Package ‘consICA’

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Type Package

biocViews Technology, StatisticalMethod, Sequencing, RNASEq, Transcriptomics, Classification, FeatureExtraction

Title consensus Independent Component Analysis

Version 2.0.0

Description consICA implements a data-driven deconvolution method – consensus independent component analysis (ICA) to decompose heterogeneous omics data and extract features suitable for patient diagnostics and prognostics.

The method separates biologically relevant transcriptional signals from technical effects and provides information about the cellular composition and biological processes.

The implementation of parallel computing in the package ensures efficient analysis of modern multicore systems.

BugReports https://github.com/biomod-lih/consICA/issues

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Encoding UTF-8

LazyData false

Imports fastICA (>= 1.2.1), sm, org.Hs.eg.db, GO.db, stats, SummarizedExperiment, BiocParallel, graph, ggplot2, methods, Rfast, pheatmap, survival, topGO, graphics, grDevices

Depends R (>= 4.2.0)

Suggests knitr, BiocStyle, rmarkdown, testthat, Seurat

VignetteBuilder knitr

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anovaIC ANOVA test for independent component across factors

Description

ANOVA (ANalysis Of VAriance) test produced for specific independent component across each
(clinical) factor as 'aov(IC ~ factor)'. Plot distributions of samples’ weight for top 9 significant
factors.

Usage

anovaIC(
cica,
Var = NULL,
icomp = 1,
plot = TRUE,
mode = "violin",
        color_by_pv = TRUE
    )

Arguments

cica list compliant to ‘consICA()’ result
Var matrix with samples’ metadata. Samples in rows and factors in columns
icomp number of component to analyse
plot if plot weights distributions for top factors
mode type of plot. Can be ‘violin’ or ‘box’
color_by_pv if TRUE plots will be colored by p-value ranges

Value

a data.frame with

factor name of factor
p.value p-value for ANOVA test for factor
p.value_disp string for p-value printing

Examples

data("samples_data")
Var <- data.frame( SummarizedExperiment::colData(samples_data))
cica <- consICA(samples_data, ncomp=10, ntry=1, ncores=1, show.every=0)
# Run ANOVA for 4th independent component
anova <- anovaIC(cica, Var=Var, icomp = 4)

consICA **Calculate consensus Independent Component Analysis**

description

calculate consensus independent component analysis (ICA) Implements efficient ICA calculations.

Usage

consICA(
    X,
    ncomp = 10,
    ntry = 1,
    show.every = 1,
    filter.thr = NULL,
    ncores = 1,
    bpparam = NULL,
consICA

```r
reduced = FALSE,
fun = "logcosh",
alg.typ = "deflation",
verbose = FALSE,
assay_string = NULL
)
```

**Arguments**

- **X**: input data with features in rows and samples in columns. Could be a ‘SummarizedExperiment’ object, matrix or ‘Seurat’ object. For ‘SummarizedExperiment’ with multiple assays or ‘Seurat’ pass the name with ‘assay_string’ parameter, otherwise the first will be taken. See `SummarizedExperiment-class`
- **ncomp**: number of components
- **ntry**: number of consensus runs. Default value is 1
- **show.every**: numeric logging period in iterations (disabled for ‘ncore’s > 1). Default value is 1
- **filter.thr**: Filter out genes (rows) with max value lower than this value from ‘X’
- **ncores**: number of cores for parallel calculation. Default value is 4
- **bpparam**: parameters from the ‘BiocParallel’
- **reduced**: If TRUE returns reduced result (no ‘X’, ‘i.best’, see ‘return’)
- **fun**: the functional form of the G function used in the approximation to neg-entropy in fastICA. Default value is "logcosh"
- **alg.typ**: parameter for fastICA(). If alg.typ == "deflation" the components are extracted one at a time. If alg.typ == "parallel" the components are extracted simultaneously. Default value is "deflation"
- **verbose**: logic TRUE or FALSE. Use TRUE for print process steps. Default value is FALSE
- **assay_string**: name of assay for ‘SummarizedExperiment’ or ‘Seurat’ input object ‘X’. Default value is NULL

