

mgsa

October 25, 2011

MgsaGoSets-class *Gene Ontology annotations*

Description

This class represents gene ontology annotations.

Details

For now, it is identical to the parental class [MgsaSets](#).

Extends

[MgsaSets](#)

See Also

[readGAF](#)

MgsaMcmcResults-class
MgsaMcmcResults-class

Description

Instances of this class are used to hold the additional information that was provided by running (possibly multiple times) an MCMC algorithm.

Slots

steps: (*numeric*) how many steps per MCMC run

restarts: (*numeric*) how many MCMC runs

alphaMcmcPost: (*matrix*) posterior estimates for each MCMC run of the parameter alpha

betaMcmcPost: (*matrix*) posterior estimates for each MCMC run of the parameter beta

pMcmcPost: (*matrix*) posterior estimates for each MCMC run of the parameter p

setsMcmcPost: ([matrix](#)) posterior estimates for each MCMC run of the sets marginal posterior probabilities

The columns of the matrices `alphaMcmcPost`, `betaMcmcPost`, `pMcmcPost` and `setsMcmcPost` stores the posterior estimates for each individual MCMC run. The row order matches the one of the slot `alphaPost`, `betaPost`, `pPots`, and `setsResults` respectively.

Accessor methods exist for each slot.

Extends

[MgsaResults](#)

Methods

[steps](#)

[restarts](#)

[alphaMcmcPost](#)

[betaMcmcPost](#)

[pMcmcPost](#)

[setsMcmcPost](#)

See Also

[mgsa](#)

MgsaResults-class *Results of an MGSA analysis*

Description

The results of an MGSA analysis.

Slots

populationSize: ([numeric](#)) The number of items in the population.

studySetSizeInPopulation: ([numeric](#)) The number of items both in the study set and in the population.

alphaPost: ([data.frame](#)) with columns `value`, `estimate` and `std.error`.

betaPost: ([data.frame](#)) with columns `value`, `estimate` and `std.error`.

pPost: ([data.frame](#)) with columns `value`, `estimate` and `std.error`.

setsResults: ([data.frame](#)) with columns `inPopulation`, `inStudySet`, `estimate` and `std.error`.

The columns of the slots `alphaPost`, `betaPost`, and `pPost` contains a realization value, its posterior estimate and standard error for the parameters `alpha`, `beta` and `p` respectively.

The columns of the slot `setsResults` contains the number of items of the set in the population, the number of items of the set in the study set, the estimate of its marginal posterior probability and its standard error. The [rownames](#) are the names of the sets if available.

Because an `MgsaResults` is the outcome of an MGSA analysis (see [mgsa](#)), accessors but no replacement methods exist for each slot.

Methods

[populationSize](#)
[studySetSizeInPopulation](#)
[alphaPost](#)
[betaPost](#)
[pPost](#)
[setsResults](#)
[show](#) signature(= "MgsaResults"): Show an [MgsaResults](#).
[plot](#) signature(= "MgsaResults"): Plot method for [MgsaResults](#) objects

See Also

[mgsa](#)

MgsaSets-class

Sets of items and their annotations

Description

This class describes sets, items and their annotations.

Details

Internally, the method [mgsa](#) indexes all elements of the sets before fitting the model. In case [mgsa](#) must be run on several observations with the same gene sets, computations can be speeded up by performing this indexing once for all. This can be achieved by building a [MgsaSets](#). In order to ensure consistency of the indexing, no replace method for any slot is provided. Accessors are available.

The data frames `setAnnotations` and `itemAnnotations` allow to store annotations. No constraint is imposed on the number and names of their columns.

Slots

sets: ([list](#)) A list whose elements are vector of item indices.
itemName2ItemIndex: ([integer](#)) The mapping of item names to index.
numberOfItems: ([integer](#)) How many items?
setAnnotations: ([data.frame](#)) Annotations of the sets. The [rownames](#) are set names.
itemAnnotations: ([data.frame](#)) Annotations of the items. The [rownames](#) are item names.

