

Package ‘sesame’

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Type Package

Title Tools For Analyzing Illumina Infinium DNA Methylation Arrays

Description

Tools For analyzing Illumina Infinium DNA methylation arrays. SeSAmE provides utilities to support analyses of multiple generations of Infinium DNA methylation BeadChips, including pre-processing, quality control, visualization and inference. SeSAmE features more accurate detection calling, intelligent inference of ethnicity, sex and advanced quality control routines.

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Depends R (>= 4.0), sesameData, methods

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URL <https://github.com/zwdzwd/sesame>

BugReports <https://github.com/zwdzwd/sesame/issues>

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'channel_inference.R' 'cnv.R' 'controls.R' 'deidentify.R'
'detection.R' 'differential_methylation.R' 'dye_bias.R'
'fileSet.R' 'open.R' 'sesamize.R' 'simulate_data.R' 'snp.R'
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sesame-package	<i>Analyze DNA methylation data</i>
----------------	-------------------------------------

Description

Sensible and step-wise analysis of DNA methylation data

Details

This package complements array functionalities that allow processing >10,000 samples in parallel on clusters.

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See Also

Useful links:

- <https://github.com/zwdzwd/sesame>
- Report bugs at <https://github.com/zwdzwd/sesame/issues>

Examples

```
sset <- readIDATpair(sub('_Grn.idat','',system.file(
  'extdata','4207113116_A_Grn.idat',package='sesameData'))))

## The OpenSesame pipeline
betas <- openSesame(sset)
```

`as.data.frame.sesameQC`*Coerce a sesameQC into a dataframe*

Description

Coerce a sesameQC into a dataframe

Usage

```
## S3 method for class 'sesameQC'  
as.data.frame(x, row.names = NULL, optional = FALSE, ...)
```

Arguments

<code>x</code>	a sesameQC object
<code>row.names</code>	see <code>as.data.frame</code>
<code>optional</code>	see <code>as.data.frame</code>
<code>...</code>	see <code>as.data.frame</code>

Value

a data.frame

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset  
qc <- sesameQC(sset)  
df <- as.data.frame(qc)
```

`BetaValueToMValue` *Convert beta-value to M-value*

Description

Logit transform a beta value vector to M-value vector.

Usage

```
BetaValueToMValue(b)
```

Arguments

<code>b</code>	vector of beta values
----------------	-----------------------

Details

Convert beta-value to M-value (aka logit transform)

Value

a vector of M values

Examples

```
BetaValueToMValue(c(0.1, 0.5, 0.9))
```

binSignals	<i>Bin signals from probe signals</i>
------------	---------------------------------------

Description

require GenomicRanges

Usage

```
binSignals(probe.signals, bin.coords, probe.coords)
```

Arguments

probe.signals	probe signals
bin.coords	bin coordinates
probe.coords	probe coordinates

Value

bin signals

bisConversionControl	<i>Compute internal bisulfite conversion control</i>
----------------------	--

Description

Compute GCT score for internal bisulfite conversion control. The function takes a SigSet as input. The higher the GCT score, the more likely the incomplete conversion.

Usage

```
bisConversionControl(sset, use.median = FALSE)
```

Arguments

sset	signal set
use.median	use median to compute GCT instead of mean

Value

GCT score (the higher, the more incomplete conversion)

Examples

```
sset <- makeExampleSeSAMEDataSet('HM450')
bisConversionControl(sset)
```

```
buildControlMatrix450k
```

Build control summary matrix

Description

The function takes a SigSet as input and outputs the control matrix summary vector. This vector summarizes one single QC metric for the array control. This includes bisulfite control, stain signal extension efficiency and more.

Usage

```
buildControlMatrix450k(sset)
```

Arguments

sset an object of class SigSet

Value

a vector with control summaries

Examples

```
sset <- makeExampleSeSAMEDataSet()
control.summary <- buildControlMatrix450k(sset)
```

```
chipAddressToSignal    Lookup address in one sample
```

Description

Lookup address and transform address to probe

Usage

```
chipAddressToSignal(dm, manifest, controls = NULL)
```

Arguments

dm data frame in chip address, 2 columns: cy3/Grn and cy5/Red
manifest a data frame with columns Probe_ID, M, U and col
controls a data frame with columns Address and Name. This is optional but might be necessary for some preprocessing methods that depends on these control probes. This is left for backward compatibility. Updated version should have controls consolidated into manifest.

Details

Translate data in chip address to probe address. Type I probes can be separated into Red and Grn channels. The methylated allele and unmethylated allele are at different addresses. For type II probes methylation allele and unmethylated allele are at the same address. Grn channel is for methylated allele and Red channel is for unmethylated allele. The out-of-band signals are type I probes measured using the other channel.

Value

a SigSet, indexed by probe ID address

cnSegmentation	<i>Perform copy number segmentation</i>
----------------	---

Description

Perform copy number segmentation using the signals in the signal set. The function takes a SigSet for the target sample and a set of normal SigSet for the normal samples. An optional arguments specifies the version of genome build that the inference will operate on. The function outputs an object of class CNSegment with signals for the segments (seg.signals), the bin coordinates (bin.coords) and bin signals (bin.signals).

Usage

```
cnSegmentation(sset, ssets.normal, refversion = c("hg19", "hg38"))
```

Arguments

sset	SigSet
ssets.normal	SigSet for normalization
refversion	hg19 or hg38

Value

an object of CNSegment

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
ssets.normal <- sesameDataGet('EPIC.5.normal')
seg <- cnSegmentation(sset, ssets.normal)
```

createUCSCtrack	<i>Turn beta values into a UCSC browser track</i>
-----------------	---

Description

Turn beta values into a UCSC browser track

Usage

```
createUCSCtrack(betas, output = NULL, platform = "HM450", refversion = "hg38")
```

Arguments

betas	a named numeric vector
output	output file name
platform	HM450, EPIC etc.
refversion	hg38, hg19 etc.

Value

when output is null, return a data.frame, otherwise NULL

Examples

```
betas.tissue <- sesameDataGet('HM450.1.TCGA.PAAD')$betas
## add output to create an actual file
df <- createUCSCtrack(betas.tissue)

## to convert to bigBed
## sort -k1,1 -k2,2n output.bed >output_sorted.bed
## bedToBigBed output_sorted.bed hg38.chrom output.bb
```

ctl	<i>ctl getter generic</i>
-----	---------------------------

Description

ctl getter generic
Get ctl slot of SigSet class

Usage

```
ctl(x)

## S4 method for signature 'SigSet'
ctl(x)
```

Arguments

x object of SigSet

Value

The ctl slot of SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
head(ctl(sset))
```

ctl<- *ctl replacement generic*

Description

ctl replacement generic

Replace ctl slot of SigSet class

Usage

```
ctl(x) <- value
```

```
## S4 replacement method for signature 'SigSet'
ctl(x) <- value
```

Arguments

x object of SigSet

value new value

Value

a new SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
df <- ctl(sset)
df[1,1] <- 10
ctl(sset) <- df
```

deIdentify	<i>De-identify IDATs by removing SNP probes</i>
------------	---

Description

Mask SNP probe intensity mean by zero.

Usage

```
deIdentify(path, out_path = NULL, snps = NULL, mft = NULL, randomize = FALSE)
```

Arguments

path	input IDAT file
out_path	output IDAT file
snps	SNP definition, if not given, default to SNP probes
mft	sesame-compatible manifest if non-standard
randomize	whether to randomize the SNPs. if TRUE, randomize the signal intensities. one can use set.seed to reidentify the IDAT with the secret seed (see examples). If FALSE, this sets all SNP intensities to zero.

