Package 'mSigAct'

March 4, 2021

```
Title mutational Signature Activity analysis ('mSigAct')
Version 2.1.0.9006
Author Steve Rozen, Alvin Wei Tian Ng, Arnoud Boot
Maintainer Steve Rozen < steverozen@gmail.com>
Description Analyze the ``activities" of mutational signatures in one or more mutational spectra.
      'mSigAct' stands for mutational Signature Activity. mSigAct can estimate (conservatively)
      whether there is evidence that a particular set of mutational signatures is present in a
      spectrum. It can also determine a *minimal* subset of signatures needed to plausibly
     reconstruct an observed spectrum. This sparse assign signatures functionality is
      *deliberately biased* toward using as few signatures as possible. This package does not
      provide all-purpose estimation for signature attribution.
License GPL-3
URL https://github.com/steverozen/mSigAct
BugReports https://github.com/steverozen/mSigAct/issues
Encoding UTF-8
LazyData true
Language en-US
Depends R (>= 4.0),
RoxygenNote 7.1.1
VignetteBuilder knitr
biocViews
Imports dplyr,
     ICAMS (>= 2.3.5.9002),
     ICAMSxtra,
     lsa,
     nloptr,
     PCAWG7 (>= 0.0.3.9006),
     philentropy,
     quadprog,
```

rlang, stats, sets, tibble, utils 2 AddSigActivity

Remotes github::steverozen/ICAMS@master,	
github::steverozen/ICAMSxtra@master,	
github::steverozen/PCAWG7@master	
Suggests BSgenome. Hsapiens. 1000 genomes. hs37d5,	
devtools,	
htmlwidgets,	
knitr,	
profvis,	
rmarkdown,	
testthat ($>= 2.1.0$),	
usethis	

R topics documented:

AddSigActivity
CancerTypes
cossim
DefaultManyOpts
ExposureProportions
g_ineq_for_ObjFnBinomMaxLH2
LLHSpectrumMAP
LLHSpectrumNegBinom
LLHSpectrumNegBinomDebug
MAPAssignActivity
MAPAssignActivity1
ObjFnBinomMaxLHMustRound
ObjFnBinomMaxLHNoRoundOK
ObjFnBinomMaxLHRound
OneMAPAssignTest
OptimizeExposure
OptimizeExposureQP
OptimizeExposureQPBootstrap
PCAWGMAPTest
PossibleArtifacts
RareSignatures
ReconstructSpectrum
ShowSigActivity
SignaturePresenceTest
SignaturePresenceTest1
SparseAssignActivity
23

AddSigActivity

Add contributing signature activity information for multiple spectra

Description

Index

Add contributing signature activity information for multiple spectra

AddSigActivity 3

Usage

```
AddSigActivity(spectra, exposure, sigs, sigs.presence.prop, nbinom.size = 5)
```

Arguments

The spectra (multiple spectra) to be reconstructed. spectra Exposures as a numerical matrix (or data.frame) with signatures in rows and exposure samples in columns. Rownames are taken as the signature names and column names are taken as the sample IDs. The signatures with which we are trying to reconstruct spectra. A numerical sigs matrix, possibly an ICAMS catalog. The column names of sigs should be a superset of row names of exposure. sigs.presence.prop The proportions of samples that contain each signature. A numerical vector (values between 0 and 1), with names being a subset of colnames (sigs). See ExposureProportions for more details. The dispersion parameter for the negative binomial distribution; smaller is more nbinom.size dispersed. See NegBinomial.

Details

This function calls ReconstructSpectrum and LLHSpectrumNegBinom.

Value

A list of lists containing output for each sample in spectra.

