## Package 'neuroSCC'

March 27, 2025

Type Package

Title Bridging Simultaneous Confidence Corridors and PET Neuroimaging

Version 1.0.0

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**Description** Tools for the structured processing of PET neuroimaging data in preparation for the estimation of Simultaneous Confidence Corridors (SCCs) for one-group, two-group, or single-patient vs group comparisons. The package facilitates PET image loading, data restructuring, integration into a Functional Data Analysis framework, contour extraction, identification of significant results, and performance evaluation. It bridges established packages (e.g., 'oro.nifti') with novel statistical methodologies (e.g., 'ImageSCC') and enables reproducible analysis pipelines, including comparison with Statistical Parametric Mapping ('SPM').

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URL https://iguanamarina.github.io/neuroSCC/,

https://github.com/iguanamarina/PhD-2023-SCC-vs-SPM-Group-vs-Group, https://github.com/iguanamarina/PhD-2024-SCC-vs-SPM-SinglePatient-vs-Group

BugReports https://github.com/iguanamarina/neuroSCC/issues

**Depends** R (>= 4.2.0)

**Imports** contoureR, dplyr, graphics, memisc, oro.nifti, stats, tidyr, utils

Suggests BPST, ImageSCC, knitr, rcmdcheck, remotes, rmarkdown, roxygen2 (>= 7.1.0), Triangulation

Additional\_repositories https://iguanamarina.github.io/neuroSCC-drat

VignetteBuilder knitr

Encoding UTF-8

LazyData true

RoxygenNote 7.3.2

NeedsCompilation no

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**Repository** CRAN

Date/Publication 2025-03-27 18:00:00 UTC

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neuroSCC-package

Wrappers for Neuroimaging Functional Data Preparation and SCC Analysis

### Description

The neuroSCC package provides tools to preprocess and structure neuroimaging data for functional data analysis using Simultaneous Confidence Corridors (SCCs). It wraps external packages to prepare data from PET images, extract contours, generate meshes, and evaluate regions of statistical significance.

The methods implemented support both group comparisons and single-subject vs. group inference, following the methodology described in Wang et al. (2020) and the author's PhD thesis.

#### calculateMetrics

#### Details

This package serves as a bridge between neuroimaging file formats (e.g., NIfTI) and advanced statistical tools like ImageSCC::scc.image. It includes the following key components.

- Loading and cleaning PET image data.
- Extracting ROIs and constructing functional data matrices.
- Generating synthetic Poisson clones for 1-vs-group settings.
- Extracting SCC-detected points and evaluating detection metrics.

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#### See Also

neuroCleaner, databaseCreator, getPoints

calculateMetrics Evaluate SCC or SPM Detection Performance

#### Description

Computes Sensitivity, Specificity, Positive Predictive Value (PPV), and Negative Predictive Value (NPV) by comparing detected points with ground truth ROI points. This function is used to assess the accuracy of SCC- or SPM-based detection in neuroimaging analysis.

#### Usage

```
calculateMetrics(detectedPoints, truePoints, totalCoords, regionName)
```

#### Arguments

detectedPoints	A data frame containing detected coordinates (x, y). SCC-detected points should be obtained using getPoints. SPM-detected points should be obtained using getSPMbinary.
truePoints	A data frame with ground truth ROI coordinates $(x, y)$ , extracted via processROIs.
totalCoords	A list with the full voxel grid dimensions, created by getDimensions. Must include named elements xDim and yDim.
regionName	A character string used to label the output region.

This function requires three precomputed objects

- detectedPoints: SCC-detected points from getPoints or SPM-detected points from getSPMbinary.
- truePoints: Ground truth ROIs extracted using processROIs.
- totalCoords: Full voxel coordinate grid from getDimensions.

#### Value

A data frame with the following evaluation metrics

- region: Name of the analyzed region.
- sensitivity: True positive rate (TP / (TP + FN) \* 100).
- specificity: True negative rate (TN / (TN + FP) \* 100).
- PPV: Positive predictive value (TP / (TP + FP) \* 100).
- NPV: Negative predictive value (TN / (TN + FN) \* 100).

#### See Also

getPoints for SCC-detected regions. getSPMbinary for binary SPM-detected points. processROIs for defining ground truth ROIs. getDimensions for generating the coordinate grid.

