. In my research, I develop novel magnetic resonance methods and technology to establish optimized tools for neuroscientific and clinical applications. My long-term goal as a physicist is to realize the full potential of magnetic resonance spectroscopy (MRS) as a diagnostic tool. My clinical long-term goal is to understand the role that neurochemicals play in the protection of the human central nervous system (CNS) or, alternatively, how dysfunction promotes vulnerability towards neurodegenerative and neuro-immunological diseases.

Reliable quantification of biochemicals with *in vivo* MRS relies on excellent spectral data quality, processing and quantification capability. In my research, I have optimized critical aspects of the measurement pipeline including experimental conditions (e.g. B0 shimming), MRS sequences (e.g. 'ghost' removal) and quantification (e.g. simulation of realistic basis spectra for linear combination modeling). I furthermore developed and disseminated the freeware INSPECTOR user-tool as a one-stop-shop solution for processing, quantification and quality assessment of *in vivo* MRS data (200+ downloads since 2017).

I have 20 years of experience in developing and conducting *in vivo* MR experiments at 3.0-11.7 Tesla field in humans and animal models and have demonstrated my scientific expertise in 50+ publications, book chapters and patents, as reviewer for 20+ scientific journals and as grant reviewer for 8 national research societies. I served as Co-Director of Yale University's 7T Brain MR Spectroscopy Core, Chair of the ISMRM Engineering Study group, member of the ISMRM Annual Meeting Program Committee (AMPC). I serve on the editorial board of *NMR in Biomedicine* and as Secretary of the ISMRM MR Spectroscopy Study Group, to be Chair in 2023.

- Landheer, K., Swanberg, K.M., Juchem, C. (2021). Magnetic resonance spectrum simulator (MARSS), a novel software package for fast and computationally efficient basis set simulation, *NMR Biomed.* 34:e4129. PMID: 3131877 (*Cover Image of Special Issue*)
- Landheer, K., Juchem, C. (2021). Are Cramer-Rao lower bounds an accurate estimate for standard deviations in in vivo magnetic resonance spectroscopy? NMR Biomed. 34:e4129. PMID: 33876459 (<u>Cover</u> <u>Image</u>)

- Landheer, K., Gajdosik, M, Treacy, M, Juchem, C. (2020). Concentration and effective T2 relaxation times of macromolecules at 3T, *Magn. Reson. Med.* 84:2327-2337. PMID: 32430924 (*Editor's Pick*)
- Juchem, C., Theilenberg, S, Kumaragamage, C., Mullen, M., DelaBarre, L., Adriany, G., Brown, P.B., McIntyre, S., Nixon, T.W., Garwood, M., de Graaf, R.A. (2020). Dynamic Multi-Coil Technique (DYNAMITE) MRI on human brain. *Magn Reson Med* 84:2953-2963. PMCID: PMC8168279 (*Editor's Pick*)

B. Positions and Honors

Positions and Employment

1995-1997	Studies of Physics, Rheinische Friedrich-Wilhelms University of Bonn, Germany
1997-1998	Studies of Physics, Universidad Autónoma de Madrid, Spain
1998-2001	Studies of Physics, Rheinische Friedrich-Wilhelms University of Bonn, Germany
2000-2001	Night operator at the particle accelerator of the University of Bonn, Bonn Isochron-Cyclotron
2002-2006	Doctoral Studies, Max-Planck Institute for Biological Cybernetics and
	Eberhard-Karls University of Tübingen, Tübingen, Germany
2007-2008	Postdoctoral Associate, Yale MR Research Center, Dept. Diagnostic Radiology
2008-2011	Associate Research Scientist, Yale MR Research Center, Dept. Diagnostic Radiology
2010-2016	System manager human 7 Tesla MR scanner, Yale MRRC, Dept. Diagnostic Radiology
2010-2016	Co-director of the 7 Tesla brain MR spectroscopy core at Yale MRRC, Dept. Diagn. Radiology
2012-2016	Assistant Professor, Departments of Radiology & Neurology, Yale University
2016-Present	Associate Professor (Tenure Track), Departments of Biomedical Engineering & Radiology,
	Columbia University
2017	Visiting Professor, Comprehensive Heart Failure Center, University Hospital Würzburg, Germany
2018-Present	Adjunct Faculty, Icahn School of Medicine at Mount Sinai (ISMMS), New York
2019-Present	Affiliate Member, Zuckerman Mind Brain Behavior Institute (ZMBBI), Columbia University