Value

NULL, changes made to the IDAT files

Examples

```
my_secret <- 13412084
set.seed(my_secret)
temp_out <- tempfile("test")
deIdentify(system.file(
  "extdata", "4207113116_A_Grn.idat", package = "sesameData"),
  temp_out, randomize = TRUE)
unlink(temp_out)
```

detectionMask	<i>Mask Sigset by detection p-value</i>
---------------	---

Description

Mask Sigset by detection p-value

Usage

```
detectionMask(sset, pval.method = NULL, pval.threshold = 0.05)
```

Arguments

sset a SigSet
 pval.method which method to use in calculating p-values
 pval.threshold the p-value threshold

Value

a filtered SigSet

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
sset.masked <- detectionMask(sset)
```

detectionPfixedNorm *Detection P-value based on normal fitting with gived parameters*

Description

The function takes a SigSet as input, computes detection p-value using negative control probes parametrized in a normal distribution and returns a new SigSet with an updated pval slot.

Usage

```
detectionPfixedNorm(  

  sset,  

  muG = 500,  

  sdG = 200,  

  muR = 500,  

  sdR = 200,  

  force = FALSE  

)
```

Arguments

sset a SigSet
 muG mean of background in Grn channel
 sdG SD of background in Grn channel
 muR mean of background in Red channel
 sdR SD of background in Red channel
 force force rerun even if result already exists

Details

Background of Grn and Red are estimated separately from a fixed normal distribution. p-value is taken from the minimum of the p-value of the two alleles (color depends on probe design).

Value

detection p-value

Examples

```
sset <- makeExampleSeSAMEDataSet()
sset <- detectionPfixedNorm(sset)
```

detectionPnegEcdf *Detection P-value based on ECDF of negative control*

Description

The function takes a SigSet as input, computes detection p-value using negative control probes' empirical distribution and returns a new SigSet with an updated pval slot.

Usage

```
detectionPnegEcdf(sset, force = FALSE)
```

Arguments

sset	a SigSet
force	force rerun even if result already exists

Value

detection p-value

Examples

```
sset <- makeExampleSeSAMEDataSet()
sset <- detectionPnegEcdf(sset)
```

detectionPnegNorm *Detection P-value based on normal fitting the negative controls*

Description

The function takes a SigSet as input, computes detection p-value using negative control probes parametrized in a normal distribution and returns a new SigSet with an updated pval slot.

Usage

```
detectionPnegNorm(sset, force = FALSE)
```

Arguments

sset	a SigSet
force	force rerun even if result already exists

Details

Background of Grn and Red are estimated separately from negative control probes-parameterized normal distribution. p-value is taken from the minimum of the p-value of the two alleles (color depends on probe design).

Value

detection p-value

Examples

```
sset <- makeExampleSeSAMEDataSet()
sset <- detectionPnegNorm(sset)
```

detectionPnegNormGS *Detection P-value emulating Genome Studio*

Description

The function takes a SigSet as input, computes detection p-value using negative control probes parametrized in a normal distribution a la Genome Studio and returns a new SigSet with an updated pval slot.

Usage

```
detectionPnegNormGS(sset, force = FALSE)
```

Arguments

sset	a SigSet
force	force rerun even if result already exists

Details

P-value is calculated using negative control probes as the estimate of background where Grn channel and Red channel are merged. But when estimating p-value the Red and Grn are summed (non-ideal).

Value

detection p-value

Examples

```
sset <- makeExampleSeSAMEDataSet()
sset <- detectionPnegNormGS(sset)
```

detectionPnegNormTotal

Detection P-value based on normal fitting the negative controls, channels are first summed

Description

The function takes a SigSet as input, computes detection p-value using negative control probes parametrized in a normal distribution with the two channels summed first and returns a new SigSet with an updated pval slot. The SD is summed to emulate the SD of the summed signal (not the most accurate treatment).

Usage

```
detectionPnegNormTotal(sset, force = FALSE)
```

Arguments

sset	a SigSet
force	force rerun even if result already exists

Value

detection p-value

Examples

```
sset <- makeExampleSeSAMEDataSet()
sset <- detectionPnegNormTotal(sset)
```

detectionPoobEcdf

Detection P-value based on ECDF of out-of-band signal

Description

aka pOOBAH (p-vals by Out-Of-Band Array Hybridization)

Usage

```
detectionPoobEcdf(sset, force = FALSE)
```

```
pOOBAH(sset, force = FALSE)
```

Arguments

sset	a SigSet
force	force rerun even if result already exists

Details

The function takes a SigSet as input, computes detection p-value using out-of-band probes empirical distribution and returns a new SigSet with an updated pval slot.

Value

detection p-value

Examples

```
sset <- makeExampleSeSAMEDataSet()
sset <- detectionPoobEcdf(sset)

sset <- makeExampleSeSAMEDataSet()
sset <- pOOBAH(sset)
```

detectionZero

Detection P-value set to all zero

Description

Detection P-value set to all zero

Usage

```
detectionZero(sset, force = FALSE)
```

Arguments

sset	a SigSet
force	force rerun even if result already exists

Value

detection p-value set to all zero

Examples

```
sset <- makeExampleSeSAMEDataSet()
sset <- detectionZero(sset)
```

diffRefSet	<i>Restrict refset to differentially methylated probes use with care, might introduce bias</i>
------------	--

Description

The function takes a matrix with probes on the rows and cell types on the columns and output a subset matrix and only probes that show discordant methylation levels among the cell types.

Usage

```
diffRefSet(g)
```

Arguments

`g` a matrix with probes on the rows and cell types on the columns

Value

`g` a matrix with a subset of input probes (rows)

Examples

```
g <- diffRefSet(getRefSet(platform='HM450'))
```

DML	<i>Test differential methylation on each locus</i>
-----	--

Description

The function takes a beta value matrix with probes on the rows and samples on the columns. It also takes a sample information data frame (`sample.data`) and formula for testing. The function outputs a list of coefficient tables for each factor tested.

Usage

```
DML(  
  betas,  
  sample.data,  
  formula,  
  se.lb = 0.06,  
  balanced = FALSE,  
  cf.test = NULL  
)
```

Arguments

betas	beta values
sample.data	data frame for sample information, column names are predictor variables (e.g., sex, age, treatment, tumor/normal etc) and are referenced in formula. Rows are samples.
formula	formula
se.lb	lower bound to standard error of slope, lower this to get more difference of small effect size.
balanced	whether design is balanced or not. default to FALSE, when unbalanced will use Welch's method to estimate standard error. balance=TRUE is faster.
cf.test	factors to test (default to all factors in formula except intercept). Use "all" for all factors.

Value

cf - a list of coefficient tables for each factor

Examples

```
data <- sesameDataGet('HM450.76.TCGA.matched')
cf <- DML(data$betas, data$sampleInfo, ~type)
```

DMR

Find Differentially Methylated Region (DMR)

Description

This subroutine uses Euclidean distance to group CpGs and then combine p-values for each segment. The function performs DML test first if cf is NULL. It groups the probe testing results into differential methylated regions in a coefficient table with additional columns designating the segment ID and statistical significance (P-value) testing the segment.

Usage

```
DMR(
  betas,
  sample.data = NULL,
  formula = NULL,
  cf = NULL,
  dist.cutoff = NULL,
  seg.per.locus = 0.5,
  platform = c("EPIC", "HM450"),
  refversion = c("hg38", "hg19"),
  ...
)
```

Arguments

betas	beta values for distance calculation
sample.data	data frame for sample information, column names are predictor variables (e.g., sex, age, treatment, tumor/normal etc) and are referenced in formula. Rows are samples.
formula	formula
cf	coefficient table from diffMeth, when NULL will be computed from beta. If cf is given, sample.data and formula are ignored.
dist.cutoff	distance cutoff (default to use dist.cutoff.quantile)
seg.per.locus	number of segments per locus higher value leads to more segments
platform	EPIC or HM450
refversion	hg38 or hg19
...	additional parameters to DML

Value

coefficient table with segment ID and segment P-value

Examples

