

SQUID Technology for Biomedical Applications

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Plan for the talk

- SQUID
- The burden of brain disorders
- MEG and MEG-MRI
- Challenges
 - Ultra-low-noise field-tolerant SQUID sensors
 - As strong pulsed prepolarization as possible (~ 150 mT in 10 ms)
 - Avoiding magnetization of the superconducting wire

SQUID Sensor

- Based on the Josephson effect, which was invented in 1962



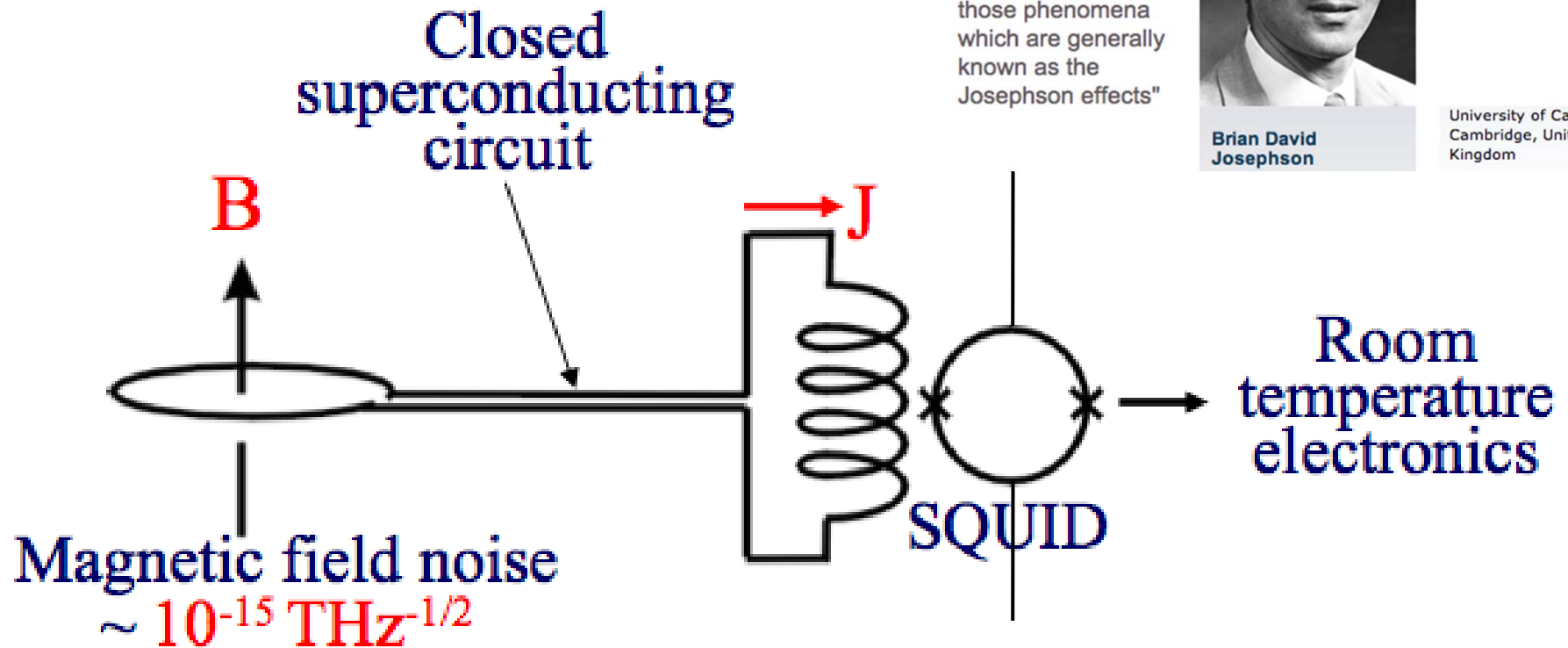
The Nobel Prize in Physics 1973

"for his theoretical predictions of the properties of a supercurrent through a tunnel barrier, in particular those phenomena which are generally known as the Josephson effects"



Brian David Josephson

University of Cambridge, United Kingdom



- The response is independent of frequency

The burden of brain diseases

1) Human suffering

- Depression: 150 million patients
- Schizophrenia: 25 million
- Dementias: 40 million
- Epilepsy: 40 million
- Stroke: 50 million

2) Cost to society: 800 billion € / year in Europe alone

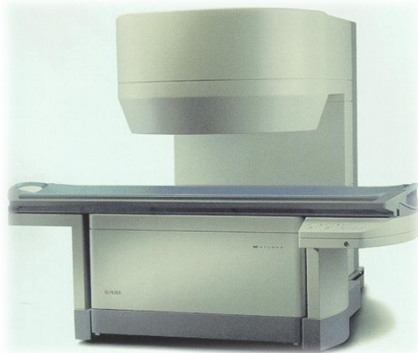
The burden is increasing with the aging population

Major Brain Research Initiatives

- **European Union: Human Brain Project**
 - to model the brain
 - 1000 M€ /10 years
- **USA: Connectome**
 - to determine the brain's wiring diagram
- **USA: BRAIN Initiative**
 - to advance innovative neurotechnology
 - 100 M\$ in 2014
- **China: BRAINNETOME**
 - to identify and understand brain networks
 - <http://www.brainnetome.org/consortium/resources.html>

Brain imaging developed in Helsinki

MRI



EEG



NIRS



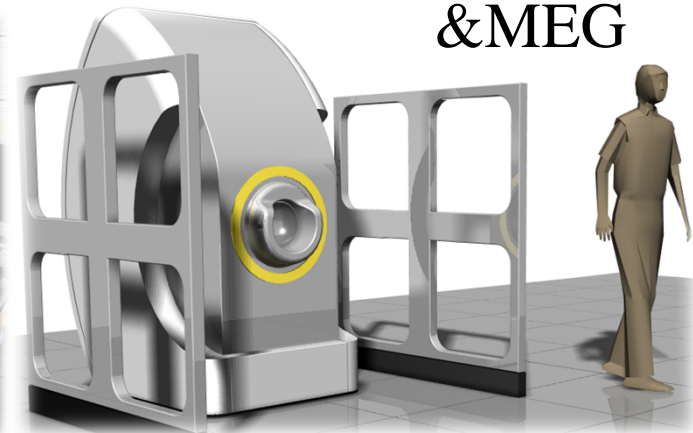
MEG



Navigated TMS



Ultra-Low-Field MRI
& MEG

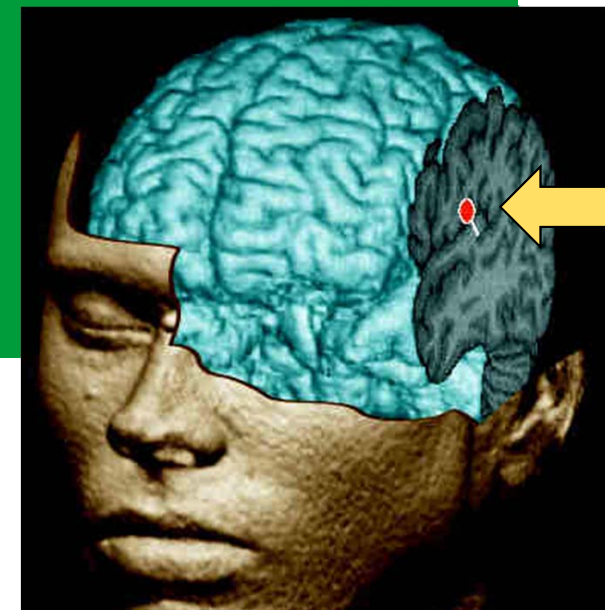
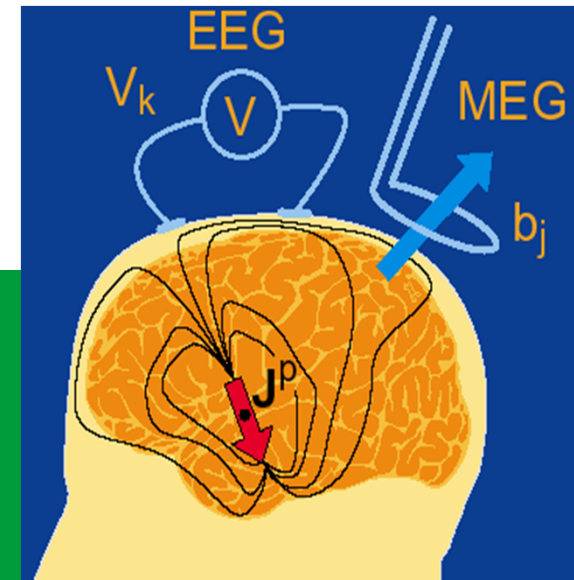


