

A Brief History of fMRI

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&

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Neurol ma e

Editor-in-Chief Peter Bandettini

Special Issue 20 Years of fMRI: **The Science** and the Stories

Available online at www.sciencedirect.com **SciVerse ScienceDirect**

Functional Magnetic Resonance Imaging in **Medicine and Physiology**

CHRIT T. W. MOONEN, PETER C. M. VAN ZIJL, JOSEPH A. FRANK, DENIS LE BIHAN, EDWIN D. BECKER

(1990) *Science,* **250, 53-61.**

angiography

Gadolinium perfusion

Diffusion

magnetization transfer

metabolic imaging (NAA)

IVIM: Intravoxel Incoherent Motion

Denis Le Bihan, MD, PhD . Eric Breton, MS . Denis Lallemand, MD • Marie-Louise Aubin, MD • Jacqueline Vignaud, MD • Maurice Laval-Jeantet, MD

Radiology 1988; 168:497-505

Separation of Diffusion and Perfusion in Intravoxel Incoherent Motion MR Imaging¹

Robert Turner, PhD . Denis Le Bihan, MD, PhD . Joseph Maier, MS (EEE) . Robert Vavrek, MS (EEE) . L. Kyle Hedges, PhD . James Pekar, PhD

Echo-Planar Imaging of Intravoxel Incoherent Motion¹ Radiology 1990; 177:407-414

Gradient factor b (s/mm2)

How it all came together…

Five Key Factors For The Emergence of Functional MRI

- **1. Magnetic properties of red blood cells**
- **2. Activation related hemodynamic changes**
- **3. Spatial scale of brain activation**
- **4. Echo Planar Imaging**
- **5. Prevalence of MRI scanners**

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Magnetic Properties of Blood

L. Pauling, C. D. Coryell, *Proc.Natl.Acad. Sci. USA 22, 210-216*, **1936**.

K.R.Thulborn, J. C.Waterton, et al., *Biochim. Biophys.Acta. 714: 265-270*, **1982**.

S. Ogawa,T. M. Lee,A. R. Kay, D.W.Tank, *Proc. Natl.Acad. Sci. USA 87, 9868-9872*, **1990**.

Turner, R., Lebihan, D., Moonen, C. T. W., Despres, D. & Frank, J. *Magnetic Resonance in Medicine,* 22, 159-166, **1991.**

red blood cells

oxygenated

deoxygenated

BOLD contrast investigation started in 1936...or even 1845.

210

PROC. N. A. S. CHEMISTRY: PAULING AND CORYELL

THE MAGNETIC PROPERTIES AND STRUCTURE OF HEMOGLOBIN, OXYHEMOGLOBIN AND CARBONMONOX YHEMOGLOBIN

BY LINUS PAULING AND CHARLES D. CORYELL

GATES CHEMICAL LABORATORY, CALIFORNIA INSTITUTE OF TECHNOLOGY

Communicated March 19, 1936

Over ninety years ago, on November 8, 1845, Michael Faraday investigated the magnetic properties of dried blood and made a note "Must try recent fluid blood." If he had determined the magnetic susceptibilities of arterial and venous blood, he would have found them to differ by a large amount (as much as twenty per cent for completely oxygenated and completely deoxygenated blood); this discovery without doubt would have excited much interest and would have influenced appreciably the course of research on blood and hemoglobin.¹

Continuing our investigations of the magnetic properties and structure of hemoglobin and related substances,² we have found oxyhemoglobin and carbonmonoxyhemoglobin to contain no unpaired electrons, and ferrohemoglobin (hemoglobin itself) to contain four unpaired electrons per heme. The description of our experiments and the interpretation and discussion of the results are given below.

Biochimica et Biophysica Acta, 714 (1982) 265-270 **Elsevier Biomedical Press**

BBA 20122

OXYGENATION DEPENDENCE OF THE TRANSVERSE RELAXATION TIME OF WATER PROTONS IN WHOLE BLOOD AT HIGH FIELD

KEITH R. THULBORN, JOHN C. WATERTON *, PAUL M. MATTHEWS and GEORGE K. RADDA Department of Biochemistry, University of Oxford, South: Parks Road, Oxford OX1 3QU (U.K.)