#### Examples

```
# Load precomputed inputs for the example
data("calculateMetricsExample", package = "neuroSCC")
# Evaluate SCC and SPM detection performance
with(calculateMetricsExample, {
    metricsSCC <- calculateMetrics(detectedSCC, trueROI, totalCoords, "Region2_SCC")
    metricsSPM <- calculateMetrics(detectedSPM, trueROI, totalCoords, "Region2_SPM")
    print(metricsSCC)
    print(metricsSPM)
})
```

calculateMetricsExample

Precomputed Inputs for SCC vs. SPM Performance Evaluation

#### databaseCreator

#### Description

A dataset containing all necessary inputs for demonstrating calculateMetrics. It enables reproducible and fast example code that compares SCC-detected and SPM-detected points against a known ground truth ROI.

These inputs were generated using sample PET and ROI files included in the neuroSCC package.

#### Usage

```
data("calculateMetricsExample")
```

#### Format

A set of four objects:

detectedSCC Data frame of SCC-detected coordinates (from getPoints). detectedSPM Data frame of SPM-detected coordinates (from getSPMbinary). trueROI Ground truth ROI voxel data (from processROIs). totalCoords List with full image grid dimensions (from getDimensions).

#### Source

Simulated PET neuroimaging study for testing SCC and SPM detection accuracy.

#### See Also

calculateMetrics, getPoints, getSPMbinary, processROIs, getDimensions

databaseCreator Create a Database of Processed PET Image Data

#### Description

Processes multiple PET image files matching a specified filename pattern. Each file is processed using neuroCleaner, and the results are aggregated into a unified data frame for functional data analysis. This function serves as a key step in the neuroSCC workflow, bridging raw image data and Simultaneous Confidence Corridors (SCC) computation.

#### Usage

```
databaseCreator(
   pattern,
   control = TRUE,
   useSequentialNumbering = FALSE,
   demo = NULL,
   quiet = FALSE
)
```

#### Arguments

pattern	character. A regular expression defining the file pattern to match. Subject identifiers are extracted from filenames based on this pattern.
control	logical. If TRUE, files are treated as control group data; if FALSE, as pathological group data. Default is TRUE.
useSequentialNumbering	
	logical. If TRUE, assigns sequential subject numbers instead of extracting them from filenames. Default is FALSE.
demo	data.frame, optional. If provided, demographic information is included for each file. Default is NULL.
quiet	logical. If TRUE, suppresses progress messages. Default is FALSE.

#### Details

The function performs the following steps

- 1. Identifies image files matching the given pattern.
- 2. Processes each file using neuroCleaner, optionally merging demographic data.
- 3. Adds a subject identifier column (CN\_number or AD\_number).
- 4. Aggregates all results into a single data frame.

If no files are successfully processed, an empty data frame is returned with a warning.

This function is typically followed by matrixCreator, which converts the output into a matrix format for functional analysis.

#### Value

A data.frame combining processed voxel-level data from all matched files. Each row represents a voxel (3D pixel). The column structure depends on input

- For the control group: CN\_number, z, x, y, pet
- For the pathological group: AD\_number, z, x, y, pet
- If demographics are included: additional columns PPT, Group, Sex, Age

#### See Also

neuroCleaner for the underlying image processing function.

matrixCreator for the next step in the workflow that converts the database to a matrix format for SCC analysis.

#### Examples

- # NOTE: To keep runtime below CRAN limits, this example processes only 1 subject.
- # You can expand the pattern to include all subjects for real use.

```
# Example: Create a database from a single synthetic PET image (control group)
controlPattern <- "^syntheticControl1\\.nii\\.gz$"</pre>
```

#### generatePoissonClones

```
databaseControls <- databaseCreator(pattern = controlPattern, control = TRUE, quiet = TRUE)
head(databaseControls)</pre>
```

generatePoissonClones Generate Synthetic Poisson Clones for PET Data

#### Description

Generates synthetic clones of a PET data matrix by adding Poisson-distributed noise to each nonzero voxel. This approach helps address the limitations of functional data analysis (FDA) in singlesubject versus group (1 vs. Group) setups, where a single subject lacks sufficient variability to reliably estimate Simultaneous Confidence Corridors (SCCs).

