

# Package ‘newFocus’

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**Type** Package

**Title** True Discovery Guarantee by the New Focus Level Procedure

**Version** 1.0

**Date** 2020-07-09

**Author** Ningning Xu

**Maintainer** Ningning Xu <n.xu@lumc.nl>

**Description** A new focus level procedure is developed based on the old focus level procedure of Goeman and Mansmann (2008) <doi:10.1093/bioinformatics/btm628> and the closed testing procedure with globaltest of Xu and Goeman (2020) <arXiv:2001.01541>. It produces the lower bound for the number of true discoveries in any gene set or GO (Gene Ontology) term. It is more powerful for the focus level GO terms chosen by user before seeing the data than the non-focus level GO terms or gene sets chosen after seeing the data.

**License** GPL (>= 2)

**Depends** ctgt

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 newFocus-package

*True Discovery Guarantee by the New Focus Level Procedure*


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## Description

A new focus level procedure is developed based on the old focus level procedure of Goeman and Mansmann (2008) <doi:10.1093/bioinformatics/btm628> and the closed testing procedure with globaltest of Xu and Goeman (2020) <arXiv:2001.01541>. It produces the lower bound for the number of true discoveries in any gene set or GO (Gene Ontology) term. It is more powerful for the focus level GO terms chosen by user before seeing the data than the non-focus level GO terms or gene sets chosen after seeing the data.

## Details

The DESCRIPTION file:

```

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```

Index of help topics:

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newFocus-package	True Discovery Guarantee by the New Focus Level Procedure
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For the GO (Gene Ontology) terms chosen as focus level nodes, [newFocus](#) function will return the number. For GO terms that are non-focus level nodes, we use [pick](#) to count the number of true discoveries based on the result of [newFocus](#).

## Author(s)

Ningning Xu

Maintainer: Ningning Xu <n.xu@lumc.nl>

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choosepath	<i>A set of focus set index</i>
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**Description**

The function aims to find out the focus set index for which the true discoveries is the most and all other focus sets that are disjoint with it .

**Usage**

```
choosepath(startingindex = 1, fsets, lowdv)
```

**Arguments**

startingindex	The index of focus set that has the first largest number of true discoveries
fsets	A list of focus level gene sets, or GO (Gene Ontology) terms
lowdv	A non-negative integer vector, which are the number of true discoveries, the length of the vector is the same as the list of focus level sets

**Value**

The function will return an integer or a numeric vector.

**Author(s)**

Ningning Xu

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ctbab	<i>Closed testing with branch and bound</i>
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**Description**

Closed testing with branch and bound algorithm specifically for globaltest

**Usage**

```
ctbab(y, Cm, Tm, upnode, level, lownode, tmin, ctrue, lf, ls, alpha, count = 0, maxIt = 0)
```

**Arguments**

y	The response variable
Cm	The matrix for calculating critical values of globaltest
Tm	The matrix for calculating test statistics of globaltest
upnode	The upper node that is used to bound critical values
level	The level that the GO term of interest
lownode	The lower node that is used to bound critical values
tmin	The minimum test statistic
ctrue	The true critical value corresponding to the minimum test statistic
lf	The lambda vector corresponding to the upper node
ls	The lambda vector corresponding to the lower node
alpha	The significance level
count	An integer stores the repetitions of the branch and bound, i.e. how many time branch and bound is implemented
maxIt	The maximal number of repetitions prespecified by user

**Value**

It will return the rejection indicator by closed testing with branch and bound algorithm.

**Author(s)**

Ningning Xu

**References**

Xu, N., & Goeman, J. (2020). Closed testing with Globaltest with applications on metabolomics data. arXiv preprint arXiv:2001.01541.

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discov

*True discoveries*

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**Description**

True discoveries calculated by the partial closed testing

**Usage**

```
discov(response, alternative, null, data, maxit = 0, alpha)
```

**Arguments**

response	The response variable
alternative	The alternative hypothesis, which is a character vector, i.e. a set of genes
null	The null hypothesis
data	A data frame with response and all covariates included
maxit	The maximal number of repetitions prespecified by user
alpha	The significance level

**Value**

It will return a non-negative integer: the lower bound for the number of true discoveries of the alternative gene set.

**Author(s)**

Ningning Xu

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newFocus	<i>The new focus level procedure</i>
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**Description**

The new focus level procedure for calculating true discoveries for focus level nodes

**Usage**

```
newFocus(response, fsets, null, data, maxit = 0, alpha = 0.05, adj = 0)
```

**Arguments**

response	The response variable
fsets	A list of focus level sets
null	The null hypothesis
data	The data frame with response and all covariates included
maxit	The maximal number of repetitions prespecified by user
alpha	The significance level
adj	The number of focus sets that are fully rejected by partial closed testing, which is used to adjust the number of focus sets, The default value is 0.

**Value**

The function will return a focus subject with the lower bound for each focus level node and the focus level nodes

**Author(s)**

Ningning Xu

**References**

Goeman, J. J., & Mansmann, U. (2008). Multiple testing on the directed acyclic graph of gene ontology. *Bioinformatics*, 24(4), 537-544.

**Examples**

```
## example data set
n= 100
m = 5
X = matrix(0, n, m,byrow = TRUE )
for ( i in 1:n){
  set.seed(1234+i)
  X[i,] = as.vector(arima.sim(model = list(order = c(1, 0, 0), ar = 0.2), n = m) )
}
y = rbinom(n,1,0.6)
X[which(y==1),1:3] = X[which(y==1),1:3] + 0.8
xs = paste("x",seq(1,m,1),sep="")
colnames(X) = xs

mydata = as.data.frame(cbind(X,y))

## focus level sets
f1 = list(c("x1", "x2"), c("x3", "x4"), "x5")
names(f1) = c("12", "34", "5")

## get td for focus level sets
focus_subject = newFocus(response = y, fsets = f1, data = mydata)

## get td for any set of interest given the focus subject
setofinterest = c("x1", "x2","x3", "x4")
pick(focus_subject, setofinterest)
```

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pick

*True discoveries for non-focus level node*

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**Description**

The number of true discoveries for the non-focus level GO terms is calculated given the focus subject.

**Usage**

```
pick(focus_obj, setofinterest)
```

**Arguments**

focus\_obj      The focus subject from function [newFocus](#)  
setofinterest    A gene set or GO term of interest

**Value**

It will return an integer: the lower bound for the number of true discoveries in the set of interest

**Author(s)**

Ningning Xu

**Examples**

```
## example data set
n= 100
m = 5
X = matrix(0, n, m,byrow = TRUE )
for ( i in 1:n){
  set.seed(1234+i)
  X[i,] = as.vector(arima.sim(model = list(order = c(1, 0, 0), ar = 0.2), n = m) )
}
y = rbinom(n,1,0.6)
X[which(y==1),1:3] = X[which(y==1),1:3] + 0.8
xs = paste("x",seq(1,m,1),sep="")
colnames(X) = xs

mydata = as.data.frame(cbind(X,y))

## focus level sets
f1 = list(c("x1", "x2"), c("x3", "x4"), "x5")
names(f1) = c("12", "34", "5")

## get td for focus level sets
focus_subject = newFocus(response = y, fsets = f1, data = mydata)

## get td for any set of interest given the focus subject
setofinterest = c("x1", "x2","x3", "x4")
pick(focus_subject, setofinterest)
```

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