

Package ‘coGPS’

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Type Package

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Description Gene Set Enrichment Analysis of P-value based statistics
for outlier gene detection in dataset merged from multiple
studies

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Contents

| | |
|-----------------------------------|---|
| coGPS-package | 2 |
| coGPS internal | 2 |
| PatientSpecificGeneList | 2 |
| PCOPA | 4 |
| permCOPA | 5 |
| PlotTopPCOPA | 7 |
| SampleData | 8 |

Index **10**

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|---------------|-----------------------------------------|
| coGPS-package | <i>Cancer Outlier Gene Profile Sets</i> |
|---------------|-----------------------------------------|

Description

Gene Set Enrichment Analysis of P-value based statistics for outlier gene detection in dataset merged from multiple studies

Author(s)

Yingying Wei, Michael Ochs Maintainer: Yingying Wei <ywei@jhsph.edu>

References

Wei, Y., Hennessey, P., Gaykalova, D., Califano, J.A., Ochs, M.F., (2011) Cancer Outlier Gene Profile Sets Elucidate Pathways in Head and Neck Squamous Cell Carcinoma.

| | |
|----------------|-----------------------------------------|
| coGPS internal | <i>coGPS package internal function.</i> |
|----------------|-----------------------------------------|

Description

These functions are not part of the package application programming interface and are not recommended to be used by the users.

Usage

plotCOPA

| | |
|-------------------------|-------------------------------------------|
| PatientSpecificGeneList | <i>Patient Specific outlier gene list</i> |
|-------------------------|-------------------------------------------|

Description

Generate an outlier gene list for each patient restricted to the top PCOPA scored genes

Usage

PatientSpecificGeneList(exprslist, alpha, side, type, TopGeneNum)

Arguments

| | |
|-------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <code>exprslist</code> | Each element of <i>exprslist</i> is a list with the first element being <i>exprs</i> and the second element being <i>classlab</i> . Each row of <i>exprs</i> represents one gene and each column represents one sample. <i>classlab</i> is a zero-one vector indicating the status of samples. We use 0 for the baseline group, usually the normal group, and 1 for the comparison group, usually the tumor group. |
| <code>alpha</code> | Significance level for P-value. |
| <code>side</code> | A vector specifying the definition of P-value in each of the study, which could be either <i>up</i> , <i>down</i> , or <i>twosided</i> . |
| <code>type</code> | A vector specifying whether the outlier pattern is <i>subtype</i> or <i>uniform</i> . |
| <code>TopGeneNum</code> | a number specifying the top number of outlier genes scored by PCOPA to be included in the generation of individual outlier gene list for each patient. |

Value

| | |
|------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <code>outliergene_bypatient</code> | a list whose length equals the number of tumor samples (patients). each element of the list is a list of length equaling to the length of <i>exprslist</i> , in other words the number of studies(or data type), showing the outlier gene for each patient in each study (or data type) |
|------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|

Author(s)

Yingying Wei

References

Wei, Y., Hennessey, P., Gaykalova, D., Califano, J.A., Ochs, M.F., (2011) Cancer Outlier Gene Profile Sets Elucidate Pathways in Head and Neck Squamous Cell Carcinoma.

Examples

```
#read in data
data(Exon_exprs_matched)
data(Methy_exprs_matched)
data(CNV_exprs_matched)
data(Exon_classlab_matched)
data(Methy_classlab_matched)
data(CNV_classlab_matched)
head(Exon_exprs_matched)

#exprslist[[i]]$exprs should be in matrix format
Exon_exprs<-as.matrix(Exon_exprs_matched)
Methy_exprs<-as.matrix(Methy_exprs_matched)
CNV_exprs<-as.matrix(CNV_exprs_matched)

#exprslist[[i]]$classlab should be in vector format
Exon_classlab<-unlist(Exon_classlab_matched)
Methy_classlab<-unlist(Methy_classlab_matched)
```

```

CNV_classlab<-unlist(CNV_classlab_matched)

#make an exprslist consisting 3 studies
trylist<-list()
trylist[[1]]<-list(exprs=Exon_exprs,classlab=Exon_classlab)
trylist[[2]]<-list(exprs=Methy_exprs,classlab=Methy_classlab)
trylist[[3]]<-list(exprs=CNV_exprs,classlab=CNV_classlab)

#generate an outlier gene list for each patient restricted to the top PCOPA scored genes
IndividualList7<-PatientSpecificGeneList(trylist,0.05,side=c("up","down","up"),type="subtype",TopGeneNum=100)

```

PCOPA

P-value based outlier gene detection

Description

Calculate P-value based statistics for outlier gene detection in dataset merged from multiple studies and give out outlier gene list for each patient.