#### Usage

```
generatePoissonClones(originalMatrix, numClones, lambdaFactor)
```

#### Arguments

originalMatrix	A numeric matrix where each row represents a flattened PET image.
numClones	An integer specifying the number of synthetic clones to generate.
lambdaFactor	A positive numeric value that scales the magnitude of Poisson noise.

#### Details

- Values equal to 0 remain unchanged to preserve background regions.
- NA values are replaced with 0 before adding noise.
- Poisson noise is applied only to positive values, scaled by lambdaFactor.
- Enables valid SCC estimation in single-subject settings by artificially increasing sample size.

#### Value

A numeric matrix with numClones rows, each representing a noisy version of originalMatrix with Poisson noise added.

#### Examples

```
# Load example input matrix for Poisson cloning
data("generatePoissonClonesExample", package = "neuroSCC")
# Select 10 random voxel positions for display
set.seed(123)
sampledCols <- sample(ncol(generatePoissonClonesExample), 10)
# Generate 1 synthetic clone
clones <- generatePoissonClones(generatePoissonClonesExample, numClones = 1, lambdaFactor = 0.25)
# Show voxel intensity values after cloning
clones[, sampledCols]
```

generatePoissonClonesExample

Example Input for Poisson Clone Generation

#### Description

A full single-subject PET matrix used to demonstrate generatePoissonClones. This matrix was extracted from simulated neuroimaging data included in the neuroSCC package.

The example avoids long runtime by generating only one synthetic clone.

#### Usage

```
data("generatePoissonClonesExample")
```

#### Format

A numeric matrix named generatePoissonClonesExample, with 1 row and all voxel columns.

#### Source

Simulated PET neuroimaging dataset included with neuroSCC.

#### See Also

generatePoissonClones

getDimensions

Get Dimensions from a Neuroimaging File

#### Description

Extracts voxel dimension information from a NIfTI or similar neuroimaging file. This function is designed to work with neuroCleaner, but it can also be used independently to inspect image dimensions.

#### Usage

getDimensions(file)

#### Arguments

file

A NIfTI file object or a file path pointing to a NIfTI image.

#### getPoints

#### Details

The function accepts either a file path or a preloaded nifti object. If a file path is provided, it uses oro.nifti::readNIfTI() to load the image. This function ensures consistent dimension extraction across the neuroSCC pipeline.

#### Value

A named list with the following elements

- xDim Number of voxels along the X axis.
- yDim Number of voxels along the Y axis.
- zDim Number of slices along the Z axis.
- dim Total number of voxels in a 2D slice (calculated as xDim \* yDim).

#### Examples

```
# Get the file path for a sample NIFTI file
niftiFile <- system.file("extdata", "syntheticControl1.nii.gz", package = "neuroSCC")
# Extract dimensions from the NIFTI file
dimensions <- getDimensions(niftiFile)
# Display the extracted dimensions
print(dimensions)
```

getPoints

Extract Significant SCC Points from an SCC Comparison Object

#### Description

Identifies and extracts coordinates where differences fall outside the simultaneous confidence corridors (SCCs), indicating statistically significant regions. This function processes the results from ImageSCC::scc.image() and returns the voxel locations that represent either hypo- or hyperactivity.

Interpretation depends on the order of inputs in the SCC computation. If SCC was computed as  $scc.image(Ya = Y_AD, Yb = Y_CN, ...)$  (i.e., the Control group is the second argument).

- positivePoints Regions where Control minus Pathological is significantly above the SCC. These correspond to areas where the Pathological group (AD) is *hypoactive* relative to Controls.
- negativePoints Regions where Control minus Pathological is significantly below the SCC. These correspond to areas where the Pathological group is *hyperactive* relative to Controls.

Always confirm the order of Ya and Yb in the SCC computation to interpret the directionality correctly.

#### Usage

getPoints(sccResult)

#### Arguments

sccResult	A list of SCC computation results, as returned by ImageSCC::scc.image. Must
	include the following components
	• Z.band — A matrix specifying grid positions.

- ind.inside.cover Indices of grid points within the confidence band.
- scc A 3D array containing the computed SCC values.

#### Value

A named list with two elements

- positivePoints A data frame with coordinates where the **first group** (Ya) shows significantly lower activity than the **second group** (Yb).
- negativePoints A data frame with coordinates where the **first group** (Ya) shows significantly higher activity than the **second group** (Yb).

#### See Also

ImageSCC::scc.image for SCC computation.