**Value**

A list with

- **X**: input object
- **nsamples, nfeatures**: dimension of X
- **S, M**: consensus metagene and weight matrix
- **ncomp**: number of components
- **X_num**: input data in matrix format
- **mr2**: mean R2 between rows of M
- **stab**: stability, mean R2 between consistent columns of S in multiple tries. Applicable only for ‘ntry’ > 1
- **i.best**: number of best iteration
coreICA

Author(s)
Petr V. Nazarov

See Also
fastICA

Examples

data("samples_data")
# Deconvolve into independent components
cica <- consICA(samples_data, ncomp=15, ntry=10, ncores=1, show.every=0)
# X = S * M, where S - independent signals matrix, M - weights matrix
dim(samples_data)
dim(cica$S)
dim(cica$M)

Description

Adaptation of fastICA for quick multiple-run calculations for consensus Independent Component Analysis (ICA)

Usage

coreICA(
  X,
  n.comp,
  preICA = NULL,
  alg.typ = c("parallel", "deflation"),
  fun = c("logcosh", "exp"),
  w.init = NULL,
  alpha = 1,
  row.norm = FALSE,
  maxit = 200,
  tol = 1e-04,
  verbose = FALSE
)

Arguments

X matrix with features in rows and samples in columns
n.comp number of components.
preICA output of ‘outICA()‘. Default is NULL
alg.typ parameter for fastICA(). If alg.typ == "deflation" the components are extracted one at a time. If alg.typ == "parallel" the components are extracted simultaneously. Default value is "deflation"

fun the functional form of the G function used in the approximation to neg-entropy in fastICA. Default value is "logcosh"

w.init initial weights

alpha default is 1

row.norm set TRUE if the normalization by rows is needed. Default is FALSE

maxit default is 200

tol default is 1e-04

verbose logic TRUE or FALSE. Use TRUE for print process steps. Default value is FALSE

Value

a list with (compliant to ‘fastICA()’ output)

X pre-processed data matrix

K pre-whitening matrix that projects data onto the first ‘n.comp’ principal components

W estimated un-mixing matrix

A estimated mixing matrix

S estimated source matrix

Author(s)

Maryna Chepeleva

enrichGO

Enrichment analysis of GO-terms based on Ensembl IDs

Description

Enrichment analysis of GO-terms for independent components with Ensembl IDs based on topGO package

Usage

enrichGO(
genesis, fdr = NULL, fc = NULL, ntop = NA, thr.fdr = 0.05,
The method estimates the variance explained by the model and by each independent component. We used the coefficient of determination (R2) between the normalized input (X-mean(X)) and (S*M)
Usage

estimateVarianceExplained(cica, X = NULL)

Arguments

cica       list compliant to 'consICA()' result
X          a 'SummarizedExperiment' object. Assay used for the model. Will be used if consICA$X is NULL, ignore otherwise.

Value

a list of:

R2          total variance explained by the model
R2_ic       Amount of variance explained by the each independent component

Examples

data("samples_data")
cica <- consICA(samples_data, ncomp=15, ntry=10, show.every=0)
var_ic <- estimateVarianceExplained(cica)

-----------------------------

getFeatures     Get features from consICA deconvolution result

-----------------------------

Description

Extract names of features (rows in 'X' and 'S' matrices) and their false discovery rates values

Usage

getFeatures(cica, alpha = 0.05, sort = FALSE)

Arguments

cica       list compliant to 'consICA()' result
alpha      value in [0,1] interval. Used to filter features with FDR < 'alpha'. Default value is 0.05
sort       sort features decreasing FDR. Default is FALSE

Value

list of dataframes 'pos' for positive and 'neg' for negative affecting features with columns:
features    names of features
fdr         false discovery rate value
getGO

Author(s)

Petr V. Nazarov

Examples

data("samples_data")
# Get deconvolution of X matrix
cica <- consICA(samples_data, ncomp=10, ntry=1, show.every=0)
# Get features names and FDR for each component
features <- getFeatures(cica)
# Positive affecting features for first components are
ic1_pos <- features$ic.1$pos

getGO Assigns IC signatures to Gene Ontologies

Description

Assigns extracted independent components to Gene Ontologies and rotate independent components (‘S’ matrix) to set most significant Gene Ontologies as positive affecting features. Set ‘ncores’ param for paralleled calculations.

