Package 'consICA'

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

Description consICA implements a data-driven deconvolution method – consensus

extract features suitable for patient diagnostics and prognostics.

The method separates biologically relevant transcriptional signals from

independent component analysis (ICA) to decompose heterogeneous omics data and

Title consensus Independent Component Analysis

Type Package

Version 2.8.0

```
technical effects and provides information about the cellular composition and biological pro-
      The implementation of parallel computing in the package ensures efficient
      analysis of modern multicore systems.
BugReports https://github.com/biomod-lih/consICA/issues
License MIT + file LICENSE
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
RoxygenNote 7.3.2
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Author Petr V. Nazarov [aut, cre] (ORCID:
       <https://orcid.org/0000-0003-3443-0298>),
      Tony Kaoma [aut] (ORCID: <a href="https://orcid.org/0000-0002-1269-4826">https://orcid.org/0000-0002-1269-4826</a>),
      Maryna Chepeleva [aut] (ORCID: <a href="https://orcid.org/0000-0003-3036-4916">https://orcid.org/0000-0003-3036-4916</a>)
Maintainer Petr V. Nazarov <petr.nazarov@lih.lu>
```

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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 \sim 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

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```
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)</pre>
```

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,
    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 'Sum-

marizedExperiment' 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

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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 simultane-

ously. 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'. De-

fault 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

Author(s)

Petr V. Nazarov

See Also

fastICA

```
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)</pre>
```

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coreICA

Fast Independent Component Analysis for multi-run mode

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

Χ	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 "log $\cos h$ "
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

6 enrichGO

K	pre-whitening matrix that projects data onto the first 'n.comp' principal components
W	estimated un-mixing matrix
Α	estimated mixing matrix
S	estimated source matrix

Author(s)

Maryna Chepeleva

enrichG0

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(
   genes,
   fdr = NULL,
   fc = NULL,
   ntop = NA,
   thr.fdr = 0.05,
   thr.fc = NA,
   db = "BP",
   genome = "org.Hs.eg.db",
   id = c("entrez", "alias", "ensembl", "symbol", "genename"),
   algorithm = "weight",
   do.sort = TRUE,
   randomFraction = 0,
   return.genes = FALSE
)
```

Arguments

```
genes
                  character vector with list of ENSEBML IDs
                  numeric vector of FDR for each gene
fdr
                  numeric vector of logFC for each gene
fc
                  number of first taken genes
ntop
                  significance threshold for FDR
thr.fdr
thr.fc
                  significance threshold for absolute logFC
                  name of GO database: "BP", "MF", "CC"
db
                  R-package for genome annotation used. For human - 'org.Hs.eg.db'
genome
id
algorithm
                  algorithm for 'runTest()'
```

do.sort if TRUE - resulted functions sorted by p-value

randomFraction for testing only, the fraction of the genes to be randomized return.genes If TRUE include genes in output. Default value is FALSE

Value

list with terms and stats

Author(s)

Petr V. Nazarov

estimateVarianceExplained

Estimate the variance explained by the model

Description

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_ics Amount of variance explained by the each independent component

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

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get	Features	:

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

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</pre>
```

getG0

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.

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Usage

```
getGO(
  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 auto-

matically 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' matricies in 'cica' object to set most significant

Gene Ontologies as positive effective features. Default is TRUE

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:

GO. ID id of Gene Ontology term

Term name of term

Annotated number of annotated genes

Significant number of significant genes

Expected estimate of the number of annotated genes if the significant genes would be randomly selected from the gene universe classisFisher

F-test

FDR false discovery rate value

Score genes score

Author(s)

Petr V. Nazarov

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Examples

```
data("samples_data")
# Calculate ICA (run with ntry=1 for quick test, use more in real analysis)
#cica <- consICA(samples_data, ncomp=2, ntry=1, ncores=1, show.every=0)
# cica <- consICA(samples_data, ncomp=40, ntry=20, show.every=0)

# Annotate independent components with gene ontoligies
# cica <- getGO(cica, db = "BP", ncores=4)
## Positively affected GOs for 2nd independent component
#head(cica$GO$GOBP$ic02$pos)</pre>
```