Magnetoencephalography (MEG)

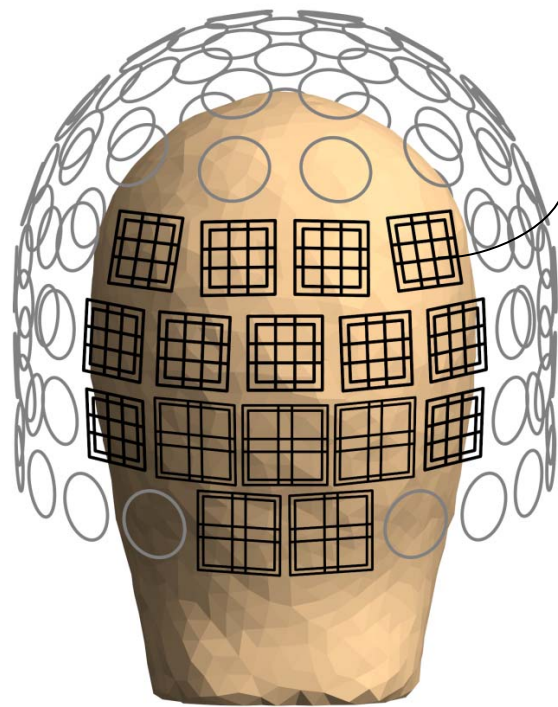
MEG is the measurement of magnetic fields produced by neuronal activity in the brain.

MEG allows one to determine locations of brain activity.

The source locations are often displayed on top of anatomical MRI images.

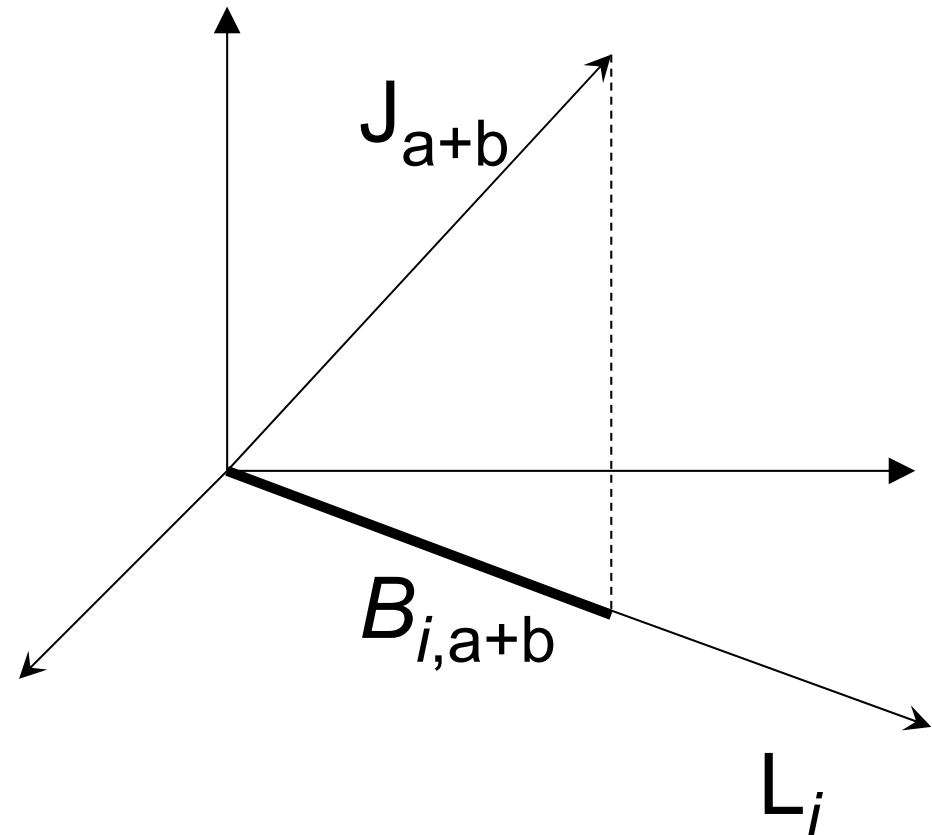


MEG: a projection of brain state



MEG sensor i

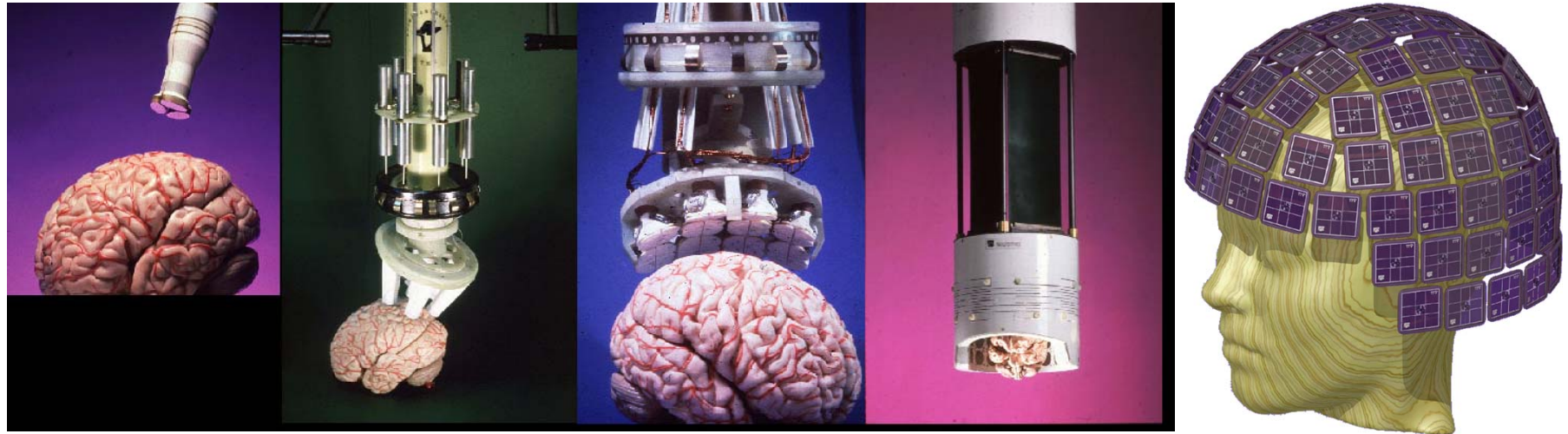
Current space



$$B_i \approx \sum a_{ik} f_k$$

Ilmoniemi 2006.

Multi-SQUID systems developed in Helsinki



	1983	1987	1989	1993	1999
Number of SQUIDs	4	7	24	122	306

Ilmoniemi et al., "A four-channel SQUID magnetometer for brain research", *Electroenceph. Clin. Neurophysiol.* 58, 467–473 (1984).

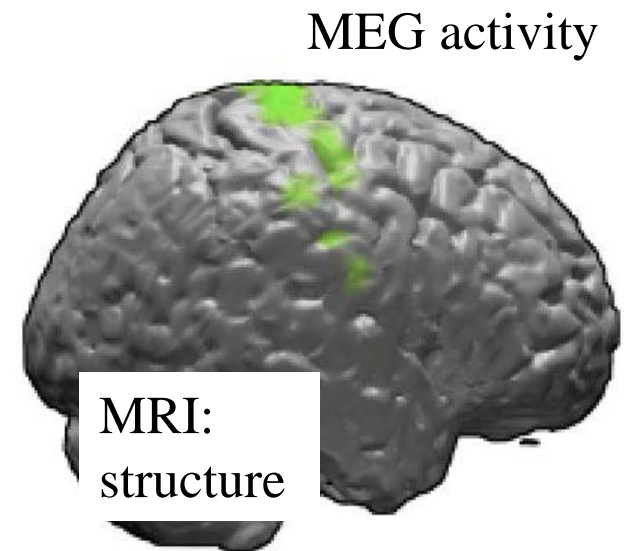
Knuutila et al., *A large-area low-noise seven-channel dc SQUID magnetometers for brain research*, *Rev. Sci. Instrum* 58, 2145–2156 (1987).

