



High density EEG

Role in epilepsy surgery evaluation, with case demonstrations

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Prelude

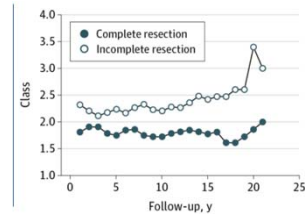
- The epileptogenic zone cannot be localized by a single method

Disclosures

- None

MRI-Negative Epilepsy

- Without potentially epileptogenic lesion regardless of final histopathology



Objectives

- Recognize the value of hdEEG in epilepsy surgery evaluation
- Describe visual hdEEG analysis using illustrative cases
- Describe EEG source imaging using illustrative cases

Implications beyond outcome

- No anatomical structure to guide surgical resection
- Location & extent of implantation not readily apparent
- Risk of complications by 40% for every 20 additional electrodes

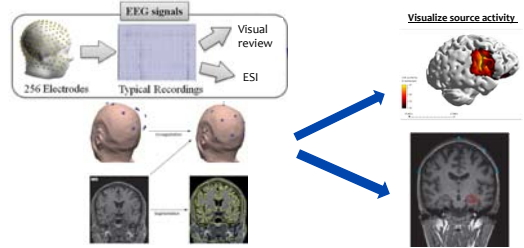
Affordable and accessible means to localize the epileptogenic zone

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Localizing the epileptogenic zone

- Advanced MRI
- SISCOM/SPM
- PET MRI
- MEG/MSI
- MR spectroscopy
- EEG-fMRI
- DTI/DSI (Connectomes)

Work flow



Modified from Hassan et al., 2016, *PLoS One* 9:e015041 and Kaboriboon et al., 2012, *Nat Rev Neurol* 8:498-507.

Pros & Cons

Advantages

- Non-invasive
- Non-lesional to lesional (PET MRI)
- Prognostication (PET/SISCOM)
- Detection of mild MTS (MRS)

	FDG PET	SISCOM/SPM	MEG/MSI*
Concordance with icEEG, nonlesional	19%-69%	64-82%	20-100%
Prediction of Engel class I outcome	75-88%*	55%-70%	38-86%
Cost (interpretation)	>\$1500*	> \$3,600	>\$2000
Availability	Ubiquitous	Limited	Limited

How about adding more electrodes?

Deo et al., 2011, *Neurology* 76:41-48.
Zheng et al., 2016, *NeuroImage Clin* 14:199-205.
Su and Bayliss, 2015, *Clinical Neurophysiology* 126:38-42.

E So, 2002, *Mayo Clin Proc* 77:1151-1156.
Knowlton et al., 2008, *Ann Neurol* 64:55-64.

How many electrodes?

- Max 2-3 cm of IE distance = less distortion of potential distributions
- Higher numbers of electrodes = smaller dipole-localization errors
- Precision of source localization is nonlinear, plateau at 100
- Consider placing more electrodes over the region of interest

Lantz et al., 2003, *Clin Neurophysiol* 114:63-69.
Ding et al., 2009, *J Clin Neurophysiol* 20:195-196.
Brodbeck et al., 2008, *Epilepsia* 49:183-191.

High density EEG (hdEEG)

- As many as 256 channels (1-2cm)
- Improves spatial sampling by reducing distance btn. electrodes
- Computer-assisted EEG source analysis methods (ESI)

Rothman, 2008, *Epilepsia* 49 (Suppl 3): 3-14.
Kaboriboon et al., 2012, *Nat Rev Neurol* 8:498-507.

Visual analysis

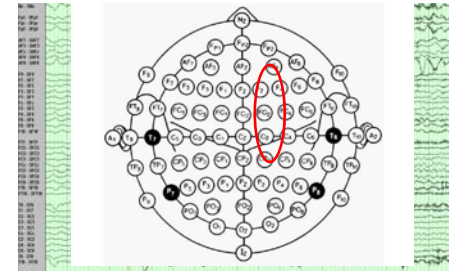
- Commonly used in routine clinical practice
- Simple unlike mathematically complex and non-intuitive-ESI
- Does not require additional software
- Not a precise tool for the identification of epileptogenic zone
- Limited insight into the extent of the involved cortical network

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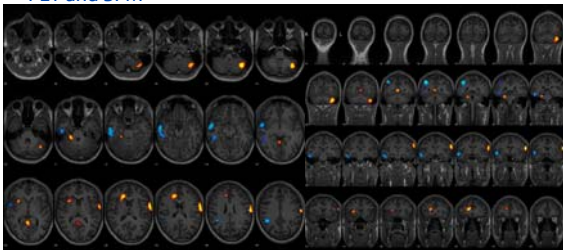
Case 1

