



# Brain Recurrence Analysis: Not Your Father's EEG

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# The EEG



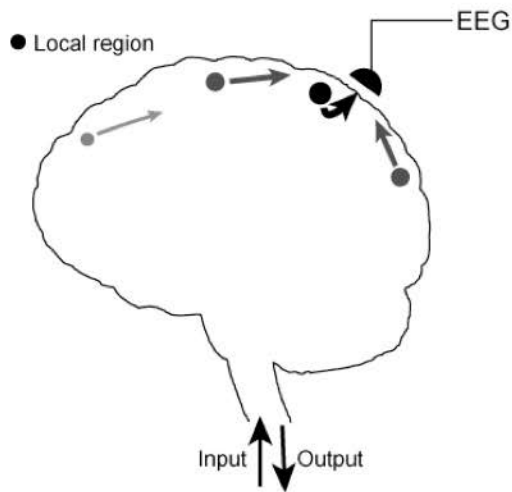
What it is: Time-dependent voltages measured between pairs of standardized locations on the scalp

Traditional clinical use: Diagnosing brain diseases: largely supplanted by CT and MRI

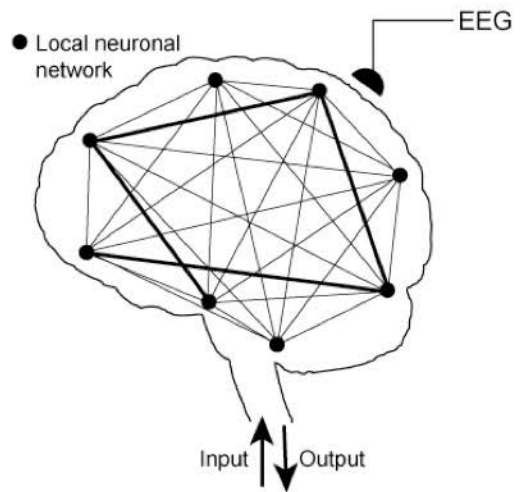
Present-day uses:

- Sleep studies
- Epilepsy
- Brain death

# Recent Developments: The Cognitive Perspective

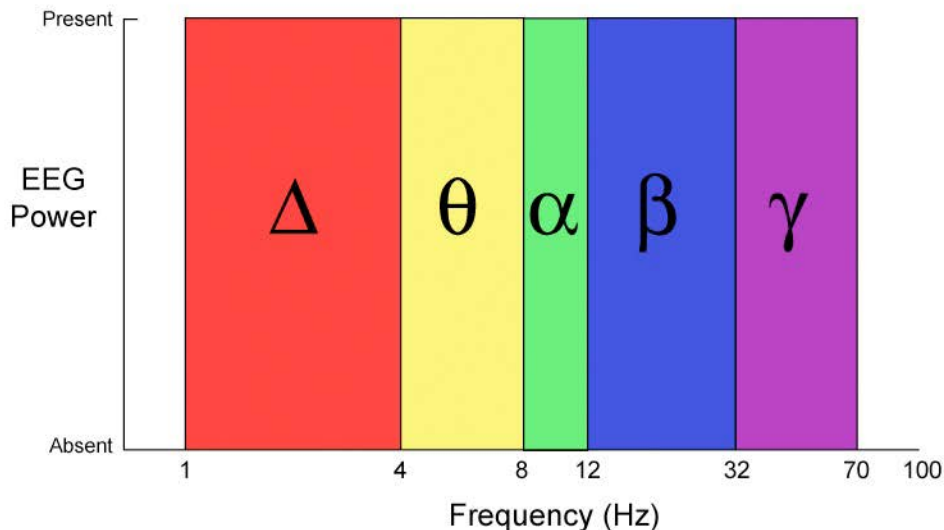


Traditional View  
(linear, quasi-static)