Methods

initialize signature(= "MgsaSets"): Initializes the mapping when some parameters are not specified
[itemAnnotations](#)
[itemAnnotations](#)
[setAnnotations](#)

```

setAnnotations
length signature( = "MgsaSets"): Length (number of sets) of MgsaSets.
itemIndices
itemIndices
show signature( = "MgsaSets"): Show an MgsaSets.
subMgsaSets
mgsa

```

See Also

[MgsaGoSets](#), [readGAF](#), [mgsa](#)

Examples

```
new("MgsaSets", sets=list(set1=c("a", "b"), set2=c("b", "c")))
```

alphaMcmcPost-methods

posterior estimates of the parameter alpha for each MCMC run

Description

Posterior estimates of the parameter alpha for each MCMC run.

Usage

```
## S4 method for signature 'MgsaMcmcResults'
alphaMcmcPost(x)
```

Arguments

x a [MgsaMcmcResults](#).

Value

alphaMcmcPost-methods: matrix: Posterior estimates of the parameter alpha for each MCMC run.

alphaPost-methods *Posterior for alpha*

Description

Realization values, posterior estimate and standard error for the parameter alpha.

Usage

```
## S4 method for signature 'MgsaResults'  
alphaPost(x)
```

Arguments

x a *MgsaResults*.

Value

alphaPost-methods: data.frame: realization values, posterior estimate and standard error for the parameter alpha.

betaMcmcPost-methods
posterior estimates of the parameter beta for each MCMC run

Description

Posterior estimates of the parameter beta for each MCMC run.

Usage

```
## S4 method for signature 'MgsaMcmcResults'  
betaMcmcPost(x)
```

Arguments

x a *MgsaMcmcResults*.

Value

betaMcmcPost-methods: matrix: Posterior estimates of the parameter beta for each MCMC run.

betaPost-methods *Posterior for beta*

Description

Realization values, posterior estimate and standard error for the parameter beta.

Usage

```
## S4 method for signature 'MgsaResults'  
betaPost(x)
```

Arguments

x a [MgsaResults](#).

Value

betaPost-methods: data.frame: realization values, posterior estimate and standard error for the parameter beta.

example-o-data *Example objects for mgsa*

Description

This are example objects for mgsa.

example-go-data *Example GO sets for mgsa*

Description

This are example GO sets for mgsa.

itemAnnotations-methods

Item annotations of a MgsaSets

Description

Item annotations of a [MgsaSets](#).

Usage

```
## S4 method for signature 'MgsaSets,missing'  
itemAnnotations(sets, items)  
## S4 method for signature 'MgsaSets,character'  
itemAnnotations(sets, items)
```

Arguments

sets an instance of class [MgsaSets](#).
items character an optional vector specifying the items of interest.

Value

itemAnnotations-methods: a `data.frame`: the item annotations.

itemIndices-methods

Item indices of a MgsaSets

Description

Returns the indices corresponding to the items

Usage

```
## S4 method for signature 'MgsaSets,character'  
itemIndices(sets, items)  
## S4 method for signature 'MgsaSets,numeric'  
itemIndices(sets, items)
```

Arguments

sets an instance of class [MgsaSets](#).
items character or numeric the items of interest.

Value

itemIndices-methods: a integer: the item indices.

length-methods *Length of a MgsaSets.*

Description

Length (number of sets) of [MgsaSets](#).

Usage

```
## S4 method for signature 'MgsaSets'
length(x)
```

Arguments

x an instance of class [MgsaSets](#).

Value

integer vector.

mgsa-methods *Performs an MGSA analysis*

Description

Estimate marginal posterior of the MGSA problem with an MCMC sampling algorithm.

Usage

```
mgsa(
o, sets, population=NULL,
alpha=seq(0.01,0.3, length.out=10), beta=seq(0.1,0.8, length.out=10),
p=seq( min(0.1, 1/length(sets)), min(0.3, 20/length(sets)), length.out=10),
steps=1e6, restarts=5, threads=0
)
```

Arguments

o The observations. It can be a numeric, integer, character or logical. See details.

sets The sets. It can be an [MgsaSets](#) or a list. In this case, each list entry is a vector of type numeric, integer, character. See details.

population The total population. Optional. A numeric, integer or character vector. Default to NULL. See details.

alpha Grid of values for the parameter alpha. Values represent probabilities of false-positive events and hence must be in [0,1]. numeric.

beta Grid of values for the parameter beta. Values represent probabilities of false-negative events and hence must be in [0,1]. numeric.