Ahlfors et al., "A 24-SQUID gradiometer for magnetoencephalography", *Physica B* 165 & 166, 97–98 (1990).

Ahonen et al., "122-channel SQUID instrument for investigating the magnetic signals from the human brain". *Phys. Scr.* T49, 198–205, (1993)

MEG needs MRI

MEG informs about function, MRI provides anatomy



MEGMRI project (FP7, 2008–12)

Goal: A **hybrid brain scanner** that can measure **MEG** and **MRI** at the same time

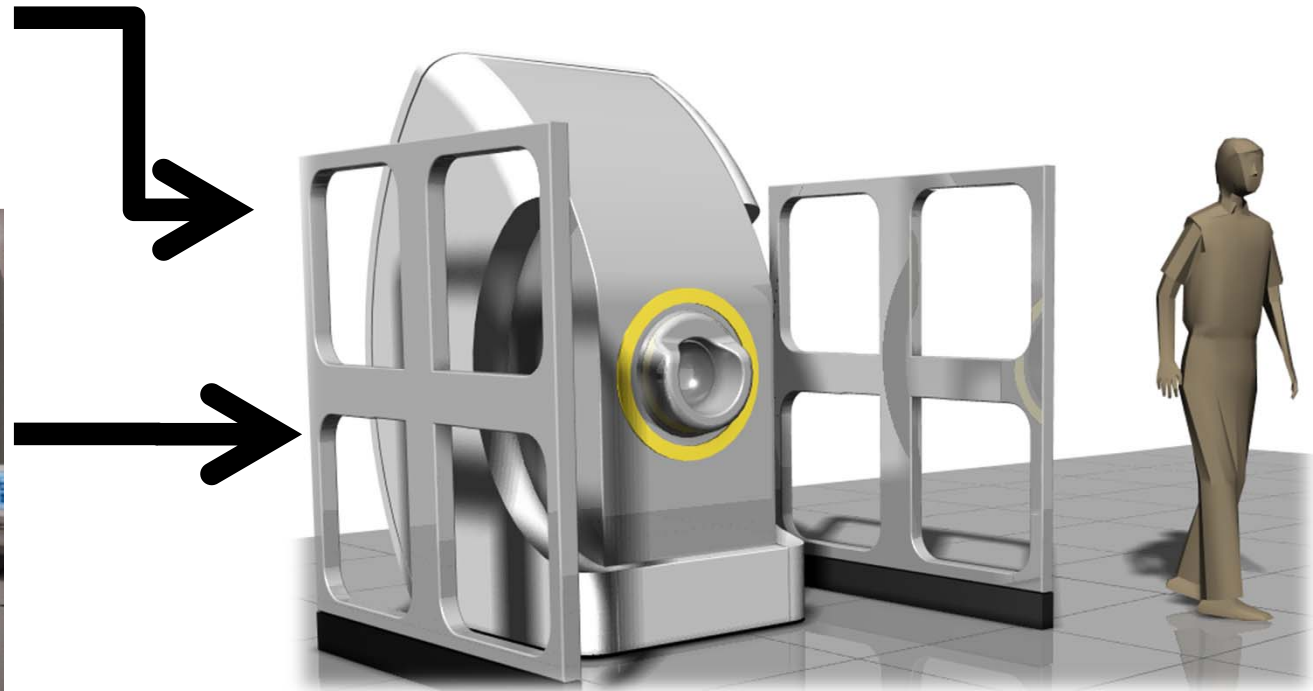
These techniques reveal information about the state of the brain



MEG

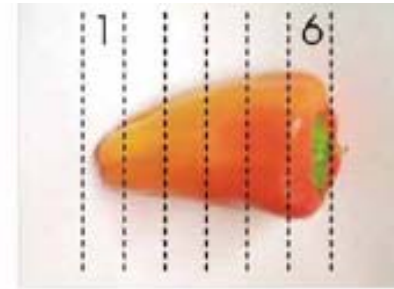


MRI



How is low-field MRI possible?

By using (as demonstrated by John Clarke et al.)

