

# Package ‘coGPS’

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**Type** Package

**Title** cancer outlier Gene Profile Sets

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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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**Suggests** limma

**Imports** graphics, grDevices

**License** GPL-2

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**biocViews** Microarray, DifferentialExpression

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

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coGPS internal	<i>coGPS package internal function.</i>
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**Description**

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

**Usage**

plotCOPA

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PatientSpecificGeneList	<i>Patient Specific outlier gene list</i>
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**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)
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**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)

```

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PCOPA

*P-value based outlier gene detection*


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### 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.
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**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

<code>exprslist</code>	Each element of $exprslist$ is a list with the first element being $exprs$ and the second element being $classlab$ . Each row of $exprs$ represents one gene and each column represents one sample. $classlab$ 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>PCOPAResult</code>	Output of <i>PCOPA</i> .
<code>topcut</code>	Cutoff of top ranked gene list.
<code>typelist</code>	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.

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## \* **Microarray, Bioinformatics, DifferentialExpression**

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