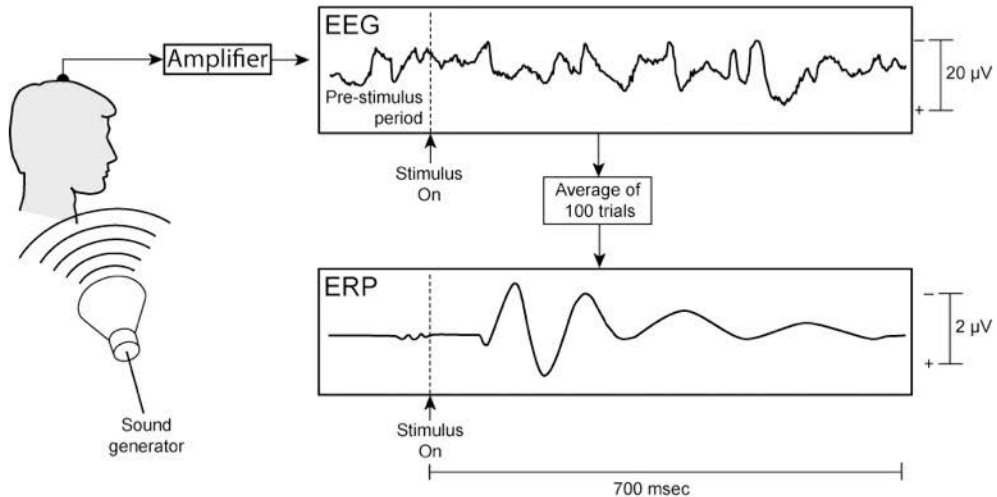
New View  
(nonlinear, dynamic)

# Linear Method: Spectral Analysis



Assumption: EEG is composed of parts

# Linear Method: Time Averaging



Assumption: Response is time locked

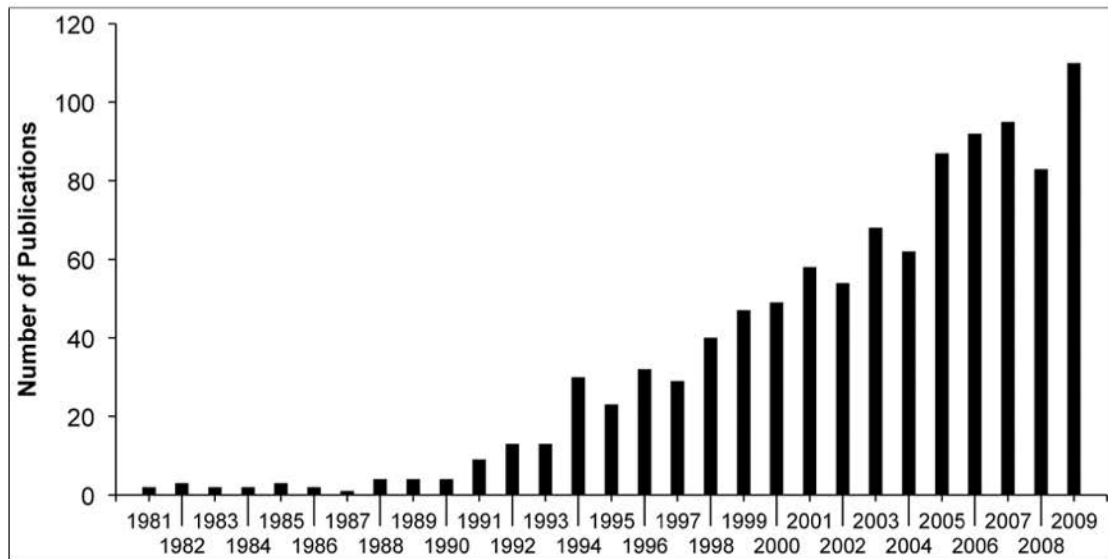
# Basic Ideas Behind Nonlinear Analytical Methods