Other Experience and Professional Memberships

2002-Present Member, DS-ISMRM (German Section of the ISMRM)
2004-Present Member, International Society for Magnetic Resonance in Medicine (ISMRM)
2007-Present Member, German Physical Society (DPG)
2008-Present Member, MR Engineering Study Group, ISMRM
2014-2015 Member, Workshop and Study Group Review Committee, ISMRM
2016-Present Member, Institute of Electrical and Electronics Engineers (IEEE)
2016-2019 Member, ISMRM Annual Meeting Program Committee (AMPC)

Journal Reviewer: 1) European Journal of Neuroscience, 2) Magnetic Resonance in Medicine, 3) Journal of Magnetic Resonance, 4) Nuclear Magnetic Resonance in Biomedicine, 5) Concepts in Magnetic Resonance A, 6) Biological Psychiatry, 7) Neurobiology of Aging, 8) Analytical Biochemistry, 9) Public Library of Science, 10) Magnetic Resonance Materials in Physics, Biology and Medicine, 11) IEEE Transactions on Biomedical Engineering, 12) IEEE Transactions on Applied Superconductivity, 13) Human Brain Mapping, 14) Magnetic Resonance Imaging, 15) Tomography, 16) Journal of MR Imaging, 17) Medical Physics, 18) Multiple Sclerosis Journal, 19) NeuroImage, 20) Neuropsychopharmacology, 21) Neurology

Editorial Board: 1) NMR in Biomedicine

<u>*Grant Reviewer*</u>. 1) Dutch National Research Society (NWO), 2) Dutch Technology Foundation (STW), 3) Research Foundation Flanders (FWO), 4) National Multiple Sclerosis Society (NMSS) / USA, 5) Multiple Sclerosis Society / United Kingdom, 6) Israel Science Foundation (ISF), 7) German Science Foundation (DFG), 8) National Institutes of Health (NIH, *Ad hoc*: EITN, SBIB-T, ITD, IGIS)

<u>Honors</u>

1997-1998	Erasmus Exchange Fellowship of the European Union
2004-06	3 ISMRM educational stipends, Annual Meetings: Kyoto/Japan, Miami Beach/FL, Seattle/WA

2006	Doctoral stipend of the Max-Planck Society
2007-2008	J.H. Brown & A. Brown Coxe postdoctoral fellowship in the medical sciences, Yale University
2010	Best scientific poster in engineering, ISMRM Annual Meeting, Stockholm, Sweden
2011	Stipend of the German Scholars Organization (GSO)
2011	I.I. Rabi Young Investigator Award finalist, ISMRM Annual Meeting, Montreal, Canada
2012-2013	Secretary, MR Engineering Study Group, ISMRM
2013	Clinical and Translational Science Award (CTSA), Yale Center for Clinical Investigation (YCCI)
2013-2014	Vice-Chair, MR Engineering Study Group, ISMRM
2014-2015	Chair, MR Engineering Study Group, ISMRM
2014-2018	Distinguished Reviewer (5x), Magnetic Resonance in Medicine (MRM)
2021-2022	Secretary, MR Spectroscopy Study Group, ISMRM

C. Contribution to Science

1. Optimized experimental conditions and methods for clinical MR spectroscopy.

The best experimental conditions and MRS methods are a necessity for excellent spectral quality and meaningful biochemical profiling with MRS - not a choice. In my research, I developed innovative B₀ shimming techniques tailored to MRS applications in the animal and human brain. My lab was the first to show that spectra free of 'ghosting' artifacts can be predictably obtained for virtually any experiment type with numerically optimized suppression of unwanted coherence pathways without the need for experimental (trial-and-error) optimization. To the best of our knowledge, we were the first to demonstrate the benefits of the MEGA semi-LASER sequence for J-difference editing of the antioxidant glutathione (GSH) in the human brain at 7T. Taken together, the experimental and methodological work of my laboratory set the stage for high-level metabolic studies in clinical populations.

- Landheer, K., Gajdosik, M, Juchem, C. (2020). A semi-LASER, single-voxel spectroscopic sequence with a minimal echo time of 20.1 ms in the human brain at 3 T, *NMR Biomed*. 33:e4324. PMID: 32557880
- b. Juchem, C., Cudalbu, C., de Graaf, R.A., Gruetter, R., Henning, A., Hetherington, H.P., Boer, V.O. (2020). B0 shimming for in vivo magnetic resonance spectroscopy: Experts' consensus recommendations. *NMR Biomed.* 34:e4350. PMID: 32596978
- Landheer, K., Juchem, C. (2019). Dephasing optimization through coherence order pathway selection (DOTCOPS) for improved crusher schemes in MR spectroscopy, *Magn. Reson. Med.* 81:2209-2222.
 PMID: 30390346
- d. Prinsen, H., de Graaf, R.A., Mason, G.F., Pelletier, D., Juchem, C. (2017). Towards Pathoneurochemical profiling of multiple sclerosis: Single-session measurement of glutathione, GABA and glutamate with MR spectroscopy at 7 Tesla, *J Magn. Reson. Imaging* 45:187-198. PMCID: PMC5167659
- e. **Juchem, C.**, Logothetis, N.K., Pfeuffer, J. (2007). 1H-MRS of the macaque monkey primary visual cortex at 7T: Strategies and pitfalls of shimming at the brain surface. *Magn Reson Imag.* 26:902-12. PMDI: 17467220