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)
```

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

get_X_num

Convert input object as numeric matrix

Description

Convert input object as numeric matrix

Usage

```
get_X_num(obj, assay_string = NULL)
```

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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

oneICA

Runs fastICA

Description

Runs fastICA once and store in a consICA manner

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Usage

```
oneICA(
   X,
   ncomp = 10,
   filter.thr = NULL,
   reduced = FALSE,
   fun = "logcosh",
   alg.typ = "deflation",
   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' param-

eter, 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 simultane-

ously. Default value is "deflation"

assay_string name of assay for 'SummarizedExperiment' or 'Seurat' input object 'X'. De-

fault 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

```
data("samples_data")
res <- oneICA(samples_data)</pre>
```

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outICA	Outside part of multiple run Independent Component Analysis

Description

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

Usage

```
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 compo-

nents

Author(s)

Maryna Chepeleva

overlapGO	Similarity of two gene ontologies lists	
-----------	---	--

Description

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

Usage

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

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Arguments

G01	list of GOs for each independent component got from 'getGO()'
G02	list of GOs for each independent component got from 'getGO()'
method	can be 'cosine' for non-parametric cosine similarity or 'jaccard' for Jaccadr 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 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 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

```
## Not run:
data("samples_data")
# Calculate ICA (run with ntry=1 for quick test, use more in real analysis)
cica1 <- consICA(samples_data, ncomp=5, ntry=1, show.every=0)</pre>
# Search enriched gene ontologies
cica1 <- getGO(cica1, db = "BP", ncores = 1)
# Calculate ICA and GOs for another dataset
cica2 <- consICA(samples_data[,1:100], ncomp=4, ntry=1, show.every=0)</pre>
cica2 <- getGO(cica2, db = "BP", ncores = 1)</pre>
# Compare two lists of enriched GOs
# Jaccard index
jc <- overlapGO(GO1 = cica1$GO$GOBP, GO2 = cica2$GO$GOBP,</pre>
method = "jaccard", fdr = 0.01)
# Cosine similarity
cos_sim <- overlapGO(GO1 = cica1$GO$GOBP, GO2 = cica2$GO$GOBP,</pre>
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

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

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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))
)
```

Arguments

cica list compliant to 'consICA()' result. May include GO list with enrichment anal-

ysis appended with 'getGO()' function

Genes features list compilant to 'getFeatures' output (list of dataframes 'pos' for pos-

itive and 'neg' for negative affecting features with names of features false dis-

covery rates columns). If NULL will generated automatically

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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 discribes the component

show.components

which compont will be shown

Value

TRUE when successfully generate report

Author(s)

Petr V. Nazarov

Examples

```
if(FALSE){
  data("samples_data")
  cica <- consICA(samples_data, ncomp=15, ntry=10, ncores = 2, show.every=0)
  if(FALSE){
    cica <- getGO(cica, db = "BP")
  }
  saveReport(cica, Var=samples_data$Var, surv = samples_data$Sur)
}</pre>
```

setOrientation

Set orientation for independent components

Description

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

Usage

```
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.

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Author(s)

Petr V. Nazarov

Examples

```
## Not run:
data("samples_data")
# Get deconvolution of X matrix
#cica <- consICA(samples_data, ncomp=10, ntry=1, show.every=0)</pre>
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")</pre>
\# 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)</pre>
# Rotate components
cica <- setOrientation(cica, verbose = T)</pre>
# 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)
```

Arguments

ncores number of processors

BPPARAM bpparameter from bpparam

Value

BAPPARAM settings

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sortDataFrame

Sort dataframe

Description

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

Usage

```
sortDataFrame(x,key, ...)
```

Arguments

```
x a data.framekey sort by this column... other parameters for 'order' function (e. g. 'decreasing')
```

Value

sorted dataframe

Examples

```
df \leftarrow 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

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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)
```

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)

```
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")])</pre>
```

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