# Package 'imsig'

July 10, 2018

| Type Package   |
|--|
| Title Immune Cell Gene Signatures for Profiling the Microenvironment<br>of Solid Tumours   |
| Version 1.0.0  |
| Author Ajit Johnson Nirmal   |
| Maintainer Ajit Johnson Nirmal <a jit="" johnson.n@gmail.com=""></a>   |
| <b>Description</b> Estimate the relative abundance of tissue-<br>infiltrating immune subpopulations abundances using gene expression data. |
| License GPL-3  |
| <pre>URL https://github.com/ajitjohnson/imsig/</pre>   |
| BugReports https://github.com/ajitjohnson/imsig/issues   |
| Encoding UTF-8   |
| LazyData true  |
| <b>Imports</b> HiClimR (>= 1.2), RColorBrewer (>= 1.1), igraph (>= 1.2), ggplot2 (>= 2.2), gridExtra (>= 2.3), survival (>= 2.4)           |
| RoxygenNote 6.0.1  |
| Suggests testthat  |
| NeedsCompilation no  |
| Repository CRAN  |
| Date/Publication 2018-07-10 20:50:03 UTC   |

# R topics documented:

| orr_matrix    | 2 |
|---------------|---|
| xample_cli    | 3 |
| xample_data   | 3 |
| eature_select | 4 |
| ene_stat      | 4 |
| nsig          | 5 |
| nsig_survival | 6 |

12

| plot_abundance | 7  |
|----------------|----|
| plot_network   | 8  |
| plot_survival  | 9  |
| pp_exp         | 10 |
| pp_sig         | 10 |
| sig            | 11 |
|                |    |

## Index

corr\_matrix Correlation matrix

# Description

Creates a correlation matrix of ImSig signature genes.

# Usage

corr\_matrix(exp, r)

## Arguments

| exp | Dataframe of transcriptomic data (natural scale) containing genes as rows and samples as columns. Note: Gene names should be set as row names and duplicates are not allowed. Missing values are not allowed within the expression matrix. Check example-head(example_data): example_data.   |
|-----|--|
| r   | Use a value between 0 and 1. Default is 0.6. This is a user defined correlation cut-off to perform feature selection (feature_select). Feature selection aids to enrich the prediction of relative abundance of immune cells by filtering off poorly correlated ImSig genes. To get an idea of what cut-off to use check the results of (gene_stat) and choose a cut-off that displays high median correlation and maintains a high proportion of genes after feature selection. |

## Value

Gene-gene correlation matrix of ImSig genes.

example\_cli

## Description

An example clinical data file. Minimum required informations are the sample name (same as that of the expression matrix), event (dead or alive) and time to event (days, months or years).

#### Usage

example\_cli

## Format

dataframe

example\_data

Example transcriptomics data

## Description

Example expression data matrix. The data is preffered to be in natural scale with genes as rows and samples as columns.Note: Gene names should be set as row names and duplicates are not allowed. Missing values are not allowed within the expression matrix. Check example-head(example\_data)

#### Usage

example\_data

## Format

dataframe

feature\_select

## Description

ImSig genes were designed to be co-expressed in tissue transcriptomic data. However, depending on the dataset some of the genes may not co-express with the dominant module. In order to remove such deviant genes, a feature selection can be carried out based on correlation. This function removes genes that exhibit a poor correlation (less than the defined r value) with the dominant ImSig module. This step of feature selection is recommended to enrich the prediction of relative abundance of immune cells.

#### Usage

feature\_select(exp, r = 0.6)

#### Arguments

| exp | Dataframe of transcriptomic data (natural scale) containing genes as rows and samples as columns. Note: Gene names should be set as row names and duplicates are not allowed. Missing values are not allowed within the expression matrix. Check example-head(example_data): example_data.   |
|-----|--|
| r   | Use a value between 0 and 1. Default is 0.6. This is a user defined correla-<br>tion cut-off to perform feature selection. To get an idea of what cut-off to use<br>check the results of (gene_stat) and choose a cut-off that displays high median<br>correlation and maintains a high proportion of genes after feature selection. |

## Value

Returns a list of 'feature selected' genes based on the set r value.

#### Examples

```
feature_select (exp = example_data, r = 0.7)
```

gene\_stat

General stastitics of ImSig analysis

#### Description

[Total genes in ImSig]: The total number of genes in ImSig list. [No. of ImSig genes in user dataset]: The number of ImSig genes found in user's dataset. Like all signatures, ImSig works best when this overlap is high, preferably over 75

## imsig

## Usage

gene\_stat(exp, r = 0.6)

#### Arguments

| exp | Dataframe of transcriptomic data (natural scale) containing genes as rows and samples as columns. Note: Gene names should be set as row names and duplicates are not allowed. Missing values are not allowed within the expression matrix. Check example-head(example_data): example_data.   |
|-----|--|
| r   | Use a value between 0 and 1. Default is 0.6. This is a user defined correlation cut-off to perform feature selection (feature_select). Feature selection aids to enrich the prediction of relative abundance of immune cells by filtering off poorly correlated ImSig genes. To get an idea of what cut-off to use check the results of (gene_stat) and choose a cut-off that displays high median correlation and maintains a high proportion of genes after feature selection. |

## Value

Dataframe of general statistics of ImSig analysis.

