

## Localizing epileptogenic network from SEEG recordings using Epileptogenicity Ranking method

Epileptogenicity Rank (ER) is a modified method of EI for quantifying epileptogenicity of brain structures in epilepsy patients. ER was calculated as the normalized values of the product of spatio-temporal parameter and energy of the signal. The spatial parameter was added along with the existing temporal domain based index calculation (EI) (Bartolomei et al., 2008) to bring the new epileptogenicity rank (ER). We set the range of ER from 1 to 10, 'ER=1' being highly epileptogenic and normal brain ranked as 'ER=10'. ER classifies the seizure onset from the propagation using abrupt frequency change in the time domain and the spatial domain by the anatomical distance from the brain structure that initiated the seizure discharges.

The EZ localization was partially automated by converting the time-series data to the frequency spectrum and applied a threshold over the mean activity to detect the seizure onset. Page and Hinkely's algorithm was implemented for seizure onset detection (Hinkley, 1971; Page, 1954). ER needs two parameters to localize epileptogenic zone (EZ) 1) SEEG and 2) The 3D location of SEEG electrode contacts.

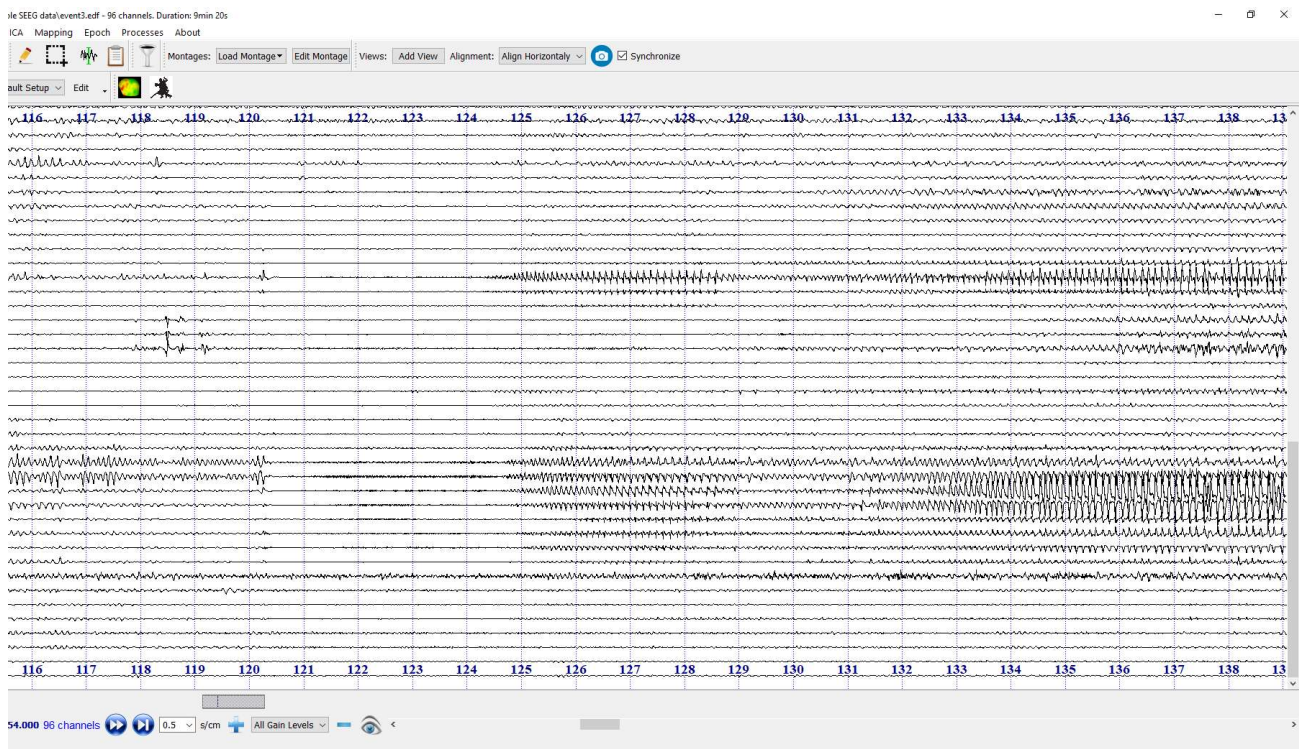
### Prerequisites

1. **MATLAB Runtime R2021a** - Download and install the windows version of MATLAB Runtime R2021a (9.10). from: <https://www.mathworks.com/products/compiler/matlab-runtime.html>
2. **Sample data** - Sample SEEG data and implantation images can be downloaded from publically available database of Brainstorm download. <https://neuroimage.usc.edu/bst/download.php>  
preprocessed data at: [click here](#)
3. **SEEG electrode location and naming using GARDEL** – GARDEL is a computational tool for automatic segmentation and labeling of SEEG electrode contacts (Medina Villalon et al., 2018). We