(Received August 4th, 1981)

Key words: Oxygenation dependence; Transverse relaxation time; Water proton; High field NMR; (Whole blood)

At high and medium magnetic field, the transverse NMR relaxation rate (T_2^{-1}) of water protons in blood is determined predominantly by the oxygenation state of haemoglobin. T_2^{-1} depends quadratically on the field strength and on the proportion of haemoglobin that is deoxygenated. Deoxygenation increases the volume magnetic susceptibility within the erythrocytes and thus creates local field gradients around these cells. From volume susceptibility measurements and the dependence of T_2^{-1} on the pulse rate in the Carr-Purcell-Meiboom-Gill experiment, we show that the increase in T_2^{-1} with increasing blood deoxygenation arises from diffusion of water through these field gradients.

Blochimica et Biophysica Acta, 714 (1982) 265-270 **Elsevier Biomedical Press**

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Department of Biochemistry, University of Oxford, South Parks Road, Oxford OX1 3QU (U.K.)

(Received August 4th, 1981)

Blood R2 proportional to Oxygenation

R2 effect is due to bulk susceptibility and not dipole-dipole interaction

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 λ

... Six years later...

Oxygenation-Sensitive Contrast in Magnetic Resonance Image of Rodent Brain at High Magnetic Fields

SEIJI OGAWA, TSO-MING LEE, ASHA S. NAYAK,* AND PAUL GLYNN

AT&T Bell Laboratories, Murray Hill, New Jersey 07974

Received November 30, 1988; accepted June 20, 1989

At high magnetic fields $(7 \text{ and } 8.4 \text{ T})$, water proton magnetic resonance images of brains of live mice and rats under pentobarbital anesthetization have been measured by a gradient echo pulse sequence with a spatial resolution of 65×65 -um posel size and 700 - μ m slice thickness. The contrast in these images depicts anatomical details of the brain by numerous dark lines of various sizes. These lines are absent in the image taken by the usual spin echo sequence. They represent the blood vessels in the image slice and appear when the deoxyhemoglobin content in the red cells increases. This contrast is most pronounced in an anoxy brain but not present in a brain with diamagnetic oxy or carbon monoxide hemoglobin. The local field induced by the magnetic susceptibility change in the blood due to the paramagnetic deoxyhemoglobin causes the intra voxel dephasing of the water signals of the blood and the surrounding tissue. This oxygenation-dependent contrast is appreciable in high field images with high spatial resolution. © 1990 Academic Press Inc.

in vivo

in vitro

20% O2

100% O₂

S. Ogawa, T.-M. Lee, A. S. Nayak, P. Glynn, Magn. Reson. Med*,* **14, 68-78 (1990)**

Susceptibility-Induced Field Distortion in the Vicinity of a Microvessel \perp to B₀.

Magnetic Resonance in Medicine "30 papers that shaped the field."

Echo-Planar Time Course MRI of Cat Brain Oxygenation Changes

ROBERT TURNER, * + DENIS LE BIHAN, † CHRIT T. W. MOONEN, § DARYL DESPRES, § AND JOSEPH FRANK‡

* Laboratory of Cardiac Energetics, ‡Diagnostic Radiology Department, and §In Vivo NMR Research Center, National Institutes of Health, Bethesda, Maryland 20892

Received June 25, 1991; revised August 7, 1991

When deoxygenated, blood behaves as an effective susceptibility contrast agent. Changes in brain oxygenation can be monitored using gradient-echo echo-planar imaging. With this technique, difference images also demonstrate that blood oxygenation is increased during periods of recovery from respiratory challenge. © 1991 Academic Press, Inc.

R. Turner, D. LeBihan, C.T.W. Moonen, D. Despres, J. Frank, Magn. Reson. Med*,* **22, 159-166 (1991)**

Ogawa predicted fMRI but got the sign wrong…

"...we expect this oxygenation-sensitive contrast could be used to monitor regional oxygen usages in the brain. When some region in a brain is much more active than other regions, the active region could show darker lines in the image because of the increased level of deoxyhemoglobin resulting from higher oxygen consumption."