## See Also

feature\_select

#### Examples

gene\_stat (exp = example\_data, r = 0.7)

| imsig | Estimate the relative abundance of tissue-infiltrating immune subpop- |
|-------|---|
|       | ulations abundances using gene expression data                        |

## Description

Estimates the relative abundance of immune cells across patients/samples.

#### Usage

imsig(exp, r = 0.6)

#### Arguments

exp

Dataframe of transcriptomic data (natural scale) containing genes as rows and samples as columns. Note: Gene names should be set as row names and duplicates are not allowed. Missing values are not allowed within the expression matrix. Check example-head(example\_data): example\_data.

r Use a value between 0 and 1. Default is 0.6. This is a user defined correlation cut-off to perform feature selection (feature\_select). Feature selection aids to enrich the prediction of relative abundance of immune cells by filtering off poorly correlated ImSig genes. To get an idea of what cut-off to use check the results of (gene\_stat) and choose a cut-off that displays high median correlation and maintains a high proportion of genes after feature selection.

## Value

Relative abundance of immune cells across samples. Returns a dataframe.

## See Also

feature\_select, example\_data

#### Examples

```
cell_abundance = imsig (exp = example_data, r = 0.7)
head(cell_abundance)
```

| imsig_survival | Survival analysis based on relative abundance of immune infiltration |
|----------------|--|
|                | estimated by ImSig   |

#### Description

Patients are split into two groups based on their immune cell abundance (median aundance value) and a regular survival analysi is carried out.

#### Usage

```
imsig_survival(exp, cli, time = "time", status = "status", r = 0.6)
```

#### Arguments

| exp    | Dataframe of transcriptomic data (natural scale) containing genes as rows and samples as columns. Note: Gene names should be set as row names and duplicates are not allowed. Missing values are not allowed within the expression matrix. Check example-head(example_data): example_data. |
|--------|--|
| cli    | Clinical metadata containting the event data (dead or alive) and time to event data. Samples names should be in rownames and same as that in the expression file. Check head() of example_cli for an example clinical data.  |
| time   | Column name of time-to-event parameter.  |
| status | Column name of event (dead or alive) parameter.  |

#### plot\_abundance

r

Use a value between 0 and 1. Default is 0.6. This is a user defined correlation cut-off to perform feature selection (feature\_select). Feature selection aids to enrich the prediction of relative abundance of immune cells by filtering off poorly correlated ImSig genes. To get an idea of what cut-off to use check the results of (gene\_stat) and choose a cut-off that displays high median correlation and maintains a high proportion of genes after feature selection.

## Value

Hazard Ratio

#### See Also

feature\_select, example\_data, example\_cli

### Examples

survival = imsig\_survival (exp = example\_data)
head(survival)

plot\_abundance Plot relative abundance of immune cells

#### Description

Barplots of relative abundance of immune cells across samples. The order of the samples are the same as that of imsig.

#### Usage

plot\_abundance(exp, r = 0.6)

#### Arguments

| exp | Dataframe of transcriptomic data (natural scale) containing genes as rows and samples as columns. Note: Gene names should be set as row names and duplicates are not allowed. Missing values are not allowed within the expression matrix. Check example-head(example_data): example_data. |
|-----|--|
| r   | Use a value between 0 and 1. Default is 0.6. This is a user defined correlation cut-off to perform feature selection (feature_select). Feature selection aids to enrich the prediction of relative abundance of immune cells by filtering off  |

to enrich the prediction of relative abundance of immune cells by filtering off poorly correlated ImSig genes. To get an idea of what cut-off to use check the results of (gene\_stat) and choose a cut-off that displays high median correlation and maintains a high proportion of genes after feature selection.

#### Value

ggplot

#### See Also

feature\_select, example\_data

## Examples

```
plot_abundance (exp = example_data, r = 0.7)
```

plot\_network

## Network graph of ImSig genes

#### Description

A Network visualization displays undirected graph structures and highlights the relationships between entities. The nodes are ImSig genes and the edges represent the correlation between them. The nodes are coloured based on cell type. Try using a correlation cut-off of '0' to get a complete picture.