#### Examples

```
# Load precomputed SCC example
data("SCCcomp", package = "neuroSCC")
# Extract significant SCC points
significantPoints <- getPoints(SCCcomp)
# Show extracted points (interpretation depends on SCC setup; see description)
head(significantPoints$positivePoints) # Pathological hypoactive vs. Control
head(significantPoints$negativePoints) # Pathological hyperactive vs. Control</pre>
```

getSPMbinary

Extract SPM-Detected Significant Points from a Binary NIfTI File

#### Description

Extracts voxel coordinates where pet = 1 (i.e., statistically significant points) from a binary NIfTI file produced by an external SPM analysis. Only voxels from a specific brain slice (z = paramZ) are retained.

The output data frame is structured identically to that of getPoints, allowing direct comparison between SCC- and SPM-detected regions via calculateMetrics.

#### matrixCreator

#### Usage

```
getSPMbinary(niftiFile, paramZ = 35)
```

#### Arguments

niftiFile	character. The path to the binary NIfTI file generated by SPM.
paramZ	integer. The specific z-slice to extract. Default is 35.

#### Details

This function converts externally generated SPM results into a format compatible with SCC analysis tools in neuroSCC. Use getDimensions to inspect the full coordinate space if needed.

#### Value

A data frame with the following columns:

• x, y – Coordinates of significant voxels at the specified slice.

#### See Also

getPoints for SCC-based detection. getDimensions for obtaining full coordinate grids. calculateMetrics for evaluating SCC vs. SPM detection performance.

#### Examples

```
# Load a sample binary NIFTI file (SPM result)
niftiFile <- system.file("extdata", "binary.nii.gz", package = "neuroSCC")
detectedSPM <- getSPMbinary(niftiFile, paramZ = 35)</pre>
```

# Show detected points
head(detectedSPM)

matrixCreator Convert Database to Functional Data Matrix Format

#### Description

Converts a PET image database (created via databaseCreator) into a matrix format suitable for functional data analysis. Each row of the resulting matrix corresponds to a subject, and each column to a voxel's PET intensity values at a specified brain slice.

#### Usage

```
matrixCreator(
   database,
   paramZ = 35,
   useSequentialNumbering = FALSE,
   quiet = FALSE
)
```

#### Arguments

database	A data frame created by databaseCreator, containing voxel-level PET image data, including subject identifiers, coordinates, and intensity values.
paramZ	An integer specifying the z-coordinate (slice) to extract. Default is 35.
useSequentialNumbering	
	logical. If TRUE, assigns sequential subject IDs instead of extracting them from filenames. Not currently used inside this function. Default is FALSE.
quiet	logical. If TRUE, suppresses progress messages. Default is FALSE.

#### Details

This function performs the following steps

- 1. Verifies that the specified z-slice exists in the database.
- 2. Identifies the correct subject grouping column (CN\_number or AD\_number).
- 3. Determines the matrix dimensions using x and y coordinates.
- 4. Extracts PET intensities per subject at the given slice.
- 5. Replaces any NaN values with 0 to ensure numerical stability.

This function typically follows databaseCreator and precedes meanNormalization in the neuroSCC workflow.

#### Value

A numeric matrix where

- Each row represents one subject's PET values at the selected z-slice.
- Each column corresponds to a voxel (flattened as a 1D row).

#### See Also

databaseCreator for generating the input database. meanNormalization for scaling matrix data prior to SCC computation.

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#### meanNormalization

#### Examples

```
# NOTE: To keep example runtime short, only one synthetic PET file is used.
# For full analysis, expand the filename pattern accordingly.
# Step 1: Generate a database for a single subject
controlPattern <- "^syntheticControl1\\.nii\\.gz$"
databaseControls <- databaseCreator(pattern = controlPattern, control = TRUE, quiet = TRUE)
# Step 2: Convert the database into a matrix format
matrixControls <- matrixCreator(databaseControls, paramZ = 35, quiet = TRUE)
# Display dimensions of the matrix
dim(matrixControls)
```

meanNormalization Mean Normalization for Matrix Data

#### Description

Normalizes each row of a matrix by dividing its elements by the row mean, ignoring NA values. This step is commonly used to adjust for global intensity differences across subjects before applying statistical comparisons or functional data analysis.