"Therefore, in addition to the anatomy of the brain, one aspect of its physiology can be studied by the MRI of water"

Oxygenation-Sensitive Contrast in Magnetic Resonance Image of Rodent Brain at High Magnetic Fields, Seiji Ogawa, Tso-Ming Lee, Asha S. Nayak, and Paul Glynn**.Magnetic Resonance in Medicine 14, 68-78 (1990).**

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The First Functional MRI Results (MGH)

Susceptibility Contrast agent bolus injection and time series collection of T2 - weighted images

The First Functional MRI Results (MGH)

Susceptibility Contrast agent bolus injection and time series collection of T2 - weighted images

The MGH Gang

Jack Belliveau, Explorer of the Brain Using M.R.I., Dies at 55

By BENEDICT CAREY MARCH 9, 2014

Jack Belliveau

Jack Belliveau, a Harvard scientist whose quest to capture the quicksilver flare of thought inside a living brain led to the first magnetic resonance image of human brain function, died on Feb. 14 in San Mateo, Calif. He was 55.

The cause was complications of a gastrointestinal disorder, said his wife, Brigitte Poncelet-Belliveau, a researcher who worked with him at the Athinoula A. Martinos Center for Biomedical Imaging at Massachusetts General Hospital. He lived in Boston. His wife said he died suddenly while visiting an uncle at his childhood home, which he owned.

Dr. Belliveau was a 30-year-old graduate student at the Martinos Center when he hatched a scheme to "see" the neural trace of brain activity. Doctors had for decades been taking X-rays and other images of the brain to look for tumors and other lesions and

to assess damage from brain injuries. Researchers had also mapped blood flow using positron emission tomography scans, but that required making and handling radioactive trace chemicals, whose signature vanished within minutes. Very few research centers had the technical knowledge or the machinery to pull it off.

Dr. Belliveau tried a different approach. He had developed a technique to track blood flow, called dynamic susceptibility contrast, using an M.R.I. scanner that took split-second images, faster than was usual at the time. This would become a standard technique for assessing blood perfusion in

Jack Belliveau, seated for an experimental EEG, with his colleague Gregory Simpson. Harvard Medical School

stroke patients and others, but Dr. Belliveau thought he would try it to spy on a normal brain in the act of thinking or perceiving.

Proc. Natl. Acad. Sci. USA
Vol. 83, pp. 1140-1144, February 1986 Neurobiology

Focal physiological uncoupling of cerebral blood flow and oxidative metabolism during somatosensory stimulation in human subjects

(positron emission tomography)

PETER T. FOX*^{†‡} AND MARCUS E. RAICHLE*[†]

*Department of Neurology and Neurological Surgery (Neurology), †Department of Radiology (Radiation Sciences), and The McDonnell Center for Studies of
Higher Brain Function, Washington University School of Medicine. St. Lou

Communicated by Oliver H. Lowry, October 7, 1985

FIG. 1. Physiological uncoupling of brain blood flow and metabolism. (Left) Resting-state measurements. (Right) Stimulated-state measurements (unilateral vibrotactile stimulation of the fingers). All images are from a single subject's scanning session and pass through the same brain plane. Color scales are linear with the maxima set at a fixed multiple (1.6) of the global average, to facilitate visual comparisons (16). During specific somatosensory stimulation a marked focal increase in CBF (29% of mean, nine subjects, three trials per subject) was produced in the contralateral sensorimotor cortex. The observed increase in the CMRo₂ was much smaller (5% of mean, nine subjects, three trials ner subject) and failed to attain sig-

nificance. This physiological uncoupling of CBF and CMRo₂ flow produced a highly significant decrease in the local OEF $(-19\% \text{ of mean})$, indicating that tissue Po. (and probably pH) rose during stimulation.

as contralateral/ipsilateral ratios (see text and Tables $1-4$, the disparity between blood flow and metabolism was evident from the raw data and was not dependent on a particular strategy of analysis.

The video that started it all. Aug 12, 1991: Tom Brady Plenary Lecture at SMRM, San Francisco

K. K. Kwong, et al, (1992) "Dynamic magnetic resonance imaging of human brain activity during primary sensory stimulation." Proc. Natl. Acad. Sci. USA. 89, 5675-5679.

