

# Package ‘DiffNetFDR’

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**Type** Package

**Title** DiffNetFDR: Differential Network Analysis with False Discovery Rate Control

**Version** 1.0

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**Description** DiffNetFDR is an R package which implements two differential network analysis methods. The methods included in DiffNetFD control the overall false discovery rate at a desired level using a multiple testing procedure.

**License** GPL (>= 2)

**Encoding** UTF-8

**NeedsCompilation** no

**Depends** R (>= 3.0), igraph, glmnet, parallel

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DiffNetFDR-package	<i>DiffNetFDR: Differential Network Analysis with False Discovery Rate Control</i>
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## Description

DiffNetFDR is an R package for differential network analysis with false discovery rate control.

## Details

We develop a user-friendly R package, named DiffNetFDR, to implement two differential network analysis methods. Compared methods implemented in existing tools, the methods included in DiffNetFDR have the following features: (i) they are based on Gaussian graphical models which can capture the conditional dependencies; (ii) they can determine the tuning parameters in a data-driven manual; (iii) they take a multiple testing method to control the overall false discovery rate; and (iv) one of them defines the differential network based on partial correlations so that the spurious differential edges caused by variants of conditional variances can be excluded. We also develop a Shiny application to provide easier analysis and visualization.

## Author(s)

Xiao-Fei Zhang

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## References

Xiao-Fei Zhang, Le Ou-Yang, Shuo Yang, Xiaohua Hu and Hong Yan, DiffNetFDR: Differential network analysis with false discovery rate control, 2018.  
 Yin Xia, Tianxi Cai and T. Tony Cai, Testing differential networks with applications to the detection of gene-gene interactions, *Biometrika*, 102(2): 247-266, 2015.  
 Weidong Liu, Structural similarity and difference testing on multiple sparse Gaussian graphical models, *Annals of Statistics*, 45(6): 2680-2707, 2017.

## See Also

[DiffNet.FDR](#), [TCGA.BRCA](#), [GSE13159.AML](#)

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DiffNet.FDR

*Differential network analysis with false discovery rate control*

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## Description

This function is implemented to infer differential network based on Gaussian graphical models. The differential network can be determined as the difference of partial correlations or the difference of precision matrices. The false discovery rate can be controlled using a multiple testing procedure.

## Usage

```
DiffNet.FDR(X, group, alpha = 0.2, test.type = "pcor", parallel = F, nCpus = 4)
```

## Arguments

X	data matrix for which the rows represent the samples and the columns represent the variables.
group	vector which defines two groups of samples under comparison.
alpha	desired FDR level ( $0 \leq \alpha \leq 1$ ), default to 0.2.

<code>test.type</code>	parameter to decide which approach is selected to define the differential network, default to "pcor". If <code>test.type = "pcor"</code> , the differential network will be inferred based on the difference of partial correlations. If <code>test.type = "pmat"</code> , the differential network will be inferred based on the difference of precision matrices.
<code>parallel</code>	logical value to indicate if the process should be run parallelly in multiple threads, default to FALSE.
<code>nCpus</code>	number of (maximum) cores to use for parallel execution, default to 4.

### Details

For details, please refer to Supplementary Information for the article: DiffNetFDR: Differential network analysis with false discovery rate control.

### Value

<code>Diff.edge</code>	the adjacency matrix of the estimated differential network.
<code>W</code>	the values of the statistics.
<code>t</code>	the threshold level such that $H_0$ is rejected.
<code>Diff.net</code>	the estimated differential network over all nodes.
<code>Diff.net.connected</code>	the estimated differential network over only the connected nodes.

### Author(s)

Xiao-Fei Zhang Maintainer: Xiao-Fei Zhang <zhangxf@mail.ccnu.edu.cn>

### References

Xiao-Fei Zhang, Le Ou-Yang, Shuo Yang, Xiaohua Hu and Hong Yan, DiffNetFDR: Differential network analysis with false discovery rate control, 2018  
 Yin Xia, Tianxi Cai and T. Tony Cai, Testing differential networks with applications to the detection of gene-gene interactions, *Biometrika*, 102(2): 247-266, 2015  
 Weidong Liu, Structural similarity and difference testing on multiple sparse Gaussian graphical models, *Annals of Statistics*, 45(6): 2680-2707, 2017.

### See Also

[TCGA.BRCA](#), [GSE13159.AML](#)