Each sublist has the following elements:

- original.spect: The original spectrum with total mutation counts added to its column name. An additional attribute "exposure" from exposure is also added.
- reconstructed.spect: The reconstructed spectrum using sigs and exposure. Its column name has the total mutation counts and cosine similarity with the original spectrum.
- contributing.sigs: The contributing signatures to the original spectrum. The column names of each contributing signature has mutation counts attributed to this signature, its contribution proportion and proposed etiology(if the etiology is unknown, then will be blank.)
- distances: Various distances and similarities between the original spectrum and reconstructed. spect.

Remark

The column names of spectra should be the same as the column name of exposure.

Examples

4 DefaultManyOpts

CancerTypes

Return a character vector of some common cancer types

Description

Return a character vector of some common cancer types

Usage

```
CancerTypes()
```

cossim

Cosine similarity with useful argument types..

Description

Calls cosine.

Usage

```
cossim(v1, v2)
```

Arguments

v1 A vector or single-column matrix v2 A vector or single-column matrix

DefaultManyOpts

Set default options for many functions, especially nloptr.

Description

Set default options for many functions, especially nloptr.

Usage

```
DefaultManyOpts()
```

Value

A list with the following elements

global.opts A sub-list with several options for nloptr, q.v., for the global optimization phase, including eval_f, the objective function.

local.opts A sub-list with several options for nloptr, q.v., for the local optimization phase, including eval_f, the objective function and the inequality constraint function eval_g_ineq

nbinom.size The dispersion parameter for the negative binomial distribution; smaller is more dispersed. See NegBinomial.

trace If > 0 print progress messages.

ExposureProportions 5

```
ExposureProportions
```

Return the proportions of tumors of a given cancer type that have a particular signature

Description

Return the proportions of tumors of a given cancer type that have a particular signature

Usage

```
ExposureProportions(
  mutation.type,
  cancer.type,
  all.sigs = NULL,
  drop.sigs.no.info = TRUE,
  must.include = character(),
  must.include.prop = 0.1
)
```

Arguments

```
mutation.type
                 A character string, one of "SBS96", "SBS192", "ID", "DBS78".
cancer.type
                 A character string. For some common cancer types, see CancerTypes for
                 more details.
                 An optional matrix of known signatures, with column names being signatures
all.sigs
                 ids. Only used to drop signatures not present in all.sigs.
drop.sigs.no.info
                 If TRUE, drop signatures not present in the column names of all.sigs.
must .include A character vector of signature IDs that must be included, even if they have
                 not previously been observed in that cancer type. The associated proportion is
                 specified by must.include.prop.
must.include.prop
                 The value used for the expected proportion of signatures in must.include
                 but not previously observed in the given cancer.type.
```

Value

A numerical vector of the proportion of tumors of type cancer.type with each signature for those signatures observed in cancer.type. The names are the signature ids.

Examples

6 LLHSpectrumMAP

```
g_ineq_for_ObjFnBinomMaxLH2
```

Function to constrain the sum of estimated exposures to the number of mutations in the spectrum.

Description

See nloptr to understand how this function is used.

Usage

```
g_ineq_for_ObjFnBinomMaxLH2(exp, spectrum, sigs, nbinom.size)
```

Arguments

exp	A numeric vector of exposures.
spectrum	The observed spectrum we are trying to reconstruct.
sigs	The signatures with which we are trying to reconstruct the spectrum. (Ignored in this function but used by $nloptr$.)
nbinom.size	Dispersion parameter. (Ignored in this function but used by nloptr.)

Likelihood that 1 observed spectrum was generated from a vector of expected mutation counts using prior information of the signature

of expected mutation counts using prior information of the presence proportions

Description

Likelihood that 1 observed spectrum was generated from a vector of expected mutation counts using prior information of the signature presence proportions

```
LLHSpectrumMAP(
   spectrum,
   expected.counts,
   nbinom.size,
   model,
   sigs.presence.prop,
   verbose = FALSE
)
```

Arguments

An observed spectrum (a numeric vector). spectrum

expected.counts

A vector of (integer) expected mutation counts, one expected count for each mutation type. We want to know the likelihood that this model generated the observed spectrum, assuming each mutational types generates counts according to a negative binomial distribution with the given expected.counts (argument mu to NegBinomial) and dispersion parameter nbinom. size.

nbinom.size

The dispersion parameter for the negative binomial distribution; smaller is more

dispersed. See NegBinomial.

model

Names of sigs present in the MAP exposure. Do not use indices. sigs.presence.prop

> The proportions of samples that contain each signature. A numerical vector (values between 0 and 1), with names being a superset of model.