#### Usage

```
plot_network(exp, r = 0.6, pt.cex = 2, cex = 1, inset = 0,
    x.intersp = 2, vertex.size = 3, vertex.label = NA,
    layout = layout_with_fr)
```

#### Arguments

| exp       | Dataframe of transcriptomic data (natural scale) containing genes as rows and samples as columns. Note: Gene names should be set as row names and duplicates are not allowed. Missing values are not allowed within the expression matrix. Check example-head(example_data): example_data.   |
|-----------|--|
| r         | Use a value between 0 and 1. Default is 0.6. This is a user defined correlation cut-off to perform feature selection (feature_select). Feature selection aids to enrich the prediction of relative abundance of immune cells by filtering off poorly correlated ImSig genes. To get an idea of what cut-off to use check the results of (gene_stat) and choose a cut-off that displays high median correlation and maintains a high proportion of genes after feature selection. |
| pt.cex    | expansion factor(s) for the points.  |
| cex       | character expansion factor relative to current par("cex"). Used for text, and provides the default for pt.cex.   |
| inset     | inset distance(s) from the margins as a fraction of the plot region when legend is placed by keyword.  |
| x.intersp | character interspacing factor for horizontal (x) spacing.  |

## plot\_survival

| vertex.size  | Node size of network graph   |
|--------------|--|
| vertex.label | Add gene names to the network graph. Default set to NA.  |
| layout       | Layout algorithm to be used for building network. Default set to force-directed layout algorithm by Fruchterman and Reingold. Read documentation of 'igraph' for other available algorithms. |

## Value

Network graph

## See Also

feature\_select

## Examples

plot\_network (exp = example\_data, r = 0.7)

plot\_survival

## Forest plot of survial analysis by ImSig

## Description

Patients are split into two groups based on their immune cell abundance (median aundance value) and a regular survival analysi is carried out. Raw values can be obtained from imsig\_survival.

## Usage

```
plot_survival(exp, cli, time = "time", status = "status", r = 0.6)
```

## Arguments

| exp    | Dataframe of transcriptomic data (natural scale) containing genes as rows and samples as columns. Note: Gene names should be set as row names and duplicates are not allowed. Missing values are not allowed within the expression matrix. Check example- head(example_data): example_data.  |
|--------|--|
| cli    | Clinical metadata containting the event data (dead or alive) and time to event data. Samples names should be in rownames and same as that in the expression file. Check head() of example_cli for an example clinical data.  |
| time   | Column name of time-to-event parameter.  |
| status | Column name of event (dead or alive) parameter.  |
| r      | Use a value between 0 and 1. Default is 0.6. This is a user defined correlation cut-off to perform feature selection (feature_select). Feature selection aids to enrich the prediction of relative abundance of immune cells by filtering off poorly correlated ImSig genes. To get an idea of what cut-off to use check the results of (gene_stat) and choose a cut-off that displays high median correlation and maintains a high proportion of genes after feature selection. |

## Value

Forest plot

## See Also

feature\_select, example\_data, example\_cli

### Examples

```
plot_survival (exp = example_data, r = 0.7, cli = example_cli, time = 'time', status= 'status')
```

pp\_exp

Pre-processing expression matrix

#### Description

Subsets the user's dataset based on the genes that are common to the users dataset and ImSig.

## Usage

pp\_exp(exp)

## Arguments

exp Dataframe of transcriptomic data (natural scale) containing genes as rows and samples as columns. Note: Gene names should be set as row names and duplicates are not allowed. Missing values are not allowed within the expression matrix. Check example- head(example\_data): example\_data.

## Value

Expression dataframe

| pp_sig | Pre-processing ImSig file |  |
|--------|---------------------------|--|
|--------|---------------------------|--|

## Description

Subsets ImSig genes based on the genes that are common to the users dataset and ImSig

## Usage

pp\_sig(exp)

## Arguments

| exp | Dataframe of transcriptomic data (natural scale) containing genes as rows and  |
|-----|--|
|     | samples as columns. Note: Gene names should be set as row names and du-        |
|     | plicates are not allowed. Missing values are not allowed within the expression |
|     | matrix. Check example- head(example_data): example_data.                       |

# Value

ImSig dataframe

sig ImSig genes

# Description

ImSig signature genes and the cell type they represent

# Usage

sig

## Format

dataframe

# Index

\*Topic datasets example\_cli, 3  $example_data, 3$ sig, 11 corr\_matrix, 2 example\_cli, 3, 6, 7, 9, 10 example\_data, 2, 3, 4–11 feature\_select, 2, 4, 5-10 gene\_stat, 2, 4, 4, 5–9 imsig, 5, 7 imsig\_survival, 6, 9 plot\_abundance, 7 plot\_network, 8  $\texttt{plot\_survival}, 9$ pp\_exp, 10  $\texttt{pp\_sig}, 10$ 

sig, 11