- **Sensitivity to initial conditions**



- **Recurrence: The future resembles (but is not identical to) the past**

# PubMed Publications on Nonlinear Analysis of EEG

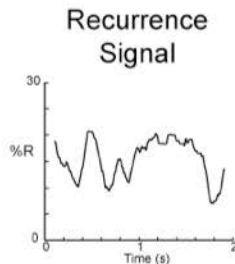
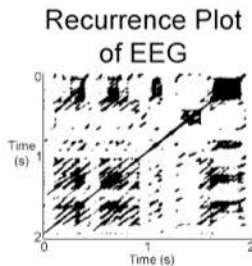
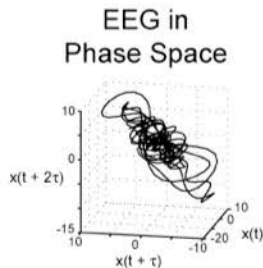
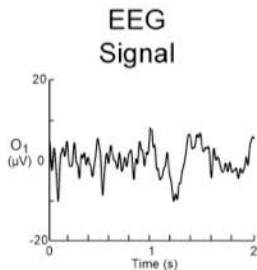


# Characteristics of Nonlinear Methods

- General method (includes linear)
- Symbolic representation of dynamics
- Units are unfamiliar and nonphysiological
- ~100 specific techniques; no consensus yet



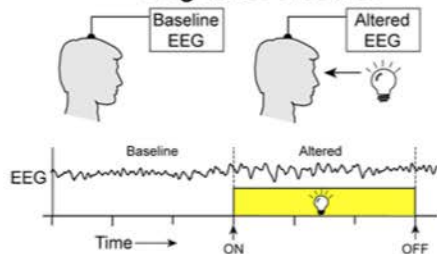
# A Representative Technique: Recurrence Analysis



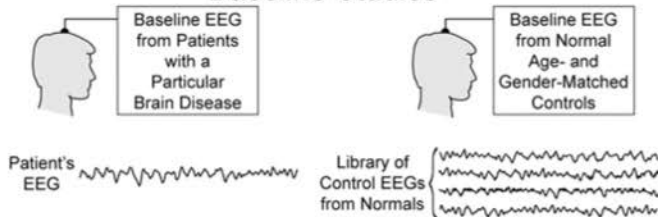
S. Carrubba, C. Frlot, A. Chesson, Jr., A.A. Marino: J. Neurosci. Meth. 157:39–47, 2006.  
S. Carrubba, C. Frlot, A.L. Chesson, Jr., A.A. Marino: J. Neurosci. Meth. 173:41–46, 2008.

