# Package 'mgmtstp27'

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<b>Title</b> DNA methylation of MGMT promoter based on Infinium Platform (27k, 450K and EPIC) from Illumina
Version 0.8
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<b>Depends</b> R (>= 4.2.1), minfi, lumi, ade4,methylumi,MASS
Suggests boot
<b>Description</b> This package contains R functions for the prediction of the DNA methylation status of MGMT promoter with data from Infinium platforms (HM-27K, HM450K, EPIC and EPICv2).
License GPL (>= 2)
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mgmtstp27-package Additional tools for the analysis of methylation data

# Description

Type Package

Additional tools for the analysis of methylation data

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#### **Details**

Package: mgmtstp27 Type: Package Version: 0.1

Date: 2014-09-11 License: GPL (>= 2)

# Author(s)

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#### References

Bady, P., D. Sciuscio, A.-C. Diserens, J. Bloch, M. J. van den Bent, C. Marosi, P.-Y. Dietrich, M. Weller, L. Mariani, F. L. Heppner, D. R. McDonald, D. Lacombe, R. Stupp, M. Delorenzi, and M. E. Hegi. 2012. MGMT methylation analysis of glioblastoma on the Infinium methylation BeadChip identifies two distinct CpG regions associated with gene silencing and outcome, yielding a prediction model for comparisons across datasets, tumor grades, and CIMP-status. Acta Neuropathologica 124:547-560.

gbm

DNA methylation of MGMT promoter region from Infinium HM-450k and HM-27K platforms

# Description

DNA methylation of MGMT promoter region from Infinium HM-450k and HM-27K platforms (table S3 and S4 described in details Bady et al. 2012).

# Usage

data("gbm")

# **Format**

two data.frames corresponding to the tables S3 and S4 described in details Bady et al. (2012)

## **Details**

The results published here are in part based upon data generated by The Cancer TCGA Genome Atlas pilot project established by the NCI and NHGRI. Information about TCGA and the investigators and institutions who constitute the TCGA research network can be found at (http://cancergenome.nih.gov). The dbGaP accession number to the specific version of the TCGA data set is phs000178.v8.p7.

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#### Source

This two datasets correspond to the tables S3 and S4 from Bady et al. (2012).

#### References

Bady, P., D. Sciuscio, A.-C. Diserens, J. Bloch, M. J. van den Bent, C. Marosi, P.-Y. Dietrich, M. Weller, L. Mariani, F. L. Heppner, D. R. McDonald, D. Lacombe, R. Stupp, M. Delorenzi, and M. E. Hegi. 2012. MGMT methylation analysis of glioblastoma on the Infinium methylation BeadChip identifies two distinct CpG regions associated with gene silencing and outcome, yielding a prediction model for comparisons across datasets, tumor grades, and CIMP-status. Acta Neuropathologica 124:547-560.

The Cancer Genome Atlas Consortium Comprehensive genomic characterization defines human glioblastoma genes and core pathways. Nature. 2008;455:1061-1068.

Noushmehr, H., D. Weisenberger, K. Diefes, H. Phillips, K. Pujara, B. Berman, F. Pan, C. Pelloski, E. Sulman, and K. Bhat. 2010. Identification of a CpG island methylator phenotype that defines a distinct subgroup of glioma. Cancer Cell 17:510 - 522.

# **Examples**

```
# table S3 (bady et al 2012)
data(NCHgbm450)
head(NCHgbm450)
# table S4 (bady et al 2012)
data(TCGAgbm27)
head(TCGAgbm27)
```

mgmt

set of tools related to prediction of the DNA methylation of MGMT promoter.

# **Description**

set of tools related to prediction of the DNA methylation of MGMT promoter.

# Usage

```
MGMTpredict(x,level = 0.05,dispersion=FALSE,transpose=FALSE,ic.distrib="normal",cutoff=1,...)
MGMTsim(n=1000,proba=NULL,newdata=NULL,...)
MGMTqc.pop(object,sim=FALSE,n=1000,which.plot=1:6,mfrow=c(3,3),...)
MGMTqc.single(object,nsample=NULL,sim=FALSE,n=1000,which.plot=1:4,mfrow=c(2,3),...)
```

# **Arguments**

x a data.frame containing contining the M-values for the probes 'cg12981137' and 'cg12434587' (in row and columns).

object an object of the class 'mgmt'.

level a numeric value corresponding to level used to compute confidence intervals

(level=0.05, by default).

dispersion a logical value. If dispersion=TRUE, dispersion correction was used to compute

Confidence intervals.