Usage

g etGO(
cica,
 alpha = 0.05,
genenames = NULL,
genome = "org.Hs.eg.db",
db = c("BP", "CC", "MF"),
ncores = 4,
rotate = TRUE
)

Arguments

cica list compliant to ‘consICA()’ result
alpha value in [0,1] interval. Used to filter features with FDR < ‘alpha’. Default value is 0.05
genenames alternative names of genes. If NULL we use rownames of ‘S’ matrix. We automatically identify type of gene identifier, you can use Ensembl, Symbol, Entrez, Alias, Genename IDs.
genome R-package for genome annotation used. For human - ‘org.Hs.eg.db’
db name of GO database: "BP","MF","CC"
ncores number of cores for parallel calculation. Default value is 4
rotate rotate components in ‘S’ and ‘M’ matrices in ‘cica’ object to set most significant Gene Ontologies as positive effective features. Default is TRUE
get_score

**Value**

rotated (if need) ‘cica’ object with added ‘GO’ - list for each db chosen (BP, CC, MM), with dataframes ‘pos’ for positive and ‘neg’ for negative affecting features for each component:

<table>
<thead>
<tr>
<th>GO.ID</th>
<th>id of Gene Ontology term</th>
</tr>
</thead>
<tbody>
<tr>
<td>Term</td>
<td>name of term</td>
</tr>
<tr>
<td>Annotated</td>
<td>number of annotated genes</td>
</tr>
<tr>
<td>Significant</td>
<td>number of significant genes</td>
</tr>
<tr>
<td>Expected</td>
<td>estimate of the number of annotated genes if the significant genes would be randomly selected from the gene universe classisFisher F-test</td>
</tr>
<tr>
<td>FDR</td>
<td>false discovery rate value</td>
</tr>
<tr>
<td>Score</td>
<td>genes score</td>
</tr>
</tbody>
</table>

**Author(s)**

Petr V. Nazarov

**Examples**

data("samples_data")
# Calculate ICA (run with ntry=1 for quick test, use more in real analysis)
cica <- consICA(samples_data, ncomp=4, ntry=1, ncores=1, show.every=0)
# cica <- consICA(samples_data, ncomp=40, ntry=20, show.every=0)

# Annotate independent components with gene ontologies
cica <- getGO(cica, db = "BP", ncores=1)
# Positively affected GOs for 2nd independent component
head(cica$GO$GOBP$ic02$pos)

---

get_score Create score depending on threshold and paradigm

**Description**

Create score depending on threshold and paradigm

**Usage**

get_score(genes, fc, thr.fc, fdr, thr.fdr, ntop)
get_X_num

Arguments

genes character vector with list of ENSEBML IDs
fc numeric vector of logFC for each gene
thr.fc significance threshold for absolute logFC
fdr numeric vector of FDR for each gene
thr.fdr significance threshold for FDR
ntop number of first taken genes

Value
numeric score vector

Description
Convert input object as numeric matrix

Usage
get_X_num(obj, assay_string = NULL)

Arguments
obj input data with features in rows and samples in columns. Could be a ‘SummarizedExperiment’ object, matrix or ‘Seurat’ object. For ‘SummarizedExperiment’ with multiple assays or ‘Seurat’ pass the name with ‘assay_string’ parameter, otherwise the first will be taken. See SummarizedExperiment-class
assay_string name of assay for ‘SummarizedExperiment’ or ‘Seurat’ input object ‘obj’. Default value is NULL

Value
matrix
is.consICA  

Is the object is consensus ICA compliant?