<code>p</code>	Grid of values for the parameter <code>p</code> . Values represent probabilities of term activity and therefore must be in <code>[0,1]</code> . <code>numeric</code> .
<code>steps</code>	The number of steps of each run of the MCMC sampler. <code>integer</code> of length 1. A recommended value is <code>1e6</code> or greater.
<code>restarts</code>	The number of different runs of the MCMC sampler. <code>integer</code> of length 1. Must be greater or equal to 1. A recommended value is 5 or greater.
<code>threads</code>	The number of threads that should be used for concurrent restarts. A value of 0 means to use all available cores. Default to 0.

Details

`mgsa-methods`: The function can handle items (such as genes) encoded as `character` or `integer`. For convenience `numeric` items can also be provided but these values should essentially be integers. The type of items in the observations `o`, the sets and in the optional population should be consistent. In the case of `character` items, `o` and population should be of type `character` and sets can either be an `MgsaSets` or a list of `character` vectors. In the case of `integer` items, `o` should be of type `integer`, `numeric` (but essentially with integer values), or `logical` and entries in sets as well as the population should be `integer`. When `o` is `logical`, it is first coerced to `integer` with a call on `which`. Observations outside the population are not taken into account. If population is `NULL`, it is defined as the union of all sets.

The default grid value for `p` is such that between 1 and 20 sets are active in expectation. The lower limit is constrained to be lower than 0.1 and the upper limit lower than 0.3 independently of the total number of sets to make sure that complex solutions are penalized. Marginal posteriors of activity of each set are estimated using an MCMC sampler as described in Bauer et al., 2010. Because convergence of an MCM sampler is difficult to assess, it is recommended to run it several times (using `restarts`). If variations between runs are too large (see `MgsaResults`), the number of steps (`steps`) of each MCMC run should be increased.

Value

`mgsa-methods`: An `MgsaMcmcResults` object.

References

Bauer S., Gagneur J. and Robinson P. GOing Bayesian: model-based gene set analysis of genome-scale data. *Nucleic Acids Research* (2010) <http://nar.oxfordjournals.org/content/38/11/3523.full>

See Also

[MgsaResults](#), [MgsaMcmcResults](#)

Examples

```
## observing items A and B, with sets {A,B,C} and {B,C,D}
mgsa(c("A", "B"), list(set1 = LETTERS[1:3], set2 = LETTERS[2:4]))

## same case with integer representation of the items and logical observation
mgsa(c(TRUE,TRUE,FALSE,FALSE), list(set1 = 1:3, set2 = 2:4))

## a small example with gene ontology sets and plot
data(example)
```

```
fit = mgsa(example_o, example_go)
## Not run:
plot(fit)
## End(Not run)
```

mgsa-package

Model-based gene set analysis

Description

Model-based Gene Set Analysis (MGSA) is a Bayesian modeling approach for gene set enrichment. The package mgsa implements MGSA and tools to use MGSA together with the Gene Ontology.

Author(s)

Sebastian Bauer <Sebastian.Bauer@charite.de>, Julien Gagneur <julien.gagneur@embl.de>

References

S. Bauer, J. Gagneur and P. N. Robinson. GOing Bayesian: model-based gene set analysis of genome-scale data. Nucleic acids research, 2010.

pMcmcPost-methods

posterior estimates of the parameter p for each MCMC run

Description

Posterior estimates of the parameter p for each MCMC run.

Usage

```
## S4 method for signature 'MgsaMcmcResults'
pMcmcPost(x)
```

Arguments

x a [MgsaMcmcResults](#).

Value

pMcmcPost-methods: matrix: Posterior estimates of the parameter p for each MCMC run.

populationSize-methods

Size of the population of a MgsaResults

Description

The size of the population on which the analysis was run.

Usage

```
## S4 method for signature 'MgsaResults'
populationSize(x)
```

Arguments

x a *MgsaResults*.

Value

populationSize-methods: integer: the size of the population.

readGAF

Read a Gene Ontology annotation file

Description

Creates a *MgsaGoSets* using gene ontology annotations provided by a file in GAF 1.0 or 2.0 format.

Usage

```
readGAF(filename, evidence=NULL, aspect=c("P", "F", "C"))
```

Arguments

filename	The name of the Gene Ontology annotation file. It must be in the GAF 1.0 or 2.0 format. It may be gzip-compressed.
evidence	character or NULL. Only annotations with evidence code in <i>evidence</i> are returned. If NULL (default), annotations of all evidence codes are returned.
aspect	character with values in P, C or F. Only annotations of the listed GO namespaces P (biological process), F (molecular function) or C (cellular component) are returned. By default, annotations of the three namespaces are returned.

