Package ‘mgsa’

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Author Sebastian Bauer <mail@sebastianbauer.info>, Julien Gagneur <gagneur@genzentrum.lmu.de>
Maintainer Sebastian Bauer <mail@sebastianbauer.info>
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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

alphaMcmcPost

posterior estimates of the parameter alpha for each MCMC run

Description

Posterior estimates of the parameter alpha for each MCMC run.

Usage

alphaMcmcPost(x)

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

Arguments

x a MgsaMcmcResults.

Value

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

alphaPost

Posterior for alpha

Description

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

Usage

alphaPost(x)

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

Arguments

x a MgsaResults.

Value

data.frame: realization values, posterior estimate and standard error for the parameter alpha.
**betaMcmcPost**

*posterior estimates of the parameter beta for each MCMC run*

**Description**

Posterior estimates of the parameter beta for each MCMC run.

**Usage**

```r
betaMcmcPost(x)
```

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

**Arguments**

- `x` a `MgsaMcmcResults`.

**Value**

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

---

**betaPost**

*Posterior for beta*

**Description**

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

**Usage**

```r
betaPost(x)
```

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

**Arguments**

- `x` a `MgsaResults`.

**Value**

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

This function takes a 1:1 mapping of go.ids to items and returns a full MgsaGOSets instance. The structure of GO is gathered from GO.db. It is sufficient to specify just the directly asserted mapping (or annotation), i.e., the most specific ones. The true path rule is taken account, that is, if an item is annotated to a term then it will be also annotated to more general terms (some people prefer to say that just the transitive closure is calculated).

**Description**

This function takes a 1:1 mapping of go.ids to items and returns a full MgsaGOSets instance. The structure of GO is gathered from GO.db. It is sufficient to specify just the directly asserted mapping (or annotation), i.e., the most specific ones. The true path rule is taken account, that is, if an item is annotated to a term then it will be also annotated to more general terms (some people prefer to say that just the transitive closure is calculated).

**Usage**

`createMgsaGoSets(go.ids, items)`

**Arguments**

- **go.ids**
  - a character vector of GO ids (GO:00001234)

- **items**
  - a vector of identifiers that are annotated to the term in the corresponding position of the go.ids vector.

**Example go**

Example GO sets for mgsa

**Description**

This data is an example GO set for mgsa.

**Example o**

Example objects for mgsa

**Description**

This data is an example objects for mgsa.
itemAnnotations

*Item annotations of a MgsaSets*

**Description**
Item annotations of a MgsaSets.

**Usage**

```r
itemAnnotations(sets, items)
```

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

```r
## 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**
a data.frame: the item annotations.

---

itemIndices

*Item indices of a MgsaSets*

**Description**
Returns the indices corresponding to the items.

**Usage**

```r
itemIndices(sets, items)
```

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

```r
## 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

- a integer: the item indices.

---

**Description**

Length (number of sets) of `MgsaSets`.

**Usage**

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

Arguments

- x: an instance of class `MgsaSets`.

Value

- integer vector.

---

**mgsa**

*Performs an MGSA analysis*

**Description**

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

mgsa(o, sets, population = NULL,
   p = seq(min(0.1, 1/length(sets)), min(0.3, 20/length(sets)), length.out = 10),
   ...)

## S4 method for signature 'integer,list'
mgsa(o, sets, population = NULL,
   p = seq(min(0.1, 1/length(sets)), min(0.3, 20/length(sets)), length.out = 10),
   ...)

## S4 method for signature 'numeric,list'
mgsa(o, sets, population = NULL,
   p = seq(min(0.1, 1/length(sets)), min(0.3, 20/length(sets)), length.out = 10),
   ...)

## S4 method for signature 'character,list'
mgsa(o, sets,
   population = NULL,
   p = seq(min(0.1, 1/length(sets)), min(0.3, 20/length(sets)), length.out = 10),
   ...)

## S4 method for signature 'logical,list'
mgsa(o, sets, population = NULL,
   p = seq(min(0.1, 1/length(sets)), min(0.3, 20/length(sets)), length.out = 10),
   ...)

## S4 method for signature 'character,MgsaSets'
mgsa(o, sets,
   population = NULL,
   p = seq(min(0.1, 1/length(sets)), min(0.3, 20/length(sets)), length.out = 10),
   ...)

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.

- **p**
  Grid of values for the parameter p. Values represent probabilities of term activity and therefore must be in [0,1].

- **...**
  Optional arguments that are passed to the methods. Supported parameters are
  - **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.
steps  The number of steps of each run of the MCMC sampler. integer of length 1. A recommended value is 1e6 or greater.

burnin  The number of burn-in MCMC steps, until sample collecting begins. integer of length 1. A recommended value is half of total MCMC steps.

thin  The sample collecting period. An integer of length 1. A recommended value is 100 to reduce autocorrelation of subsequently collected samples.

flip.freq  The frequency of MCMC Gibbs step that randomly flips the state of a random set from active to inactive or vice versa. numeric from (0,1].

restarts  The number of different runs of the MCMC sampler. integer of length 1. Must be greater or equal to 1. A recommended value is 5 or greater.

threads  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

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 \texttt{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 \texttt{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 \texttt{MgsaResults}), the number of steps (steps) of each MCMC run should be increased.