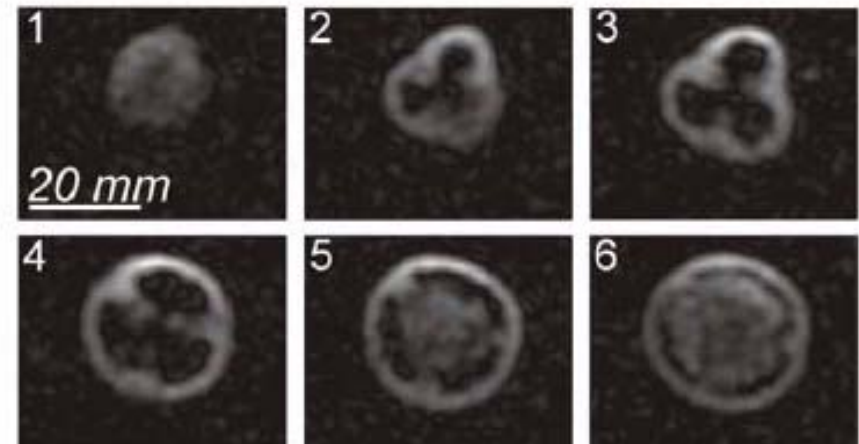


1. SQUIDS

- Response is independent of frequency

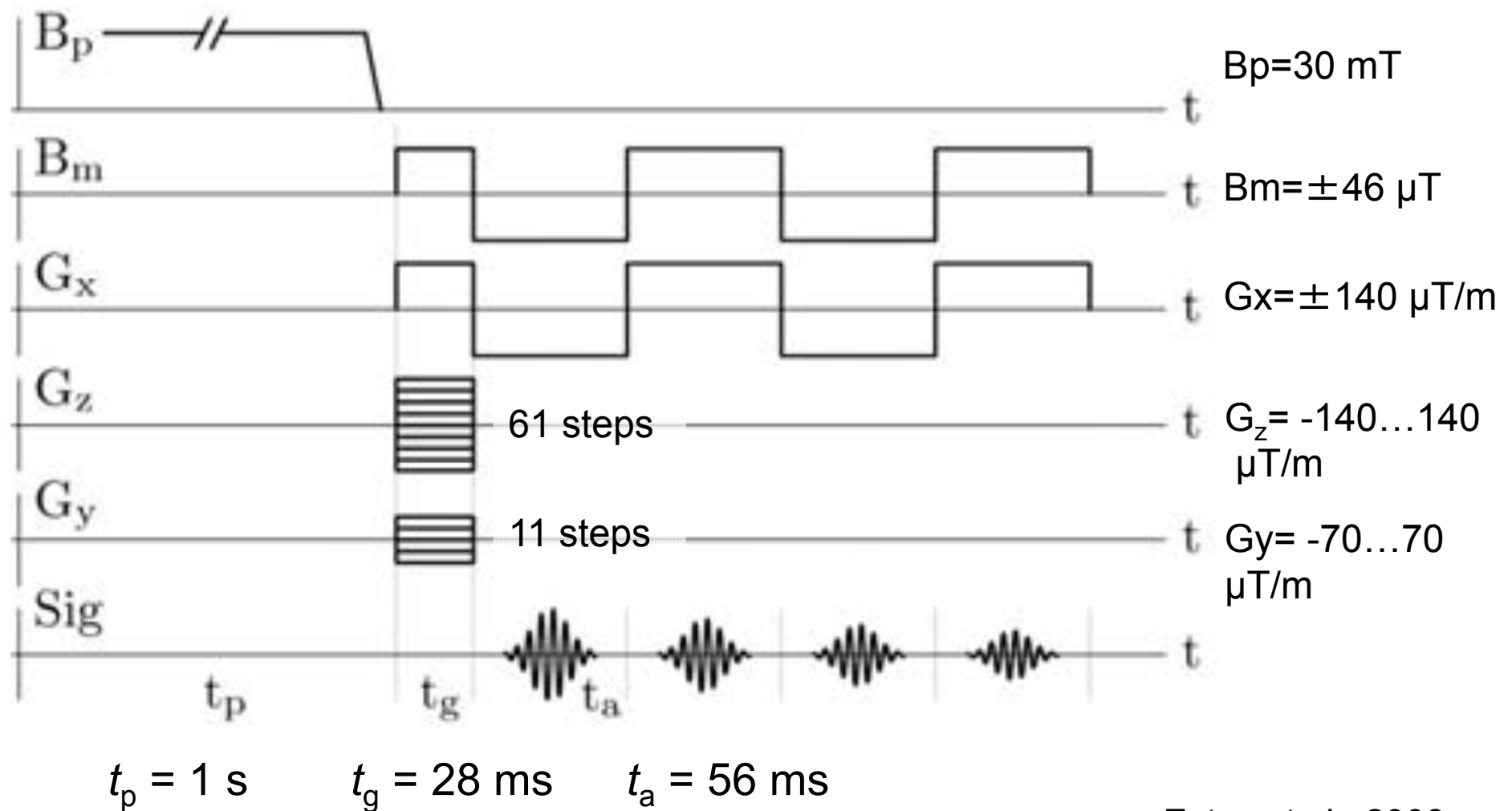
2. Prepolarization

- Polarization is independent of measurement field



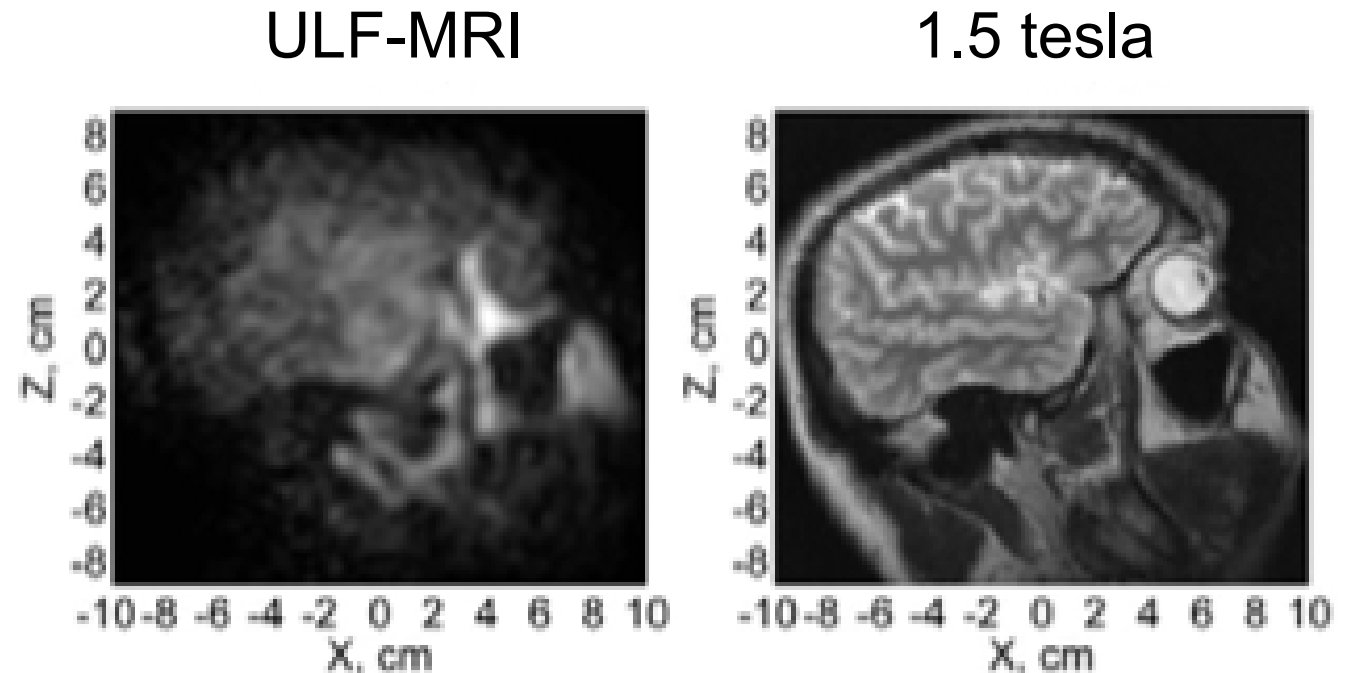
- Add second phase encoding sequence
- Polarizing field 60 mT
- Imaging field 132 μ T, gradients 150 μ T/m
- Resolution 1.2 mm x 1.2 mm

Prepolarization (& other fields)

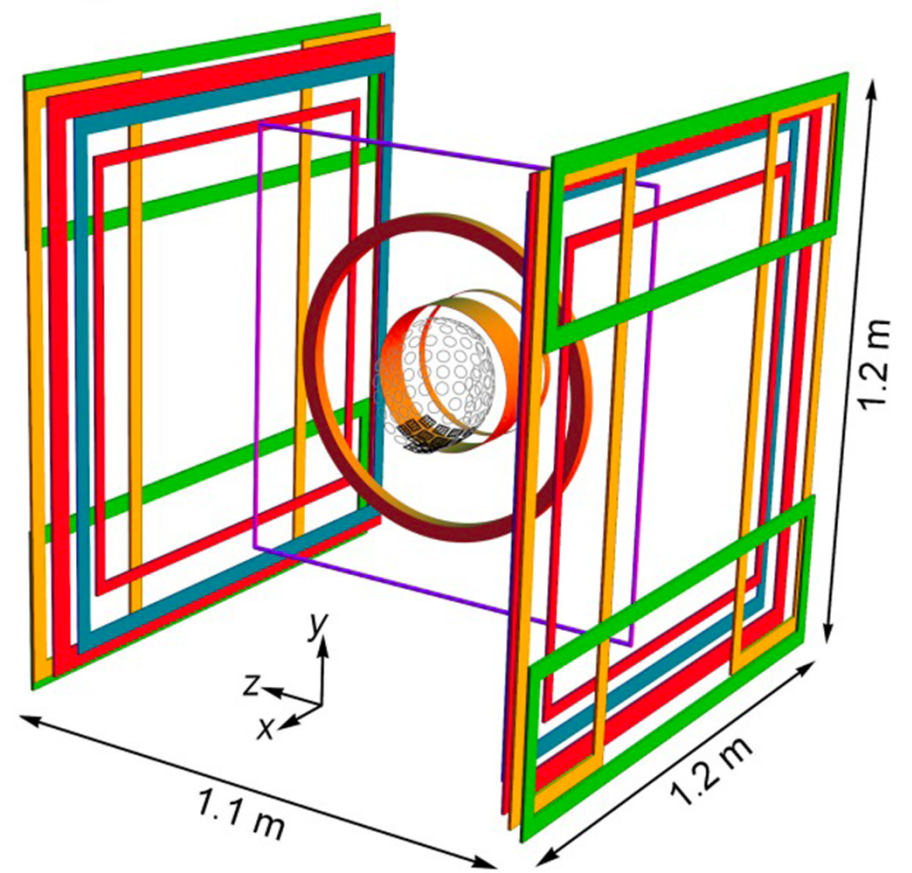
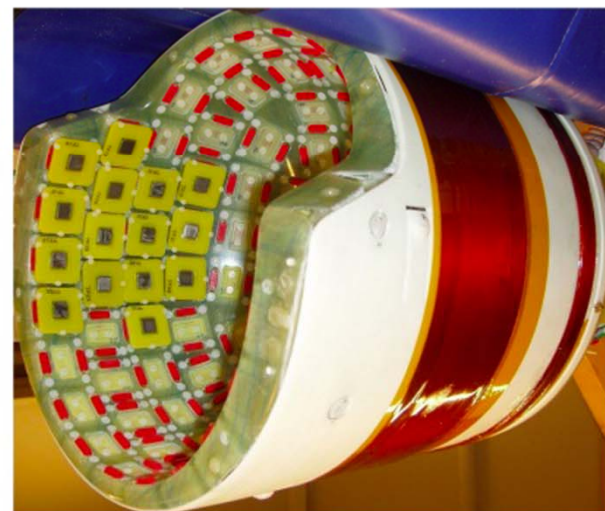