# Experimental Design

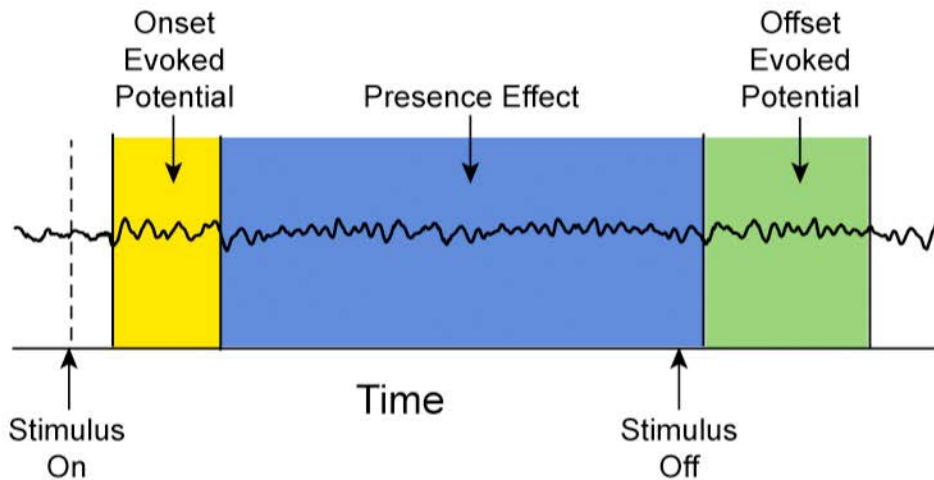
## Cognitive Studies



## Baseline Studies

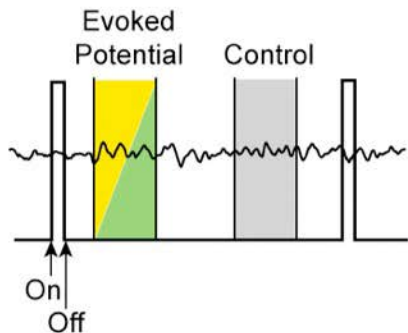


# Distinct Brain Electrical Responses

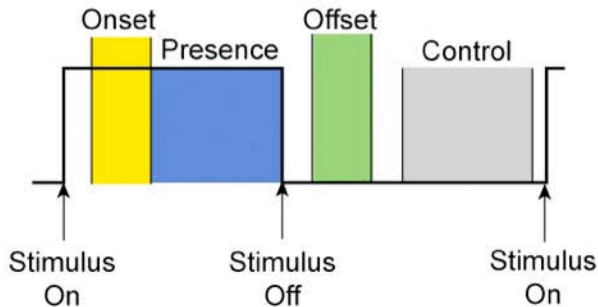


# Experimental Procedures

## Evoked-Potential Study



## Onset EP, Offset EP, and Presence Study



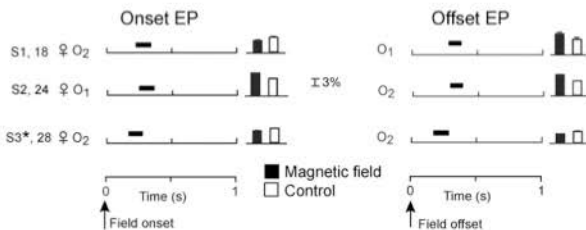
# Discovery of Human Magnetic Sense

**Rationale:** Sensory perception entails evoked potentials

**Procedure:**



**Typical Results**  
(N=17 subjects):



**Summary:**

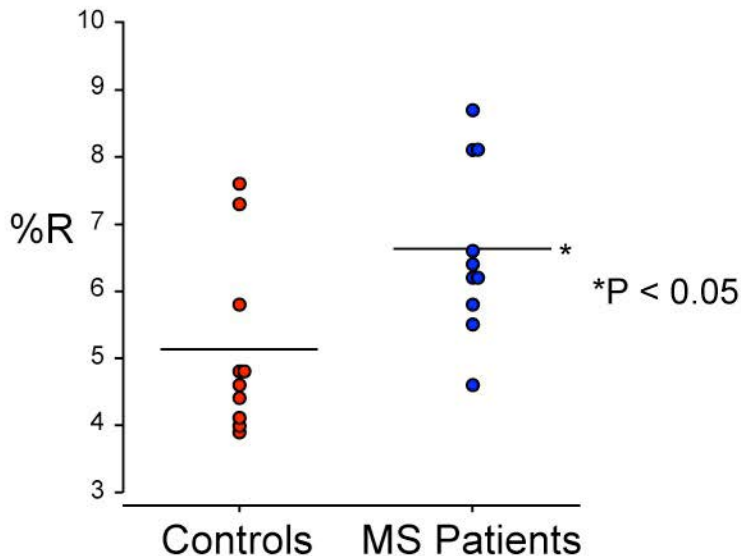
- Each subject detected the magnetic field ( $P < 0.05$ )
- Latency and direction of effect (relative to control) varied with subject
- Effect not bilateral