used GARDEL for 1) to coregister and localize SEEG electrodes in post-op MRI and 2) to export 3d coordinates of SEEG electrode contacts and brain model.

GARDEL can be downloaded from: <https://meg.univ-amu.fr/wiki/GARDEL:presentation>

- 4. Data preprocessing - AnyWave :** multi-platform software for visualizing and processing EEG/SEEG/MEG/XMG data (Colombet et al., 2015). We used AnyWave for preprocessing and excluding the channels other than SEEG / bad channels. The preprocessed data exported in EDF format. Later this file was imported in EPI-rank. Seizure 3 in the sample dataset will look like the below plot.



AnyWave can be downloaded from : <https://meg.univ-amu.fr/wiki/AnyWave>

# EPI-rank interface

After downloading the EPI-rank from <https://github.com/Brain-Mapping/EPI-rank>.

On a windows machine/computer, unzip, then double click on “epi\_rank.exe”. This open up the tool interface. EPI-rank tool interface has eight steps/sub-menus to localize EZ from SEEG.

epi\_rank

**1. Upload SEEG signal and electrode contact coordinates**  
Upload pruned SEEG a few minutes before the seizure initiation (15 mins). as .edf ; Upload the SEEG electrode MRI coordinates generated using GARDEL.

**2. Validate the channel details**

**3. Set Parameters**  
Filter power noise at 50Hz and 60 Hz

Theta low: 4 Hz, Alpha low: 8 Hz, Beta low: 13 Hz, Gamma low: 31 Hz  
Theta high: 7 Hz, Alpha high: 12 Hz, Beta high: 30 Hz, Gamma high: 90 Hz

**4. Select 'er' calculation**  
☐ er = sum(Beta low : Gamma high) ☐ er = sum(Theta low : Gamma high)  
☒ er = sum(Beta low : Gamma high) / sum (Alpha low : Alpha high)  
☐ er = sum(Beta high : Gamma high) / sum (Beta low: Beta high)

**5.** Slope: 15 (1 to 20) Threshold: 0.2  
Time window: 1 in seconds, value valid from 0.4 to 1.0  
er adaptation from: 1 to 100 in seconds.