10 cm. 5/ice cardsac $5mGp$ gree photo sti. ! TR:2.55 $T8=y8 (MP9)91$ C_1 T T $A = 109$
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Re 7A gestim. avec 33-70 (38) and $C - C$ B 70 ser shin $disdeg-z$ 30 Pre 40 prot $2s$ $T1 = 1.058$ IR $3.5EB$ disdag $TR = 3s$ $TT = 1100ms$ $TF = 42$ 40 pc 40 post $z \sim$ $\sqrt{\frac{29}{30}}$ 40 80 IR Image 66 20 30.50 W $\partial \overline{\partial}$ $90 \mu c$ itation (40) \mathcal{F} $irst$; Trst:n. $2-\sigma$ 20 irstim 79 10 cr. spie cardiac. $irstim .su$ removing $\int x \, dx$ in. Are 23 $5 - 8$ 13 14 16 18 19 25 - 21 $(16 + \sqrt{3}L)$ irstim.s $44:160$ $(44 + pH)$ Irafim. And lag get mirstim. avg (savernom).

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Gepre. av.g. gest. 35 yR IR 370 (0 ν $77 = 1.05$ $9.$ $3.5EB$ disdag $TR = 35$ $TT = 1100ms$ $T_{t} = 40$ 40 pc 40 post z_{ν} 80 **IR (CBF) Contrast**20 20 $\partial \overline{\partial}$ light of 30 in ages i tstin p μ 3-30 (28) $90 \mu c$ 40° \mathcal{G} $84 - 33 - 68$ $LT57$ $67 - 80$ (47) Trstin. sub looks good. only -270. $2-\sigma$ 757.80
 47.80
 $4-30$
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 67.80 **The original block design** $\frac{1}{1}$ rstim.sub (73) (subtractif d'un 4) $\frac{10.05}{11.00}$ paradigm *semany* $(16 + \sqrt{3}L)$ irstim.s $2-5$ 7 10-17 19-50 $44:1100$ $\{r_1\}$: 1, pro $34-38$ 40-49-465 67-80
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S. Ogawa, et al., (1992) "Intrinsic signal changes accompanying sensory stimulation: functional brain mapping with magnetic resonance imaging." Proc. Natl. Acad. Sci. USA. 89, 5951-5955.

What preceded the results from the Medical College of Wisconsin...

MAGNETIC RESONANCE IN MEDICINE 21, 39-48 (1991)

Coil Optimization for MRI by Conjugate Gradient Descent

ERIC C. WONG,* A. JESMANOWICZ, AND JAMES S. HYDE

Biophysics Section, Department of Radiology, Medical College of Wisconsin, Milwaukee, Wisconsin 53226

Received April 30, 1990; revised June 29, 1990

Local head gradient coils: Window(s) of opportunity

Fig. 1 GUI for gradient descent gradient coil design tool. The design shown is one octant of the X gradient coil designed and built in August 1991. The program was written in Objective C and ran on a NeXT Cube computer.

NeuroImage, Volume 62, Issue 2, 2012, 660 - 664

http://dx.doi.org/10.1016/j.neuroimage.2012.01.025

August, 1991

Initially could only do one slice…

2.5 cm !

TR = 2 sec TE = 50 ms One slice In plane 3.75 x 3.75

One little known fact…

We didn't even need a gradient coil:

EPI at 5mm x 5mm x 5mm was quite possible using 100 amp gradient amplifiers and the whole body gradient coils…

Every scanner in the world in 1991 could have performed EPI-based fMRI at perfectly reasonable resolution.

P. A. Bandettini, et al., (1992) "Time course EPI of human brain function during task activation." Magn. Reson. Med 25, 390-397.

Blamire, A. M., et al. (1992). "Dynamic mapping of the human visual cortex by high-speed magnetic resonance imaging." Proc. Natl. Acad. Sci. USA 89: 11069-11073.

MAGNETIC RESONANCE IN MEDICINE 23, 37-45 (1992).