```
data <- sesameDataGet('HM450.76.TCGA.matched')
cf <- DMR(data$betas, data$sampleInfo, ~type)
```

dyeBiasCorr *Correct dye bias in by linear scaling.*

Description

The function takes a SigSet as input and scale both the Grn and Red signal to a reference (ref) level. If the reference level is not given, it is set to the mean intensity of all the in-band signals. The function returns a SigSet with dye bias corrected.

Usage

```
dyeBiasCorr(sset, ref = NULL)
```

Arguments

sset	a SigSet
ref	reference signal level

Value

a normalized SigSet

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
sset.db <- dyeBiasCorr(sset)
```

dyeBiasCorrMostBalanced

Correct dye bias using most balanced sample as the reference

Description

The function chose the reference signal level from a list of SigSet. The chosen sample has the smallest difference in Grn and Red signal intensity as measured using the normalization control probes. In practice, it doesn't matter which sample is chosen as long as the reference level does not deviate much. The function returns a list of SigSets with dye bias corrected.

Usage

```
dyeBiasCorrMostBalanced(ssets)
```

Arguments

ssets a list of normalized SigSets

Value

a list of normalized SigSets

Examples

```
ssets <- sesameDataGet('HM450.10.TCGA.BLCA.normal')
ssets.db <- dyeBiasCorrMostBalanced(ssets)
```

dyeBiasCorrTypeINorm *Dye bias correction by matching green and red to mid point*

Description

This function compares the Type-I Red probes and Type-I Grn probes and generates and mapping to correct signal of the two channels to the middle. The function takes one single SigSet and returns a SigSet with dye bias corrected.

Usage

```
dyeBiasCorrTypeINorm(sset)
```

Arguments

sset a SigSet

Value

a SigSet after dye bias correction.

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
sset.db <- dyeBiasCorrTypeINorm(sset)
```

 estimateCellComposition

Estimate cell composition using reference

Description

This is a reference-based cell composition estimation. The function takes a reference methylation status matrix (rows for probes and columns for cell types, can be obtained by getRefSet function) and a query beta value measurement. The length of the target beta values should be the same as the number of rows of the reference matrix. The method assumes one unknown component. It outputs a list containing the estimated cell fraction, the error of optimization and methylation status of the unknown component.

Usage

```
estimateCellComposition(g, q, refine = TRUE, dichotomize = FALSE, ...)
```

Arguments

g	reference methylation
q	target measurement: length(q) == nrow(g)
refine	to refine estimate, takes longer
dichotomize	to dichotomize query beta value before estimate, this relieves unclean background subtraction
...	extra parameters for optimization, this includes temp - annealing temperature (0.5) maxIter - maximum iteration to stop after converge (1000) delta - delta score to reset counter (0.0001) verbose - output debug info (FALSE)

Value

a list of fraction, min error and unknown component methylation state

estimateLeukocyte

Estimate leukocyte fraction using a two-component model

Description

The method assumes only two components in the mixture: the leukocyte component and the target tissue component. The function takes the beta values matrix of the target tissue and the beta value matrix of the leukocyte. Both matrices have probes on the row and samples on the column. Row names should have probe IDs from the platform. The function outputs a single numeric describing the fraction of leukocyte.

Usage

```
estimateLeukocyte(
  betas.tissue,
  betas.leuko = NULL,
  betas.tumor = NULL,
  platform = c("EPIC", "HM450", "HM27")
)
```

Arguments

betas.tissue tissue beta value matrix (#probes X #samples)
 betas.leuko leukocyte beta value matrix, if missing, use the SeSAmE default by infinium platform
 betas.tumor optional, tumor beta value matrix
 platform "HM450", "HM27" or "EPIC"

Value

leukocyte estimate, a numeric vector

Examples

```
betas.tissue <- sesameDataGet('HM450.1.TCGA.PAAD')$betas
estimateLeukocyte(betas.tissue)
```

 extra

extra getter generic

Description

extra getter generic
 Get extra slot of SigSet class

Usage

```
extra(x)

## S4 method for signature 'SigSet'
extra(x)
```

Arguments

x object of SigSet

Value

The extra slot of SigSet

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
head(extra(sset))
```

extra<- *extra replacement generic*

Description

extra replacement generic
 Replace extra slot of SigSet class

Usage

```
extra(x) <- value

## S4 replacement method for signature 'SigSet'
extra(x) <- value
```

Arguments

x	object of SigSet
value	new value

Value

a new SigSet

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
df <- extra(sset)
extra(sset) <- list(pval=numeric(0))
```

formatVCF *Convert SNP from Infinium array to VCF file*

Description

Convert SNP from Infinium array to VCF file

Usage

```
formatVCF(sset, vcf = NULL, refversion = "hg19", annoS = NULL, annoI = NULL)
```

Arguments

sset	SigSet
vcf	output VCF file path, if NULL output to console
refversion	reference version, currently only support
annoS	SNP variant annotation, download if not given
annoI	Infinium-I variant annotation, download if not given hg19 and hg38 in human

Value

VCF file. If vcf is NULL, a data.frame is output to console. The data.frame does not contain VCF headers.

Note the vcf is not sorted. You can sort with `awk '$1 ~ /^#/ print $0;next print $0 | "sort -k1,1 -k2,2n"'`

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset

annoS <- sesameDataPullVariantAnno_SNP('EPIC', 'hg19')
annoI <- sesameDataPullVariantAnno_InfiniumI('EPIC', 'hg19')
## output to console
head(formatVCF(sset, annoS=annoS, annoI=annoI))
```

getAFTypeIbySumAlleles

Get allele frequency treating type I by summing alleles

Description

Takes a SigSet as input and returns a numeric vector containing extra allele frequencies based on Color-Channel-Switching (CCS) probes. If no CCS probes exist in the SigSet, then an numeric(0) is returned.

Usage

```
getAFTypeIbySumAlleles(sset, known.ccs.only = TRUE)
```

Arguments

```
sset          SigSet
known.ccs.only  consider only known CCS probes
```

Value

beta values

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
betas <- getAFTypeIbySumAlleles(sset)
```

getBetas	<i>Get beta Values</i>
----------	------------------------

Description

sum.typeI is used for rescuing beta values on Color-Channel-Switching CCS probes. The function takes a SigSet and returns beta value except that Type-I in-band signal and out-of-band signal are combined. This prevents color-channel switching due to SNPs.

Usage

```
getBetas(sset, mask = TRUE, sum.TypeI = FALSE)
```

Arguments

sset	SigSet
mask	whether to use mask
sum.TypeI	whether to sum type I channels

Value

a numeric vector, beta values

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset  
betas <- getBetas(sset)
```

getBinCoordinates	<i>Get bin coordinates</i>
-------------------	----------------------------

Description

requires GenomicRanges, IRanges

Usage

```
getBinCoordinates(seqInfo, gapInfo, probe.coords)
```

Arguments

seqInfo	chromosome information object
gapInfo	chromosome gap information
probe.coords	probe coordinates

Value

bin.coords

getNormCtrls	<i>get normalization control signal</i>
--------------	---

Description

get normalization control signal from SigSet. The function optionally takes mean for each channel.