If TRUE print messages under some circumstances. verbose

Value

log(likelihood(spectrum | expected.counts)) + log(probability(model | sigs.presence.prop)), or, in more detail, the sum of the negative binomial likelihoods that each element of the spectrum (i.e., the count for each mutation type e.g. ACT > AAT) was generated from the expected count for that mutation type plus the probability of the signature model used in the reconstruction given the prior sigs.presence.prop.

LLHSpectrumNegBinom

Likelihood that 1 observed spectrum was generated from a vector of expected mutation counts.

Description

Likelihood that 1 observed spectrum was generated from a vector of expected mutation counts.

Usage

LLHSpectrumNegBinom(spectrum, expected.counts, nbinom.size, verbose = FALSE)

Arguments

spectrum An observed spectrum (a numeric vector)

expected.counts

A vector of (integer) expected mutation counts, one expected count for each mutation type. We want to know the likelihood that this model generated the observed spectrum, assuming each mutational types generates counts according to a negative binomial distribution with the given expected. counts (argument mu to NegBinomial) and dispersion parameter nbinom.size.

nbinom.size

The dispersion parameter for the negative binomial distribution; smaller is more

dispersed. See NegBinomial.

If TRUE print messages under some circumstances. verbose

Value

log(likelihood(spectrum | expected.counts)), or, in more detail, the sum of the negative binomial likelihoods that each element of the spectrum (i.e., the count for each mutation type e.g. ACT > AAT) was generated from the expected count for that mutation type.

LLHSpectrumNegBinomDebug

A verbose version of LLHSpectrumNegBinom for testing

Description

We use a separate function so as not to slow down the heavily used LLHSpectrumNegBinom and to provide more information in the output

Usage

```
LLHSpectrumNegBinomDebug(
   spectrum,
   expected.counts,
   nbinom.size,
   verbose = FALSE
)
```

Arguments

spectrum (a numeric vector)

expected.counts

A vector of (integer) expected mutation counts, one expected count for each mutation type. We want to know the likelihood that this model generated the observed spectrum, assuming each mutational types generates counts according to a negative binomial distribution with the given expected.counts (argument mu to NegBinomial) and dispersion parameter nbinom.size.

 $\verb|nbinom.size| The dispersion parameter for the negative binomial distribution; smaller is more$

dispersed. See NegBinomial.

verbose If TRUE print messages under some circumstances.

Value

A tibble with self-explanatory columns and rows.

MAPAssignActivity 9

MAPAssignActivity Find Maximum A Posteriori (MAP) assignment of signature exposures that explain multiple spectra

Description

Find Maximum A Posteriori (MAP) assignment of signature exposures that explain multiple spectra

Usage

```
MAPAssignActivity(
  spectra,
  sigs,
  sigs.presence.prop,
  output.dir,
  max.level = 5,
  p.thresh = 0.05,
  m.opts = DefaultManyOpts(),
  num.parallel.samples = 5,
  mc.cores.per.sample = min(20, 2^max.level),
  progress.monitor = NULL,
  seed = NULL
)
```

Arguments

The spectra (multiple spectra) to be reconstructed. spectra A numerical matrix, possibly an ICAMS catalog. sigs sigs.presence.prop The proportions of samples that contain each signature. A numerical vector (values between 0 and 1), with names being a subset of colnames (sigs). See ExposureProportions for more details. Directory path to save the output file. output.dir The maximum number of signatures to try removing. max.level If the p value for a better reconstruction with as opposed to without a set of p.thresh signatures is > than this argument, then we can use exposures without this set. See DefaultManyOpts. m.opts num.parallel.samples The (maximum) number of samples to run in parallel. On Microsoft Windows machines it is silently changed to 1. Each sample in turn can require multiple cores, as governed by mc.cores.per.sample. mc.cores.per.sample