Perfusion Imaging

JOHN A. DETRE, * † JOHN S. LEIGH, * DONALD S. WILLIAMS, ‡ AND ALAN P. KORETSKY 1.6

* Metabolic Magnetic Resonance Research Center and Department of Biochemistry and Biophysics. University of Pennsylvania School of Medicine. Philadelphia. Pennsylvania 19104: and ‡Pittsburgh NMR Center for Biomedical Research and §Department of Biological Sciences, Carnegie Mellon University, Pittsburgh, Pennsylvania 15213

Received July 2, 1990; revised January 3, 1991

Measurement of tissue perfusion is important for the functional assessment of organs in vivo. Here we report the use of H NMR imaging to generate perfusion maps in the rat brain at 4.7 T. Blood water flowing to the brain is saturated in the neck region with a sliceselective saturation imaging sequence, creating an endogenous tracer in the form of proximally saturated spins. Because proton T_1 times are relatively long, particularly at high field strengths, saturated spins exchange with bulk water in the brain and a steady state is created where the regional concentration of saturated spins is determined by the regional blood flow and regional T_1 . Distal saturation applied equidistantly outside the brain serves as a control for effects of the saturation pulses. Average cerebral blood flow in normocapnic rat brain under halothane anesthesia was determined to be 105 ± 16 cc $\cdot 100$ g⁻¹ \cdot min⁻¹ (mean \pm SEM, $n = 3$), in good agreement with values reported in the literature, and was sensitive to increases in arterial $pCO₂$. This technique allows regional perfusion maps to be measured noninvasively, with the resolution of ¹H MRI, and should be readily applicable to human studies. @ 1992 Academic Press. Inc.

Proc. Natl. Acad. Sci. USA Vol. 89, pp. 212-216, January 1992 **Biophysics**

Magnetic resonance imaging of perfusion using spin inversion of arterial water

(cerebral blood flow/adiabatic fast passage/hypercarbia/rat brain/cold injury)

DONALD S. WILLIAMS*, JOHN A. DETRE^{†‡}, JOHN S. LEIGH[†], AND ALAN P. KORETSKY^{*§}

*Pittsburgh Nuclear Magnetic Resonance Center for Biomedical Research, and ^{\$}Department of Biological Sciences, Carnegie Mellon University, Pittsburgh, PA 15213; and [†]Metabolic Magnetic Resonance Research Center, Department of Radiology, and [‡]Department of Neurology, University of Pennsylvania School of Medicine, Philadelphia, PA 19104

Communicated by Mildred Cohn, September 19, 1991

FIG. 2. (A) Coronal image of a rat head. The resonance planes for radiofrequency used for spin inversion by AFP for control and inversion images are indicated by 1 and 3, respectively, and plane 2 is the detection plane. (B) Control transverse image from the detection plane (plane 2 in A). (C) Difference image between control and inversion images. (D) T_{lapp} image.

FIG. 5. Comparison of conventional MRI and perfusion imaging of a rat brain subjected to a regional cold injury. (A) Conventional T_2 -weighted image $(TE = 60$ ms, $TR = 2$ s). The injured region shows up as hyperintensity due to a longer T_2 . (B) Perfusion image of the same slice. The grey scale is from 0 to 6 ml·g⁻¹·min⁻¹. The injured region is dark due to low flow.

Magnetic Resonance in Medicine "30 papers that shaped the field."

Perfusion Contrast EPISTAR FAIR

TI (ms) FAIR EPISTAR

- **K. K. Kwong, et al, (1992)** "Dynamic magnetic resonance imaging of human brain activity during primary sensory stimulation." **Proc. Natl. Acad. Sci. USA. 89, 5675-5679.**
- **S. Ogawa, et al., (1992)** "Intrinsic signal changes accompanying sensory stimulation: functional brain mapping with magnetic resonance imaging." **Proc. Natl. Acad. Sci. USA. 89, 5951-5955.**
- **P. A. Bandettini, et al., (1992)** "Time course EPI of human brain function during task activation." **Magn. Reson. Med 25, 390-397.**
- **Blamire, A. M., et al. (1992).** "Dynamic mapping of the human visual cortex by high-speed magnetic resonance imaging." **Proc. Natl. Acad. Sci. USA 89: 11069-11073.**
- **Frahm, J., et al (1992)** "Dynamic MR Imaging of Human Brain Oxygenation During Rest and Photic-Stimulation." **Journal of Magnetic Resonance Imaging, 2, 501-505.**

