

# Package ‘MUTSCOT’

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**Title** MutScot

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**Description** The MutScot package.

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**Imports** stats, utils, MASS, dndscv

**License** GPL-2

**NeedsCompilation** yes

**Depends** R (>= 2.10)

## R topics documented:

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MutScot-package	<i>MutScot</i>
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## Description

The MutScot package for identifying driver genes

## Details

Identifying cancer driver genes is essential for understanding mechanisms of carcinogenesis and designing therapeutic strategies. Consequently, a set of driver genes has been identified for each cancer types, assumed to be identical across subtypes. This assumption may not hold, and the sets of driver genes are possibly distinct across cancer subtypes. The MutScot package includes a statistical framework that identifies driver genes and utilizes patient information to investigate subtype heterogeneity of driver genes. The function `MutScot` identifies driver genes, and the function `MutScotPatientHeter` is used for conducting a subtype heterogeneity test.

**Author(s)**

Ho-Hsiang Wu and Bin Zhu <bin.zhu@nih.gov>

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data	<i>Data for the MutScot example</i>
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**Description**

Data for the MutScot example.

**Details**

The object `data` is a data frame with columns for the sample id, chromosome, position (hg19), reference allele, and alternate allele.

**Examples**

```
data(data, package="MUTSCOT")
data[1:5, ]
```

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getBenchmarkGenes	<i>Benchmark Genes</i>
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**Description**

Get a vector of gene names given by Bailey et al (2018)

**Usage**

```
getBenchmarkGenes()
```

**Details**

This is the list of driver gene names given by Bailey et al (2018).

**Value**

A character vector of gene names.

**Examples**

```
getBenchmarkGenes()
```

MutScot

*MutScot***Description**

MutScot

**Usage**

```
MutScot(data, MutSigCV.out=NULL, options=NULL)
```

**Arguments**

<code>data</code>	Data frame containing five columns for sample id, chromosome, position (hg19), reference allele, and alternate allele. The columns must be in the order given above. No default
<code>MutSigCV.out</code>	Information provided by MutSigCV algorithm (see details). The default is NULL.
<code>options</code>	List of options (see details). The default is NULL.

**Details**

See `data(MutSigCV)` for an example format of `MutSigCV.out`. Currently, the only columns of this object that are used are: `gene`, `X`, `x`, and `q`.

**Options**

Option	Description	Default
<code>maxMutPerPat</code>	maximum number of mutations per patient	300
<code>multitest</code>	method used for multiple tests (see <code>p.adjust</code> )	"fdr"
<code>alpha.q</code>	significance level	0.1
<code>benchmarkGenes</code>	vector of genes for summary	genes returned from <code>getBenchmarkGenes</code>

**Value**

A list of class "MutScot" with the following names and descriptions:

- `driverGenes` Vector of identified driver genes.
- `significantGenes` Data frame of significant genes among the methods determined by `options$alpha.q`.
- `MutScotGenes` Data frame of p-values and adjusted p-values for the MutScot method.
- `countMat` Matrix of TP, FP, FI counts by method.
- `dndsout` Output from the dndscv algorithm.
- `patientIds` Vector of patient ids.
- `genes` Vector of gene names.
- `mutationRates` List containing relative SNV mutation rates of patients (`PatRat`), weight parameter of patients w.r.t SNV mutations (`wt`), relative Indel mutation rates of patients (`PatRatI`), weight parameter of patients w.r.t Indel mutations (`wtI`), annotated Indel mutations (`Indels_record`), annotated observed/expected Indel mutations (`geneindels`), IVW estimated gene-specific background mutation rates (`Lambda_g`).

- obsExp List containing gene-specific observed and expected missense mutation counts (miss), gene-specific observed and expected nonsense mutation counts (nsen), gene-specific observed and expected splicing site mutation counts (splc), gene-specific observed and expected Indel mutation counts (indel).

Author(s)

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See Also

MutScotPatientHeter

Examples

```
data(data, package="MUTSCOT")
data(MutSigCV.out, package="MUTSCOT")
# Not run due to the time it takes
#MutScot(data, MutSigCV.out=MutSigCV.out)
```

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MutScotPatientHeter
<i>Test for heterogeneity</i>

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Description

Test genes for subtype heterogeneity

Usage

```
MutScotPatientHeter(obj, patientInfo, testGenes, options=NULL)
```

Arguments

obj	Object returned from MutScot. No default.
patientInfo	Data frame containing sample ids and numeric columns of adjusted continuous covariates. The first column must be the ids and there must be at least one covariate (see details). No default.
testGenes	Character vector of gene names to test. No default.
options	List of options (see details). The default is NULL.

Details

The first column of patient ids in the patientInfo data frame are matched with obj\$patientIds.

Options

Option	Description	Default
MCT	number of Monte-Carlo simulations	1e4

**Value**

A list with the following names and descriptions:

- `pvalues` Data frame of gene-specific p-values for missense, nonsense, splicing, indel, and the test of subtype heterogeneity. A row of NAs will appear for any gene not found in `obj$genes`.
- `glm_miss` List of the glm fitting of missense mutation for each gene.
- `glm_nsen` List of the glm fitting of nonsense mutation for each gene.
- `glm_splic` List of the glm fitting of splicing site mutation for each gene.
- `glm_indel` List of the glm fitting of indel mutation for each gene.

**Author(s)**

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**See Also**

`MutScot`

**Examples**

```
data(data, package="MUTSCOT")
data(MutSigCV.out, package="MUTSCOT")
data(PatInfo, package="MUTSCOT")

# Not run due to the time it takes
#retM <- MutScot(data, MutSigCV.out=MutSigCV.out)
#x <- retM$significantGenes
#tmp <- (x[, "Benchmark"]==TRUE) & (x[, "MutScot"]==TRUE)
#TestSet <- as.vector(x[tmp, "Gene"])
#retH <- MutScotPatientHeter(retM, PatInfo, TestSet)
```

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MutSigCV.out

*Example of the MutSigCV.out data object*

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**Description**

Example of the `MutSigCV.out` data object

**Details**

The object is a data frame named `MutSigCV.out`. Currently, the only columns of this data that are used are: `gene`, `X`, `x`, and `q`.

**Examples**

```
# Look at the first five rows
data(MutSigCV.out, package="MUTSCOT")
MutSigCV.out[1:5, ]
```

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`PatInfo`*Data for the `MutScotPatientHeter` example*

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**Description**

Data for the `MutScotPatientHeter` example.

**Details**

The object `data` is a data frame with columns for the sample id and ER status.

**Examples**

```
data(PatInfo, package="MUTSCOT")
PatInfo[1:5, ]
```