Description

Check if the object is a list in the same format as the result of `consICA()`

Usage

is.consICA(cica)

Arguments

cica  list

Value

TRUE or FALSE

Examples

# returns TRUE
is.consICA(list("ncomp" = 2, "nsples" = 2, "nfeatures" = 2,
    "S" = matrix(0,2,2),"M" = matrix(0,2,2)))

oneICA  

Runs fastICA

Description

Runs fastICA once and store in a consICA manner

Usage

oneICA(
    X,  
ncomp = 10,  
filter.thr = NULL,  
reduced = FALSE,  
fun = "logcosh",  
alg.typ = "deflation",  
assay_string = NULL
)
oneICA

Arguments

- **X**
  input data with features in rows and samples in columns. Could be a ‘SummarizedExperiment’ object, matrix or ‘Seurat’ object. For ‘SummarizedExperiment’ with multiple assays or ‘Seurat’ pass the name with ‘assay_string’ parameter, otherwise the first will be taken. See `SummarizedExperiment-class`

- **ncomp**
  number of components. Default value is 10

- **filter.thr**
  filter rows in input matrix with max value > `filter.thr`. Default value is `NULL`

- **reduced**
  If TRUE returns reduced result (no X, see ‘return’)

- **fun**
  the functional form of the G function used in the approximation to neg-entropy in fastICA. Default value is “logcosh”

- **alg.typ**
  parameter for fastICA(). if alg.typ == "deflation" the components are extracted one at a time. if alg.typ == "parallel" the components are extracted simultaneously. Default value is "deflation"

- **assay_string**
  name of assay for ‘SummarizedExperiment’ or ‘Seurat’ input object ‘X’. Default value is `NULL`

Value

a list with

- **X**
  input ‘SummarizedExperiment’ object

- **nsamples, nfeatures**
  dimension of X assay

- **S, M**
  consensus metagene and weight matrix

- **ncomp**
  number of components

Author(s)

Petr V. Nazarov

See Also

- `fastICA`

Examples

```r
data("samples_data")
res <- oneICA(samples_data)
```
outICA

*Outside part of multiple run Independent Component Analysis*

**Description**

Calculate a common part for consensus Independent Component Analysis (ICA)

**Usage**

```r
outICA(X, n.comp, row.norm = FALSE, verbose = FALSE)
```

**Arguments**

- `X`: matrix with features in rows and samples in columns
- `n.comp`: number of components
- `row.norm`: rows normalization flag. Default value is FALSE
- `verbose`: logic TRUE or FALSE. Use TRUE for print process steps. Default value is FALSE

**Value**

a list with

- `X`: input matrix
- `X1`: interim calculated matrix
- `K`: pre-whitening matrix that projects data onto the first `n.comp` principal components

**Author(s)**

Maryna Chepeleva

overlapGO

*Similarity of two gene ontologies lists*

**Description**

Calculate similarity matrix of gene ontologies (GOs) of independent components. The measure could be cosine similarity or Jaccard index (see details)

**Usage**

```r
overlapGO(GO1, GO2, method = c("cosine", "jaccard"), fdr = 0.01)
```
Arguments

- **GO1**: list of GOs for each independent component got from `getGO()`
- **GO2**: list of GOs for each independent component got from `getGO()`
- **method**: can be 'cosine' for non-parametric cosine similarity or 'jaccard' for Jaccard index. See details
- **fdr**: FDR threshold for GOs that would be used in measures. Default value is 0.01

Details

Jaccard index is a measure of the similarity between two sets of data. It is calculated as intersection divided by union

\[ J(A, B) = \frac{|A \cap B|}{|A \cup B|} \]

Results are from 0 to 1.

Cosine similarity here is calculated in a non-parametric way: for two vectors of gene ontologies, the space is created as a union of GOs in both vectors. Then, two rank vectors in this space are created, most enriched GOs get the biggest rank and GOs from space not included in the GO vector get 0. Cosine similarity is calculated between two scaled rank vectors. Such approach allows to take the order of enriched GO into account. Results are from -1 to 1. Zero means no similarity.

Value

A similarity matrix of cosine or Jaccard values, rows corresponds to independent components in 'GO1', columns to independent components in 'GO2'.