Functional MRI of the Brain

A Report on the SMRM/SMRI Workshop held in Arlington, Virginia

June 17–19, 1993

MRM 30:405-408 (1993)

Society of Magnetic Resonance in Medicine Society for Magnetic Resonance Imaging

FUNCTIONAL MRI OF THE BRAIN

Syllabus

A Workshop Presented by the **Society of Magnetic Resonance in Medicine** and the **Society for Magnetic Resonance Imaging**

June 17-19, 1993 The Ritz-Carlton, Pentagon City Arlington, Virginia

Denis Le Bihan National Institutes of Health Diagnostic Radiology Department Building 10, Room 1C-660 Bethesda, Maryland 20892

Robert Turner National Institutes of Health Laboratory of Cardiac Energetics Building 10, Room B1D-161 Bethesda, Maryland 20892

Michael E. Moseley Department of Radiology **Stanford University** Stanford, California 94305-5488

James S. Hyde **Biophysics Research Institute** Medical College of Wisconsin 8701 Waterton Plank Road Milwaukee, Wisconsin 53226

Functional Neuroimaging with EPI: Sequence Issues

Robert Turner, Peter Jezzard, #Lucie Hertz-Pannier, #Denis Le Bihan, *David Feinberg

Laboratory of Cardiac Energetics, National Heart, Lung, and Blood Institute, and #Diagnostic Radiology Department, Clinical Center, NIH, Bethesda, MD 20892 *Department of Radiology, NYU Medical Center, New York, NY

ABSTRACT

Freedom from motion artifact, comparatively good SNR, rapid multi-slice capability, and excellent time resolution make Echo-Planar Imaging an excellent choice for BOLD contrast MR functional neuroimaging. However, when the gradient echo version of EPI is used for this purpose, problems arise regarding image quality and interpretation. Large draining veins distant from active neural regions are the major confusing factor. At high enough static magnetic fields, spin-echo EPI can be used to obtain images showing local changes of blood oxygenation related to brain activation, in which draining veins have less effect. The idea MRFN equence will combine gradient-recalled echo and spin echo features, and thus will be some variant of GRASE (GRAdient echo and Spin Echo).

A proposed acronym...

The earliest successful magnetic resonance functional neuroimaging (MRFN) studies with BOLD contrast were made using a gradient-echo version of echo-planar imaging (EPI). The EPI technique, proposed by Mansfield in 1977 (4), allows the capture of a complete MR image in under 100 ms. Thus most motions in the body are frozen and motion artifact rarely appears. EPI relies on a very rapidly switched magnetic field gradient of large amplitude, and a fast data capture rate. Since these features were not considered necessary by most manufacturers of commercial MR systems until recently, the technique has been available only in a few pioneering laboratories. The technique normally uses a full 90 degree rf pulse for spin excitation, and hence provides a comparatively high single-shot signal/noise ratio (SNR), considering the large receiver bandwidth required. For brain imaging, with equal voxel size, an EPI image with 40 ms acquisition time has been found to have the same SNR as a FLASH image with optimized bandwidth taking 2 seconds to acquire. Faster FLASH images will have a poorer SNR than EPI. Low flip-angle variants of EPI (5) can of course provide much higher values of SNR/unit time, though this sacrifices SNR in each

Functional Mapping of the Human Visual Cortex at 4 and 1.5 Tesla Using Deoxygenation Contrast EPI

R. Turner, P. Jezzard, H. Wen, K. K. Kwong, D. Le Bihan, T. Zeffiro, R. S. Balaban

MRM 29:277-279 (1993)

FIG. 2. Plot of fractional change in 4 T (squares) and 1.5 T (triangles) EPI image intensity versus time in the eight-voxel regions of interest in the visual cortex shown in Fig. 1, for a volunteer experiencing alternate 30-s periods of rest and photic stimulation. Details of acquisition for the 4 and 1.5 T data are described in the

Local Gradient Coil

NIH_{4T}

Siemens' new 7T

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Visual Cortex Organization

Right

Ocular Dominance Column Mapping

Menon, R. S., S. Ogawa, et al. (1997). J Neurophysiol 77(5): 2780-7. 0.54×0.54 in plane resolution