The maximum number of cores to use for each sample. On Microsoft Windows machines it is silently changed to 1.

```
progress.monitor
```

Function called at the start of each new level (number of signatures to try excluding). Must take named arguments value and detail, and no others. Designed for a AsyncProgress progress bar function.

10 MAPAssignActivity1

seed

Random seed; set this to get reproducible results. (The numerical optimization is in two phases; the first, global phase might rarely find different optima depending on the random seed.)

Value

A list with the elements:

- proposed.assignment: Proposed signature assignment for spectra with the highest MAP found.
- proposed.reconstruction: Proposed reconstruction of spectra based on MAP.
- reconstruction.distances: Various distances and similarities between spectra and proposed.reconstruction.
- error.messages: Only appearing if there are errors running MAPAssignActivity.
- results.details: Detailed results for each sample in spectra.

These elements will be NULL if the algorithm could not find the optimal reconstruction or there are errors coming out.

Examples

```
## Not run:
# This is a long running example unless multiple CPU cores are available
indices <- grep("Lung-AdenoCA", colnames(PCAWG7::spectra$PCAWG$SBS96))</pre>
spectra <- PCAWG7::spectra$PCAWG$SBS96[, indices[1:2], drop = FALSE]</pre>
sigs <- PCAWG7::COSMIC.v3.1$signature$genome$SBS96</pre>
sigs.prop <- ExposureProportions (mutation.type = "SBS96",</pre>
                                   cancer.type = "Lung-AdenoCA")
MAP.out <- MAPAssignActivity(spectra = spectra,
                              sigs = sigs,
                              sigs.presence.prop = sigs.prop,
                              output.dir = file.path(tempdir(), "Lung-AdenoCA"),
                              \max.level = length(sigs.prop) - 1,
                              p.thresh = 0.05 / ncol(spectra),
                              num.parallel.samples = 2,
                              mc.cores.per.sample = 10)
## End(Not run)
```

MAPAssignActivity1 Find a Maximum A Posteriori (MAP) assignment of signature exposures that explain one spectrum.

Description

Find a Maximum A Posteriori (MAP) assignment of signature exposures that explain one spectrum.

MAPAssignActivity1 11

Usage

```
MAPAssignActivity1(
  spect,
  sigs,
  sigs.presence.prop,
  max.level = 5,
  p.thresh = 0.05,
  m.opts = DefaultManyOpts(),
  max.mc.cores = min(20, 2^max.level),
  progress.monitor = NULL,
  seed = NULL
)
```

Arguments

A single spectrum. spect A numerical matrix, possibly an ICAMS catalog. sigs sigs.presence.prop

> The proportions of samples that contain each signature. A numerical vector (values between 0 and 1), with names being a subset of colnames (sigs). See ExposureProportions for more details.

max.level The maximum number of signatures to try removing.

If the p value for a better reconstruction with as opposed to without a set of p.thresh

signatures is > than this argument, then we can use exposures without this set.

m.opts See DefaultManyOpts.

max.mc.cores The maximum number of cores to use. On Microsoft Windows machines it is silently changed to 1.

progress.monitor

Function called at the start of each new level (number of signatures to try excluding). Must take named arguments value and detail, and no others. Designed for a AsyncProgress progress bar function.

Random seed; set this to get reproducible results. (The numerical optimization is in two phases; the first, global phase might rarely find different optima

depending on the random seed.)