Author(s)

Maryna Chepeleva

Examples

```r
# Not run:
data("samples_data")
cica1 <- consICA(samples_data, ncomp=5, ntry=1, show.every=0)
cica1 <- getGO(cica1, db = "BP", ncores = 1)
cica2 <- consICA(samples_data[,1:100], ncomp=4, ntry=1, show.every=0)
cica2 <- getGO(cica2, db = "BP", ncores = 1)
cj <- overlapGO(GO1 = cica1$GO$GOBP, GO2 = cica2$GO$GOBP,
    method = "jaccard", fdr = 0.01)
ccos_sim <- overlapGO(GO1 = cica1$GO$GOBP, GO2 = cica2$GO$GOBP,
    method = "cosine", fdr = 0.01)
```

## End(Not run)
plotICVarianceExplained

Barplot variance explained by each IC

Description

Method to plot variance explained (R-squared) by the MOFA model for each view and latent factor. As a measure of variance explained for gaussian data we adopt the coefficient of determination (R2). For details on the computation see the help of the estimateVarianceExplained function

Usage

plotICVarianceExplained(
  cica,
  sort = NULL,
  las = 2,
  title = "Variance explained per IC",
  x.cex = NULL,
  ...
)

Arguments

cica | consICA compliant list
sort | specify the arrangement as 'asc'/'desc'. No sorting if NULL
las | orientation value for the axis labels (0 - always parallel to the axis, 1 - always horizontal, 2 - always perpendicular to the axis, 3 - always vertical)
title | character string with title of the plot
x.cex | specify the size of the tick label numbers/text with a numeric value of length 1
... | extra arguments to be passed to barplot

Value

A numeric vector compliant to ‘barplot’ output

Examples

data("samples_data")
cica <- consICA(samples_data, ncomp=15, ntry=10, show.every=0)
p <- plotICVarianceExplained(cica, sort = "asc")
samples_data

Samples of gene expression

Description
A dataset containing the expression of 2454 genes for 472 samples from skin cutaneous melanoma (SKCM) TCGA cohort, their metadata such as age, gender, cancer type etc. and survival time-to-event data

Usage
data(samples_data)

Format
A SummarizedExperiment object:

- **assay** expression matrix with genes by rows and samples by columns
- **colData** data frame with sample metadata (clinical variables)

saveReport
Save PDF report with analysis of each independent component

Description
Save PDF report with description of each independent component (IC) consists of most affected genes, significant Go terms, survival model for the component, ANOVA analysis for samples characteristics and stability

Usage
saveReport(
cica,
Genes = NULL,
Var = NULL,
surv = NULL,
genenames = NULL,
file = sprintf("report_ICA_%d.pdf", ncol(IC$S)),
main = "Component # %d (stability = %.3f)",
show.components = seq.int(1, ncol(cica$S))
)
setOrientation

Arguments

- **cica**: list compliant to ‘consICA()’ result. May include GO list with enrichment analysis appended with ‘getGO()’ function.
- **Genes**: features list compliant to ‘getFeatures’ output (list of dataframes ‘pos’ for positive and ‘neg’ for negative affecting features with names of features false discovery rates columns). If NULL will generated automatically.
- **Var**: matrix with samples metadata.
- **surv**: dataframe with time and event values for each sample.
- **genenames**: alternative gene names for printing in the report.
- **file**: report filename, ends with ".pdf".
- **main**: title for each list describes the component.
- **show.components**: which component will be shown.

Value

TRUE when successfully generate report.

Author(s)

Petr V. Nazarov.

Examples

```r
data("samples_data")
cica <- consICA(samples_data, ncomp=40, ntry=10, show.every=0)
if(FALSE){
cica <- getGO(cica, db = "BP")
}
saveReport(cica, Var=samples_data$Var, surv = samples_data$Sur)
```

Description

Set orientation for independent components as positive in most enriched direction. Use first element of ‘GOs’ for direction establishment.