Value

An \texttt{MgsaMcmcResults} object.

References


See Also

\texttt{MgsaResults,MgsaMcmcResults}
Examples

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

## same case with integer representation of the items and logical observation
msga(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)
```

---

### MgsaGoSets

**Gene Ontology annotations**

**Description**

This class represents gene ontology annotations.

**Details**

For now, it is identical to the parental class `MgsaSets`.

**See Also**

`readGAF`

---

### MgsaMcmcResults

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

**Description**

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

Slots

- nsamplshow many samples collected per MCMC run
- stepshow many steps per MCMC run
- restarthow many MCMC runs
- alphanmcmcPostposterior estimates for each MCMC run of the parameter alpha
- betanmcmcPostposterior estimates for each MCMC run of the parameter beta
- pmnmcmcPostposterior estimates for each MCMC run of the parameter p
- setsnmcPostposterior estimates for each MCMC run of the sets marginal posterior probabilities

The columns of the matrices alphanmcmcPost, betanmcmcPost, pmnmcmcPost and setsnmcPost 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.

See Also

mgsa

MgsaResults

Results of an MGSA analysis

Description

The results of an MGSA analysis.

Slots

- populationSizeThe number of items in the population.
- studySetSizeInPopulationThe number of items both in the study set and in the population.
- alphapostwith columns value, estimate and std.error.
- betapostwith columns value, estimate and std.error.
- ppostwith columns value, estimate and std.error.
- setsresultswith 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.

See Also

mgsa
MgsaSets

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: A list whose elements are vector of item indices.
- itemName2ItemIndex: The mapping of item names to index.
- numberOfItems: How many items?
- setAnnotations: Annotations of the sets. The rownames are set names.
- itemAnnotations: Annotations of the items. The rownames are item names.

See Also

MgsaGoSets, readGAF, mgsa

Examples

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

nsamples

How many samples per MCMC run collected

Description

how many samples collected per MCMC run.

Usage

nsamples(x)

### S4 method for signature 'MgsaMcmcResults'
nsamples(x)
Arguments

- `x`: a `MgsaMcmcResults`.

Value

integer: how many samples per MCMC run collected.

---

Plot method for `MgsaResults` objects

### Usage

```r
plotMgsaResults(x, y, ...)```

Arguments

- `x`: a `MgsaResults`.
- `y`: unused
- `...`: unused

---

Posterior estimates of the parameter `p` for each MCMC run

### Usage

```r
pMcmcPost(x)
```

Arguments

- `x`: a `MgsaMcmcResults`.

Value

matrix: Posterior estimates of the parameter `p` for each MCMC run.
populationSize  
*Size of the population of a MgsaResults*

**Description**

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

**Usage**

```r
populationSize(x)
```

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

**Arguments**

- `x` a `MgsaResults`.

**Value**

integer: the size of the population.

---

pPost  
*Posterior for beta*

**Description**

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

**Usage**

```r
pPost(x)
```

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

**Arguments**

- `x` a `MgsaResults`.

**Value**

data.frame: realization values, posterior estimate and standard error for the parameter p.
**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**

```r
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 evidence 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**


**See Also**

- `MgsaGoSets`
- `mgsa`

**Examples**

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

**How many MCMC runs**

**Description**

How many MCMC runs.

**Usage**

```r
restarts(x)
```

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

**Arguments**

- `x` a `MgsaMcmcResults`.

**Value**

integer: how many MCMC runs.

---

### setAnnotations

**Set annotations of a MgsaSets**

**Description**

Set annotations of a `MgsaSets`.

**Usage**

```r
setAnnotations(sets, names)
```

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

```R
## 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**

a `data.frame`: the set annotations.
setsMcmcPost

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

**Usage**
```r
setsMcmcPost(x)
```

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

**Arguments**
- `x`: a `MgsaMcmcResults`.

**Value**
`matrix`: Posterior estimates of the set marginal probabilities for each MCMC run.

setsResults

**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**
```r
setsResults(x)
```

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

**Arguments**
- `x`: a `MgsaResults`.

**Value**
`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

---

**Description**

How many steps per MCMC run.

**Usage**

```r
steps(x)
```

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

**Arguments**

- `x` a `MgsaMcmcResults`.

**Value**

- integer: how many steps per MCMC run.

---

studySetSizeInPopulation

---

**Description**

Size of the study set of a MgsaResults

**Usage**

```r
studySetSizeInPopulation(x)
```

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

**Arguments**

- `x` a `MgsaResults`.

**Value**

- integer: the size of the study set.
subMgsaSets  

 Subset of an MgsaSets

Description

Returns a subset of an MgsaSets that contains only the specified items. Empty sets are removed.

Usage

```r
subMgsaSets(sets, items)
```

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

Arguments

- `sets` an MgsaSets.
- `items` character. The items to restrict on.

Value

an MgsaSets.
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