Usage

```
PCOPA(exprslist, alpha, side, type)
```

Arguments

| | |
|------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <code>exprslist</code> | Each element of <i>exprslist</i> is a list with the first element being <i>exprs</i> and the second element being <i>classlab</i> . Each row of <i>exprs</i> represents one gene and each column represents one sample. <i>classlab</i> is a zero-one vector indicating the status of samples. We use 0 for the baseline group, usually the normal group, and 1 for the comparison group, usually the tumor group. |
| <code>alpha</code> | Significance level for P-value. |
| <code>side</code> | A vector specifying the definition of P-value in each of the study, which could be either <i>up</i> , <i>down</i> , or <i>twosided</i> . |
| <code>type</code> | A vector specifying whether the outlier pattern is <i>subtype</i> or <i>uniform</i> . |

Value

| | |
|------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <code>PCOPAstatistics</code> | the P-value based outlier gene detection statistics |
| <code>outliergene_bypatient</code> | a list whose length equals the number of tumor samples (patients). each element of the list is a list of length equaling to the length of <i>exprslist</i> , in other words the number of studies(or data type), showing the outlier gene for each patient in each study (or data type) |

Author(s)

Yingying Wei

References

Wei, Y., Hennessey, P., Gaykalova, D., Califano, J.A., Ochs, M.F., (2011) Cancer Outlier Gene Profile Sets Elucidate Pathways in Head and Neck Squamous Cell Carcinoma.

Examples

```
#read in data
data(Exon_exprs_matched)
data(Methy_exprs_matched)
data(CNV_exprs_matched)
data(Exon_classlab_matched)
data(Methy_classlab_matched)
data(CNV_classlab_matched)
head(Exon_exprs_matched)

#exprslist[[i]]$exprs should be in matrix format
Exon_exprs<-as.matrix(Exon_exprs_matched)
Methy_exprs<-as.matrix(Methy_exprs_matched)
CNV_exprs<-as.matrix(CNV_exprs_matched)

#exprslist[[i]]$classlab should be in vector format
Exon_classlab<-unlist(Exon_classlab_matched)
Methy_classlab<-unlist(Methy_classlab_matched)
CNV_classlab<-unlist(CNV_classlab_matched)

#make an exprslist consisting 3 studies
trylist<-list()
trylist[[1]]<-list(exprs=Exon_exprs,classlab=Exon_classlab)
trylist[[2]]<-list(exprs=Methy_exprs,classlab=Methy_classlab)
trylist[[3]]<-list(exprs=CNV_exprs,classlab=CNV_classlab)

#calculate P-value based statistics for outlier gene detection and output the outlier gene list for each patient
a7<-PCOPA(trylist,0.05,side=c("up","down","up"),type="subtype")
```

permCOPA

Calculate PCOPA value for permutations

Description

Run permutations by randomly shuffling the sample class labels and calculate a vector of PCOPA values for each permutation.

Usage

```
permCOPA(exprslist, alpha=0.05, side, type, perms=100)
```

Arguments

| | |
|------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <code>exprslist</code> | Each element of <i>exprslist</i> is a list with the first element being <i>exprs</i> and the second element being <i>classlab</i> . Each row of <i>exprs</i> represents one gene and each column represents one sample. <i>classlab</i> is a zero-one vector indicating the status of samples. We use 0 for the baseline group, usually the normal group, and 1 for the comparison group, usually the tumor group. |
| <code>alpha</code> | Significance level for P-value. |
| <code>side</code> | A vector specifying the definition of P-value in each of the study, which could be either <i>up</i> , <i>down</i> , or <i>twosided</i> . |
| <code>type</code> | A vector specifying whether the outlier pattern is <i>subtype</i> or <i>uniform</i> . |
| <code>perms</code> | Number of permutations to run. |

Value

| | |
|-------------------------|---------------------------------------------------------------------------------------------|
| <code>permResult</code> | A matrix where each row correspond to a gene and each column correspond to one permutation. |
|-------------------------|---------------------------------------------------------------------------------------------|

Author(s)

Michael Ochs

References

Wei, Y., Hennessey, P., Gaykalova, D., Califano, J.A., Ochs, M.F., (2011) Cancer Outlier Gene Profile Sets Elucidate Pathways in Head and Neck Squamous Cell Carcinoma.

Examples

```
#read in data
data(Exon_exprs_matched)
data(Methy_exprs_matched)
data(CNV_exprs_matched)
data(Exon_classlab_matched)
data(Methy_classlab_matched)
data(CNV_classlab_matched)
head(Exon_exprs_matched)

#exprslist[[i]]$exprs should be in matrix format
Exon_exprs<-as.matrix(Exon_exprs_matched)
Methy_exprs<-as.matrix(Methy_exprs_matched)
CNV_exprs<-as.matrix(CNV_exprs_matched)

#exprslist[[i]]$classlab should be in vector format
Exon_classlab<-unlist(Exon_classlab_matched)
Methy_classlab<-unlist(Methy_classlab_matched)
CNV_classlab<-unlist(CNV_classlab_matched)

#make an exprslist consisting 3 studies
trylist<-list()
```

```

trylist[[1]]<-list(exprs=Exon_exprs,classlab=Exon_classlab)
trylist[[2]]<-list(exprs=Methy_exprs,classlab=Methy_classlab)
trylist[[3]]<-list(exprs=CNV_exprs,classlab=CNV_classlab)

#run 2 permutations
perma7<-permCOPA(trylist,0.05,side=c("up","down","up"),type="subtype",perms=2)

```

PlotTopPCOPA

Plot expression patterns of top ranked genes.