Usage

```r
setOrientation(cica, verbose = FALSE)
```

Arguments

- **cica**: list compliant to ‘consICA()’ result. Must contain GO, see ‘getGO()’.
- **verbose**: logic TRUE or FALSE. Use TRUE for print process steps. Default is FALSE.
Value

cica object after rotation, with rotated ‘S’, ‘M’ and added ‘compsign’ which is vector defined rotation: ‘S_rot = S * compsign, M_rot = M * compsign, GO_rot = GO * compsign’

Note

Implemented inside ‘getGO()’ in version >= 1.1.1.

Author(s)

Petr V. Nazarov

Examples

```r
## Not run:
data("samples_data")
# Get deconvolution of X matrix
#cica <- consICA(samples_data, ncomp=10, ntry=1, show.every=0)
cica <- consICA(samples_data, ncomp=2, ntry=1, show.every=0) # timesaving example
GOs <- getGO(cica, db = "BP")
# Get already rotated S matrix and Gene Ontologies
cica <- getGO(cica, db = "BP")

# Get Gene Ontologies without rotation (actually, you don’t need to do this)
# This may used for GO generated with version < 1.1.1. Add GO to cica list.
cica <- getGO(cica, db = "BP", rotate = FALSE)
# Rotate components
cica <- setOrientation(cica, verbose = T)
# Which components was rotated
which(cica$compsign == -1)

## End(Not run)
```

---

**set_bpparam**

Set up for the parallel computing for biocParallel Adapt from ‘FEAST’

This function sets up the environment for parallel computing.

**Description**

Set up for the parallel computing for biocParallel Adapt from ‘FEAST’ This function sets up the environment for parallel computing.

**Usage**

```
set_bpparam(ncores = 0, BPPARAM = NULL)
```
sortDataFrame

Arguments

ncores number of processors
BPPARAM bpparameter from bpparam

Value

BAPPARAM settings

Description

Sort dataframe, adapted from http://snippets.dzone.com/user/r-fanatic

Usage

sortDataFrame(x,key, ...)

Arguments

x a data.frame
key sort by this column
... other parameters for ‘order’ function (e.g. ‘decreasing’)

Value

sorted dataframe

Examples

df <- data.frame("features" = c("f1", "f2", "f3"), fdr = c(0.02, 0.002, 1))
sortDataFrame(df, "fdr")
sortFeatures  Sort Genes of consICA object

Description

Sort Genes for independent components

Usage

sortFeatures(Genes)

Arguments

Genes  list compilant to 'getFeatures' output

Value

sorted list

Examples

#features <- list(ic1 = list(  # "pos" = data.frame("features" = c("f1", "f2", "f3"),  # "fdr" = c(0.0043, 0.4, 0.04)),  # "neg" = data.frame("features" = c("f1", "f2", "f3"),  # "fdr" = c(0, 0.1, 0.9))))
#sortFeatures(features)

survivalAnalysis  Survival analysis based on significant IC

Description

Cox regression (based on R package 'survival') on the weights of independent components with significant contribution in individual risk model. For more see Nazarov et al. 2019 In addition the function plot Kaplan-Meier diagram.

Usage

survivalAnalysis(cica, surv = NULL, time = NULL, event = NULL, fdr = 0.05)
survivalAnalysis

Arguments

- `cica`: list compliant to 'consICA()' result
- `surv`: dataframe with time and event values for each sample. Use this parameter or 'time' and 'event'
- `time`: survival time value for each sample
- `event`: survival event factor for each sample (TRUE or FALSE)
- `fdr`: false discovery rate threshold for significant components involved in final model. Default value is 0.05

Value

a list with

- `cox.model`: an object of class ‘coxph’ representing the fit. See ‘coxph.object’ for details
- `hazard.score`: hazard score for significant components (fdr < ‘fdr’ in individual cox model)

Examples

data("samples_data")
# Get deconvolution of X matrix
cica <- consICA(samples_data, ncomp=10, ntry=1, show.every=0)
surv <- survivalAnalysis(cica,
  surv = SummarizedExperiment::colData(samples_data)[,c("time", "event")])
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