- 21 yo RH M with refractory stereotyped sleep-related seizures
 - Grunts and might hold his laughter like a "child in a church"
 - Body turning and some unusual mouth movements
 - Symmetric bilateral arm and leg extension and stiffening
- 10-20 EEG: non-lateralizing (midline)
- MRI-ve

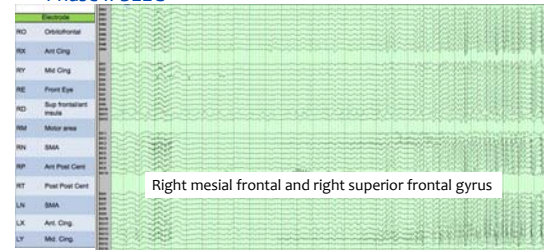
Ictal EEG



PET and SPM



Phase II-SEEG



Right mesial frontal and right superior frontal gyrus

Seizures abated with lifestyle changes and add on CBZ

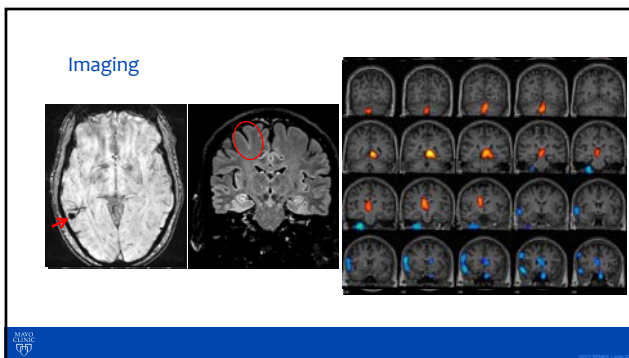
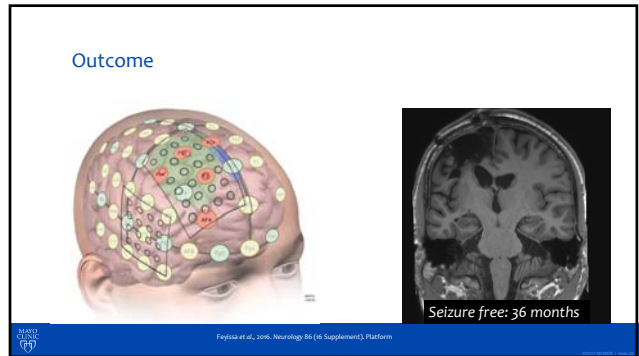
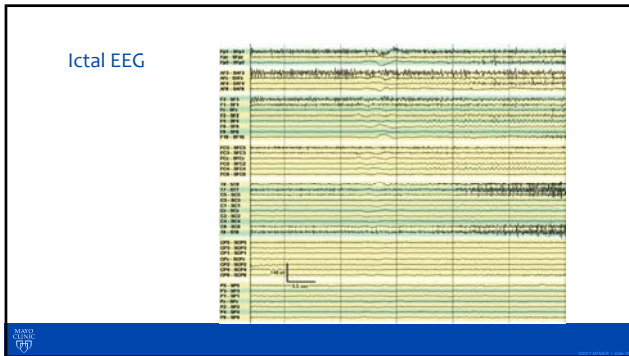
Interictal EEG



Case 2

- A 34 years old right handed man with drug resistant focal epilepsy
 - An aura consisting of an out-of-body kind of experience + deja vu
 - Staring blankly, loss of awareness, unresponsiveness,
 - Suddenly laughter, and lastly hypermotor behavior
 - With GTCS

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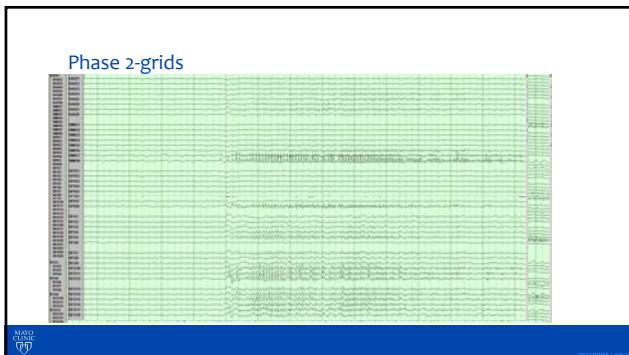


hdEEG in FLE

- The 10-20 system in FLE
 - short seizure duration
 - fast cortical spread
 - motion and muscle artifact
 - suboptimal spatial distribution
 - false lateralization from MF

Nonlocalizing, pseudodiffuse, or wo correlate

Basaldua et al., 1998, Neurology 51:757-761
Catalano et al., 2001, Epilepsy Res 49:147-155



Visual hdEEG in suspected midline FLE

	Lobar concordance with IEEG*	Lateralization concordance with IEEG**
hdEEG (10-10 sys)	12/14	14/14
Conventional EEG	3/14	9/14
SISCOM/SPM	2/12	3/12

All cases with diffuse hdEEG ictal onset also had diffuse IEEG onset!