First ULF-MRI Images of the Brain

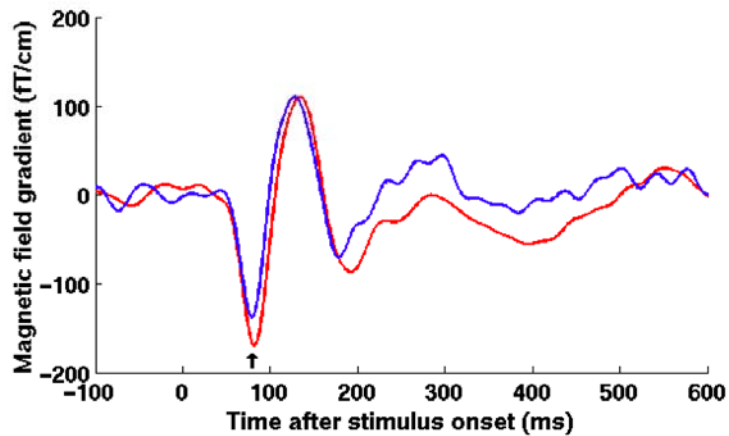
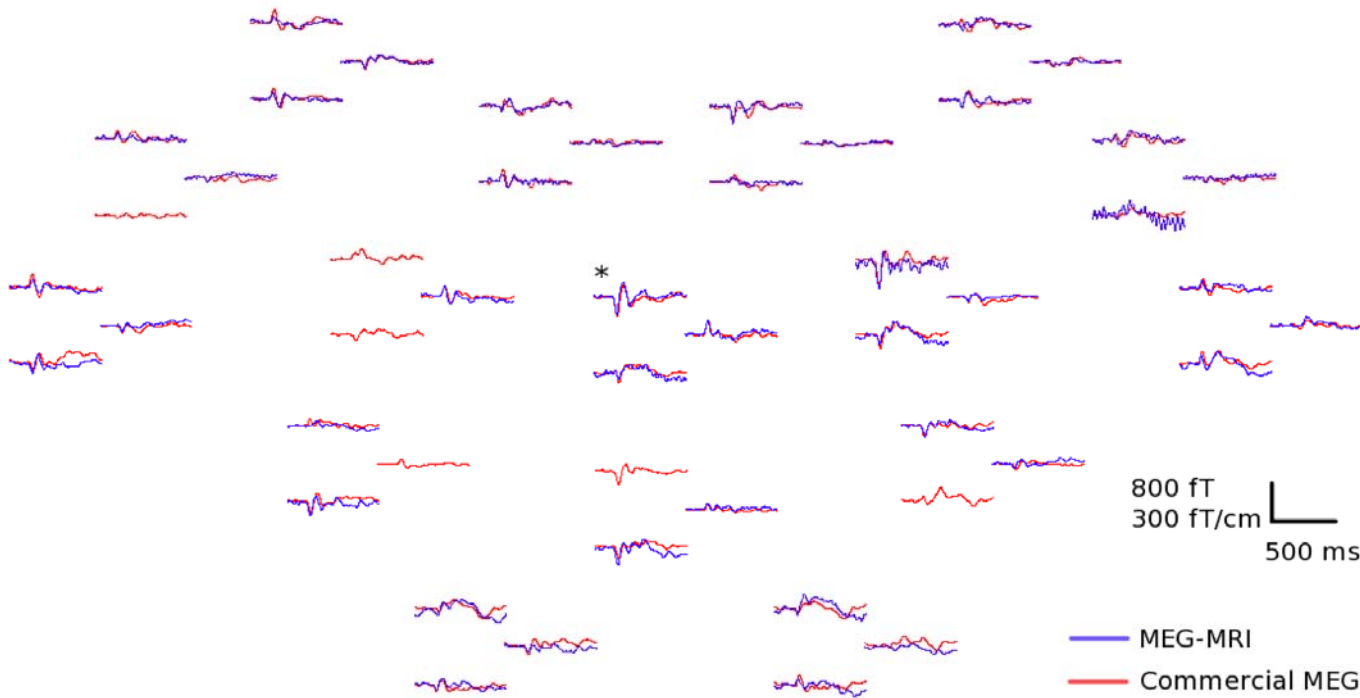
- 7 SQUIDs in parallel
- $B_p = 30 \text{ mT}$
- $B_0 = 46 \text{ } \mu\text{T}$
- 90-minute measurement



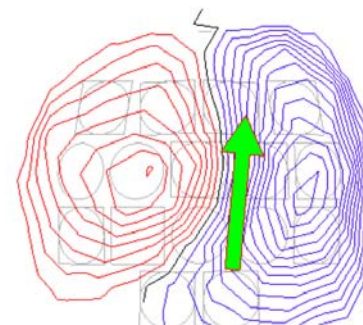
Zotev et al., IEEE/CSC & ESAS European Superconductivity News Forum, No. 4, April 2008