Usage

```
getNormCtrls(sset, average = FALSE)
```

Arguments

sset	a SigSet
average	whether to average

Value

a data frame of normalization control signals

Examples

```
sset <- readIDATpair(file.path(system.file(
  'extdata', '', package='sesameData'), '4207113116_B'))

df.ctl <- getNormCtrls(sset)
```

getProbesByGene	<i>Get Probes by Gene</i>
-----------------	---------------------------

Description

Get probes mapped to a gene. All transcripts for the gene are considered. The function takes a gene name as appears in UCSC RefGene database. The platform and reference genome build can be changed with 'platform' and 'refversion' options. The function returns a vector of probes that falls into the given gene.

Usage

```
getProbesByGene(
  geneName,
  platform = c("EPIC", "HM450"),
  upstream = 0,
  downstream = 0,
  refversion = c("hg38", "hg19")
)
```

Arguments

geneName	gene name
platform	EPIC or HM450
upstream	number of bases to expand upstream of target gene
dwestream	number of bases to expand downstream of target gene
refversion	hg38 or hg19

Value

probes that fall into the given gene

Examples

```
probes <- getProbesByGene('CDKN2A', upstream=500, dwestream=500)
```

getProbesByRegion *Get probes by genomic region*

Description

The function takes a genomic coordinate and output the a vector of probes on the specified platform that falls in the given genomic region.

Usage

```
getProbesByRegion(
  chrm,
  beg = 1,
  end = -1,
  platform = c("EPIC", "HM450"),
  refversion = c("hg38", "hg19")
)
```

Arguments

chrm	chromosome
beg	begin, 1 if omitted
end	end, chromosome end if omitted
platform	EPIC or HM450
refversion	hg38 or hg19

Value

probes that fall into the given region

Examples

```
getProbesByRegion('chr5', 135413937, 135419936,
  refversion = 'hg19', platform = 'HM450')
```

getProbesByTSS	<i>Get Probes by Gene Transcription Start Site (TSS)</i>
----------------	--

Description

Get probes mapped to a TSS. All transcripts for the gene are considered. The function takes a gene name as appears in UCSC RefGene database. The platform and reference genome build can be changed with 'platform' and 'refversion' options. The function returns a vector of probes that falls into the TSS region of the gene.

Usage

```
getProbesByTSS(
  geneName,
  upstream = 1500,
  dwestream = 1500,
  platform = c("EPIC", "HM450"),
  refversion = c("hg38", "hg19")
)
```

Arguments

geneName	gene name
upstream	the number of base pairs to expand upstream the TSS
dwestream	the number of base pairs to expand dwestream the TSS
platform	EPIC or HM450
refversion	hg38 or hg19

Value

probes that fall into the given gene

Examples

```
probes <- getProbesByTSS('CDKN2A')
```

getRefSet	<i>Retrieve reference set</i>
-----------	-------------------------------

Description

The function retrieves the curated reference DNA methylation status for a set of cell type names under the Infinium platform. Supported cell types include "CD4T", "CD19B", "CD56NK", "CD14Monocytes", "granulocytes", "scFat", "skin" etc. See package sesameData for more details. The function output a matrix with probes on the rows and specified cell types on the columns. 0 suggests unmethylation and 1 suggests methylation. Intermediate methylation and nonclusive calls are left with NA.

Usage

```
getRefSet(cells = NULL, platform = c("EPIC", "HM450"))
```

Arguments

cells	reference cell types
platform	EPIC or HM450

Value

g, a 0/1 matrix with probes on the rows and specified cell types on the columns.

Examples

```
betas <- getRefSet('CD4T', platform='HM450')
```

getSegment	<i>Select segment from coefficient table</i>
------------	--

Description

This function takes a coefficient table and returns a subset of the table targeting only the specified segment using segment ID.

Usage

```
getSegment(cf1, seg.id)
```

Arguments

cf1	coefficient table of one factor from DMR
seg.id	segment ID

Value

coefficient table from given segment

Examples

```
data <- sesameDataGet('HM450.76.TCGA.matched')
cf <- DMR(data$betas, data$sampleInfo, ~type)
getSegment(cf[[1]], cf[[1]][['Seg.ID']][1])
```

getSexInfo	<i>Get sex-related information</i>
------------	------------------------------------

Description

The function takes a `SigSet` and returns a vector of three numerics: the median intensity of chrY probes; the median intensity of chrX probes; and fraction of intermediate chrX probes. chrX and chrY probes excludes pseudo-autosomal probes.

Usage

```
getSexInfo(sset)
```

Arguments

sset a `SigSet`

Value

medianY and medianX, fraction of XCI, methylated and unmethylated X probes, median intensities of auto-chromosomes.

Examples

```
sset <- makeExampleSeSAMEDataSet()
getSexInfo(sset)
```

IG	<i>IG getter generic</i>
----	--------------------------

Description

IG getter generic
Get IG slot of `SigSet` class

Usage

```
IG(x)

## S4 method for signature 'SigSet'
IG(x)
```

Arguments

x object of `SigSet`

Value

The IG slot of `SigSet`

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
head(IG(sset))
```

 IG<-

IG replacement generic

Description

IG replacement generic
 Replace IG slot of SigSet class

Usage

```
IG(x) <- value

## S4 replacement method for signature 'SigSet'
IG(x) <- value
```

Arguments

x	object of SigSet
value	new value

Value

a new SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
df <- IG(sset)
df[1,1] <- 10
IG(sset) <- df
```

 II

II getter generic

Description

II getter generic
 Get II slot of SigSet class

Usage

```
II(x)

## S4 method for signature 'SigSet'
II(x)
```

Arguments

x object of SigSet

Value

The II slot of SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
head(II(sset))
```

 II<-

II replacement generic

Description

II replacement generic

Replace II slot of SigSet class

Usage

```
II(x) <- value
```

```
## S4 replacement method for signature 'SigSet'
II(x) <- value
```

Arguments

x object of SigSet

value new value

Value

a new SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
df <- II(sset)
df[1,1] <- 10
II(sset) <- df
```

inferEthnicity	<i>Infer Ethnicity</i>
----------------	------------------------

Description

This function uses both the built-in rsprobes as well as the type I Color-Channel-Switching probes to infer ethnicity.

Usage

```
inferEthnicity(sset)
```

Arguments

sset a SigSet

Details

sset better be background subtracted and dyebias corrected for best accuracy

Please note: the betas should come from sigset **without** channel inference.

Value

string of ethnicity

Examples

```
sset <- makeExampleSeSAMEDataSet("HM450")
inferEthnicity(sset)
```

inferSex	<i>Infer Sex</i>
----------	------------------

Description

Infer Sex

Usage

```
inferSex(sset)
```

Arguments

sset a SigSet

Value

'F' or 'M' We established our sex calling based on the median intensity of chromosome X, Y and the fraction of intermediately methylated probes among the identified X-linked probes. This is similar to the approach by Minfi (Aryee et al., 2014) but also different in that we used the fraction of intermediate beta value rather than median intensity for all chromosome X probes. Instead of using all probes from the sex chromosomes, we used our curated set of Y chromosome probes and X-linked probes which exclude potential cross-hybridization and pseudo-autosomal effect.

XXY male (Klinefelter's), 45,X female (Turner's) can confuse the model sometimes. Our function works on a single sample.

