



# Editorial: *In Vivo* Magnetic Resonance at Ultra High Field

He Zhu<sup>1</sup> and Ewald Moser<sup>2, 3\*</sup>

<sup>1</sup> Vanderbilt University Institute of Imaging Science, Nashville, TN, United States, <sup>2</sup> Center for Medical Physics and Biomedical Engineering, Medical University of Vienna, Vienna, Austria, <sup>3</sup> MR Center of Excellence, Medical University of Vienna, Vienna, Austria

Keywords: MRI, MRS, 1H, 13C, UHF magnets, heart, brain

### **Editorial on the Research Topic**

### In Vivo Magnetic Resonance at Ultra High Field

This Special Research Topic includes a representative collection of the topics outlined in the overview. With the editorial, we aim to set the stage for these excellent articles. We also emphasize our view that biomedical imaging continues to be the frontier of science and engineering today. The Brain Research through Advancing Innovative Neurotechnologies (BRAIN) initiative in the US [1], for example, is regarded as the highest priority of the nation. It is often referred to as the "Apollo project of the brain" and the basic approach is best described as "reverse engineering the brain" [2]. Similarly, the EU has launched its Human Brain Project, a H2020 FET flagship project "which strives to accelerate the fields of neuroscience, computing and brain-related medicine". In both efforts, neuroimaging serves as a natural foundation because seeing it is the first step of figuring out how it works. This principal applies to the entire body and differs in the specific challenges of an imaging method. It can be low sensitivity of molecular imaging, motion in cardiac imaging or inhomogeneous magnetic fields at ultra high fields and frequencies. In general, discoveries in biomedical sciences can be expected when the technology of the tools moves forward.

A frugal collection of articles cover a particularly broad range of topics. In a review focused on the history of NMR magnet technology Moser et al. addressed the driving issue of this topic that is the motivation for higher and higher magnetic fields. This review summarizes the magnet development since its early commercial availability in the 1950s in analytical NMR, up to now at 7 T or above for clinical (human) and preclinical (animal) purposes. To establish the background, the article first explained the behaviors of signal and noise versus field strengths [3]. The development in the following 60 plus years was chronicled in different stages (resistive, permanent, and superconducting magnets) to what is available today. Here we should note that Dr. Lauterbur actually never received a patent for his idea to employ NMR for imaging (which he dubbed "zeugmatography"), as in 1971 his university did not believe this was worthwhile. Concerning the current state of clinical UHF MRI, Siemens obtained CE-labeling for its 7 T Terra system in August of 2017, whereas FDA approval is still pending (M. Blasche, personal communication). The second review in this collection is focused on RF decoupling methods of <sup>13</sup>C spectroscopy (Li et al.). In contrast to the broad scope of the first one, this review goes deep into details on a small topic in NMR [4]. However, we expect to see this topic grow larger than it is because <sup>13</sup>C spectroscopy allows detection of many important organic compounds. Since the 90's, a few groups have performed <sup>13</sup>C experiments from 2.1 T to 14.1 T. One example of these efforts is to distinguish glutamate (Glu) and glutamine (Gln)in the brain [5, 6]. As Glu-Gln cycling accounts for up to 80% of the energy consumption, it is essential in understanding the brain in the clinic or in the lab [7].

The first research article by Miller et al. describes a series of unique experiments of <sup>13</sup>C spectroscopy on glucose and glycogen. The goal was to investigate the modulation effect of glucagon on glycogen and glucose in the liver [8]. <sup>13</sup>C spectroscopy was first performed on a preclinical platform at 11.7 T on perfused mouse liver. Then the same experiments were

# **OPEN ACCESS**

# Edited and reviewed by:

Alex Hansen, Norwegian University of Science and Technology, Norway

### \*Correspondence:

Ewald Moser ewald.moser@meduniwien.ac.at

### Specialty section:

This article was submitted to Biomedical Physics, a section of the journal Frontiers in Physics

Received: 24 August 2017 Accepted: 19 September 2017 Published: 28 September 2017

### Citation:

Zhu H and Moser E (2017) Editorial: In Vivo Magnetic Resonance at Ultra High Field. Front. Phys. 5:45. doi: 10.3389/fphy.2017.00045

1

<sup>1</sup>www.humanbrainproject.eu/en/

Zhu and Moser In Vivo MR at UHF

performed on non-human primates (NHP) at 7 T. Similar dynamics of glucose and glycogen concentrations were measured ex vivo and in vivo as proof of concept as well as validation. <sup>1</sup>H spectroscopy is perhaps the hallmark application at UHF as it can be included with regular although appropriately adapted MRI protocols without special hardware. An article by Li et al. [9] demonstrates this advantage in MRI guided single voxel MRS with GABA editing in the brain [10]. They applied the special editing scheme to suppress macromolecules, i.e., acquiring GABA instead of GABA+ [11]. More importantly, the placements of the voxels were automatically generated using MRI images with segmentation and registration. This procedure addresses a long-standing issue of single voxel MRS that is the inconsistency of human interpretation of brain structures and manually placing voxels. Their algorithm achieved spatial variations of approximately 1 mm or less.