**6. Set Epileptogenicity Rank (ER)**  
Epileptogenicity Rank: 7 takes value between 1 and 10

**7.** Detect Clear Quit

**8.** Save parameters Load parameters Locate EZ in patient brain model

**EPI-rank**

**Energy Ratio plot**

**Predicting seizure onset** onset order

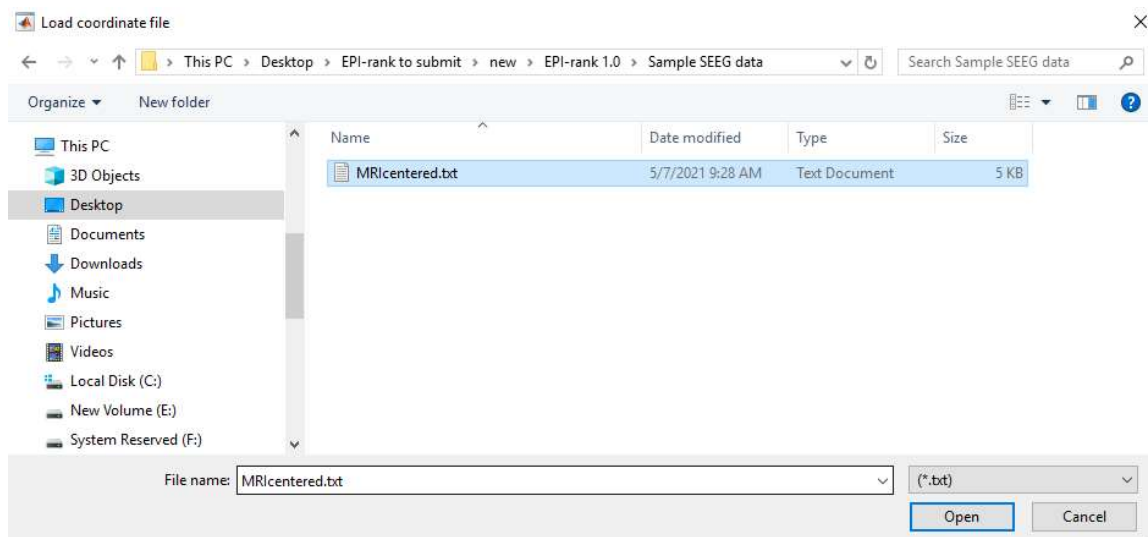
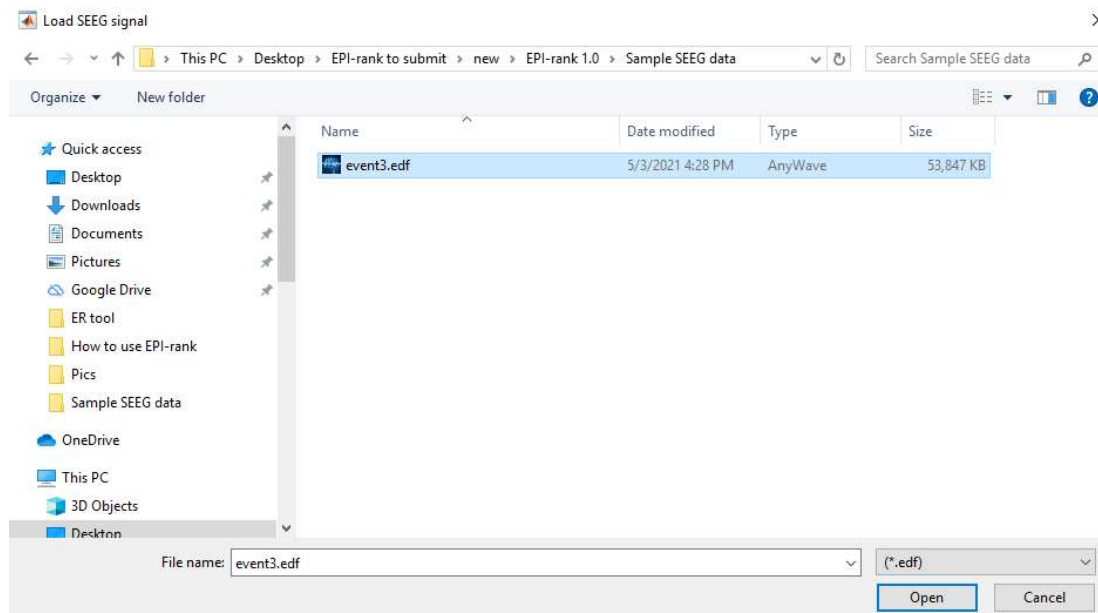
**Epileptic network**

