

## 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.

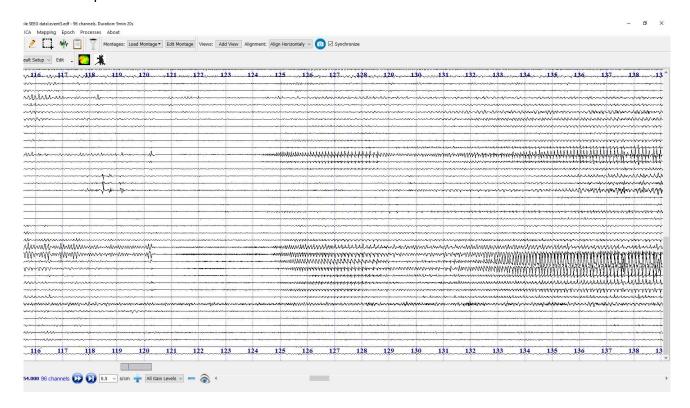
## **Prerequisites**

- **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. <a href="https://neuroimage.usc.edu/bst/download.php">https://neuroimage.usc.edu/bst/download.php</a> 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

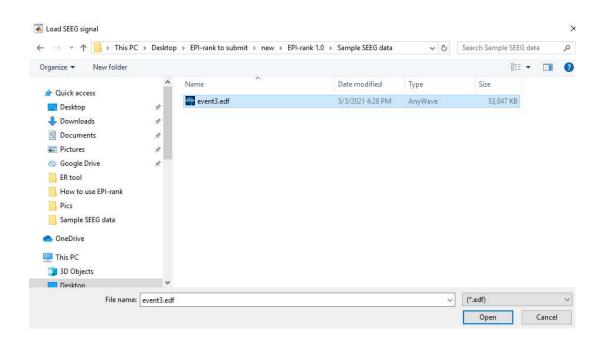
## **EPI-rank interface**

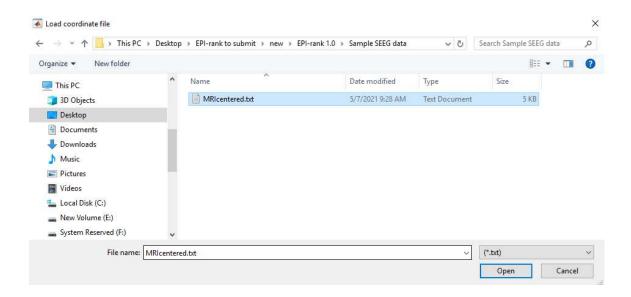
After downloading the EPI-rank from <a href="https://github.com/Brain-Mapping/EPI-rank">https://github.com/Brain-Mapping/EPI-rank</a>.

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.

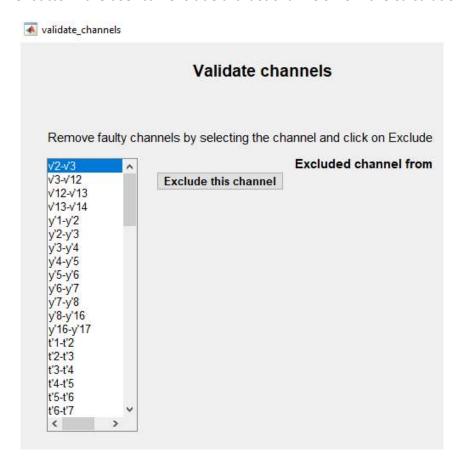
epi_rank				- □ ×
Upload	Upload SEEG signal and electrode contact coordinates pruned SEEG a few minutes before the seizure initiation (15 mins), as .edf; Upload the electrode MRI coordinates generated using GARDEL.	EPI-rank	Energy Ratio plot	
	File upload	Ci z-i din	0.6	
	Validate the channels  Set Parameters  Validate channels		0.4 -	
Theta low Theta high		Gamma low 31 Hz Gamma high 90 Hz	0 0.1 0.2 0.3 0.4 0.5 0.6 0.7  Predicting seizure onset	0.8 0.9 1
4.	Select 'er' calculation  or = sum(Beta low: Gamma high)  er = sum(Beta low: Gamma high) / sum (Alpha low: Alpha high)  er = sum(Beta low: Gamma high) / sum (Beta low: Beta high)	igh)	Epileptic network Channel Name Time ER	
5.	Slope 15 (1 to 20)  Time window 1 in seconds, value valid from 0.4 to 1.0 er adaptation from 1 to 100 in seconds.  Check detection	Threshold 0.2	Propagation network	v
6.	Set Epileptogenicity Rank (ER)		Channel Name Time ER	
7.	Epileptogenicity Rank 7 takes value between 1 and 10  Detect Clear Quit	View detection on SEEG		
8.	Save parameters Load parameters	ocate EZ in patient brain model		