2. Dynamic Shimming with Spherical Harmonic Shapes

Inhomogeneous B_0 magnetic fields cause spatial distortions and signal dropout in MRI. To date, limited B_0 homogeneity is a primary reason why the expected gains of high and ultra-high field MRI have not been fully kept. Researchers at Yale and Stanford demonstrated in the mid 1990's the gains in B_0 homogeneity that can be achieved when linear correction fields are tailored to specific slices and applied in a dynamic fashion, so-called dynamic shimming (as opposed to static shimming that considers the entire region-of-interest at once). Together with my colleagues, I further pushed the limits of spherical harmonic-based B_0 shimming with the first successful implementation of zero through third order dynamic shimming with full eddy-current and offset compensation. I carried out the research under the supervision of my PI at the time Dr. Robin A. de Graaf based on amplifier technology developed by Terence W. Nixon. Even today – 10 years later – the achieved B_0 homogeneity represents the gold standard for conventional B_0 shimming in the human brain.

- a. Juchem, C., Nixon, T.W., Diduch, P., McIntyre, S., Rothman, D.L., Starewicz, P., & de Graaf, R.A. (2010). Dynamic shimming of the human brain at 7 Tesla, *Conc. Magn. Reson.* 37B:116-128. PMCID: PMC2907895
- b. Boer, V.O., Klomp, D.W.J., Juchem, C., Luijten, P.R., de Graaf, R.A. (2011) Multi-slice ¹H MRSI of the human brain at 7 Tesla using dynamic B₀ and B₁ shimming. *Magn. Reson. Med.* 68:662-670. PMCID: PMC3306521

3. Multi-Coil B_0 Magnetic Field Control for B_0 Shimming and MR Imaging

For more than half a century B_0 magnetic correction fields were generated by dedicated, electrical coils resembling the shapes of low-order spherical harmonic terms. Although this approach does not allow fully satisfactory B_0 homogeneity in the rodent or human brain, the exclusive use of these shapes has never been questioned. I took the lead in initiating a complete paradigm shift and, together with my colleagues, developed a new and generalized method that allows the synthesis of advanced magnetic field distributions with generic (i.e. non-orthogonal) basis fields from a set of localized coils. Multi-coil B_0 shimming provides dramatically better B_0 homogeneity in the mouse, rat and human brain than standard methods and in the future should close to completely eliminate B_0 inhomogeneity as a problem.

- a. Juchem, C., Theilenberg, S., Kumaragamage, C., Mullen, M., DelaBarre, L., Adriany, G., Brown, P. B., McIntyre, S., Nixon, T. W., Garwood, M., de Graaf, R.A. (2020). Dynamic multicoil technique (DYNAMITE) MRI on human brain. *Magn. Reson. Med.* 84:2953-2963. PMID: 32544274 (*Editor's Pick*)
- Rudrapatna, U., Fluerenbrock, F., Nixon, T.W., de Graaf, R.A., Juchem, C. (2018). Combined imaging and shimming with the dynamic multi-coil technique. *Magn. Reson. Med.* 81:1424-1433. PMCID: PMC4120278
- Juchem, C.*, Rudrapatna U.*, Nixon, T.W., de Graaf, R.A. (2015). Dynamic Multi-Coil Technique (DYNAMITE) Shimming for Echo-Planar Imaging of the Human Brain at 7 Tesla. *NeuroImage.* 105:462-472. PMCID: PMC4262558
- d. **Juchem, C.**, Nixon, T.W., McIntyre, S., Rothman, D.L., & de Graaf, R.A. (2010). Magnetic field modeling with a set of individual localized coils, *J. Magn. Reson.* 204:281-289. PMCID: PMC2884296

4. Dissemination of Freeware Software Packages

To date, advanced solutions for optimized experiment preparation, analysis and quantification of MR spectroscopy are limited. Such methods are at the heart of my research program and I share MR sequences, algorithms and processing software with the MR community free of charge. I expect easy access to advanced methods to further promote both research and clinical potential of MR.