# Basis for Functional Test for Multiple Sclerosis

Patients with Multiple Sclerosis			Normal Controls		
MS participant (age)	Onset Evoked Potential	Offset Evoked Potential	Normal participant (age)	Onset Evoked Potential	Offset Evoked Potential
1 (40)	No Response	No Response	1 (51)	O <sub>2</sub>	C <sub>3</sub> C <sub>4</sub> P <sub>3</sub>
2 (34)	No Response	O <sub>1</sub> O <sub>2</sub> P <sub>4</sub>	2 (66)	O <sub>2</sub> C <sub>3</sub> P <sub>4</sub>	C <sub>4</sub> P <sub>4</sub>
3 (52)	No Response	O <sub>1</sub> C <sub>3</sub>	3 (22)	No Response	O <sub>2</sub> P <sub>3</sub>
4 (32)	O <sub>1</sub> O <sub>2</sub> C <sub>4</sub>	C <sub>3</sub> C <sub>4</sub>	4 (26)	C <sub>3</sub> C <sub>4</sub> P <sub>3</sub>	No Response
5 (19)	No Response	O <sub>2</sub> C <sub>3</sub> P <sub>3</sub>	5 (23)	C <sub>3</sub> C <sub>4</sub> P <sub>4</sub>	O <sub>2</sub> C <sub>4</sub> P <sub>3</sub>
6 (30)	O <sub>2</sub> C <sub>3</sub>	O <sub>2</sub> C <sub>3</sub> P <sub>3</sub>	6 (23)	C <sub>3</sub> C <sub>4</sub>	C <sub>4</sub> P <sub>4</sub>
7 (18)	No Response	C <sub>3</sub> C <sub>4</sub> P <sub>3</sub> P <sub>4</sub>	7 (23)	O <sub>1</sub> C <sub>3</sub> P <sub>3</sub>	O <sub>1</sub> O <sub>2</sub> C <sub>3</sub> P <sub>3</sub> P <sub>4</sub>
8 (27)	C <sub>3</sub> C <sub>4</sub> P <sub>4</sub>	O <sub>2</sub> C <sub>4</sub> P <sub>4</sub>	8 (46)	O <sub>1</sub> C <sub>3</sub>	O <sub>1</sub> C <sub>3</sub> P <sub>3</sub>
9 (50)	No Response	No Response	9 (23)	O <sub>1</sub> O <sub>2</sub> C <sub>4</sub> P <sub>3</sub> P <sub>4</sub>	C <sub>3</sub> P <sub>3</sub>
10 (31)	No Response	O <sub>1</sub> P <sub>3</sub>	10 (25)	P <sub>3</sub> P <sub>4</sub>	C <sub>3</sub> P <sub>3</sub> P <sub>4</sub>
11 (38)	No Response	O <sub>2</sub> C <sub>4</sub>			

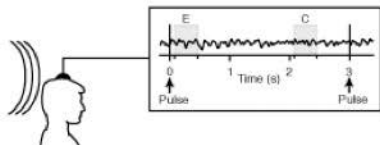
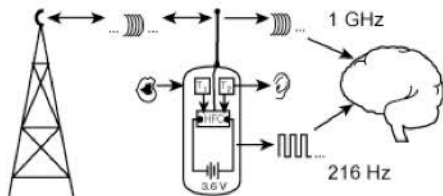
S. Carrubba, A. Minagar, E. Gonzalez-Toledo, A.L. Chesson Jr., C. Frilot II, A.A. Marino: Neurol. Res. 32:297, 2010.

# Comparison of Baseline EEGs from MS and Control Patients



S. Carrubba, A. Minagar, A.L. Chesson Jr, A.A. Marino: Submitted.

# Effect of Cell-Phone Magnetic Fields (EMFs) on Brain Electrical Activity



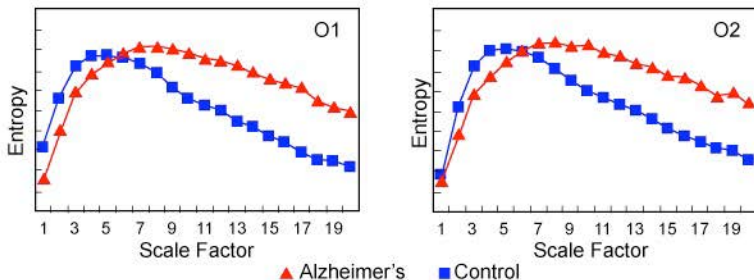
**Summary:** Typical cell phone triggers 216 evoked potentials per sec of use