Details

The function extracts from the annotation file all direct gene annotations and infers from the Gene Ontology all the indirect annotations (due to term relationships). This is done using the package *Go.db* which provides the ontology as a database and *RSQLite* for querying the database.

Value

An `MgsaGoSets` object.

References

The Gene Ontology Consortium. Gene Ontology: tool for the unification of biology. Nature Genetics, 2000. The GAF file format: <http://www.geneontology.org/GO.format.annotation.shtml> GO evidence codes: <http://www.geneontology.org/GO.evidence.shtml>

See Also

`MgsaGoSets`, `mgsa`

Examples

```
## parsing provided example file (yeast)
gofile = system.file("example_files/gene_association_head.sgd", package="mgsa")
readGAF(gofile)
## only annoations infered from experiment or a direct assay
readGAF(gofile, evidence=c("EXP", "IDA"))
```

restarts-methods *How many MCMC runs*

Description

how many MCMC runs.

Usage

```
## S4 method for signature 'MgsaMcmcResults'
restarts(x)
```

Arguments

x a `MgsaMcmcResults`.

Value

restarts-methods: integer: how many MCMC runs.

setAnnotations-methods

Set annotations of a MgsaSets

Description

Set annotations of a [MgsaSets](#).

Usage

```
## S4 method for signature 'MgsaSets,missing'
setAnnotations(sets, names)
## S4 method for signature 'MgsaSets,character'
setAnnotations(sets, names)
```

Arguments

sets an instance of class [MgsaSets](#).
 names character an optional vector specifying the names of interest.

Value

setAnnotations-methods: a `data.frame`: the set annotations.

setsMcmcPost-methods

posterior estimates of the the set marginal probabilities for each MCMC

Description

Posterior estimates of the set marginal probabilities for each MCMC run.

Usage

```
## S4 method for signature 'MgsaMcmcResults'
setsMcmcPost(x)
```

Arguments

x a [MgsaMcmcResults](#).

Value

setsMcmcPost-methods: `matrix`: Posterior estimates of the set marginal probabilities for each MCMC run.

`setsResults-methods`*Posterior for each set*

Description

Number of items of the set in the population, the number of items of the set in the study set, the estimate of its marginal posterior probability and its standard error.

Usage

```
## S4 method for signature 'MgsaResults'  
setsResults(x)
```

Arguments

`x` a [MgsaResults](#).

Value

`setsResults-methods`: `data.frame`: For each set, number of items of the set in the population, number of items of the set in the study set, estimate of its marginal posterior probability and standard error.

`show, MgsaResults-method`*Show an MgsaResults*

Description

Show an [MgsaResults](#).

Usage

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

Arguments

`object` an instance of class [MgsaResults](#).

Value

an invisible `NULL`

show, MgsaSets-method

Show an MgsaSets

Description

Show an [MgsaSets](#).

Usage

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

Arguments

object an instance of class [MgsaSets](#).

Value

an invisible NULL

steps-methods

How many steps per MCMC run

Description

how many steps per MCMC run.

Usage

```
## S4 method for signature 'MgsaMcmcResults'  
steps(x)
```

Arguments

x a [MgsaMcmcResults](#).

Value

steps-methods: integer: how many steps per MCMC run.

studySetSizeInPopulation-methods
Size of the study set of a MgsaResults

Description

The size of the study set on which the analysis was run.

Usage

```
## S4 method for signature 'MgsaResults'  
studySetSizeInPopulation(x)
```

Arguments

x a [MgsaResults](#).

Value

studySetSizeInPopulation-methods: integer: the size of the study set.

subMgsaSets-methods
Subset of an MgsaSets

Description

Returns a subset of an [MgsaSets](#) in which only the required items are kept. Empty sets are removed.

Usage

```
## S4 method for signature 'MgsaSets,numeric'  
subMgsaSets(sets, items)
```

Arguments

sets an [MgsaSets](#).
items numeric. The items to restrict on.

Value

subMgsaSets-methods: an [MgsaSets](#).

Note

TODO: do we need that method? Does it work on items or item indexes?

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