Value

seed

A list with the elements:

- proposed.assignment: Proposed signature assignment for spect with the highest MAP found.
- proposed.reconstruction: Reconstruction based on MAP.
- · reconstruction.distances: Various distances and similarities between spect and proposed.reconstruction.
- all.tested: A tibble of all the search results.
- time.for.MAP.assign: Value from system.time for running MAPAssignActivity1.
- error.messages: Only appearing if there are errors running MAPAssignActivity1.

These elements will be NULL if the algorithm could not find the optimal reconstruction or there are errors coming out.

Examples

ObjFnBinomMaxLHMustRound

A deprecated negative binomial maximum likelihood objective function.

Description

Use ObjFnBinomMaxLHRound instead.

Usage

```
ObjFnBinomMaxLHMustRound(exp, spectrum, sigs, nbinom.size)
```

Arguments

exp A vector of exposures ("activities").

spectrum The spectrum to assess.
sigs The matrix of signatures.

nbinom.size The dispersion parameter for the negative binomial distribution; smaller is more

dispersed. See NegBinomial.

Details

This function will lead to errors in some situations when the rounded reconstructed signature contains 0s for mutations classes for which the target spectrum is > 0.

ObjFnBinomMaxLHNoRoundOK

A deprecated negative binomial maximum likelihood objective function.

Description

Use ObjFnBinomMaxLHRound instead.

Usage

```
ObjFnBinomMaxLHNoRoundOK(exp, spectrum, sigs, nbinom.size)
```

Arguments

exp A vector of exposures ("activities").

spectrum The spectrum to assess.
sigs The matrix of signatures.

nbinom.size The dispersion parameter for the negative binomial distribution; smaller is more

dispersed. See NegBinomial.

Details

This function rounds sometimes, which leads to minor differences in log likelihoods of reconstructed spectra (LLHSpectrumNegBinom) compared to the value returned by this function.

ObjFnBinomMaxLHRound

The preferred negative binomial maximum likelihood objective function.

Description

Can be used as the objective function for SparseAssignActivity, SparseAssignActivity1, and SignaturePresenceTest1. (Internally used by by nloptr.)

Usage

```
ObjFnBinomMaxLHRound(exp, spectrum, sigs, nbinom.size)
```

Arguments

exp A vector of exposures ("activities").

spectrum The spectrum to assess.
sigs The matrix of signatures.

nbinom.size The dispersion parameter for the negative binomial distribution; smaller is more

dispersed. See NegBinomial.

14 OneMAPAssignTest

Value

-1 * log(likelihood(spectrum | reconstruction))

nloptr minimizes the objective function, so the lower the objective function, the better.

OneMAPAssignTest Run one test of MAPAssignActivity1.

Description

Run one test of MAPAssignActivity1.

Usage

```
OneMAPAssignTest(
   spect,
   reference.exp,
   cancer.type,
   mutation.type,
   exposure.mutation.type,
   max.subsets = 1000,
   max.level = 5,
   max.mc.cores = 100,
   m.opts = DefaultManyOpts(),
   out.dir = NULL,
   p.thresh,
   max.presence.proportion,
   sigs.prop = NULL,
   sigs = NULL
)
```

Arguments

```
spect
                 A single spectrum.
reference.exp
                 Compare the inferred exposures to this.
                 Character string from a fixed set indicating different cancer types, used to look
cancer.type
                 up the set of signatures known in that cancer type and the proportion of cancers
                 of that type that have the signature. TODO: provide information on how to find
                 the allowed cancer types.
mutation.type
                 One of "SBS96", "SBS192", "ID", "DBS78".
exposure.mutation.type
                 One of "SBS96", "ID", "DBS78".
max.subsets The maximum number of subsets that can be tested for removal from the set of
                 signatures.
                 The maximum number of signatures to try removing.
max.level
max.mc.cores The maximum number of cores to use. On Microsoft Windows machines it is
                 silently changed to 1.
```