Subject	Evoked Potentials	Family-wise Error
1 (24/M)	O1 O2 P4	0.010
2 (53/F)	C3 C4 P3	0.002
3 (22/F)	C3 P3 P3	0.001
4 (22/M)	C3 C4 P3	0.001
5 (22/F)	O1 O1	0.081
6 (43/F)	O1 C4	0.006
7 (22/F)	O2 P4	0.031
8 (50/F)	O2 P3 P4	0.062
9 (62/M)	—	—
10 (18/F)	O2 C3	0.078
11 (36/F)	O2 C3	0.043
12 (47/F)	C3 P3 P4	0.085
13 (32/F)	O1 P3	0.006
14 (24/F)	O1 O2	0.038
15 (52/M)	C3 P3	0.015
16 (22/F)	C3 C4	0.021
17 (23/F)	—	—
18 (29/M)	C4 P3	0.043
19 (23/M)	C3 P3	0.023
20 (26/M)	P3 P4	0.015



# Nonlinear Dynamics of EEG in Alzheimer's Disease

- Decreased complexity<sup>1</sup>
- A particularly promising measure<sup>2</sup>

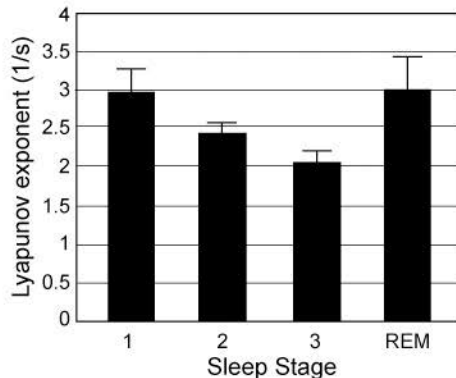
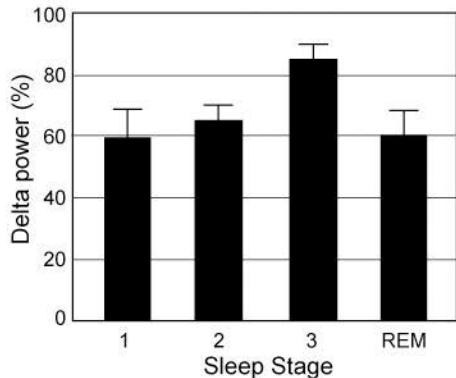


1. J. Jeong: Clin. Neurophysiol. 115:1490-1505.
2. T. Mizuno, T. Takahashi, R.Y. Cho, M. Kikuchi, T. Murata, K. Takahashi, Y. Wada: Clin. Neurophysiol. 121:1438-1446, 2010.

# Sleep Medicine: Frontier for Nonlinear EEG Analysis

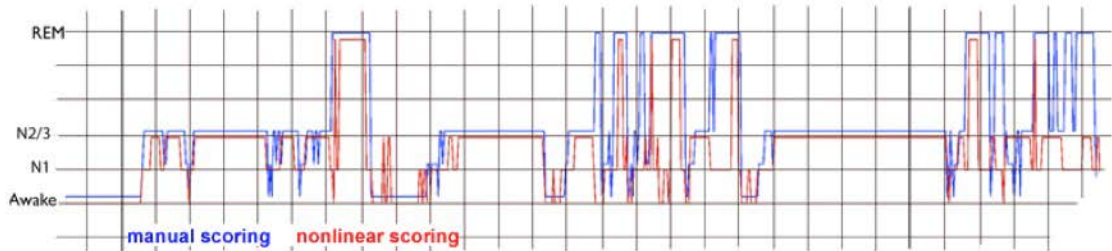
- Sleep phenomenology/pathology defined using EEG
- Goal No. 1: Automated scoring
- Goal No. 2: Fine structure of macro states