Optical Imaging

R. D. Frostig et. al, PNAS 87: 6082-6086, (1990).

Cheng, et al. (2001) Neuron,32:359-374

 0.47×0.47 in plane resolution

Cheng, et al. (2001) Neuron,32:359-374

Yacoub et al. PNAS 2008

Orientation Columns in Human V1 as Revealed by fMRI at 7T

Yacoub et al. PNAS 2008

Scalebar = 0.5 mm
Multi-sensory integration

M.S. Beauchamp et al.,

Functional Neuroimaging Techniques

after Churchland and Sejnowski, 1988

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MRI vs. fMRI

high resolution (1 mm)

one image

MRI **the contract of the contr**

many images (e.g., every 2 sec for 5 mins)

Sir Peter Mansfield conceived of Echo Planar Imaging long before it was physically possible…

Spatial Mapping of the Chemical Shift in NMR, P. Mansfield. Magnetic Resonance in Medicine 1, 370-386 (1984).

LETTER TO THE EDITOR

A target field approach to optimal coil design

R Turner Department of Physics, Nottingham University, Nottingham NG7 2RD, UK

Received 2 May 1986

Abstract. Using a Fourier-Bessel expansion of the magnetic field generated by currents flowing on a cylinder, the relationship between current and field may be inverted. Hence the current density required to generate a specified target field B_z on the surface of a cylinder of smaller radius may be evaluated. Gradient, solenoidal and shim coil designs based on this approach provide exceptionally large usable volumes while remaining compact and, if necessary, of low inductance.

the gradient is uniform to 5% is shaded. Wire positions:

Snapshot Head Imaging at 0.5 T Using the Echo Planar Technique

R. J. ORDIDGE, R. COXON, A. HOWSEMAN, B. CHAPMAN, R. TURNER. M. STEHLING, AND P. MANSFIELD

Department of Physics, University of Nottingham, Nottingham NG7 2RD, United Kingdom

Received April 22, 1988; revised June 17, 1988

The echo planar imaging (EPI) method and related variants of this technique can produce complete two-dimensional images from the data collected in a single experiment lasting a fraction of a second. EPI methods are used at 0.5 T to produce snapshot images of the human head with a spatial resolution of less than 2 mm. \degree 1988 Academic Press, Inc.

FIG. 2. The gradient and RF pulse sequence used in the MBEST experiment. An unphased image is produced by taking the magnitude of the Fourier transformed data. The largest spin echoes occur midway through the experiment, thus providing discrimination against signal with a short T_2 relaxation time.

Approximate EPI Timeline

1976-84 P. Mansfield conceives of EPI 1989 EPI of humans emerges on a handful of scanners 3 x 3 x 3-10 mm3 1989 ANMR retrofitted with GE scanners for EPI 1991 Home built head gradient coils perform EPI 1996 EPI is standard on clinical scanners 2000 Gradient performance continues to increase 2002 Parallel imaging allows for higher resolution EPI 2006 1.5 x 1.5 x 1.5 mm3 single shot EPI possible 2009 At 7T sub – mm single shot EPI for fMRI is possible

Imaging System Components

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Scopus: Articles or Reviews Published per Year

How it all came together…

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Motor (black) Primary Sensory (red) egrative Sensory (violet) Basic Cognition (green) High onition (yellow) Emotion (blue)

J. Illes, M. P. Kirschen, J. D. E. Gabrielli, Nature Neuroscience, 6 (3)m p.205

Technology

Coil arrays High field strength High resolution Novel sequences

Methodology

Paradigm design Univariate / Multivariate Multi-modal integration Real time feedback Classification

Fluctuations **Dynamics** Functional Resolution

Interpretation V Applications

Healthy Brain Organization Clinical Research Clinical Applications

A bit more history…

- **Event-Related fMRI (1993, 1996)**
- **Retinotopy (1995)**
- **Resting State fMRI (1995…2006)**
- **Ocular Dominance Columns (1997)**
- **High Field and Multi-channel Receive (2000)**
- **fMRI "Decoding" (2001)**
- **Intracortical recordings and fMRI (2001)**
- **DARPA Brain Reading Competition (2006)**
- **Approval for Pre-surgical Mapping (2007)**

Blamire, A. M., et al. (1992). "Dynamic mapping of the human visual cortex by high-speed magnetic resonance imaging." Proc. Natl. Acad. Sci. USA 89: 11069-11073.