### Examples

```
rm(list=ls())
data("TCGA.BRCA")

result = DiffNet.FDR(TCGA.BRCA$X,TCGA.BRCA$group, alpha = 0.2, test.type = "pcor")
Diff.net.connected = result$Diff.net.connected
# Visualize the estimated differential network in an interactive manner.
tkid <- tkplot(Diff.net.connected, vertex.size= degree(Diff.net.connected)*1.2, layout =layout_with_fr,
               vertex.color="red", vertex.group.cex=0.8, edge.width =1.2, edge.color="orange")
# Visualize the estimated differential network in a non-interactive manner.
# grab the coordinates from tkplot
loc <- tkplot.getcoords(tkid)
```

```

plot(Diff.net.connected, layout=loc, vertex.size= degree(Diff.net.connected)*1.2, vertex.color="red",
     vertex.group.cex=0.8, edge.width =1.2, edge.color="orange")

rm(list=ls())
data("TCGA.BRCA")
result = DiffNet.FDR(TCGA.BRCA$X,TCGA.BRCA$group, alpha = 0.2, test.type = "pmat")
Diff.net.connected = result$Diff.net.connected
# Visualize the estimated differential network in an interactive manner.
tkid <- tkplot(Diff.net.connected, vertex.size= degree(Diff.net.connected)*1.2, layout =layout_with_fr,
               vertex.color="red", vertex.group.cex=0.8, edge.width =1.2, edge.color="orange")
# Visualize the estimated differential network in a non-interactive manner.
# grab the coordinates from tkplot
loc <- tkplot.getcoords(tkid)
plot(Diff.net.connected, layout=loc, vertex.size= degree(Diff.net.connected)*1.2, vertex.color="red",
     vertex.group.cex=0.8, edge.width =1.2, edge.color="orange")

rm(list=ls())
data("GSE13159.AML")

result = DiffNet.FDR(GSE13159.AML$X,GSE13159.AML$group, alpha = 0.2, test.type = "pcor")
Diff.net.connected = result$Diff.net.connected
# Visualize the estimated differential network in an interactive manner.
tkid <- tkplot(Diff.net.connected, vertex.size= degree(Diff.net.connected)*2, layout =layout_with_fr,
               vertex.color="red", vertex.group.cex=0.8, edge.width =1.2, edge.color="orange")
# Visualize the estimated differential network in a non-interactive manner.
# grab the coordinates from tkplot
loc <- tkplot.getcoords(tkid)
plot(Diff.net.connected, layout=loc, vertex.size= degree(Diff.net.connected)*2, vertex.color="red",
     vertex.group.cex=0.8, edge.width =1.2, edge.color="orange")

rm(list=ls())
data("GSE13159.AML")
result = DiffNet.FDR(GSE13159.AML$X,GSE13159.AML$group, alpha = 0.2, test.type = "pmat")
Diff.net.connected = result$Diff.net.connected
# Visualize the estimated differential network in an interactive manner.
tkid <- tkplot(Diff.net.connected, vertex.size= degree(Diff.net.connected)*2, layout =layout_with_fr,
               vertex.color="red", vertex.group.cex=0.8, edge.width =1.2, edge.color="orange")
# Visualize the estimated differential network in a non-interactive manner.
# grab the coordinates from tkplot
loc <- tkplot.getcoords(tkid)
plot(Diff.net.connected, layout=loc, vertex.size= degree(Diff.net.connected)*2, vertex.color="red",
     vertex.group.cex=0.8, edge.width =1.2, edge.color="orange")

```

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GSE13159.AML

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*Acute myeloid leukemia data (GSE13159)*


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## Description

Gene expression datasets for acute myeloid leukemia (GSE13159). The data is obtained from <http://discern-leelab.cs.washington.edu/>. It includes expression measurements for 541 AML cancers and 73 normal samples. The data only include expression measurements of genes that overlap

with the acute myeloid leukemia pathway (hsa05221) collected from the Kyoto Encyclopedia of Genes and Genomes databas. It includes an expression matrix for which the rows represent the samples and the columns represent the genes, and a vector denoting the group membership of each sample.

**Usage**

```
data("GSE13159.AML")
```

**Author(s)**

Xiao-Fei Zhang

**References**

Xiao-Fei Zhang, Le Ou-Yang, Shuo Yang, Xiaohua Hu and Hong Yan, DiffNetFDR: An R package for differential network analysis with false discovery rate control, 2018.

Maxim Grechkin, Benjamin A. Logsdon, Andrew J. Gentles and Su-In Lee, Identifying network perturbation in cancer, PLoS Computational Biology, 12(5): e1004888, 2016.

**See Also**

[DiffNet.FDR](#)

**Examples**

```
data(GSE13159.AML)
## maybe str(GSE13159.AML) ...
```

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TCGA.BRCA

*TCGA breast cancer data*

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**Description**

The TCGA breast cancer gene expression dataset used in our case study. The dataset (level 3, Agilent G450 microarray, version: May 6 2017) is obtained from the TCGA database using the TCGA2STAT R package. It includes gene expression measurements for 231 luminal A cancers and 95 basal-like cancers. The data only includes expression measurement of genes that overlap with the breast cancer pathway (hsa05224) collected from the Kyoto Encyclopedia of Genes and Genomes database. It includes an expression matrix for which the rows represent the samples and the columns represent the genes, and a vector denoting the group membership of each sample.

**Usage**

```
data("TCGA.BRCA")
```

**Author(s)**

Xiao-Fei Zhang

**References**

Xiao-Fei Zhang, Le Ou-Yang, Shuo Yang, Xiaohua Hu and Hong Yan, DiffNetFDR: Differential network analysis with false discovery rate control, 2018.

The Cancer Genome Atlas Research Network, Comprehensive molecular portraits of human breast tumors, Nature, 490(7418): 61-70, 2012. (<http://cancergenome.nih.gov/>)

**See Also**

[DiffNet.FDR](#)

**Examples**

```
data(TCGA.BRCA)
## maybe str(TCGA.BRCA) ...
```

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