# Discrimination of Sleep Stages: Comparison Between Linear and Nonlinear Measures



J. Fell, J. Röschke, K. Mann, C. Schäffner. *Electroencephalog. Clin. Neurophysiol.* 98:401–410, 1996.

# Sleep States Derived from Manual and Nonlinear (Computational) Scoring



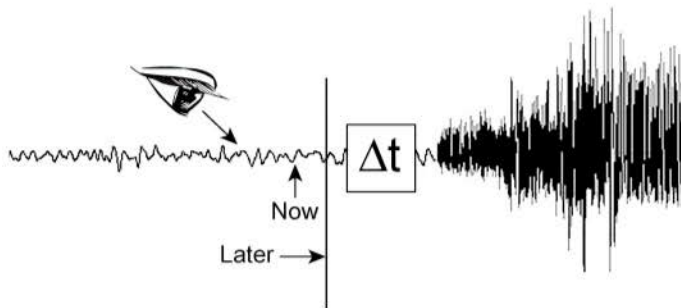
M.R. Titchener: Entropy sensitive to sleep state. NOLTA, 2006.

# Nonlinear EEG Analyses in Sleep Medicine

- Sleep states stable in minutes  $\implies$  Sleep ideal for testing computational schemes to distinguish brain states
- Expect additional (not redundant) information
- Applications:
  - Diagnosis
  - Effects of drugs
  - Aging

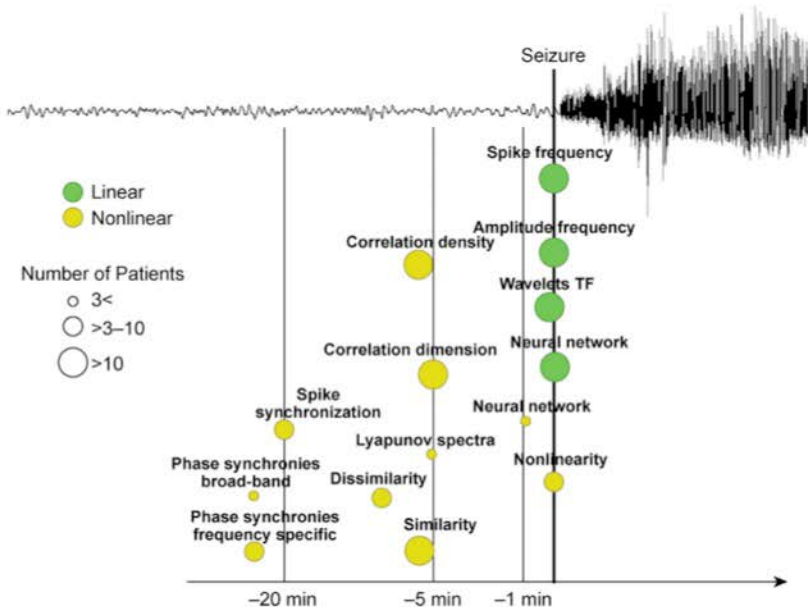
# Epilepsy: Is a Seizure Coming?

Basic Idea:



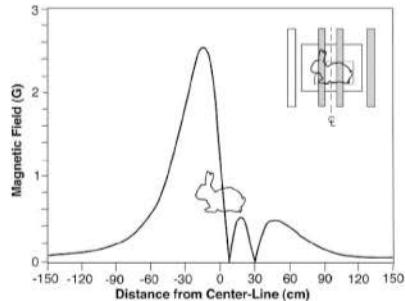
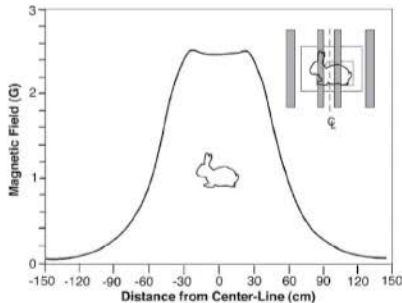
- Is such a thing possible?
- How long is  $\Delta t$ ?