We are delighted to include an article on cardiac MRI because of its difficulty [12] and worthy alternative in CT [13]. Huelnhagen et al. present a comprehensive report on the potential and challenge of myocardial  $T_2^*$  mapping using 7 T MRI. All aspects are discussed from signal mechanism, hardware requirement, available sequences to post processing and clinical merit. What is particularly valuable is the theoretical and experimental analysis of the dependence of  $T_2^*$  on spatial resolution and  $B_0$  inhomogeneity. This framework will benefit all gradient echo based sequences on 7 T as they are the workhorse due to RF restrictions. Last but not least, Orzada et al. present an analysis of an integrated 8-channel Tx/Rx body array as a body coil on 7 T MRI [14]. Due to the high frequency, and thus decreasing wavelength in tissue, whole-body Tx/Rx

# **REFERENCES**

- Weiss PS. President Obama announces the BRAIN Initiative. ACS Nano (2013) 7:2873-4. doi: 10.1021/nn401796f
- Cepelewicz J. The U.S. Government Launches a \$100-Million "Apollo Project of the Brain". Sci Am. (2016). Available online at: https://www.scientificamerican.com/article/the-u-s-government-launches-a-100-million-apollo-project-of-the-brain/
- Brunner P. Ernst RR. Sensitivity and performance time in NMR imaging. J Magn Reson. (1979) 33:83–106. doi: 10.1016/0022-2364(79)90192-6
- de Graaf RA, Mason GF, Patel AB, Behar KL, Rothman DL. *In vivo* 1H-[13C]-NMR spectroscopy of cerebral metabolism. *NMR Biomed.* (2003) 16:339–57. doi: 10.1002/nbm.847
- Shen J, Petersen KF, Behar KL, Brown P, Nixon TW, Mason GF, et al. Determination of the rate of the glutamate/glutamine cycle in the human brain by in vivo 13C NMR. Proc Natl Acad Sci USA. (1999) 96:8235–40. doi: 10.1073/pnas.96.14.8235
- Sonnay S, Duarte JM, Just N, Gruetter R. Compartmentalised energy metabolism supporting glutamatergic neurotransmission in response to increased activity in the rat cerebral cortex: A 13C MRS study in vivo at 14.1 T. J Cereb Blood Flow Metab. (2016) 36:928–40. doi: 10.1177/0271678X16629482
- Hyder F, Fulbright RK, Shulman RG, Rothman DL. Glutamatergic function in the resting awake human brain is supported by uniformly high oxidative energy. J Cereb Blood Flow Metab. (2013) 33:339–47. doi: 10.1038/jcbfm.2012.207
- Roden M, Perseghin G, Petersen KF, Hwang JH, Cline GW, Gerow K, et al. The roles of insulin and glucagon in the regulation of hepatic glycogen synthesis and turnover in humans. J Clin Invest. (1996) 97:642–8. doi: 10.1172/JCI118460
- 9. Li Y, Bian W, Larson P, Crane JC, Parvathaneni P, Nagarajan, et al. Reliable and reproducible GABA measurements using automated spectral editing MRS at ultra-high field. *Front Phys.* (2017) 5.

coils remain a challenge. The integrated 8-channel array was compared to a local 8-channel array. The bigger and wider integrated body coil demonstrates the inherent advantage of producing a more homogeneous excitation profile while similar SAR may be achieved with future optimization.

A typical scientist or engineer in biomedical research would not sensibly compare our own profession to the moon landing. Now as the analogy appears in the media for the general public, we are happy to use the APOLLO project to articulate our point. There was no real significance in putting two people on the moon for a walk. Nevertheless, it is regarded as a historic achievement because the pursuit led to new science and technology that changed the world. On a smaller scale, the pursue in imaging technology will bring progress to various biomedical fields [15]. We proposed this topic to express this view and we hope this collection shows that our aspirations resonate with fellow researchers.

## **AUTHOR CONTRIBUTIONS**

All authors listed have made a substantial, direct and intellectual contribution to the work, and approved it for publication.

# NOTE

In August 21, 2017, i.e., after the Editorial was accepted, the National High Magnetic Field Laboratory, Tallahassee, Florida State University, USA, reached 41.4 T with a dc resistive magnet (M.D. Bird, private communication). This clearly surpasses the former record of 38.5 T in Hefei, China.

- Mescher M, Merkle H, Kirsch J, Garwood M, Gruetter R. Simultaneous in vivo spectral editing and water suppression. NMR Biomed (1998) 11:266–72. doi: 10.1002/(SICI)1099-1492(199810)11:6<266::AID-NBM530>3.0. CO;2-J
- Henry PG, Dautry C, Hantraye P, Bloch G. Brain GABA editing without macromolecule contamination. *Magn Reson Med.* (2001) 45:517–20. doi: 10.1002/1522-2594(200103)45:3<517::AID-MRM1068>3.0.CO;2-6
- Lee VS, Hecht EM, Taouli B, Chen Q, Prince K, Oesingmann N. Body and cardiovascular MR imaging at 3.0 T. Radiology (2007) 244:692–705. doi: 10.1148/radiol.2443060582
- Flohr TG, Schaller S, Stierstorfer K, Bruder H, Ohnesorge BM, Schoepf UJ. Multi-detector row CT systems and image-reconstruction techniques. Radiology (2005) 235:756–73. doi: 10.1148/radiol.23530 40037
- Erturk MA, Raaijmakers AJ, Adriany G, Ugurbil K, Metzger GJ.
  A 16-channel combined loop-dipole transceiver array for 7 Tesla body MRI. Magn Reson Med. (2017) 77:884–94. doi: 10.1002/mrm. 26153
- Le Bihan D, Schild T. Human brain MRI at 500 MHz, scientific perspectives and technological challenges. Supercond Sci Technol. (2017) 30. doi: 10.1088/1361-6668/30/3/033003

**Conflict of Interest Statement:** The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Copyright © 2017 Zhu and Moser. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) or licensor are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.