# Package 'chNet'

May 10, 2021

Type Package	
<b>Title</b> Differential network analysis by simultaneously considering changes in gene interactions and gene expression.	
Version 2.0.0	
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<b>Description</b> chNet is a method designed for inferring diffferential network that satisfy hierarchichal constraints. chNet can simultaneously take into account the change of gene interactions and gene expression.	
<b>Depends</b> $R(>=3.5.1)$	
Imports MASS, Matrix, igraph, mytnorm, glmnet	
Suggests knitr, rmarkdown	
VignetteBuilder knitr	
RoxygenNote 6.1.1	
License $GPL(>=2)$	
Encoding UTF-8	
LazyData true	
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chNet	Differential network analysis by simultaneously considering changes in gene interactions and gene expression

## **Description**

The complete procedure for estimating differential network using chNet. For details, refer to method (Sections 2.4 and 2.5 in the main text).

# Usage

```
chNet (X,group,subsampling,R,lambar,parallel,nCpus)
```

## **Arguments**

X Data matrix for which the rows represent the samples and the columns represent

the genes.

group Vector which defines two groups of samples under comparison.

subsampling Logical value to indicate if the process should be run to obtain a weight matrix,

default to FALSE.

R The number of sub-sampled datasets; default is 20.

lambar The tuning (threshold) parameter controls the sparsity level.

parallel Logical value to indicate if the process should be run parallelly in multiple

threads, default to FALSE.

nCpus Number of (maximum) cores to use for parallel execution, default to 4.

## **Details**

This function is implemented to infer differential network that satisfy hierarchichal constraints. We first define the differential network as the difference of partial correlations from two different conditions, and develop a new test statistic to quantify the change of partial correlations. Then the Student's t-test statistic is used to quantify the changes in expression levels of individual genes. Finally, an optimization model is developed to combine the two different types of test statistics so that the estimated differential networks exhibit the hierarchical structures. A closed-formed solution is derived to solve the optimization model. In addition, based on sub-sampling experiments, a weighted hierarchial differential network can also be inferred.

#### Value

diff.edge.weight

the estimated weighted differential network.

diff.edge the adjacency matrix of the estimated differential network.

diff.gene the adjacency matrix of the estimated differentially expressed genes.

Diff. net the estimated differential network over all genes.

Diff.net.connected

the estimated differential network over only the connected genes.

## Author(s)

Jia-Juan Tu

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#### References

Jia-Juan Tu, Le Ou-Yang, Hong Yan, Hong Qin and Xiao-Fei Zhang (2021), Differential network analysis by simultaneously considering changes in gene interactions and gene expression.

#### See Also

```
generate.data, TCGA.BRCA, GSE13159.AML
```

#### **Examples**

```
# Simulation data
data.x= generate.data(p = 100, n = 100, umin = 0.5, umax = 1)
result = chNet(data.x$X,data.x$group, subsampling = FALSE, R= 20,
lambar = 3, parallel = FALSE, nCpus = 4)

# TCGA breast cancer data
data("TCGA.BRCA")
result = chNet(TCGA.BRCA$X,TCGA.BRCA$group,subsampling = FALSE, R= 20,
lambar = 2.825, parallel = FALSE, nCpus = 4)
# GSE13159 AML
data("GSE13159.AML")
result = chNet(TCGA.BRCA$X,TCGA.BRCA$group,subsampling = FALSE, R= 20,
lambar = 2.8, parallel = FALSE, nCpus = 4)
```

generate.data

Generate simulated data

# Description

The complete procedure for generating simulated data. For details, refer to simulation study (Supplementary Section 3.3.1).

# Usage

```
generate.data (p, n, umin, umax)
```

# **Arguments**

p The number of genes.
n The sample size.

umin The lower limits of the edge values.

The upper limits of the edge values.

# **Details**

The function is used to generate the gene expression datasets.

## Value

X A matrix of sample matrice  $(2n \times p)$  from two different conditions. A matrix of sample label matrice  $(2n \times 1)$  from two different conditions.

rho A list (length = 2) of the partical coefficients matrices  $(p \times p)$ .

hubgene A set hub genes.

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#### Author(s)

Jia-Juan Tu

#### References

Jia-Juan Tu, Le Ou-Yang, Hong Yan, Hong Qin and Xiao-Fei Zhang (2021), Differential network analysis by simultaneously considering changes in gene interactions and gene expression.

#### See Also

```
chNet, TCGA.BRCA, GSE13159.AML
```

# **Examples**

```
# Simulation data
data.x = generate.data(p = 100, n = 100, umin = 0.5, umax = 1)
```

GSE13159.AML

Acute myeloid leukemia data (GSE13159)

# **Description**

Gene expression datasets for acute myeloid leukemia (GSE13159) is download from http://discern-leelab.cs.washington.ed. There are 541 AML and 73 normal samples. The data include expression measurements of genes that overlap with the mTOR signaling pathway (hsa04150) from the Kyoto Encyclopedia of Genes and Genomes databas and 50 non-differentially expressed genes (Student's t-test, p-value > 0.05) with the highest variation AML samples. 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")
```

# **Format**

An object of class list of length 2.

## Author(s)

Jia-Juan Tu

# **Source**

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

# References

Jia-Juan Tu, Le Ou-Yang, Hong Yan, Hong Qin and Xiao-Fei Zhang (2021), Differential network analysis by simultaneously considering changes in gene interactions and gene expression.

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#### See Also

```
generate.data, chNet,TCGA.BRCA
```

#### **Examples**

data("GSE13159.AML")

TCGA.BRCA

TCGA breast cancer data

# **Description**

The TCGA breast cancer gene expression dataset used in our case study. The data (level 3, Agilent G450 microarray, version: May 6 2017) is diwnloaded 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 includes expression measurement of genes that overlap with the breast cancer pathway (hsa05224) collected from the Kyoto Encyclopedia of Genes and Genomes database and 50 non-differentially expressed genes (Student's t-test, p-value > 0.05) with the highest variation across luminal A and basal-like cancers. 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")
```

## **Format**

An object of class list of length 2.

#### Author(s)

Jia-Juan Tu

# **Source**

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

#### References

Jia-Juan Tu, Le Ou-Yang, Hong Yan, Hong Qin and Xiao-Fei Zhang (2021), Differential network analysis by simultaneously considering changes in gene interactions and gene expression.

# See Also

```
generate.data, TCGA.BRCA, GSE13159.AML
```

# **Examples**

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

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