#### Usage

```
meanNormalization(
  matrixData,
  handleInvalidRows = c("warn", "error", "omit"),
  returnDetails = FALSE,
  quiet = FALSE
)
```

#### Arguments

matrixData	A matrix where each row represents one subject's PET data, typically generated by matrixCreator.
handleInvalidRo	ws
	character. Specifies how to handle rows with invalid means (either zero or NA). Options include "warn" (default), "error", or "omit".
returnDetails	logical. If TRUE, returns a list with the normalized matrix and additional diagnostics. If FALSE (default), returns only the normalized matrix.
quiet	logical. If TRUE, suppresses console messages. Default is FALSE.

#### Details

The function performs the following steps

- 1. Computes the row means of the input matrix, ignoring NAs.
- 2. Divides each row by its corresponding mean.
- 3. Replaces NaN values (from division by 0) with 0 if applicable.
- 4. Handles problematic rows according to the selected handleInvalidRows option: "warn" (default) issues a warning, "error" stops execution, and "omit" removes the affected rows from the result.

This step is often used prior to applying SCC methods to ensure comparability across subjects.

#### Value

A normalized matrix, or a list if returnDetails = TRUE.

- normalizedMatrix The normalized matrix.
- problemRows Indices of rows that had zero or NA means.

#### See Also

matrixCreator for building the matrix input to normalize.

#### Examples

neuroCleaner

Clean and Load Data from NIfTI Neuroimaging Files

#### Description

Loads a NIfTI-format neuroimaging file and transforms it into a structured data frame, organizing voxel-level information for downstream analysis. This function is the first step in the neuroimaging processing pipeline in neuroSCC, converting raw PET data into a format suitable for functional data analysis. SCCs are later computed using functions from the ImageSCC package, such as ImageSCC::scc.image().

#### neuroCleaner

#### Usage

neuroCleaner(name, demo = NULL, demoRow = 1)

#### Arguments

name	character. The full path to the NIfTI file to process.
demo	Optional data.frame containing demographic information. If provided, it should include columns (case-insensitive): PPT, Group, Sex, and Age. If automatic matching via the PPT column fails, the row specified by demoRow is used. Default is NULL.
demoRow	integer. Row to use from the demographic table if automatic matching fails. Default is 1.

#### Details

The function performs the following steps

- 1. Loads the NIfTI file using oro.nifti::readNIfTI().
- 2. Converts the 3D image into a tidy data frame.
- 3. Adds z, x, and y voxel coordinates.
- 4. If demographic data is provided, attempts to match based on PPT (case-insensitive). If no match is found, demoRow is used.

The resulting data frame serves as input for databaseCreator, matrixCreator, and other core functions in the neuroSCC pipeline.

#### Value

A data frame where each row represents a voxel (3D pixel).

- If demographics are provided: the columns include PPT, Group, Sex, Age, z, x, y, and pet.
- If demographics are not provided: the columns include z, x, y, and pet.

The pet column contains the PET intensity value at each voxel location.

#### See Also

```
databaseCreator for batch image processing.
readNIfTI for reading NIfTI-format files.
```

#### Examples

```
# Load a sample Control NIfTI file
niftiFile <- system.file("extdata", "syntheticControl1.nii.gz", package = "neuroSCC")
# Example Without demographic data
petData <- neuroCleaner(niftiFile)
petData[sample(nrow(petData), 10), ] # Show 10 random voxels</pre>
```

```
neuroContour
```

#### Description

This function extracts contours from a neuroimaging NIFTI file where values change according to specified levels. It processes the NIFTI file with neuroCleaner to extract structured neuroimaging data, then extracts contours using contoureR::getContourLines. These contours serve as input for Triangulation::TriMesh, which is used in Simultaneous Confidence Corridors (SCCs) calculations.

While **not mandatory**, it is **highly recommended** that the input NIFTI file be pre-processed such that zero values represent the background and non-zero values represent regions of interest. The function's default behavior extracts contours at level 0, which is ideal for well-masked data.

#### Usage

```
neuroContour(niftiFile, paramZ = 35, levels = c(0), plotResult = FALSE)
```

#### Arguments

niftiFile	character, the path to the NIFTI file containing neuroimaging data. Ideally, the file should be masked so that zero values represent the background.
paramZ	integer, the specific z-slice to extract contours from. Default is 35.
levels	numeric, a vector of levels at which to draw the contours. Default is $c(0)$ .
plotResult	logical, if TRUE, plots the extracted contours. Default is FALSE.