Description

It first sorts the expression value $exprslist[[i]]$exprs[j,]$ among the baseline samples (e.g. normal ones) and comparison group (e.g. tumor ones) separately for selected gene j , and then plot the sorted expression values. The first argument *exprslist* should be the same one as for *PCOPA*; the second argument *PCOPAResult* should be an output of *PCOPA*; the third argument *topcut* determines how far we would go down the top ranked list; and the last argument *typelist* is a vector specifying the titles for each graph corresponds to a specific study.

Usage

```
PlotTopPCOPA(exprslist, PCOPAResult, topcut, typelist)
```

Arguments

| | |
|--------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <i>exprslist</i> | Each element of <i>exprslist</i> is a list with the first element being <i>exprs</i> and the second element being <i>classlab</i> . Each row of <i>exprs</i> represents one gene and each column represents one sample. <i>classlab</i> is a zero-one vector indicating the status of samples. We use 0 for the baseline group, usually the normal group, and 1 for the comparison group, usually the tumor group. |
| <i>PCOPAResult</i> | Output of <i>PCOPA</i> . |
| <i>topcut</i> | Cutoff of top ranked gene list. |
| <i>typelist</i> | A vector specifying the titles for each graph corresponds to a specific study. |

Author(s)

Michael Ochs, Yingying Wei

Examples

```

#read in data
data(Exon_exprs_matched)
data(Methy_exprs_matched)
data(CNV_exprs_matched)
data(Exon_classlab_matched)
data(Methy_classlab_matched)
data(CNV_classlab_matched)
head(Exon_exprs_matched)

```

```

#exprslist[[i]]$exprs should be in matrix format
Exon_exprs<-as.matrix(Exon_exprs_matched)
Methy_exprs<-as.matrix(Methy_exprs_matched)
CNV_exprs<-as.matrix(CNV_exprs_matched)

#exprslist[[i]]$classlab should be in vector format
Exon_classlab<-unlist(Exon_classlab_matched)
Methy_classlab<-unlist(Methy_classlab_matched)
CNV_classlab<-unlist(CNV_classlab_matched)

#make an exprslist consisting 3 studies
trylist<-list()
trylist[[1]]<-list(exprs=Exon_exprs,classlab=Exon_classlab)
trylist[[2]]<-list(exprs=Methy_exprs,classlab=Methy_classlab)
trylist[[3]]<-list(exprs=CNV_exprs,classlab=CNV_classlab)

#calculate P-value based statistics for outlier gene detection and output the outlier gene list for each patient
a7<-PCOPA(trylist,0.05,side=c("up","down","up"),type="subtype")

#plot expression patterns of top ranked genes.
PlotTopPCOPA(trylist,a7,topcut=1,typelist=c("Exon","Methy","CNV"))

```

SampleData

Sample Data for coGPS

Description

Here we present an example of coGPS analysis.

Arguments

Exon_exprs_matched
Expression data for 44 tumors and 25 normals. Each row indicates a gene with row name showing gene name and each column indicates a sample with column name showing sample name.

Exon_class_matched
A length 69 vector showing status of corresponding exon samples, 0 for normals and 1 for tumors.

Methy_exprs_matched
Methylation data for 44 tumors and 25 normals.

Methy_class_matched
A length 69 vector showing status of corresponding methylation samples, 0 for normals and 1 for tumors.

CNV_exprs_matched
Copy number data for 44 tumors and 25 normals.

CNV_class_matched
A length 69 vector showing status of corresponding copy number samples, 0 for normals and 1 for tumors.

Hs.gmt1.c1
Broad Institute C1 Positional Gene Sets.

Details

In this application, the columns of each data type are matched. In other words, the first columns of `Exon_exprs_matched`, `Methy_exprs_matched` and `CNV_exprs_matched` correspond to the same patient. And hence the `Exon_class_matched`, `Methy_class_matched` and `CNV_class_matched` are identical. However, suppose in applications that we are not concerned with the outlier gene list for each patient, we can leave with the samples (columns) unmatched.

Index

* **Microarray, Bioinformatics, DifferentialExpression**

coGPS-package, 2

CNV_classlab_matched (SampleData), 8

CNV_exprs_matched (SampleData), 8

coGPS (coGPS-package), 2

coGPS internal, 2

coGPS-package, 2

Exon_classlab_matched (SampleData), 8

Exon_exprs_matched (SampleData), 8

Hs.gmt1.c1 (SampleData), 8

Methy_classlab_matched (SampleData), 8

Methy_exprs_matched (SampleData), 8

PatientSpecificGeneList, 2

PCOPA, 4

permCOPA, 5

plotCOPA (coGPS internal), 2

PlotTopPCOPA, 7

SampleData, 8