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
inferSex(sset)
```

inferSexKaryotypes *Infer Sex Karyotype*

Description

The function takes a SigSet and infers the sex chromosome Karyotype and presence/absence of X-chromosome inactivation (XCI). chrX, chrY and XCI are inferred relatively independently. This function gives a more detailed look of potential sex chromosome aberrations.

Usage

```
inferSexKaryotypes(sset)
```

Arguments

sset a SigSet

Value

Karyotype string, with XCI

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
inferSexKaryotypes(sset)
```

inferTypeIChannel	<i>Infer and reset color channel for Type-I probes instead of using what is specified in manifest. The results are stored to sset@extra\$IGG and sset@extra\$IRR slot.</i>
-------------------	--

Description

IGG => Type-I green that is inferred to be green IRR => Type-I red that is inferred to be red

Usage

```
inferTypeIChannel(
  sset,
  switch_failed = FALSE,
  verbose = FALSE,
  summary = FALSE
)
```

Arguments

sset	a SigSet
switch_failed	whether to switch failed probes (default to FALSE)
verbose	whether to print correction summary
summary	return summarized numbers only.

Value

a SigSet, or numerics if summary == TRUE

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
inferTypeIChannel(sset)
```

initFileSet	<i>initialize a fileSet class by allocating appropriate storage</i>
-------------	---

Description

initialize a fileSet class by allocating appropriate storage

Usage

```
initFileSet(map_path, platform, samples, probes = NULL, inc = 4)
```

Arguments

map_path	path of file to map
platform	EPIC, HM450 or HM27, consistent with sset@platform
samples	sample names
probes	probe names
inc	bytes per unit data storage

Value

a sesame::fileSet object

Examples

```
fset <- initFileSet('mybetas2', 'HM27', c('s1','s2'))
```

 IR

IR getter generic

Description

IR getter generic

Get IR slot of SigSet class

Usage

```
IR(x)
```

```
## S4 method for signature 'SigSet'
IR(x)
```

Arguments

x object of SigSet

Value

The IR slot of SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
head(IR(sset))
```

IR<- *IR replacement generic*

Description

IR replacement generic
 Replace IR slot of SigSet class

Usage

```
IR(x) <- value

## S4 replacement method for signature 'SigSet'
IR(x) <- value
```

Arguments

x	object of SigSet
value	new value

Value

a new SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
df <- IR(sset)
df[1,1] <- 10
IR(sset) <- df
```

makeExampleSeSAMEDataSet
Make a simulated SeSAMEData set

Description

Constructs a simulated SigSet dataset. For the given platform, randomly simulate methylated and unmethylated allele signals. In-band signals were simulated using a N(4000, 200) normal distribution. Out-of-band signals were simulated using a N(400, 200) normal distribution. Control signals were simulated using a N(400, 300) normal distribution.

Usage

```
makeExampleSeSAMEDataSet(platform = c("HM450", "EPIC", "HM27"))
```

Arguments

platform	optional, HM450, EPIC or HM27
----------	-------------------------------

Value

Object of class SigSet

Examples

```
sset <- makeExampleSeSAMEDataSet()
```

```
makeExampleTinyEPICDataSet
```

Make a tiny toy simulated EPIC data set

Description

Construct a tiny EPIC SigSet of only 6 probes. In-band signals were simulated using a N(4000, 200) normal distribution. Out-of-band signals were simulated using a N(400, 200) normal distribution. Control signals were simulated using a N(400, 300) normal distribution.

Usage

```
makeExampleTinyEPICDataSet()
```

Value

Object of class SigSet

Examples

```
sset <- makeExampleTinyEPICDataSet()
```

```
mapFileSet
```

Deposit data of one sample to a fileSet (and hence to file)

Description

Deposit data of one sample to a fileSet (and hence to file)

Usage

```
mapFileSet(fset, sample, named_values)
```

Arguments

fset	a sesame::fileSet, as obtained via readFileSet
sample	sample name as a string
named_values	value vector named by probes

Value

a sesame::fileSet

Examples

```
## create two samples
fset <- initFileSet('mybetas2', 'HM27', c('s1','s2'))

## a hypothetical numeric array (can be beta values, intensities etc)
hypothetical <- setNames(runif(fset$n), fset$probes)

## map the numeric to file
mapFileSet(fset, 's1', hypothetical)

## get data
sliceFileSet(fset, 's1', 'cg00000292')
```

meanIntensity	<i>Mean Intensity</i>
---------------	-----------------------

Description

The function takes one single SigSet and computes mean intensity of all the in-band measurements. This includes all Type-I in-band measurements and all Type-II probe measurements. Both methylated and unmethylated alleles are considered. This function outputs a single numeric for the mean.

Usage

```
meanIntensity(sset)
```

Arguments

sset a SigSet

Value

mean of all intensities

Examples

```
sset <- makeExampleSeSAMEDataset()
meanIntensity(sset)
```

MValueToBetaValue	<i>Convert M-value to beta-value</i>
-------------------	--------------------------------------

Description

Convert M-value to beta-value (aka inverse logit transform)

Usage

```
MValueToBetaValue(m)
```

Arguments

m	a vector of M values
---	----------------------

Value

a vector of beta values

Examples

```
MValueToBetaValue(c(-3, 0, 3))
```

noob	<i>Noob background correction</i>
------	-----------------------------------

Description

The function takes a SigSet and returns a modified SigSet with background subtracted. Background was modelled in a normal distribution and true signal in an exponential distribution. The Norm-Exp deconvolution is parameterized using Out-Of-Band (oob) probes

Usage

```
noob(sset, oobRprobes = NULL, oobGprobes = NULL, offset = 15)
```

Arguments

sset	a SigSet
oobRprobes	out-of-band red probes, if not given use all oobR
oobGprobes	out-of-band grn probes, if not given use all oobG
offset	offset

Value

a new SigSet with noob background correction

Examples

```
sset <- makeExampleTinyEPICDataSet()
sset.nb <- noob(sset)
```

noobsb	<i>Background subtraction with bleeding-through subtraction</i>
--------	---

Description

The function takes a SigSet and returns a modified SigSet with background subtracted. Signal bleed-through was modelled using a linear model with error estimated from cross-channel regression. Norm-Exp deconvolution using Out-Of-Band (oob) probes.

Usage

```
noobsb(sset, offset = 15, detailed = FALSE)
```

Arguments

sset	a SigSet
offset	offset
detailed	if TRUE, return a list of SigSet and regression function

Value

a modified SigSet with background correction

Examples

```
sset <- makeExampleSeSAMEDataSet('HM450')
sset.nb <- noobsb(sset)
```

oobG	<i>oobG getter generic</i>
------	----------------------------

Description

oobG getter generic
Get oobG slot of SigSet class

Usage

```
oobG(x)

## S4 method for signature 'SigSet'
oobG(x)
```

Arguments

x	object of SigSet
---	------------------

Value

The oobG slot of SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
head(oobG(sset))
```

```
oobG<- oobG replacement generic
```

Description

oobG replacement generic
 Replace oobG slot of SigSet class

Usage

```
oobG(x) <- value

## S4 replacement method for signature 'SigSet'
oobG(x) <- value
```

Arguments

x	object of SigSet
value	new value

Value

a new SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
df <- oobG(sset)
df[1,1] <- 10
oobG(sset) <- df
```

```
oobR oobR getter generic
```

Description

oobR getter generic
 Get oobR slot of SigSet class

Usage

```
oobR(x)

## S4 method for signature 'SigSet'
oobR(x)
```

Arguments

x object of SigSet

Value

The oobR slot of SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
head(oobR(sset))
```

oobR<- *oobR replacement generic*

Description

oobR replacement generic
Replace oobR slot of SigSet class

Usage

```
oobR(x) <- value

## S4 replacement method for signature 'SigSet'
oobR(x) <- value
```

Arguments

x object of SigSet
value new value

Value

a new SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
df <- oobR(sset)
df[1,1] <- 10
oobR(sset) <- df
```

 openSesame

The openSesame pipeline

Description

This function is a simple wrapper of noob + nonlinear dye bias correction + pOOBAH masking.

Usage

