

Package ‘EZFrangility’

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Title Compute Neural Frangility for Ictal iEEG Time Series

Version 2.1.1

Description Provides tools to compute the neural frangility matrix from intracranial electrocorticographic (iEEG) recordings, enabling the analysis of brain dynamics during seizures. The package implements the method described by Li et al. (2017) <doi:10.23919/ACC.2017.7963378> and includes functions for data preprocessing ('Epoch'), frangility computation ('calcAdjFrag'), and visualization.

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Depends R (>= 4.1.0), Epoch

LazyData true

Imports stats, methods, ggplot2 (>= 3.4.0), viridis, ggtext, glue, rlang, foreach, progress, ramify, reshape2

Suggests knitr, rmarkdown, testthat (>= 3.0.0), doSNOW, gsignal

Config/testthat/edition 3

VignetteBuilder knitr

URL <https://github.com/Jiefei-Wang/EZFrangility>

NeedsCompilation no

Author Jiefei Wang [aut, cre] (ORCID: <<https://orcid.org/0000-0002-2709-5332>>),
Patrick Karas [aut],
Anne-Cecile Lesage [aut] (ORCID:
<<https://orcid.org/0000-0002-9528-4899>>),
Ioannis Malagaris [aut] (ORCID:
<<https://orcid.org/0000-0001-5126-2068>>),
Oliver Zhou [aut],
Liliana Camarillo Rodriguez [aut] (ORCID:
<<https://orcid.org/0000-0001-8288-6885>>),
Sean O'Leary [aut] (ORCID: <<https://orcid.org/0000-0003-3650-705X>>),
Yuanyi Zhang [aut]

Maintainer Jiefei Wang <szwjf08@gmail.com>

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calcAdjFrag	<i>Calculate adjacency matrices and fragility matrix from iEEG recording</i>
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Description

The function calculates the neural fragility column from an adjacency matrix in each time window

Usage

```
calcAdjFrag(
  epoch,
  window,
  step,
  lambda = NULL,
  nSearch = 100L,
  progress = FALSE,
  parallel = FALSE
)
```

Arguments

epoch	Matrix or Epoch object. iEEG data matrix or Epoch object. If matrix, the row names are the electrode names and the column names are the time points. For a matrix input, the sampling rate is assumed to be 1 Hz and the start time is 0.
window	Integer. The number of time points to use in each window
step	Integer. The number of time points to move the window each time
lambda	Numeric. The lambda value for regularization to use in the ridge regression. If NULL, the lambda will be chosen automatically ensuring that ensuring that the adjacent matrix is stable (see details)
nSearch	Integer. Number of instable eigenvalues with norm=1 to search for the minimum norm perturbation. This parameter is used only when the lambda is NULL
progress	Logical. If TRUE, print progress information. If parallel is TRUE, this option only support the doSNOW backend.
parallel	Logical. If TRUE, use parallel computing. Users must register a parallel backend with the foreach package

Details

1/ For each time window i , a discrete stable Linear time system (adjacency matrix) is computed named A_i such that $A_i x(t) = x(t + 1)$. The 'lambda' option is the regularization parameter for the ridge regression. lambda=NULL(default) will find a lambda value that ensures the stability of the estimated A_i .

2/For each stable estimated A_i , the minimum norm perturbation Γ_{ik} (k index of the electrodes) for column perturbation is computed. Each column is normalized $\frac{\max(\Gamma_i) - \Gamma_{ik}}{\max(\Gamma_i)}$

Value

A Fragility object

Source

Recreation of the method described in Li A, Huynh C, Fitzgerald Z, Cajigas I, Brusko D, Jagid J, et al. Neural fragility as an EEG marker of the seizure onset zone. Nat Neurosci. 2021 Oct;24(10):1465–74 ([pubmed](#)). We have found solutions to fill up missing details in the paper method description

Examples

```
## A dummy example with 5 electrodes and 20 time points
data <- matrix(rnorm(100), nrow = 5)
## create an Epoch object
epoch <- Epoch(data, startTime = 0, samplingRate = 1)
windowNum <- 10
step <- 5
lambda <- 0.1
calcAdjFrag(
  epoch = epoch, window = windowNum,
```

```
    step = step, lambda = lambda, progress = TRUE
  )

## A more realistic example with parallel computing

if (requireNamespace("doSNOW")) {
  ## Register a SNOW backend with 4 workers
  library(parallel)
  library(doSNOW)
  cl <- makeCluster(4, type = "SOCK")
  registerDoSNOW(cl)

  data("pt01EcoG")
  window <- 250
  step <- 125
  title <- "PT01 seizure 1"
  calcAdjFrag(
    epoch = pt01EcoG, window = window,
    step = step, parallel = TRUE, progress = TRUE
  )

  ## stop the parallel backend
  stopCluster(cl)
}
```

checkIndex

Check and keep valid index only

Description

Check and keep valid index only

Usage

```
checkIndex(indices, names)
```

Arguments

indices	Numeric or character index to check
names	Character. All names corresponding to the indices

estimateSOZ	<i>Find Seizure Onset Zone</i>
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Description

The function estimates the seizure onset zone (SOZ). For each row, it calculates the maximum, minimum, or mean of row. The rows with the highest values are considered as the SOZ.