Proc. Natl. Acad. Sci. USA Vol. 93, pp. 14878-14883, December 1996 Neurobiology

Detection of cortical activation during averaged single trials of a cognitive task using functional magnetic resonance imaging

(neuroimaging/single trial/language/prefrontal)

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fMRI of human visual cortex

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Mapping striate and extrastriate visual areas in human cerebral cortex

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Resting State Correlations

correlation with reference function seed voxel in motor cortex Activation:

Rest:

B. Biswal et al., MRM, 34:537 (1995)

Activation-based fMRI and "resting state" f

Resting Correlation (Right Hand Seed)

Resting state fMRI: Why is this area important?

- The number of papers and applications has exploded.
- Neuronal, psychiatric, and developmental disorders may relate to altered connectivity.
- Methods are in their infancy, and rapidly evolving.
- Neuronal correlates of spontaneous fluctuations are not fully understood.

Magnetic Resonance in Medicine "30 papers that shaped the field."

Distributed and Overlapping Representations of Faces and Objects in Ventral Temporal Cortex

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Logothetis et al. (2001) "Neurophysiological investigation of the basis of the fMRI signal" Nature, 412, 150-157

Parametric manipulation of brain activation demonstrated that BOLD contrast approximately followed the level of brain activation: visual system (Kwong et al., 1992), auditory system (Binder et al., 1994), and motor system (Rao et al., 1996).

The use of continuous variation of visual stimuli parameters as a function of time was proven a powerful method for fMRI-based retinotopy: (Engel et al., 1994, Deyoe et al., 1994, Sereno et al., 1995).

Event-related fMRI was first demonstrated (Blamire et al., 1992).

Application of event-related fMRI to cognitive activation was shown (Buckner et al., 1996, McCarthy et al., 1997).

Development of mixed event-related and block designs was put forward: (Donaldson et al., 2002).

Paradigms were demonstrated in which the activation timing of multiple brain systems timing was orthogonal, allowing multiple conditions to be cleanly extracted from a single run (Courtney et al., 1997).

High resolution maps were created: For spatial resolution: ocular dominance columns (Menon et al., 1997, Cheng et al., 2001) and cortical layer activation maps were created (Logothetis et al., 2002).

Extraction of information at high spatial frequencies within regions of activation was demonstrated (Haxby et al., 2001).

For temporal resolution: Timings from ms to hundreds of ms were extracted (Ogawa et al., 2000, Menon et al., 1998, Henson et al., 2002, Bellgowan et al., 2003).

The development of "deconvolution" methods allowed for rapid presentation of stimuli (Dale and Buckner, 1997).

Early BOLD contrast models were put forward: (Ogawa et al., 1993, Buxton and Frank, 1997).

More sophisticated models were published that more fully integrated the latest data on hemodynamic and metabolic changes (Buxton et al., 2004).

The development of "clustered volume" acquisition was put forth as a method to avoid scanner noise artifacts: (Edmister et al., 1999).

The findings of functionally related resting state correlations: (Biswal et al., 1995) and regions that consistently show deactivation (Binder et al., 1999, Raichle et al., 2001) were described.

Observation of the pre-undershoot in fMRI (Hennig et al., 1997, Menon et al., 1995, Hu et al., 1997) and correlation with optical imaging was reported (Malonek and Grinvald, 1996).

Simultaneous use of fMRI and direct electrophysiological recording in non-human primate brain during visual stimulation elucidated the relationship between fMRI and BOLD contrast. (Logothetis et al., 2001). Simultaneous electrophysiological recordings in animal models revealed a correlation between negative signal changes and decreased neuronal activity (Shmuel et al., 2002). Simultaneous electrophysiological recordings in animal models provided evidence that inhibitory input could cause an increase in cerebral blood flow (Matheiesen et al., 1998).