# Relative Success of Different Methods for Predicting Seizures



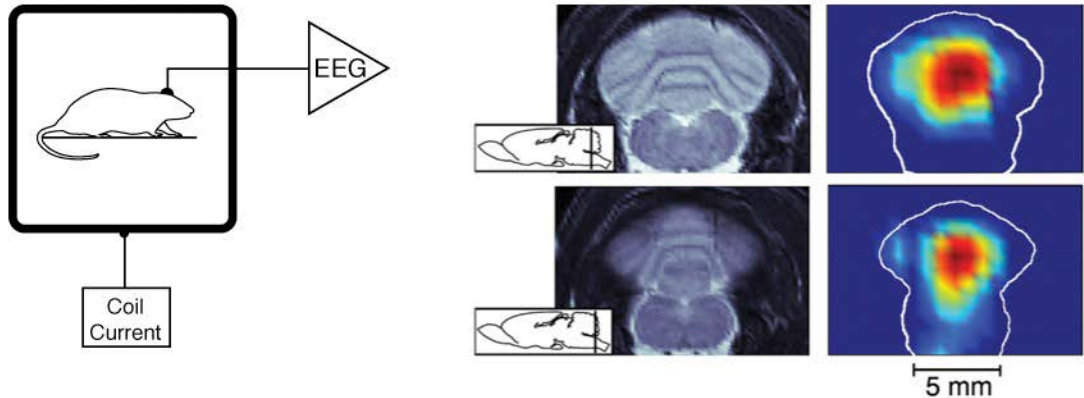
M. Le Van Quyen, J. Martinerie, V. Navarro, M. Baulac, F.J. Varela: J. Clin. Neurophysiol. 18:191–208, 2001.

# Basic-Science Studies: Anatomical Location of Field Detector



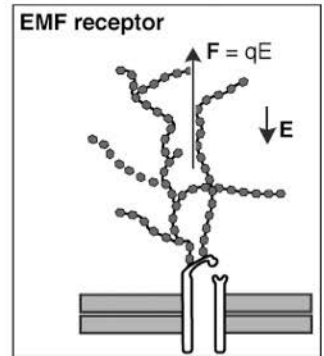
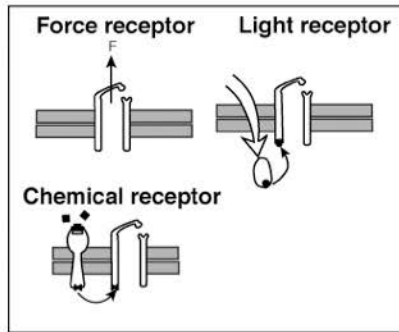
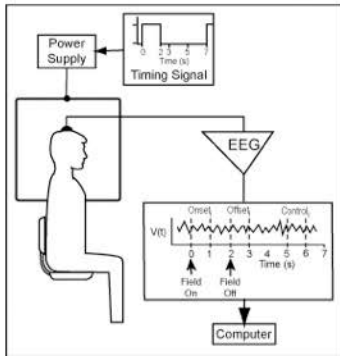


# Basic-Science Studies: Field-Activated Brain Region Assessed Using PET



PET images of field-activated region in rat hindbrain

# Basic-Science Studies: Field Detector is a Force Receptor

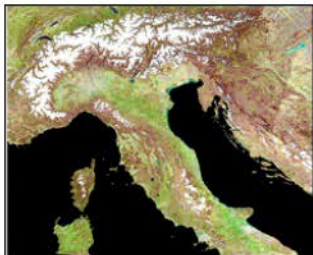


A.A. Marino, S. Carrubba, C. Frilot, A.L. Chesson Jr.: *Neurosci. Lett.* 452:119–123, 2009.

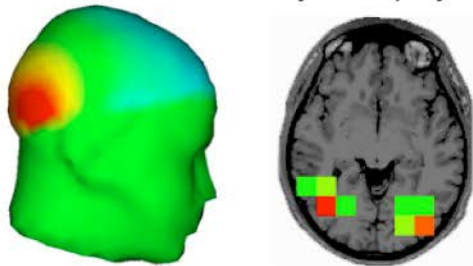
S. Carrubba, C. Frilot 2<sup>nd</sup>, F.X. Hart, A.L. Chesson Jr., A.A. Marino: *Int. J. Radiat. Biol.* 85:622–632, 2009.

# Summary

Limitation of MRI and CT



Almost limitless ability to display data



Nonlinear dynamic model can yield more neurophysiological information



Model Limitation: Not mechanistic