Usage

```
estimateSOZ(
  x,
  method = c("mean", "median", "max", "min"),
  proportion = 0.1,
  ...
)
```

Arguments

x	Fragility object
method	Character. The method to use to find the onset zone. Must be one of 'max', 'min', or "mean"
proportion	Numeric. The proportion of electrodes to consider as the onset zone. The electrode number will be rounded to the nearest integer.
...	Additional arguments

Value

A vector of electrode names, or indices if the electrode names are NULL

fragilityRow	<i>Compute the normalized fragility row for adjacency matrix A</i>
--------------	--

Description

The matrix A is used for the regression: $A * x(t) = x(t+1)$

Usage

```
fragilityRow(A, nSearch = 100, normalize = TRUE)
```

Arguments

A	Numeric. Adjacency Matrix
nSearch	Integer. Number of eigenvalues tried to find the minimum norm vector
normalize	Logical. If TRUE, the fragility row is normalized

fragStat	<i>Compute quantiles, mean and standard deviation for two electrodes groups</i>
----------	---

Description

Compute quantiles, mean and standard deviation for two electrodes groups

Usage

```
fragStat(frag, groupIndex = NULL, groupName = "SOZ", ranked = FALSE)
```

Arguments

frag	A Fragility object from calcAdjFrag
groupIndex	Integer or string. A group of electrodes to mark
groupName	Character. Name of the group of electrodes, default is "SOZ"
ranked	Logical. If TRUE, use the ranked fragility matrix from Fragility object

Value

list of 5 items with quantile matrix, mean and sdv from both electrodes groups

Examples

```
data("pt01Frag")
data("pt01EcoG")
## sozNames is the name of the electrodes we assume are in the SOZ
sozNames <- metaData(pt01EcoG)$sozNames
pt01fragstat <- fragStat(frag = pt01Frag, groupIndex = sozNames)
```

nrow,Fragility-method *Get the number of rows or columns of a Fragility object*

Description

Get the number of rows or columns of a Fragility object

Usage

```
## S4 method for signature 'Fragility'
nrow(x)

## S4 method for signature 'Fragility'
ncol(x)
```

Arguments

x A Fragility object

Value

- `nrow(x)`: The number of rows (electrodes) in the fragility matrix.
- `ncol(x)`: The number of columns (time points) in the fragility matrix.
- `dim(x)`: A vector of length 2 containing the number of rows and columns in the fragility matrix.

plot,Fragility,missing-method

Visualization functions (raw signal, fragility matrix)

Description

`plot`: plot fragility heatmaps with electrodes marked as soz colored

`plotFragQuantile`: Plot Fragility time quantiles for two electrodes groups

`plotFragQuantile`: Plot Fragility time distribution for two electrodes groups

Usage

```
## S4 method for signature 'Fragility,missing'
```

```
plot(  
  x,  
  y,  
  groupIndex = NULL,  
  maxLabels = 50,  
  ranked = FALSE,  
  x.lab.size = 10,  
  y.lab.size = 10  
)
```

```
plotFragQuantile(  
  frag,  
  groupIndex = NULL,  
  groupName = "SOZ",  
  x.lab.size = 10,  
  y.lab.size = 10  
)
```

```
plotFragDistribution(  
  frag,  
  groupIndex = NULL,  
  groupName = "SOZ",
```

```

bandType = c("SEM", "SD"),
rollingWindow = 1,
ranked = FALSE,
x.lab.size = 10,
y.lab.size = 10
)

```

Arguments

<code>x</code>	Fragility object from <code>calcAdjFrag</code>
<code>y</code>	Not used (for S4 method compatibility)
<code>groupIndex</code>	Integer or string. A group of electrodes to mark
<code>maxLabels</code>	Integer. Maximum number of labels to show on y-axis. Default is 50. The actual number of labels may be less than this value if there are too many electrodes.
<code>ranked</code>	Logical. If TRUE, use the ranked fragility matrix from Fragility object
<code>x.lab.size</code>	Numeric. Size of x-axis labels. Default is 4.
<code>y.lab.size</code>	Numeric. Size of y-axis labels. Default is 10
<code>frag</code>	Fragility object from <code>calcAdjFrag</code>
<code>groupName</code>	Character. Name of the group of electrodes, default is "SOZ"
<code>bandType</code>	Character. The type of band to use, either "SEM" or "SD". Default is "SEM".
<code>rollingWindow</code>	Integer. Window size for rolling average smoothing. Default is 1 (no smoothing).