a**b****c**

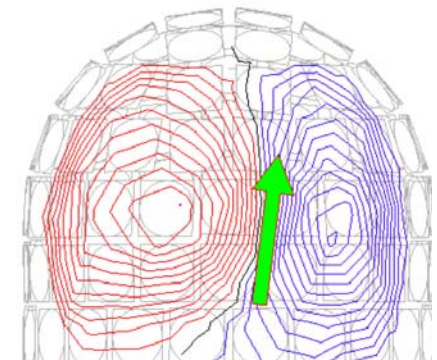
MEG signals obtained with the prototype



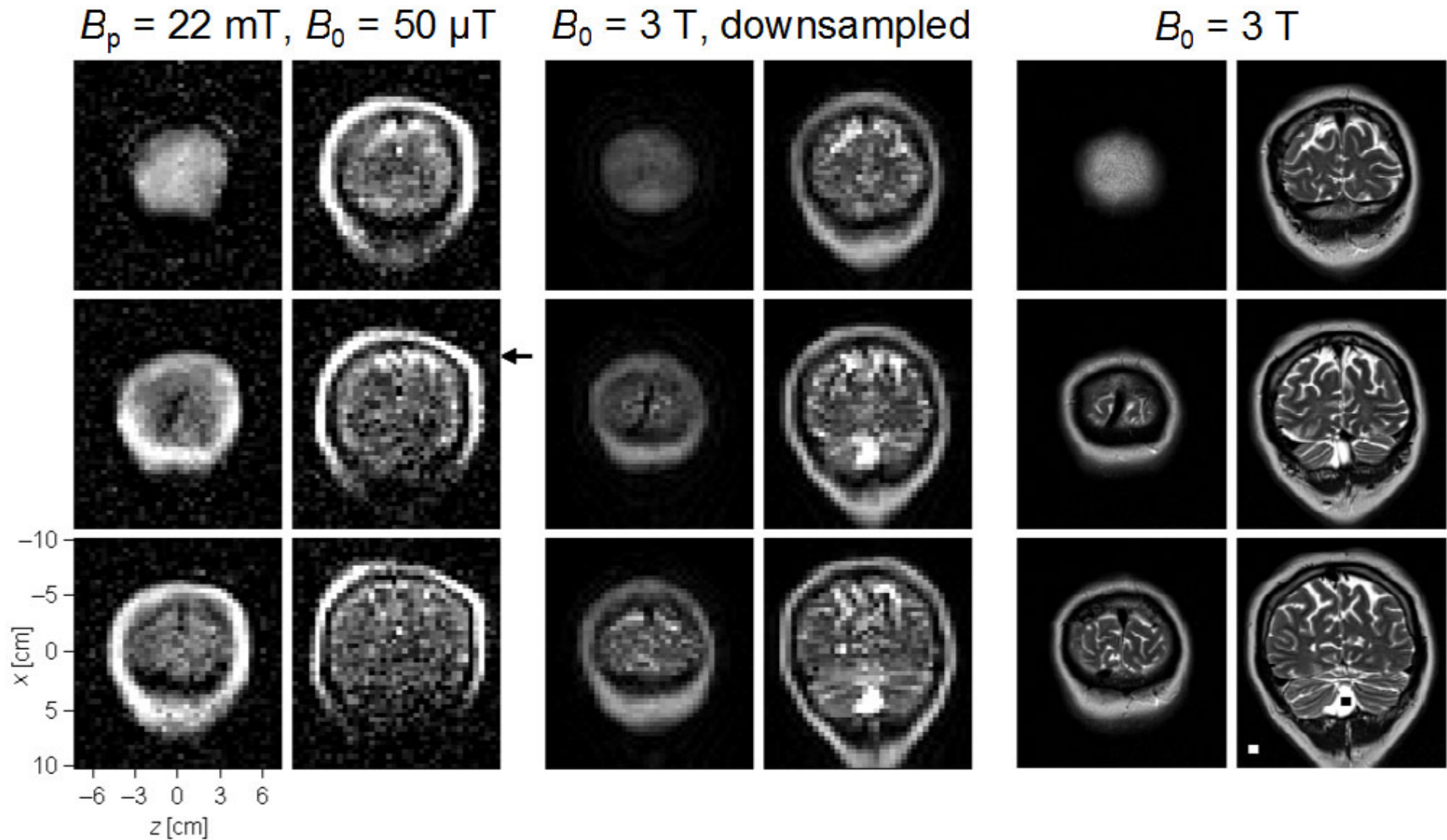
MEG-MRI



Commercial MEG



Brain Images

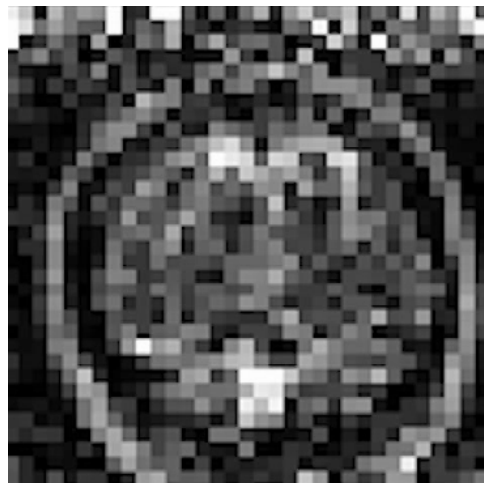


Benefits of ULF-MRI

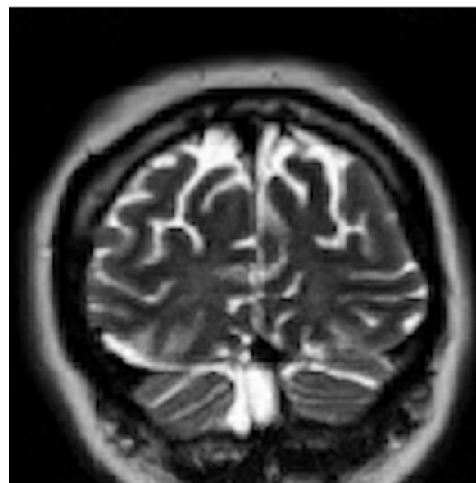
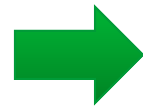
- Simultaneous MEG and MRI
 - Superb registration accuracy
 - Improved work flow
- Safe, quiet, open
 - No projectile danger, safe with pacemakers
 - Better for infants and children
- Superior T1 contrast
- Future promise
 - Superb source location accuracy

Future Prospects

- Sensor noise down by a factor of 10 (5 to 0.5 fT)
 - Prepolarization field up by a factor of 5 (22 to 110 mT)
 - Number of SQUIDs up by a factor of 6 (48 to 306)
 - More intelligent measurement sequences
- ⇒ MRI measurement time < 10 min, resolution < 2 mm
- ⇒ Clinical interest & commercial market
- + Conductivity recordings ?



Present



Next generation



Challenges

Clinical success can be expected only if the data rate is increased by more than 1000

- SQUIDs: lowest possible noise, field tolerance
- SC coils: highest possible field, no magnetization
- Dealing with fields from 1 to 10000000000000000 fT (0.1 T)

Field-Tolerant SQUIDs for ULF-MRI

All-planar SQUIDs and pickup coils for combined
MEG and MRI

J. Luomahaara¹, P. T. Vesänen², J. Penttilä³, J. O. Nieminen²,
J. Dabek², J. Simola⁴, M. Kiviranta¹, L. Grönberg¹, C. J.
Zevenhoven², R. J. Ilmoniemi² and J. Hassel¹.

Supercond. Sci. Technol. 24 075020 (2011)

VTT Technical Research Centre of Finland

Fig.1

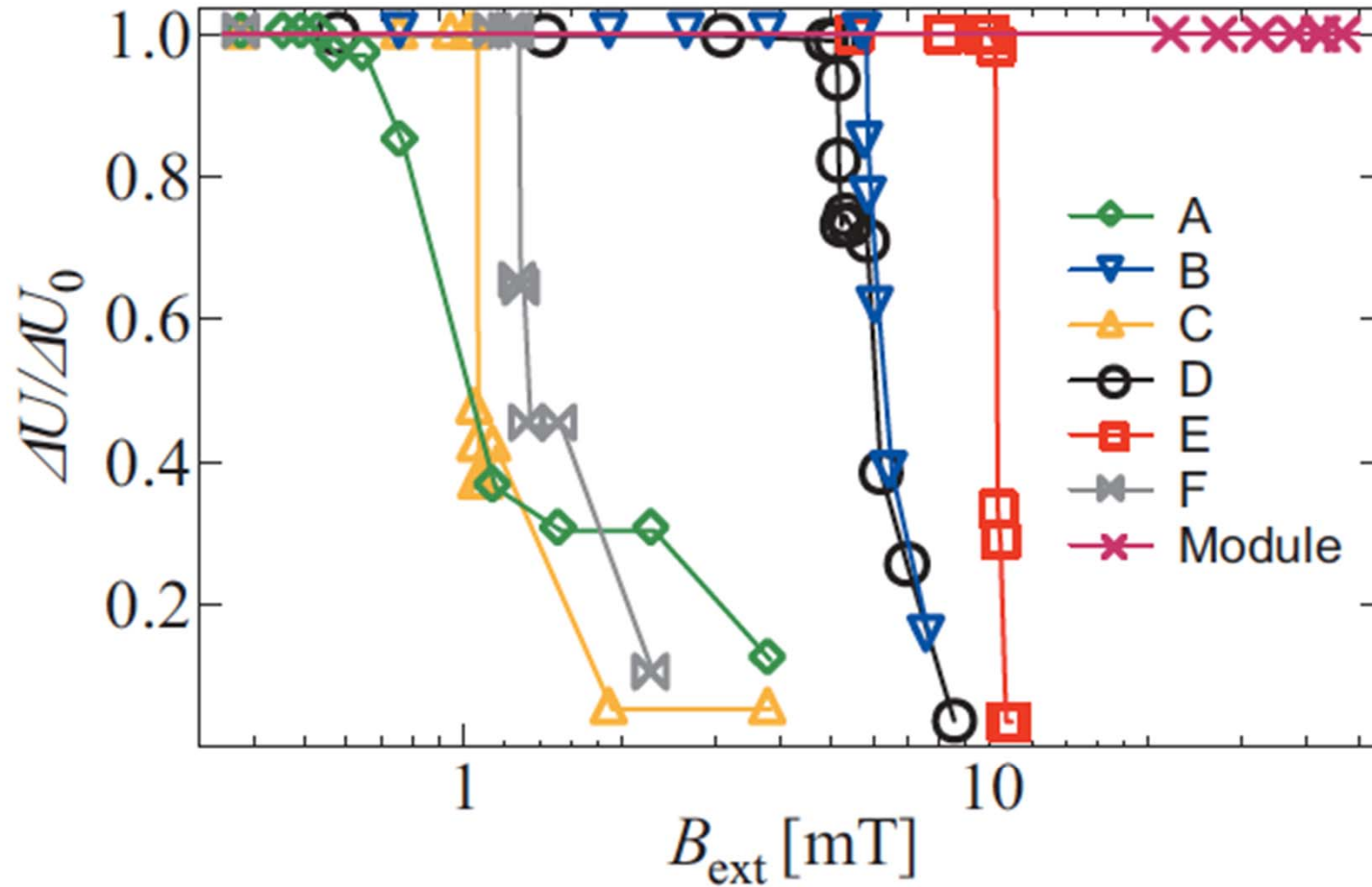


Figure 1. The relative modulation depth $\Delta U/\Delta U_0$ as a function of applied perpendicular magnetic field B_{ext} for different SQUID types and a sensor module (see next section) constructed from F type SQUIDs and $11 \times 11 \times 2 \text{ mm}^3$ Nb shields.