Channel Name	Time	ER
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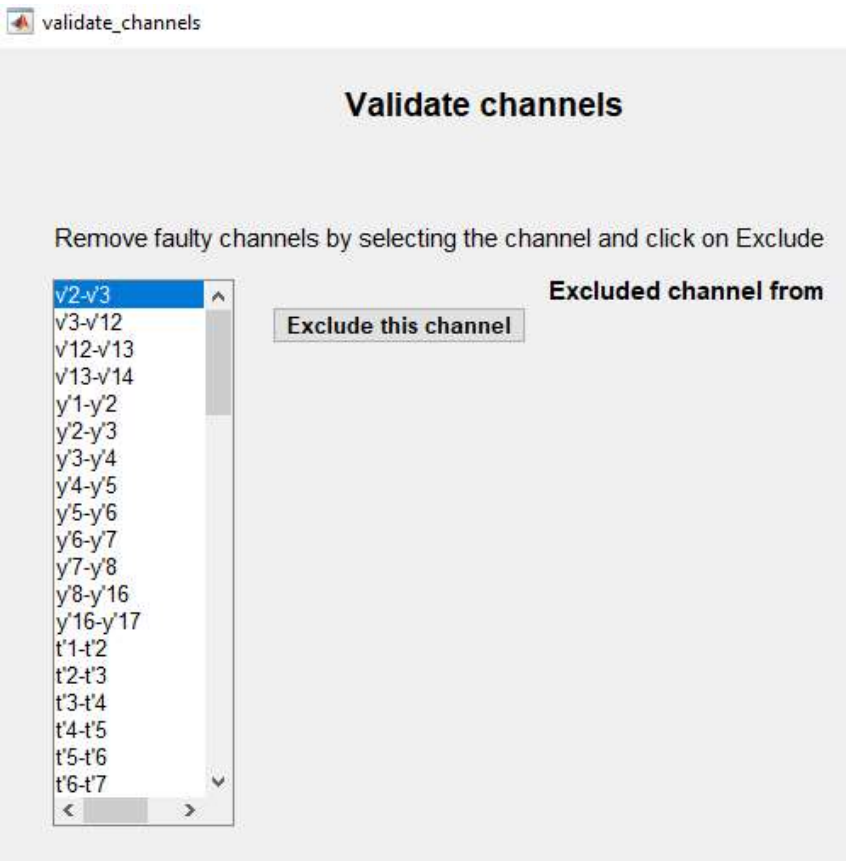
**Propagation network**

Channel Name	Time	ER
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**Step 1:** Click on “File upload” button to upload SEEG (data should be in “.edf”) and electrode location file (“MRIcentered.txt” generated using GARDEL). See how to use the GARDEL: <https://meg.univ-amu.fr/wiki/GARDEL:presentation>



**Step2:** On successful loading of SEEG and electrode location, a window/pop-up will appear “validate channels”. The SEEG channels were automatically converted to bipolar montage in this step. Using the “exclude this channel button” the user can exclude artifact channels from the calculation.



**Step 3:** This step removes the powerline noise from all SEEG channels. We can also set values for various EEG bands in this step. By clicking on “View SEEG”, the user can visualize the signal in bipolar montage.

3. Set Parameters

☒ Filter power noise at 50Hz and 60 Hz

Theta low

4

Hz

Alpha low

8

Hz

Beta low

13

Hz

Gamma low

31

Hz

Theta high

7

Hz

Alpha high

12

Hz

Beta high

30

Hz

Gamma high

90

Hz

**Step 4:** The user can select an equation for the calculation of energy ratio in this step.

4. Select 'er' calculation

☐  $er = \text{sum}(\text{Beta low} : \text{Gamma high})$  ☐  $er = \text{sum}(\text{Theta low} : \text{Gamma high})$

☒  $er = \text{sum}(\text{Beta low} : \text{Gamma high}) / \text{sum}(\text{Alpha low} : \text{Alpha high})$

☐  $er = \text{sum}(\text{Beta high} : \text{Gamma high}) / \text{sum}(\text{Beta low} : \text{Beta high})$

**Step 5:** Parameters including “time window”, “slope of energy ratio” and “energy ratio adaptation period” can be set in this step. er adaptation period usually set as an absolute period 1-2 minutes before the approximate seizure onset. Once all parameter set, then click on “Check detection” button that will compute the energy spectrum plot and each trace represents the energy spectrum of an SEEG. The interictal to ictal transition were colored in blue and red.