Value

A ggplot object

Examples

```

data("pt01EcoG")

## sozNames is the name of the electrodes we assume are in the SOZ
sozNames <- metaData(pt01EcoG)$sozNames

## precomputed fragility object
data("pt01Frag")

## plot the fragility heatmap
plot(pt01Frag, groupIndex = sozNames)

## plot the fragility quantiles
plotFragQuantile(frag = pt01Frag, groupIndex = sozNames)

## plot the fragility distribution
plotFragDistribution(frag = pt01Frag, groupIndex = sozNames)

## plot with smoothing
plotFragDistribution(frag = pt01Frag, groupIndex = sozNames, rollingWindow = 2)

```

pt01EcoG

Pt01 seizure 1 around seizure onset

Description

This data corresponds to the first seizure of patient from the Fragility Data Set. EcoG recording gathered in collaboration with the National Institute of Health. The data contains only the good channels. It has been notch filtered and common average referenced in RAVE. The time range for full data is (-10:10s). Due to the size limit of the package, The full data has been epoched -1:2s around the seizure onset. The acquisition frequency is 1000 Hz

Usage

```
## EEG data
data(pt01EcoG)
```

Format

pt01EcoG: A Matrix with 84 rows (electrodes) and 3000 columns (time points)

pt01Frag: A fragility object results of applying the main function `calcAdjFrag` to pt01EcoG with `window = 250` and `step = 125`

Source

Fragility Multi-Center Retrospective Study ([OpenNeuro](#))

ridge

fit a generalized linear model to compute adjacency matrix A

Description

$$A x(t) = x(t+1)$$

Usage

```
ridge(xt, xtp1, lambda)
```

Arguments

xt	matrix. iEEG time series for a given window, with electrodes names as rows and time points as columns
xtp1	matrix. the iEEG time serie at the next time point, with electrodes names as rows and time points as columns
lambda	Numeric Vector. A user supplied lambda sequence.

Value

adjacency matrix A

ridgeR2	<i>computes R2</i>
---------	--------------------

Description

computes R2

Usage

ridgeR2(xt, xtp1, A)

Arguments

xt	matrix. iEEG time series for a given window, with electrodes names as rows and time points as columns
xtp1	matrix. the iEEG time serie at the next time point, with electrodes names as rows and time points as columns
A	adjacency matrix

ridgeSearch	<i>Ridge Regression for Electrode Readings</i>
-------------	--

Description

Ridge regression to compute matrix adjancency matrix A such as $A \cdot xt = xtp1$ the lambda parmeter is found by dichotomy such that A is stable (all eigenvalues have a norm less than one)

Usage

ridgeSearch(xt, xtp1, lambda = NULL)

Arguments

xt	matrix. iEEG time series for a given window, with electrodes names as rows and time points as columns
xtp1	matrix. the iEEG time serie at the next time point, with electrodes names as rows and time points as columns
lambda	Numeric Vector. A user supplied lambda sequence.

Value

adjacency matrix A_{fin} with lambda as attribute

show,Fragility-method *Print the Fragility object*

Description

Print the Fragility object

Usage

```
## S4 method for signature 'Fragility'  
show(object)
```

Arguments

object A Fragility object

Value

the object itself

show,FragStat-method *Print the FragStat object*

Description

Print the FragStat object

Usage

```
## S4 method for signature 'FragStat'  
show(object)
```

Arguments

object A FragStat object

Value

the object itself

[,Fragility-method *Subset a Fragility object*

Description

Subset a Fragility object

Usage

```
## S4 method for signature 'Fragility'
x[i, j, ..., drop = FALSE]
```

Arguments

x	A Fragility object
i	A logical vector or a numeric vector of indices to subset the electrodes
j	A logical vector or a numeric vector of indices to subset the time windows
...	Additional arguments (not used)
drop	Additional arguments (not used)

Value

A new Fragility object with the subsetted data

\$.FragStat-method *Getters and Setters for S4 object*

Description

Getters and Setters for S4 object

Usage

```
## S4 method for signature 'FragStat'
x$name

## S4 replacement method for signature 'FragStat'
x$name <- value

## S4 method for signature 'Fragility'
x$name

## S4 replacement method for signature 'Fragility'
x$name <- value
```

Arguments

x	S4 object
name	Slot name
value	Value to set

Value

S4 object itself or slot value

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