**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: <a href="https://meg.univ-amu.fr/wiki/GARDEL:presentation">https://meg.univ-amu.fr/wiki/GARDEL:presentation</a>





**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.

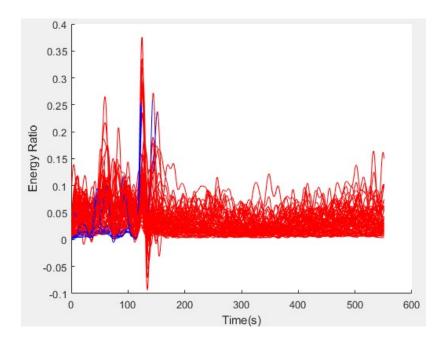


**Step 4:** The user can select an equation for the calculation of energy ratio in this step.

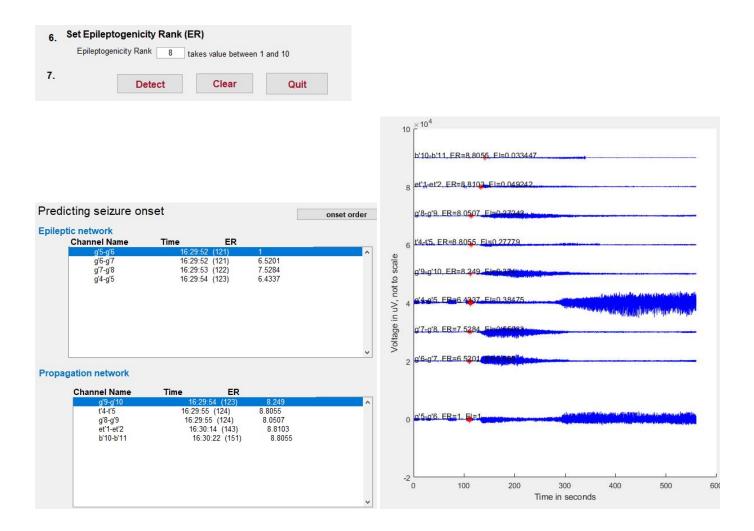
er = sum(Beta low : Gamma high) / sum (Alpha low : Alpha high)	

**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.

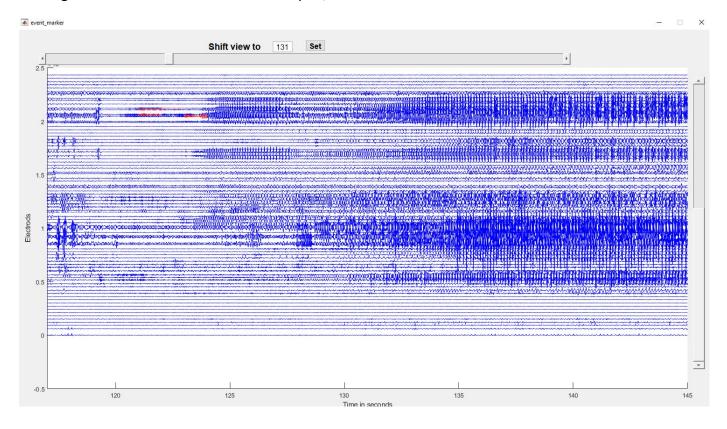
5.	Slope 15 (1 to 20) Threshold 0.2	
	Time window 1 in seconds, value valid from 0.4 to 1.0	
	er adaptation from 1 to 100 in seconds.	
	Check detection	



**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.



## Clicking on "view detection on SEEG" will open,



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



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