```
openSesame(
  x,
  platform = "",
  manifest = NULL,
  what = "beta",
  BPPARAM = SerialParam(),
  ...
)
```

Arguments

x	SigSet(s), IDAT prefix(es), minfi GenomicRatioSet(s), or RGChannelSet(s)
platform	optional platform string
manifest	optional dynamic manifest
what	either 'sigset' or 'beta'
BPPARAM	get parallel with MulticoreParam(n)
...	parameters to getBetas

Details

If the input is an IDAT prefix or a SigSet, the output is the beta value numerics. If the input is a minfi GenomicRatioSet or RGChannelSet, the output is the sesamized GenomicRatioSet.

Value

a numeric vector for processed beta values

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
IDATprefixes <- searchIDATprefixes(
  system.file("extdata", "", package = "sesameData"))
betas <- openSesame(IDATprefixes)
```

openSesameToFile *openSesame pipeline with file-backed storage*

Description

openSesame pipeline with file-backed storage

Usage

```
openSesameToFile(map_path, idat_dir, BPPARAM = SerialParam(), inc = 4)
```

Arguments

map_path	path of file to be mapped (beta values file)
idat_dir	source IDAT directory
BPPARAM	get parallel with MulticoreParam(2)
inc	bytes per item data storage. increase to 8 if precision is important. Most cases 32-bit representation is enough.

Value

a sesame::fileSet

Examples

```
openSesameToFile('mybetas',
  system.file('extdata', package='sesameData'))
```

parseGEOsignalABFile *Parse GEO signal-A/B File into a SigSet List*

Description

This function is meant to be a convenience function for parsing data from Signal_A and Signal_B file provided by GEO. In many cases, this function generates a "partial" SigSet due to lack of out-of-band signal and control probe measurement in those Signal_A/B files. The detection p-value is based on a fixed normal distribution rather than from negative control or OOB probes.

Usage

```
parseGEOsignalABFile(path, platform = "HM450", drop = TRUE, parallel = TRUE)
```

Arguments

path	path to Signal-A/B file downloaded from GEO. The file can remain gzipped.
platform	HM450, EPIC or HM27
drop	whether to reduce to SigSet when there is only one sample.
parallel	whether to use multiple cores.

Value

a SigSetList or a SigSet

Examples

```
path = system.file(
  'extdata',
  'GSE36369_NonEBV_SignalA_SignalB_3samples_1k.txt.gz',
  package='sesame')
ssets <- parseGEOSignalABFile(path)
```

predictAgeHorvath353 *Horvath 353 age predictor*

Description

The function takes a named numeric vector of beta values. The name attribute contains the probe ID (cg, ch or rs IDs). The function looks for overlapping probes and estimate age using Horvath aging model (Horvath 2013 Genome Biology). The function outputs a single numeric of age in years.

Usage

```
predictAgeHorvath353(betas)
```

Arguments

betas a probeID-named vector of beta values

Value

age in years

Examples

```
betas <- sesameDataGet('HM450.1.TCGA.PAAD')$betas
predictAgeHorvath353(betas)
```

predictAgePheno *Phenotypic age predictor*

Description

The function takes a named numeric vector of beta values. The name attribute contains the probe ID (cg, ch or rs IDs). The function looks for overlapping probes and estimate age using Horvath aging model (Levine et al. 2018 Aging, 513 probes). The function outputs a single numeric of age in years.

Usage

```
predictAgePheno(betas)
```

Arguments

betas a probeID-named vector of beta values

Value

age in years

Examples

```
betas <- sesameDataGet('HM450.1.TCGA.PAAD')$betas  
predictAgePheno(betas)
```

predictAgeSkinBlood *Horvath Skin and Blood age predictor*

Description

The function takes a named numeric vector of beta values. The name attribute contains the probe ID (cg, ch or rs IDs). The function looks for overlapping probes and estimate age using Horvath aging model (Horvath et al. 2018 Aging, 391 probes). The function outputs a single numeric of age in years.

Usage

```
predictAgeSkinBlood(betas)
```

Arguments

betas a probeID-named vector of beta values

Value

age in years

Examples

```
betas <- sesameDataGet('HM450.1.TCGA.PAAD')$betas  
predictAgeSkinBlood(betas)
```

print.fileSet *Print a fileSet*

Description

Print a fileSet

Usage

```
## S3 method for class 'fileSet'  
print(x, ...)
```

Arguments

x a sesame::fileSet
... stuff for print

Value

string representation

Examples

```
fset <- initFileSet('mybetas2', 'HM27', c('s1','s2'))  
fset
```

print.sesameQC *Print sesameQC object*

Description

Print sesameQC object

Usage

```
## S3 method for class 'sesameQC'  
print(x, ...)
```

Arguments

x a sesameQC object
... extra parameter for print

Value

print sesameQC result on screen

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
sesameQC(sset)
```

probeNames	<i>Get Probe Names of SigSet class</i>
------------	--

Description

Get Probe Names of SigSet class

Usage

```
probeNames(x)

## S4 method for signature 'SigSet'
probeNames(x)
```

Arguments

x object of Sigset

Value

A char vector

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset
head(probeNames(sset))
```

pval	<i>pval getter generic</i>
------	----------------------------

Description

pval getter generic
Get pval slot of SigSet class

Usage

```
pval(x)

## S4 method for signature 'SigSet'
pval(x)
```

Arguments

x object of SigSet

Value

The pval slot of SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset  
head(pval(sset))
```

pval<- *pval replacement generic*

Description

pval replacement generic

Replace pval slot of SigSet class

Usage

```
pval(x) <- value  
  
## S4 replacement method for signature 'SigSet'  
pval(x) <- value
```

Arguments

x	object of SigSet
value	new value

Value

a new SigSet

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset  
df <- pval(sset)  
df[1] <- 0.01  
pval(sset) <- df
```

qualityMask	<i>Mask beta values by design quality</i>
-------------	---

Description

Currently quality masking only supports three platforms

Usage

```
qualityMask(sset, mask.use.tcga = FALSE)
```

Arguments

sset	a SigSet object
mask.use.tcga	whether to use TCGA masking, only applies to HM450

Value

a filtered SigSet

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
sset.masked <- qualityMask(sset)
```

qualityRank	<i>This function looks at public data of similar nature e.g., tissue, FFPE vs non-FFPE, etc to evaluate the quality of the target data quality</i>
-------------	--

Description

This function looks at public data of similar nature e.g., tissue, FFPE vs non-FFPE, etc to evaluate the quality of the target data quality

Usage

```
qualityRank(sset, tissue = NULL, samplePrep = NULL, raw = FALSE)
```

Arguments

sset	a raw (unprocessed) SigSet
tissue	A string (blood,buccal and saliva)
samplePrep	FFPE, fresh, frozen
raw	to return the raw comparison table

Value

three numbers: 1. The number of public samples compared. 2. The fraction of public samples with more nondetection, and 3. The fraction of public samples with lower mean intensity 4. The higher the fraction, the better the sample.

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
ranks <- qualityRank(sset)
```

readFileSet	<i>Read an existing fileSet from storage</i>
-------------	--

Description

This function only reads the meta-data.

Usage

```
readFileSet(map_path)
```

Arguments

map_path path of file to map (should contain valid _idx.rds index)

Value

a sesame::fileSet object

Examples

```
## create two samples
fset <- initFileSet('mybetas2', 'HM27', c('s1','s2'))

## a hypothetical numeric array (can be beta values, intensities etc)
hypothetical <- setNames(runif(fset$n), fset$probes)

## map the numeric to file
mapFileSet(fset, 's1', hypothetical)

## read it from file
fset <- readFileSet('mybetas2')

## get data
sliceFileSet(fset, 's1', 'cg00000292')
```

readIDATpair	<i>Import a pair of IDATs from one sample</i>
--------------	---

Description

The function takes a prefix string that are shared with `_Grn.idat` and `_Red.idat`. The function returns a `SigSet`.