#### Details

This function extracts contours from a **NIFTI** file, typically a **masked** image where background values are set to zero, and regions of interest contain non-zero values. While users can specify a different boundary level, the recommended approach is to use levels = 0 for masked data.

The extracted contours are typically used as input to Triangulation::TriMesh to create a triangular mesh of the region, which is then used for Simultaneous Confidence Corridors calculations.

#### Value

A list of data frames, where each data frame contains the x and y coordinates of a contour. The first element typically represents the external boundary, while subsequent elements (if present) represent internal contours or holes. Each data frame has two columns:

- x x-coordinates of the contour points.
- y y-coordinates of the contour points.

#### See Also

getContourLines for the underlying contour extraction. Triangulation::TriMesh for the next step in the SCC calculation process.

#### processROIs

#### Examples

```
# Get the file path for a sample NIFTI file
niftiFile <- system.file("extdata", "syntheticControl1.nii.gz", package = "neuroSCC")
# Extract contours at level 0
contours <- neuroContour(niftiFile, paramZ = 35, levels = 0, plotResult = TRUE)
# Display the first few points of the main contour
head(contours[[1]])
```

processROIs

Process ROI Voxel Data from a NIfTI File

#### Description

Processes Regions of Interest (ROIs) from a binary NIfTI file by extracting voxel-level coordinates and labeling each voxel as part of the ROI or not. The function preserves the spatial structure and is typically used to prepare ground truth ROIs for comparison with SCC-detected regions via calculateMetrics.

#### Usage

```
processROIs(
   roiFile,
   region,
   number,
   save = TRUE,
   outputDir = tempdir(),
   verbose = TRUE
)
```

#### Arguments

roiFile	character. Path to the binary NIfTI file containing ROI data.
region	character. Name of the ROI region (e.g., "Region2").
number	character. Identifier for the subject or group (e.g., "18").
save	logical. If TRUE, saves the result as an .RDS file. If FALSE, returns a data frame in the console. Default is TRUE.
outputDir	character. Directory where the ROI table will be saved if save = TRUE. Default is a temporary file: tempdir().
verbose	logical. If TRUE, displays progress messages. Default is TRUE.

#### Details

The function uses neuroCleaner to load and flatten the NIfTI file into a structured data frame. All voxels are retained, with the pet column indicating which ones are part of the ROI (1) versus background ( $\emptyset$ ). An ROI label is added in the group column.

This output is used as ground truth for evaluating detection performance in SCC analyses.

#### Value

A data frame with voxel-level ROI information.

- group Combined identifier built from region and number.
- z, x, y Voxel coordinates.
- pet Binary value indicating ROI membership (1 = ROI, 0 = non-ROI).

If save = TRUE, the data frame is saved as an .RDS file and not returned to the console.

#### See Also

calculateMetrics for evaluating SCC detection performance. neuroCleaner for reading and structuring voxel data.

#### Examples

# Load and process a sample ROI NIFTI file (console output)

SCCcomp

Example SCC Computation Result

#### Description

A precomputed example of a Simultaneous Confidence Corridor (SCC) analysis comparing a group of pathological subjects against controls. This object was generated using the ImageSCC::scc.image function and represents a realistic output from SCC-based neuroimaging group comparisons.

This dataset is used in the examples of getPoints and calculateMetrics, allowing users to explore SCC outputs without needing to recompute them.

#### Usage

data("SCCcomp")

#### SCCcomp

#### Format

A named list of class "image" with the following elements

scc 3D array of SCC confidence bands, dimensions [n, 2, alpha].

Z.band Matrix of grid coordinates corresponding to evaluated locations.

ind.inside.cover Integer vector of indices for grid points inside the SCC band.

V.est.a, V.est.b Vertex matrices for triangulated domains (pathological and control groups).

Tr.est.a, Tr.est.b Triangle index matrices corresponding to the domain meshes.

alpha Vector of confidence levels used (e.g., 0.1, 0.05, 0.01).

d.est Spline degree used in mean function estimation.

r Smoothing parameter used during fitting.

#### Source

Simulated PET neuroimaging study for evaluating SCC methodology.

#### See Also

getPoints, calculateMetrics, ImageSCC::scc.image

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