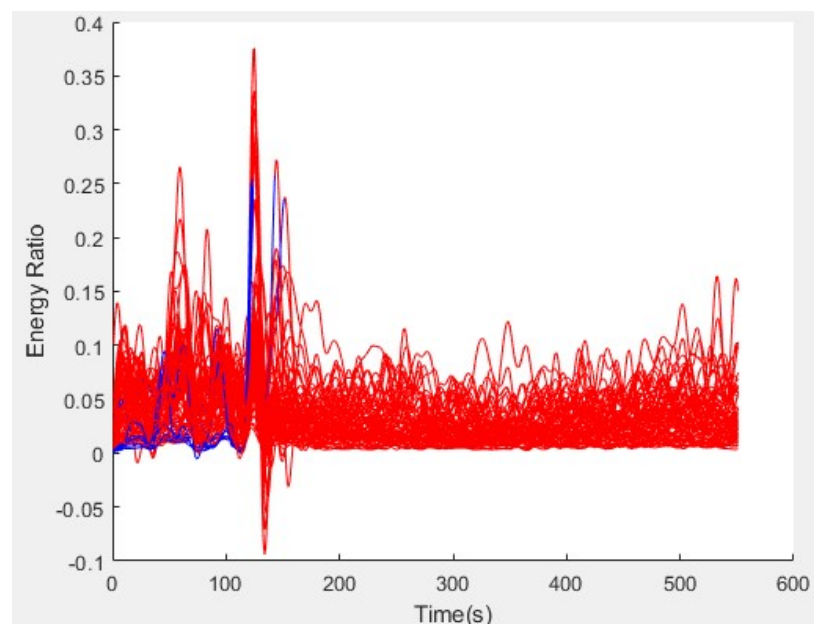
5.

Slope  (1 to 20) Threshold

Time window  in seconds, value valid from 0.4 to 1.0

er adaptation from  to  in seconds.

[Check detection](#)





**Step 6 and 7:** In this step, the user can set the threshold for epileptogenicity rank. Localize the EZ by clicking the “Detect” button. This will compute ER for all SEEG contacts and differentiate the seizure onset and from the propagation. Follow to that click on “Localize EZ in patient brain model” button to view the EZ localized in patient brain model. The user needs to load “mesh\_irm\_centered\_GARDEL.mat” file generated during SEEG electrode localization.

6. Set Epileptogenicity Rank (ER)

Epileptogenicity Rank  takes value between 1 and 10

7.

Detect

Clear

Quit

Predicting seizure onset

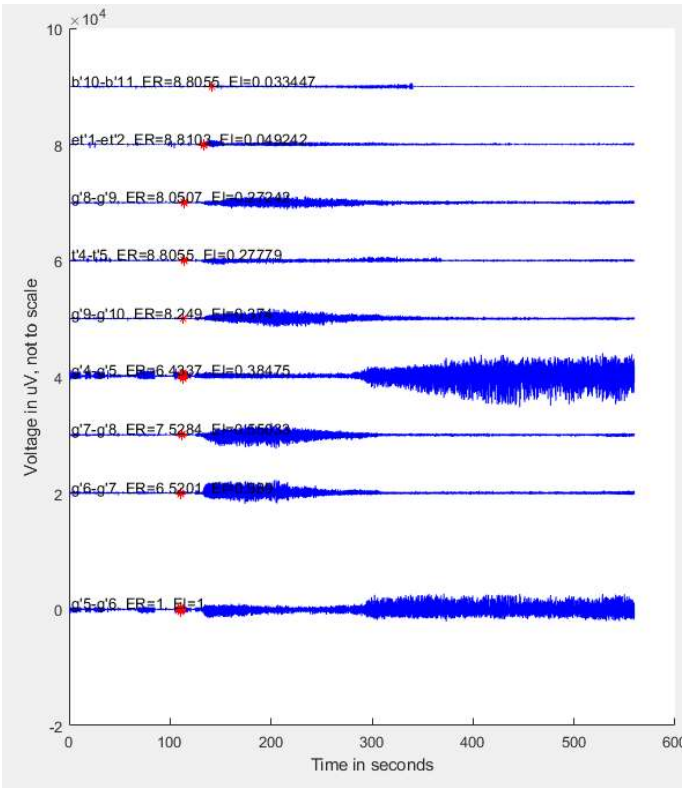
onset order

Epileptic network

Channel Name	Time	ER
g5-g6	16:29:52 (121)	1
g6-g7	16:29:52 (121)	6.5201
g7-g8	16:29:53 (122)	7.5284
g4-g5	16:29:54 (123)	6.4337

Propagation network

Channel Name	Time	ER
g9-g10	16:29:54 (123)	8.249
t4-t5	16:29:55 (124)	8.8055
g8-g9	16:29:55 (124)	8.0507
et1-et2	16:30:14 (143)	8.8103
b10-b11	16:30:22 (151)	8.8055



**Step 8:** The respective buttons save the parameters used for EZ localization by individual SEEGs.

8.

Save parameters

Load parameters

**Step9:** Reset the application by clicking “Clear” button and close the application using “Quit” button.