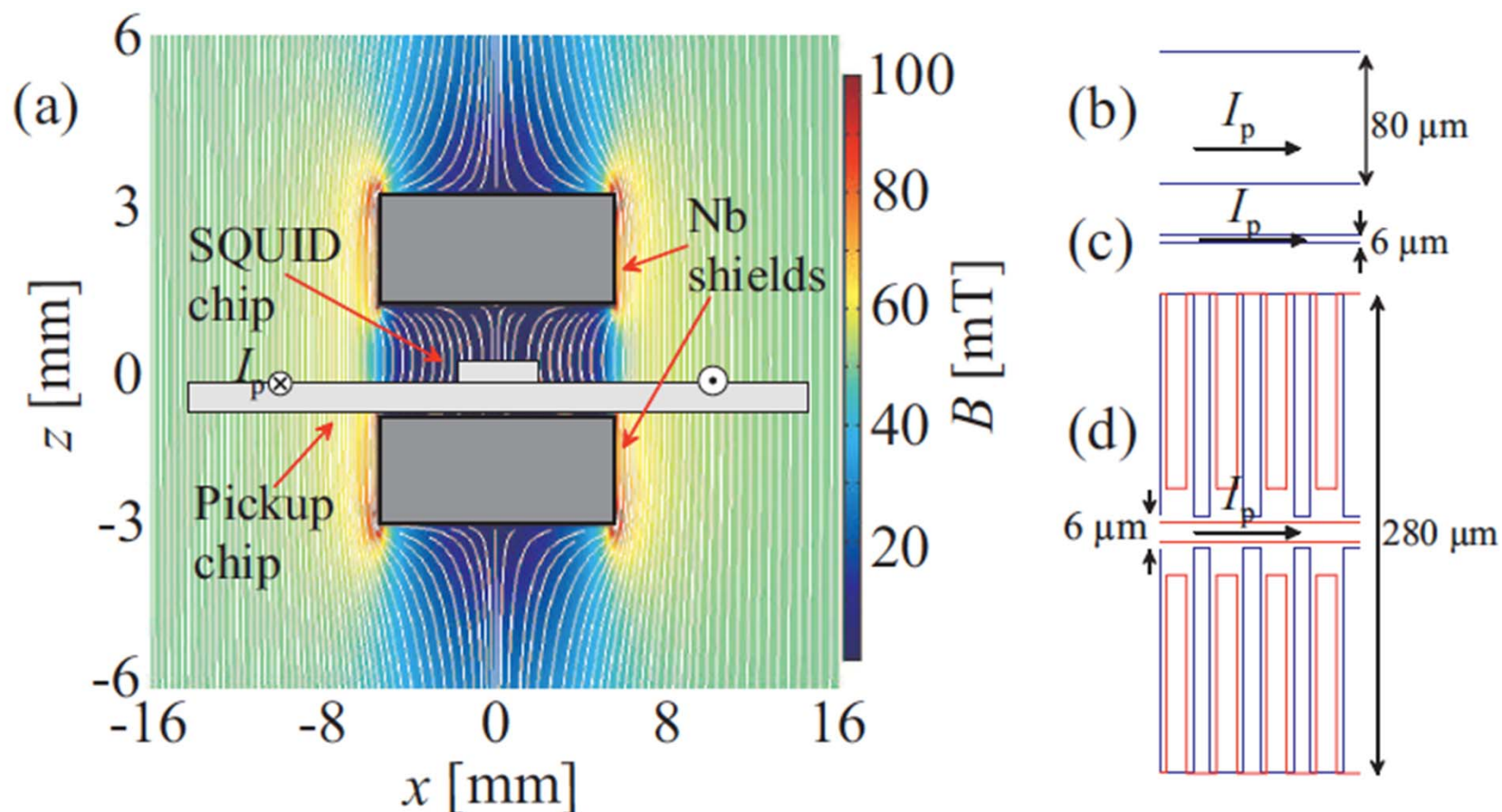


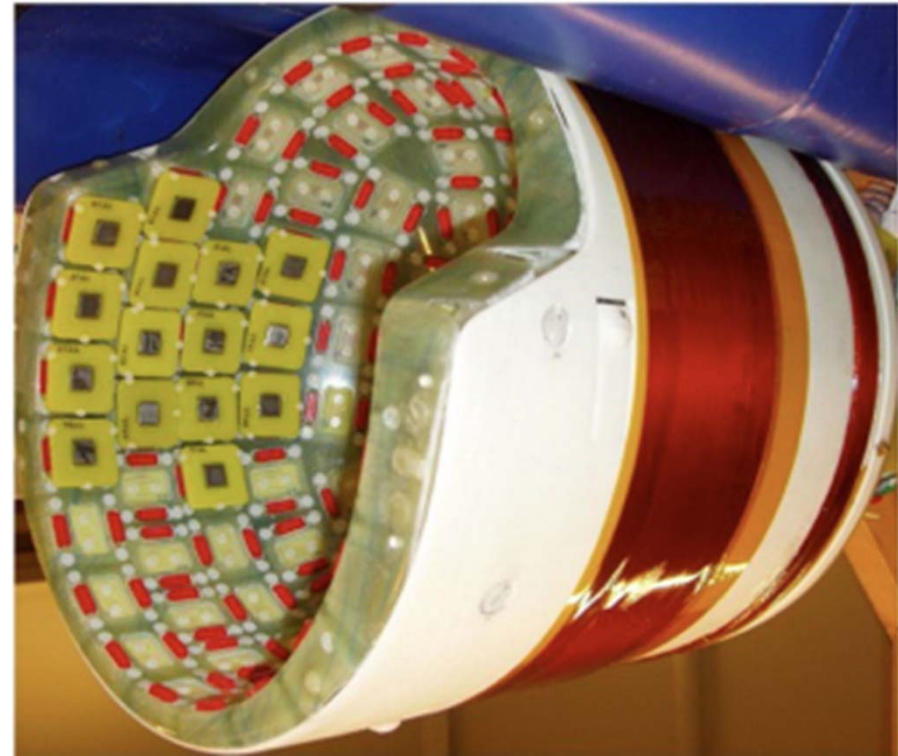
Figure 3. (a) A cross-section of the sensor module showing the pickup and SQUID chips as well as superconducting shields. The shielding effect is calculated by a finite element method showing the position dependence of the field in the ambient field of 50 mT in z -direction. With thin-film devices, the linewidth reduction from (b) to (c) decreased the sensitivity of a magnetometer. As a result, (d) a comb-like structure constructed of 6- μm -wide films was introduced. I_p flows in blue layer whereas in (d) an additional superconducting layer (denoted with red) was deposited on top of the gaps for screening purposes. The effective width of the transmission line increases as the flux generated by I_p is compelled to enclose the whole 280- μm -wide structure, thus, lowering the inductance of the pickup coil and raising the sensitivity of the sensor.