OptimizeExposure 15

m.opts	See DefaultManyOpts.
out.dir	If non-NULL create this directory if necessary and put results there.
p.thresh	If the p value for a better reconstruction with than without a set of signatures is $>$ than p.thresh, then we can use exposures without this set.
max.presence	.proportion
	The maximum value of the proportion of tumors that must have a given signature. Used so that it is possible to exclude a signature from a spectrum, e.g. perhaps all examples of tumor types have SBS5, but we want to allow a small chance that SBS5 is not present.
sigs.prop	The proportions of samples that contain each signature. A numerical vector (values between 0 and 1), with names being signature identifiers. Can be the return value from $\texttt{ExposureProportions}$.
sigs	Matrix of signatures.

OptimizeExposure

Optimize the reconstruction of a spectrum from a set of signatures.

Description

Optimize the reconstruction of a spectrum from a set of signatures.

Usage

```
OptimizeExposure(spectrum, sigs, m.opts, ...)
```

Arguments

spectrum The spectrum to be reconstructed.
sigs The available signatures.
m.opts Options that govern the numerical optimization. For documentation see DefaultManyOpts.
... Additional arguments for eval_f.

Value

```
A list with elements

loglh The log likelihood of the best solution (set of exposures) found.

exposure The vector of exposures that generated loglh, i.e. the number of mutations ascribed to each signature.

objective The final value of the objective function.

solution The optimum exposures. Deprecated.

warnings A character vector of warnings.

global.search.diagnostics Diagnostics from nloptr.

local.search.diagnostics Diagnostics from nloptr.
```

OptimizeExposureQP Quadratic programming optimization of signature activities

Description

Quadratic programming optimization of signature activities

Usage

```
OptimizeExposureQP(spectrum, signatures)
```

Arguments

Mutational signature spectrum as a numeric vector or single column data frame or matrix.

Matrix or data frame of signatures from which reconstruct spectrum. Rows are mutation types and columns are signatures. Should have column names for interpretable results. Cannot be a vector because the column names are needed.

Value

A vector of exposures with names being the colnames from signatures. Code adapted from SignatureEstimation::decomposeQP.

```
OptimizeExposureQPBootstrap
```

Bootstrap OptimizeExposureQP and filter exposures by confidence intervals

Description

Bootstrap OptimizeExposureQP and filter exposures by confidence intervals

```
OptimizeExposureQPBootstrap(
   spectrum,
   signatures,
   num.replicates = 10000,
   conf.int = 0.95,
   mc.cores = 10,
   seed = NULL
)
```

PCAWGMAPTest 17

Arguments

spectrum Mutational signature spectrum as a numeric vector or single column data frame or matrix.

01 11111111

Matrix or data frame of signatures from which reconstruct spectrum. Rows are mutation types and columns are signatures. Should have column names for interpretable results. Cannot be a vector because the column names are needed.

num.replicates

signatures

Number of bootstrap replicates.

conf.int Discard signatures with conf.int that overlaps 0.

mc.cores The maximum number of cores to use. On MS Windows machines it defaults to

1.

seed Random seed; set this to get reproducible results.

#' @return A list with elements

exposure The vector of exposures that generated log1h, i.e. the number of mutations ascribed to each signature. The names of exposure are a subset of the colnames (signatures).

euclidean.dist The final value of the objective function.

cosine.sim The cosine similarity between spectrum and the reconstruction based on signatures.

If the spectrum has 0 mutations, no bootstrapping is done, and in the return value all signaures have 0 exposures, euclidian.dist is 0, and cosine.sim is NaN.

PCAWGMAPTest

Run MAPAssignActivity1 on one sample from the PCAWG platinum data set.

Description

Run MAPAssignActivity1 on one sample from the PCAWG platinum data set.