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transpose a logical value (transpose=TRUE, when the probes are organized by rows).

ic.distrib a character value indicating the probability distribution used to compute confi-

dence intervals ("student" or "normal", by default ic.distrib="normal").

cutt-off a numeric value identifying the cut-off used to calculate the DNA methylation

state of MGMT promoter (1= better classification, 2= balance among sensitivity

and specificty). By default, cutoff is equal to 1.

n number of simulated samples

newdata an object of the class 'mgmt' containing containing the M-values for the probes

'cg12981137' and 'cg12434587' (in columns).

sim a logical value. If sim=TRUE, the reference is obtained by simulation (Multi-

variate distribution. If sim=FALSE (by default), the reference is the training

data.

nsample a numerical value identifying the sample of interest (only used in the function

MGMTqc.single, bydefault nsample=1).

which.plot a vector telling which plots to produce.

mfrow parameter for the array of figures to be drawn.

... further arguments passed to or from other methods

#### **Details**

For EPIC v2 infinium platform, the function uses the two probes "cg12434587\_BC11" and "cg12981137\_TC11" (which will be renamed as "cg12434587" and "cg12981137"). The selection of these two probes is based on the concordance of the annotation information and the sequences of the probes (see variable "AlleleA\_ProbeSeq" from manifest file). add information about simulation and QC graphical ouput.

# Value

The function 'MGMTsim' return a data.frame containing two simulated M-values corresponding to the probes 'cg12981137' and 'cg12434587'. The function 'MGMTpredict' return data.frame contianing the following information:

sample sample(row) names

cg12434587 M-value for the probe 'cg12434587' cg12981137 M-value for the probe 'cg12981137'

pred probability that MGMT promoter is methylated

lower limit of the confidence intervals for the probability upper upper limit of the confidence intervals for the probability

state DNA methylation state of MGMT promoter (U,M)using the cut-off provided in

Bady et al. (2012)

extended DNA methylation state of MGMT promoter (U,u,m,M) using the cut-off pro-

vided and confidence interval in Bady et al. (2012)

# Author(s)

P.BADY

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#### References

Bady, P., D. Sciuscio, A.-C. Diserens, J. Bloch, M. J. van den Bent, C. Marosi, P.-Y. Dietrich, M. Weller, L. Mariani, F. L. Heppner, D. R. McDonald, D. Lacombe, R. Stupp, M. Delorenzi, and M. E. Hegi. 2012. MGMT methylation analysis of glioblastoma on the Infinium methylation BeadChip identifies two distinct CpG regions associated with gene silencing and outcome, yielding a prediction model for comparisons across datasets, tumor grades, and CIMP-status. Acta Neuropathologica 124:547-560.

# **Examples**

```
data(MGMTSTP27)
training1 <- MGMTSTP27$data
pred1 <- MGMTpredict(training1)
sim1 <- MGMTsim(n=100,newdata=pred1)
qqplot(pred1[,"cg12434587"],sim1[,"cg12434587"])
MGMTqc.pop(pred1)</pre>
```

MGMTSTP27

Model to predict MGMTSTP27

### Usage

```
data("MGMTSTP27")
```

#### **Format**

an object 'glm' with additional numerical vector called 'perf' containing performance information.

perf a numeric vector containing information

# **Details**

The function MGMTpredict provides prediction of DNA methylation status of MGMT promoter as described in Bady et al (2012). The model is an object glm with an additional numerical vector called 'perf' containing performance information.

# Source

The model is presented in details in Bady et al. (2012).

#### References

Bady, P., D. Sciuscio, A.-C. Diserens, J. Bloch, M. J. van den Bent, C. Marosi, P.-Y. Dietrich, M. Weller, L. Mariani, F. L. Heppner, D. R. McDonald, D. Lacombe, R. Stupp, M. Delorenzi, and M. E. Hegi. 2012. MGMT methylation analysis of glioblastoma on the Infinium methylation BeadChip identifies two distinct CpG regions associated with gene silencing and outcome, yielding a prediction model for comparisons across datasets, tumor grades, and CIMP-status. Acta Neuropathologica 124:547-560.

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# Examples

data(MGMTSTP27)
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# maximization of good classification
MGMTSTP27\$perf1
# balance amon specificity and sensitivity
MGMTSTP27\$perf2

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