SC Coils for Prepolarization

Acceptable specs:

- $B > 100 \text{ mT}$
- Switch time $< 10 \text{ ms}$
- Remanent field $< 100 \text{ nT}$

Currently, $B < 22 \text{ mT}$
limited by magnetization
of the wire



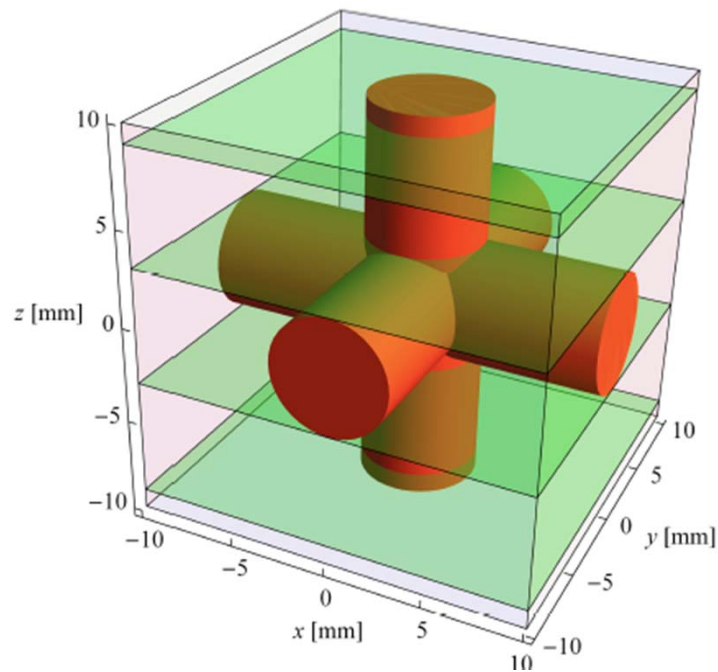
- 552 turns, diameter 30 cm
- 0.44-mm wire; 24000 1- μm filaments
bronze matrix (Supercon, Inc)
- 0.02-mm insulation
- 2 mT/A, total inductance 130 mT

If we succeed:

- Measurement of conductivity distribution with high spatial accuracy (1 mm) and with small relative error (perhaps 5% when current errors may be 50%)
=> Elimination of the geometrical uncertainty in the structure of the head from the MEG/EEG inverse problem
- MEG accuracy from 5 to 1–2 mm
- EEG accuracy from perhaps 20 mm to < 5 mm
- TMS accuracy from 5 to 1–2 mm
- Improved work flow
- New markets

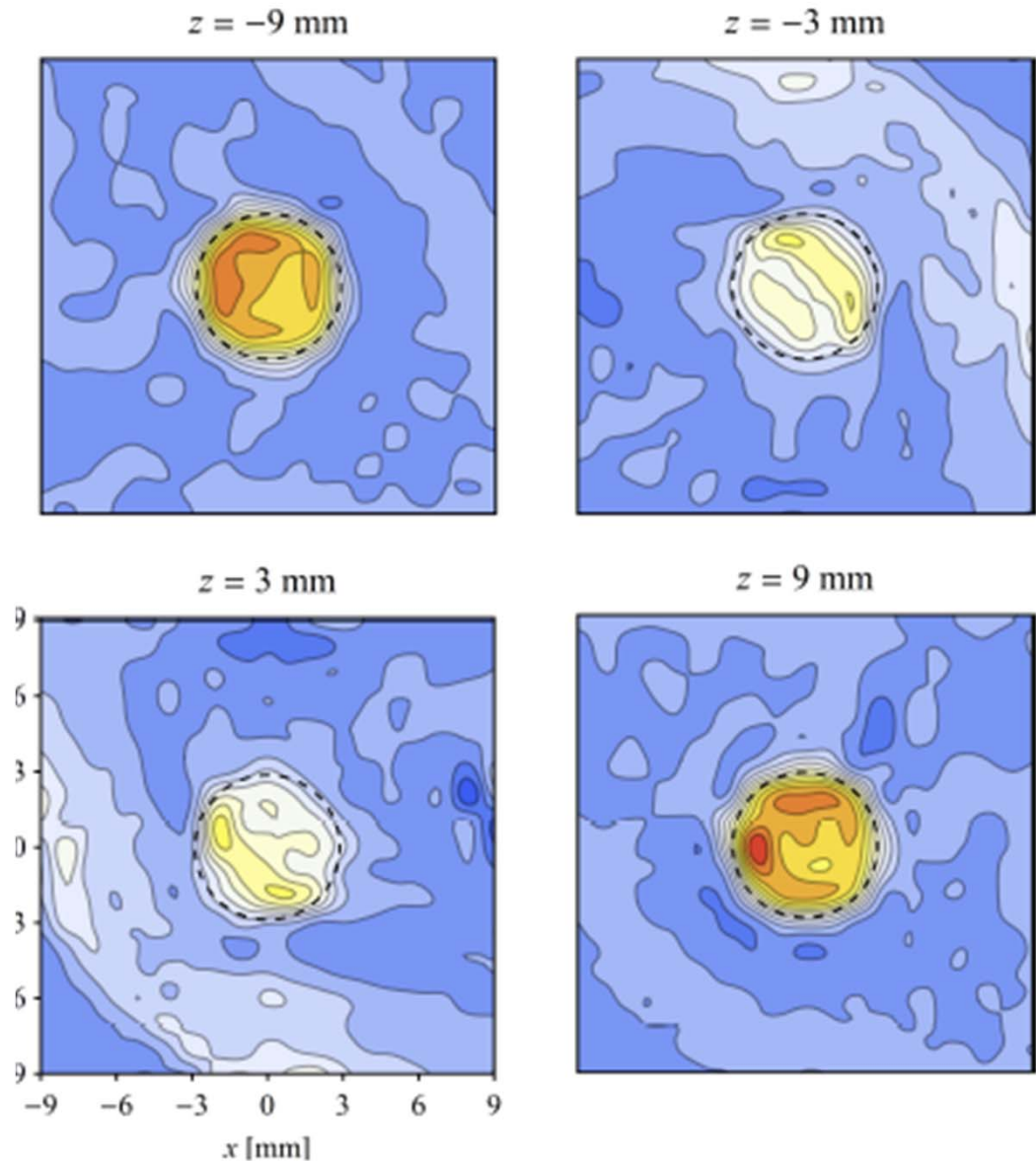
Conductivity imaging with ULF-MRI

Simulation

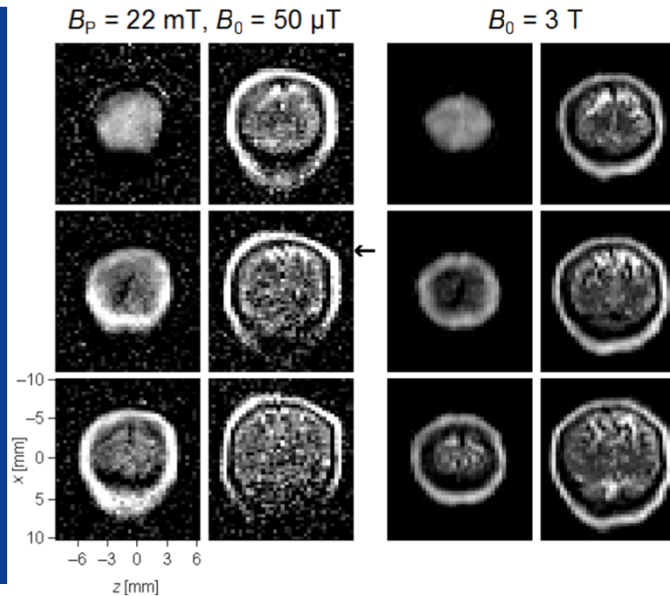
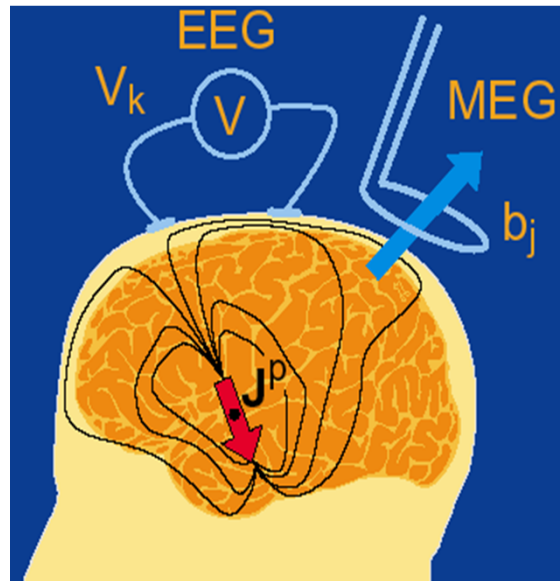


Presently, the S/N is not sufficient for human experimentation (too strong currents would be needed)

Nieminen, Vesänen et al. 2012



Thank you for your attention!



Panu Vesanen, Koos Zevenhoven, Juhani Dabek, Sarianna Alanko, Andrey Zhdanov, Mika Pollari, Fa Hsuan-Lin, Jaakko Nieminen, Tuomas Hirvonen, Juha Simola, Lauri Parkkonen, Antti Ahonen, Juho Luomahaara, Juha Hassel, Jari Penttilä, Jyrki Mäkelä, Juha Montonen, and the MEGMRI consortium; megmri.net