 $Run\ {\tt MAPAssignActivity1}\ on\ one\ sample\ from\ the\ PCAWG\ platinum\ data\ set\ with\ artifact\ signatures\ removed.$

```
PCAWGMAPTest(
  cancer.type,
  sample.index,
  mutation.type,
  max.level = 5,
  max.mc.cores,
  out.dir = NULL,
  p.thresh = 0.01,
  m.opts = DefaultManyOpts(),
  max.presence.proportion = 0.99,
  sigs.prop = NULL
)
```

18 PossibleArtifacts

```
PCAWGMAPTest(
  cancer.type,
  sample.index,
  mutation.type,
  max.level = 5,
  max.mc.cores,
  out.dir = NULL,
  p.thresh = 0.01,
  m.opts = DefaultManyOpts(),
  max.presence.proportion = 0.99,
  sigs.prop = NULL
)
```

Arguments

cancer.type A cancer type from the PCAWG exposures matrix. sample.index The index of the sample within the exposures matrix. mutation.type

One of "SBS96", "SBS192", "ID", "DBS78"

max.level The maximum number of signatures to try removing.

max.mc.cores The maximum number of cores to use. On Microsoft Windows machines it is

silently changed to 1.

out.dir If non-NULL create this directory if necessary and put results there.

p.thresh If the p value for a better reconstruction with than without a set of signatures is

> than p. thresh, then we can use exposures without this set.

max.presence.proportion

The maximum value of the proportion of tumors that must have a given signature. Used so that it is possible to exclude a signature from a spectrum, e.g. perhaps all examples of tumor types have SBS5, but we want to allow a small

chance that SBS5 is not present.

sigs.prop The proportions of samples that contain each signature. A numerical vector

(values between 0 and 1), with names being signature identifiers. Can be the

return value from ExposureProportions.

Value

See OneMAPAssignTest.

A list with two elements, each the result for one call to OneMAPAssignTest.

Possible Artifacts $Return\ a\ character\ vector\ of\ the\ IDs\ of\ possible\ SBS96\ signature\ artifacts.$

Description

Return a character vector of the IDs of possible SBS96 signature artifacts.

```
PossibleArtifacts()
```

RareSignatures 19

RareSignatures

Return a character vector of the IDs of rare SBS96 signatures.

Description

Return a character vector of the IDs of rare SBS96 signatures.

Usage

```
RareSignatures()
```

ReconstructSpectrum

Given signatures (sigs) and exposures (exp), return a spectrum or spectra

Description

Given signatures (sigs) and exposures (exp), return a spectrum or spectra

Usage

```
ReconstructSpectrum(sigs, exp, use.sig.names = FALSE)
```

Arguments

sigs Signature as a matrix or or data frame, with each row one mutation type (g.e.

CCT > CAT or CC > TT) and each column a signature.

exp The exposures for one or more samples as a matrix or data.frame, with each row

a signature and each column a sample.

use.sig.names

If ${\tt TRUE}\ check\ that\ {\tt rownames}\ ({\tt exp})\ is\ a\ subset\ of\ {\tt colnames}\ ({\tt sigs})\ ,$ and

use only the columns in sigs that are present in exp.

Details

Does not care or check if colSums (sigs) == 1. Error checking is minimal since this function is called often.

```
ShowSigActivity Show signature activity from the output generated by AddSigActivity
```

Description

Show signature activity from the output generated by AddSigActivity

Usage

```
ShowSigActivity(
   list.of.sig.activity,
   output.dir,
   plot.all.samples.in.one.pdf = TRUE,
   plot.exposure.proportion = FALSE,
   ...
)
```

Arguments

```
list.of.sig.activity
```

A list of contributing signature activity information for multiple spectra. See the return value of AddSigActivity for more details.

output.dir The directory to save the results. Create this directory if it does not exist. plot.all.samples.in.one.pdf

Whether to plot all the signature activity information within one PDF. Default is TRUE. If FALSE, then plot one PDF for each sample.

plot.exposure.proportion

Whether to plot exposure proportions rather than counts.

... Other arguments passed to PlotCatalogToPdf.

Examples

SignaturePresenceTest

Test whether a given signature is plausibly present in a catalog.