Consider placing more electrodes over the region of interest!

Fayissa et al., 2017, Epilepsy Res 159:157-166

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EEG source imaging (ESI)

- Co-registration of the electric source estimations with MRI
- Signal processing techniques to estimate current sources
- Insight into the extent of the involved cortical network
- Identify the electric sources underlying epileptogenic activity



Kalbarboon et al., 2012, Nat Rev Neurol 8:498-507.

Inverse solution

- The problem of identifying the source of a given surface field potential



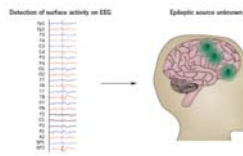
- Equivalent current dipole (ECD) vs distributed source models



Modified from: Kalbarboon et al., 2012, Nat Rev Neurol 8:498-507.

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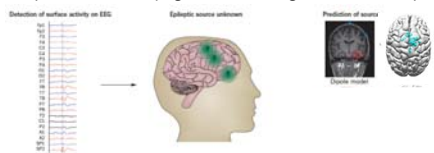
Utility of ESI in epilepsy

- Identification of the irritative zone (spikes and/or sharp waves)
- Identification of the ictal onset zone (EEG seizure onset)
- Identification of seizure networks (seizure propagation)



Inverse solution

- The problem of identifying the source of a given surface field potential



- Equivalent current dipole (ECD) vs distributed source models



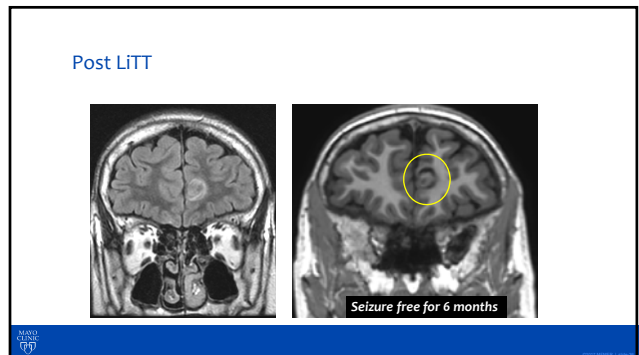
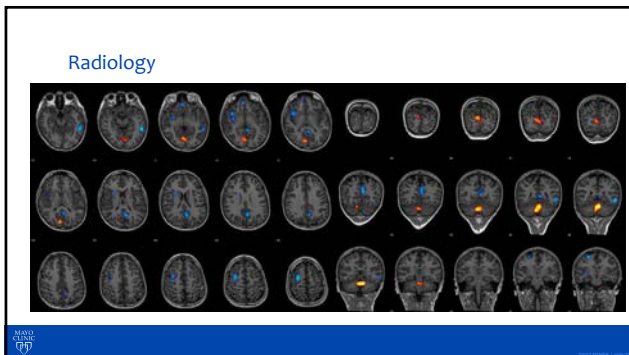
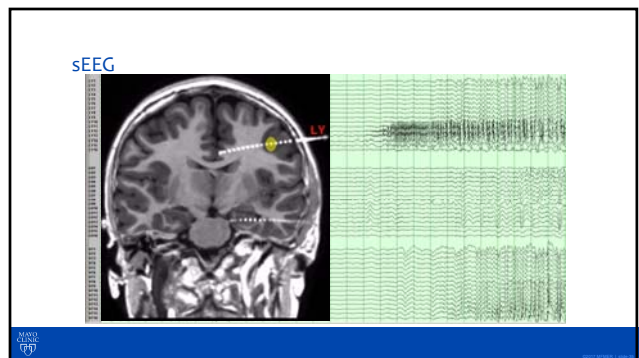
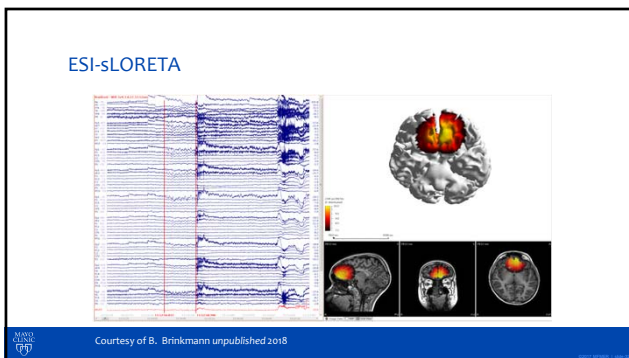
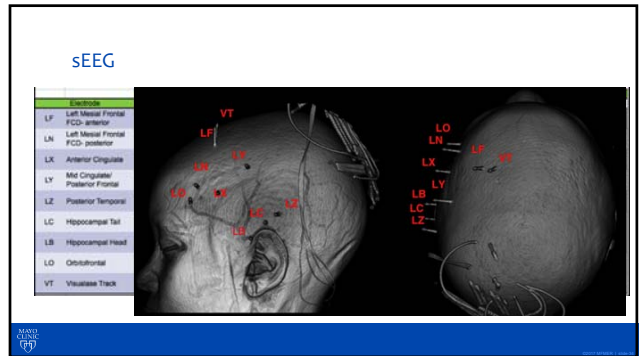
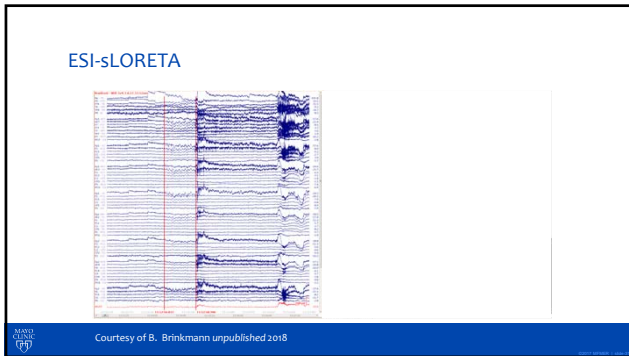
Modified from: Kalbarboon et al., 2012, Nat Rev Neurol 8:498-507.