- a. Landheer K, **Juchem C**. Constrained Optimized Water Suppression (COWS). CTV license: CU21111 (2021). <u>innovation.columbia.edu/technologies/CU21111_COWS</u>
- b. Gajdošik M, Landheer K, Juchem C. Pocket MRS: Mobile application for rapid analysis of magnetic resonance spectroscopy. CTV license: CU21108 (2021). innovation.columbia.edu/technologies/CU21108 PocketMRS
- c. Landheer K, **Juchem C**. Magnetic Resonance Spectrum Simulator (MARSS). CTV license: CU19205 (2019). innovation.columbia.edu/technologies/CU19215_MARSS
- Landheer K, Juchem C. Dephasing optimization through coherence order pathway selection (DOTCOPS) for improved crusher and phase cycling schemes in MR spectroscopy. CTV license CU18146 (2018). <u>innovation.columbia.edu/technologies/CU18406</u>
- e. Juchem C. INSPECTOR Magnetic resonance spectroscopy software for *in vivo* biomedical and clinical research. CTV license CU17130 (2017). innovation.columbia.edu/technologies/cu17130_INSPECTOR
- f. **Juchem, C**. B0DETOX: B₀ detoxification software for magnetic field shimming. CTV license CU17326 (2017), <u>innovation.columbia.edu/technologies/CU17326_B0DETOX</u>

D. Research Support Ongoing Research Support

R01 EB030560 (Juchem, C) NIH / NIBIB

Dynamic Multi-Coil B₀ Shimming for Diagnostic MRI of Frontal Brain

In this NIH-R01 project, we combine DYNAMITE B_0 shim with clinical RF technology to establish the first integrated multi-coil B_0 and radio-frequency (MC/RF) setup dedicated to clinical diagnostics and workflow. The research is significant because it is expected to fundamentally leverage the diagnostic potential of gradient-echo MRI in the ventral PFC and orbits, setting the stage for widespread clinical use of state-of-the-art B_0 shim technology and true translation from bench to bedside.

Role: Principal Investigator

R01 MH123142 (Kantrowitz, J, Javitt, D, Grinband, J) NIH / NIMH

Glutamatergic mechanisms of psychosis and target engagement

Several glutamate-targeted medications have failed in pivotal clinical trials despite robust effectiveness in preclinical models. The present project further develops ketamine biomarkers for glutamate-based treatments and explores glutamate and dopamine mechanisms of schizophrenia using brain imaging. Role: Investigator

Established Investigator Grant (Juchem, C)

General Electric, GE Healthcare

A Fastmap Shim Tool (FAMASITO) for User-Friendly B0 Field Homogenization

This research seeks to integrate advanced B0 magnetic field shimming methodology with GE's EPIC scanner environment for clinical single-voxel MR spectroscopy applications.

Role: Principal Investigator

DOD W81XWH1810221 (Martinez, D)

Department of Defense (DoD)

Investigation of the NMDA Antagonist Ketamine as a Treatment for Tinnitus

Tinnitus is a very common problem that often accompanies hearing loss. Research in humans and animals suggest that the neurotransmitters glutamate and GABA are important in the development and maintenance of tinnitus. In this project, we will test the effect of ketamine on tinnitus using a placebo controlled design. Magnetic Resonance Spectroscopy (MRS) scans will be obtained at the time of placebo/ketamine infusions to investigate the effect on GABA and glutamate levels.

Role: Investigator

Collaboratory Award (Juchem, C, Kegeles, L) Columbia University

In Vivo Magnetic Resonance Spectroscopy - from Data to Clinical Benefit

In this project, we will establish an educational course that comprises all aspects of *in vivo* MRS from theory to experiment, from data acquisition to the derivation of metabolic signatures, and from study design to clinical interpretation. The course will provide an understanding of MRS techniques and bridge the gap between theoretical concepts, hands-on training and direct experimental experience on a human 3T MR scanner. Role: Co-PI

U01 EB025153 (Garwood, M, University of Minnesota) NIH / NIBIB

Imaging Human Brain Function with Minimal Mobility Restrictions

The goal of this project is to build the first-ever human MRI scanner requiring only the head to be inside the magnet bore and having a large window for viewing outside the magnet bore. The small size, weight, and power requirements of this 1.5 Tesla MRI system will enable it to be transported and sited almost anywhere in the world and to bring the magnet to the subject rather than the other way around. Role: Co-PI of Columbia subcontract

04/07/2021 - 12/31/2024

12/01/2020 - 11/30/2025

12/01/2017 - 11/30/2021

01/01/2019 - 12/31/2021

09/01/2017 - 09/30/2022

09/01/2017 - 08/31/2022

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