Description

Test whether a given signature is plausibly present in a catalog.

SignaturePresenceTest1 21

Usage

```
SignaturePresenceTest(
   spectra,
   sigs,
   target.sig.index,
   m.opts = NULL,
   mc.cores = 10
)
```

Arguments

The catalog (matrix) to analyze. This could be an ICAMS catalog or a numerical matrix.

sigs A catalog of signatures from which to choose. This could be and ICAMS catalog or a numerical matrix.

target.sig.index
The index of the signature the presence of which we want to test.

m.opts If NULL use the return from calling DefaultManyOpts. For documentation see DefaultManyOpts.

mc.cores Number of cores to use. Always silently changed to 1 on Microsoft Windows.

```
SignaturePresenceTest1
```

Test whether a given signature is plausibly present in a spectrum.

Description

For backward compatibility. See also AnySigSubsetPresent.

Usage

```
SignaturePresenceTest1(spectrum, sigs, target.sig.index, m.opts)
```

Arguments

```
spectrum The spectrum to analyze.

sigs A catalog of signatures from which to choose.

target.sig.index
The index of the signature the presence of which we want to test.

m.opts For documentation see DefaultManyOpts.
```

22 SparseAssignActivity

```
SparseAssignActivity
```

Find known signatures that can most sparsely reconstruct each spectrum in a catalog.

Description

Find known signatures that can most sparsely reconstruct each spectrum in a catalog.

Usage

```
SparseAssignActivity(
  spectra,
  sigs,
  max.level = 5,
  p.thresh = 0.05,
  m.opts = NULL,
  num.parallel.samples = 5,
  mc.cores.per.sample = min(20, 2^max.level)
)
```

Arguments

```
The spectra (multiple spectra) to be reconstructed.
spectra
sigs
                 The known signatures to use in reconstruction.
max.level
                 The largest number of signatures to consider discarding in the reconstruction.
                 The maximum p value based on which it is decided to retain a signature in a
p.thresh
                 reconstruction.
                 For documentation see DefaultManyOpts.
m.opts
num.parallel.samples
                 The (maximum) number of samples to run in parallel; each sample in turn can
                 require multiple cores, as governed by mc.cores.per.sample.
mc.cores.per.sample
                 The maximum number of cores to use for each sample. On Microsoft Windows
```

machines it is silently changed to 1.

Value

A list with the inferred exposure matrix as element exposure.

Index

```
AddSigActivity, 2, 20
AnySigSubsetPresent, 21
AsyncProgress, 9, 11
CancerTypes, 4, 5
cosine, 4
cossim.4
DefaultManyOpts, 4, 9, 11, 15, 18, 21, 22
ExposureProportions, 3, 5, 9, 11, 15, 18
g_ineq_for_ObjFnBinomMaxLH2,6
ICAMS, 3, 9, 11, 21
LLHSpectrumMAP, 6
LLHSpectrumNegBinom, 3, 7, 8, 13
LLHSpectrumNegBinomDebug, 8
MAPAssignActivity, 9
MAPAssignActivity1, 10, 14, 17
NegBinomial, 3, 4, 7, 8, 12, 13
nloptr, 4, 6, 13-15
ObjFnBinomMaxLHMustRound, 12
ObjFnBinomMaxLHNoRoundOK, 13
ObjFnBinomMaxLHRound, 12, 13, 13
OneMAPAssignTest, 14, 18
OptimizeExposure, 15
OptimizeExposureQP, 16, 16
OptimizeExposureQPBootstrap, 16
PCAWGMAPTest, 17
PlotCatalogToPdf, 20
PossibleArtifacts, 18
RareSignatures, 19
ReconstructSpectrum, 3, 19
ShowSigActivity, 20
SignaturePresenceTest, 20
SignaturePresenceTest1, 13, 21
SparseAssignActivity, 13, 22
SparseAssignActivity1, 13
tibble, 8
```