Case 3

- A 13 yo RH boy with medically intractable focal epilepsy
 - Confused followed by generalized convulsive activity with UI
 - Nocturnal events of staring off, nonspecific vocalization with UI
 - 2-3/week
- MRI -ve (initial read)



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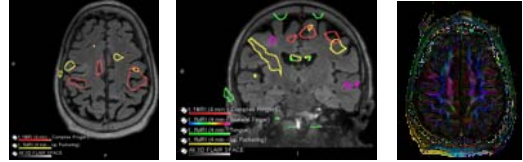
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Case 4

- A 70 yo RM with lesional drug resistant epilepsy
 - Aura of indescribable "funny feeling"
 - Then speech arrest
 - Posturing on the left and post-ictal paresis on the left
 - 1-2/month; GTC X2
- MRI negative but later fMRI revealed suspicious Rt. SMA FCD



fMRI...



- Concern for eloquent cortex seizure onset zone
- Patient offered RNS vs Chronic subthreshold cortical stimulation
- Patient still considering his options



Interictal



Courtesy of B. Brinkmann unpublished 2018



What do we know?

	# of patients	concordance with IEEG or outcome
Interictal ESI		
Brodbeck et al., 2010	10	80%
Wang et al., 2011	7	100%
Brodbeck et al., 2011	192	84%, 88%*
Migeveard et al., 2014	32	80%
Ictal ESI		
Holmes et al., 2010	10	80%
Beniczky et al., 2016	22	73%
Nemtas et al., 2017	14	93%
Visual inspection		
Feyissa et al., 2017	14	86%, 100%

Why hasn't it become a standard of care?

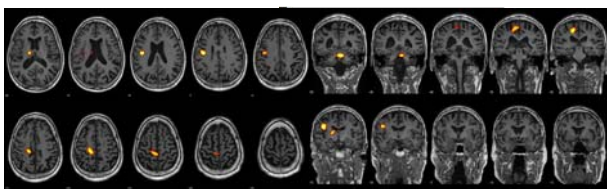
Need user-friendly and easy-to-use ESI software



Brodbeck et al., 2010. Epilepsia 51:131-136.
 Holmes et al., 2010. Neurosurgery 69:1594-1600.
 Wang et al., 2011. Clin Neurophysiol 122:1068-1075.
 Brodbeck et al., 2011. Brain 134:188-199.

Migeveard et al., 2014. J Neural Neurosurg Psychiatry 85:38-45.
 Beniczky et al., 2016. Seizure 43: 1-5.
 Nemtas et al., 2017. Epilepsia 58:2216-2226.
 Feyissa et al., 2017. Epilepsia 58:1920-1926.

Radiology



Last word

- hdEEG improves identification of potential candidates for iEEG & surgery
- Early integration may help identify cases with a low yield iEEG
- Consider placing more electrodes over the region of interest
- ESI is a promising tool and deserves a role in epilepsy surgery evaluation



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