Usage

```
readIDATpair(
  prefix.path,
  platform = "",
  manifest = NULL,
  controls = NULL,
  verbose = FALSE
)
```

Arguments

<code>prefix.path</code>	sample prefix without <code>_Grn.idat</code> and <code>_Red.idat</code>
<code>platform</code>	EPIC, HM450 and HM27 etc.
<code>manifest</code>	optional design manifest file
<code>controls</code>	optional control probe manifest file
<code>verbose</code>	be verbose? (FALSE)

Value

a `SigSet`

Examples

```
sset <- readIDATpair(sub('_Grn.idat', '', system.file(
  "extdata", "4207113116_A_Grn.idat", package = "sesameData")))
```

reIdentify	<i>Re-identify IDATs by restoring scrambled SNP intensities</i>
------------	---

Description

This requires setting a seed with a secret number that was used to de-identify the IDAT (see example). This requires a secret number that was used to de-identify the IDAT

Usage

```
reIdentify(path, out_path = NULL, snps = NULL, mft = NULL)
```

Arguments

path	input IDAT file
out_path	output IDAT file
snps	SNP definition, if not given, default to SNP probes
mft	sesame-compatible manifest if non-standard

Value

NULL, changes made to the IDAT files

Examples

```
temp_out <- tempfile("test")

set.seed(123)
reIdentify(system.file(
  "extdata", "4207113116_A_Grn.idat", package = "sesameData"), temp_out)
unlink(temp_out)
```

reopenSesame	<i>re-compute beta value for GenomicRatioSet</i>
--------------	--

Description

re-compute beta value for GenomicRatioSet

Usage

```
reopenSesame(x, naFrac = 0.2)
```

Arguments

x	GenomicRatioSet
naFrac	maximum NA fraction for a probe before it gets dropped (1)

Value

a GenomicRatioSet

RGChannelSetToSigSets *Convert RGChannelSet (minfi) to a list of SigSet (SeSAME)*

Description

Notice the colData() and rowData() is lost. Most cases, rowData is empty anyway.

Usage

```
RGChannelSetToSigSets(rgSet, manifest = NULL, BPPARAM = SerialParam())
```

Arguments

rgSet	a minfi::RGChannelSet
manifest	manifest file
BPPARAM	get parallel with MulticoreParam(n)

Value

a list of sesame::SigSet

Examples

```
if (require(FlowSorted.Blood.450k)) {
  rgSet <- FlowSorted.Blood.450k[,1:2]
  ssets <- RGChannelSetToSigSets(rgSet)
}
```

searchIDATprefixes *Identify IDATs from a directory*

Description

The input is the directory name as a string. The function identifies all the IDAT files under the directory. The function returns a vector of such IDAT prefixes under the directory.

Usage

```
searchIDATprefixes(dir.name, recursive = TRUE, use.basename = TRUE)
```

Arguments

dir.name	the directory containing the IDAT files.
recursive	search IDAT files recursively
use.basename	basename of each IDAT path is used as sample name This won't work in rare situation where there are duplicate IDAT files.

Value

the IDAT prefixes (a vector of character strings).

Examples

```
## only search what are directly under
IDATprefixes <- searchIDATprefixes(
  system.file("extdata", "", package = "sesameData"))

## search files recursively is by default
IDATprefixes <- searchIDATprefixes(
  system.file(package = "sesameData"), recursive=TRUE)
```

segmentBins	<i>Segment bins using DNACopy</i>
-------------	-----------------------------------

Description

Segment bins using DNACopy

Usage

```
segmentBins(bin.signals, bin.coords)
```

Arguments

bin.signals	bin signals (input)
bin.coords	bin coordinates

Value

segment signal data frame

sesameQC	<i>Generate summary numbers that indicative of experiment quality Please provide a raw sigset (before any preprocessing). Usually directly from readIDATpair</i>
----------	--

Description

Generate summary numbers that indicative of experiment quality Please provide a raw sigset (before any preprocessing). Usually directly from readIDATpair

Usage

```
sesameQC(sset)
```

Arguments

sset	a SigSet object
------	-----------------

Value

a sesameQC class object

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
sesameQC(sset)
```

sesamize	<i>"fix" an RGChannelSet (for which IDATs may be unavailable) with Sesame The input is an RGSet and the output is a sesamized minfi::GenomicRatioSet</i>
----------	--

Description

"fix" an RGChannelSet (for which IDATs may be unavailable) with Sesame The input is an RGSet and the output is a sesamized minfi::GenomicRatioSet

Usage

```
sesamize(
  rgSet,
  naFrac = 1,
  BPPARAM = SerialParam(),
  HDF5 = NULL,
  HDF5SEdestination = paste0(tempdir(check = TRUE), "/sesamize_HDF5_scratch"),
  replace = FALSE
)
```

Arguments

rgSet	an RGChannelSet, perhaps with colData of various flavors
naFrac	maximum NA fraction for a probe before it gets dropped (1)
BPPARAM	get parallel with MulticoreParam(n)
HDF5	is the rgSet HDF5-backed? if so, avoid eating RAM (perhaps)
HDF5SEdestination	character(1) path to where the HDF5-backed GenomicRatioSet will be stored
replace	logical(1) passed to saveHDF5SummarizedExperiment

Value

a sesamized GenomicRatioSet

Note

We employ BPRED0 for a second chance if bplapply hits an error.

show, SigSet-method *The display method for SigSet*

Description

The function outputs the number of probes in each category and the first few signal measurements.

Usage

```
## S4 method for signature 'SigSet'  
show(object)
```

Arguments

object displayed object

Value

message of number of probes in each category.

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset  
print(sset)
```

signalMU *report M and U for regular probes*

Description

report M and U for regular probes

Usage

```
signalMU(sset)
```

Arguments

sset a SigSet

Value

a data frame of M and U columns

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset  
signalMU(sset)
```

SigSet-class	<i>SigSet class</i>
--------------	---------------------

Description

This is the main data class for SeSAmE. The class holds different classes of signal intensities.

The function takes a string describing the platform of the data. It can be one of "HM27", "HM450" or "EPIC".

The function takes a string describing the platform of the data. It can be one of "HM27", "HM450" or "EPIC".

Usage

```
## S4 method for signature 'SigSet'
initialize(.Object, platform, ...)
```

```
SigSet(...)
```

Arguments

.Object	target object
platform	"EPIC", "HM450", "HM27" or other strings for custom arrays
...	additional arguments

Value

a SigSet object

a SigSet object

Slots

IG intensity table for type I probes in green channel

IR intensity table for type I probes in red channel

II intensity table for type II probes

oobG out-of-band probes in green channel

oobR out-of-band probes in red channel

ctl all the control probe intensities

pval named numeric vector of p-values

extra extra data, currently, IGG => Type-I green that is inferred to be green IRR => Type-I red that is inferred to be red pvals => list of other pvals

platform "EPIC", "HM450" or "HM27"

Examples

```
## Create an empty EPIC object.
SigSet("EPIC")
SigSet('EPIC')
```

SigSetList *constructor*

Description

constructor

Usage

```
SigSetList(...)
```

Arguments

... the SigSet objects that will be the List elements

Value

a SigSetList

Examples

```
sset1 <- readIDATpair(file.path(system.file(
  'extdata', '', package='sesameData'), '4207113116_A'))

sset2 <- readIDATpair(file.path(system.file(
  'extdata', '', package='sesameData'), '4207113116_B'))

SigSetList(sset1, sset2)
```

SigSetList-class *a List of SigSets with some methods of its own*

Description

a List of SigSets with some methods of its own

SigSetList-methods *SigSetList methods (centralized). Currently scarce... 'show' print a summary of the SigSetList.*

Description

SigSetList methods (centralized). Currently scarce...
 'show' print a summary of the SigSetList.

Usage

```
## S4 method for signature 'SigSetList'
show(object)
```

Arguments

object a SigSetList

Value

Description of SigSetList

Examples

```
SigSetListFromPath(system.file("extdata", "", package = "sesameData"))
```

SigSetListFromIDATs *read IDATs into a SigSetList*

Description

FIXME: switch from 'parallel' to BiocParallel

Usage

```
SigSetListFromIDATs(stubs, parallel = FALSE)
```

Arguments

stubs the IDAT filename stubs
parallel run in parallel? (default FALSE)

Value

a SigSetList

Examples

```
## a SigSetList of length 1
ssets <- SigSetListFromIDATs(file.path(
  system.file("extdata", "", package = "sesameData"), "4207113116_A"))
```

SigSetListFromPath *read an entire directory's worth of IDATs into a SigSetList*

Description

read an entire directory's worth of IDATs into a SigSetList

Usage

```
SigSetListFromPath(path = ".", parallel = FALSE, recursive = TRUE)
```

Arguments

path	the path from which to read IDATs (default ".")
parallel	run in parallel? (default FALSE)
recursive	whether to search recursively

Value

a SigSetList

Examples

```
## Load all IDATs from directory
ssets <- SigSetListFromPath(
  system.file("extdata", "", package = "sesameData"))
```

SigSetsToRGChannelSet *Convert sesame::SigSet to minfi::RGChannelSet*

Description

Convert sesame::SigSet to minfi::RGChannelSet

Usage

```
SigSetsToRGChannelSet(ssets, BPPARAM = SerialParam(), annotation = NA)
```

Arguments

ssets	a list of sesame::SigSet
BPPARAM	get parallel with MulticoreParam(n)
annotation	the minfi annotation string, guessed if not given

Value

a minfi::RGChannelSet

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
rgSet <- SigSetsToRGChannelSet(sset)
```

SigSetToRatioSet	<i>Convert one sesame::SigSet to minfi::RatioSet</i>
------------------	--

Description

Convert one sesame::SigSet to minfi::RatioSet

Usage

```
SigSetToRatioSet(sset, annotation = NA)
```

Arguments

sset	a sesame::SigSet
annotation	minfi annotation string

Value

a minfi::RatioSet

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset
ratioSet <- SigSetToRatioSet(sset)
```

sliceFileSet	<i>Slice a fileSet with samples and probes</i>
--------------	--

Description

Slice a fileSet with samples and probes

Usage

```
sliceFileSet(fset, samples = fset$samples, probes = fset$probes, memmax = 10^5)
```

Arguments

fset	a sesame::fileSet, as obtained via readFileSet
samples	samples to query (default to all samples)
probes	probes to query (default to all probes)
memmax	maximum items to read from file to memory, to protect from accidental memory congestion.

Value

a numeric matrix of length(samples) columns and length(probes) rows

Examples

```
## create two samples
fset <- initFileSet('mybetas2', 'HM27', c('s1','s2'))

## a hypothetical numeric array (can be beta values, intensities etc)
hypothetical <- setNames(runif(fset$n), fset$probes)

## map the numeric to file
mapFileSet(fset, 's1', hypothetical)

## get data
sliceFileSet(fset, 's1', 'cg00000292')
```

SNPcheck	<i>Check sample identity using SNP probes</i>
----------	---

Description

Check sample identity using SNP probes

Usage

```
SNPcheck(betas)
```

Arguments

betas numeric matrix (row: probes, column: samples)

Value

grid object plotting SNP clustering

Examples

```
betas <- sesameDataGet('HM450.10.TCGA.PAAD.normal')
SNPcheck(betas)
```

subsetSignal	<i>Select a subset of probes</i>
--------------	----------------------------------

Description

The function takes a SigSet as input and output another SigSet with probes from the given probe selection.

Usage

```
subsetSignal(sset, probes)
```


Arguments

sset a SigSet
probes target probes

Value

another sset with probes specified

Examples

```
sset <- sesameDataGet('EPIC.1.LNCaP')$sset  
subsetSignal(sset, rownames(slot(sset, 'IR')))
```

topLoci	<i>Top loci in differential methylation</i>
---------	---

Description

This is a convenience function to show top differential methylated segments. The function takes a coefficient table as input and output the same table ordered by the significance of the locus.

Usage

```
topLoci(cf1)
```

Arguments

cf1 coefficient table of one factor from diffMeth

Value

coefficient table ordered by p-value of each locus

Examples

```
data <- sesameDataGet('HM450.76.TCGA.matched')  
cf <- DMR(data$betas, data$sampleInfo, ~type)  
topLoci(cf[[1]])
```

topSegments	<i>Top segments in differential methylation</i>
-------------	---

Description

This is a utility function to show top differential methylated segments. The function takes a coefficient table as input and output the same table ordered by the significance of the segments.

Usage

```
topSegments(cf1)
```

Arguments

cf1 coefficient table of one factor from DMR

Value

coefficient table ordered by adjusted p-value of segments

Examples

```
data <- sesameDataGet('HM450.76.TCGA.matched')
cf <- DMR(data$betas, data$sampleInfo, ~type)
topSegments(cf[[1]])
```

totalIntensities	<i>M+U Intensities for All Probes</i>
------------------	---------------------------------------

Description

The function takes one single SigSet and computes total intensity of all the in-band measurements by summing methylated and unmethylated alleles. This function outputs a single numeric for the mean.

Usage

```
totalIntensities(sset)
```

Arguments

sset a SigSet

Value

a vector of M+U signal for each probe

Examples

```
sset <- makeExampleSeSAMeDataSet()
intensities <- totalIntensities(sset)
```

totalIntensityZscore *Calculate intensity Z-score*

Description

This function compute intensity Z-score with respect to the mean. Log10 transformation is done first. Probes of each design type are grouped before Z-scores are computed.

Usage

```
totalIntensityZscore(sset)
```

Arguments

sset a SigSet

Value

a vector of Z-score for each probe

Examples

```
sset <- sesameDataGet('HM450.1.TCGA.PAAD')$sset  
head(totalIntensityZscore(sset))
```

twoCompsEst2 *Estimate the fraction of the 2nd component in a 2-component mixture*

Description

Estimate the fraction of the 2nd component in a 2-component mixture

Usage

```
twoCompsEst2(  
  pop1,  
  pop2,  
  target,  
  use.ave = TRUE,  
  diff_1m2u = NULL,  
  diff_1u2m = NULL  
)
```

Arguments

pop1	Reference methylation level matrix for population 1
pop2	Reference methylation level matrix for population 2
target	Target methylation level matrix to be analyzed
use.ave	use population average in selecting differentially methylated probes
diff_1m2u	A vector of differentially methylated probes (methylated in population 1 but unmethylated in population 2)
diff_1u2m	A vector of differentially methylated probes (unmethylated in population 1 but methylated in population 2)

Value

Estimate of the 2nd component in the 2-component mixture

visualizeGene	<i>Visualize Gene</i>
---------------	-----------------------

Description

Visualize the beta value in heatmaps for a given gene. The function takes a gene name which is taken from the UCSC refGene. It searches all the transcripts for the given gene and optionally extend the span by certain number of base pairs. The function also takes a beta value matrix with sample names on the columns and probe names on the rows. The function can also work on different genome builds (default to hg38, can be hg19).

Usage

```
visualizeGene(
  geneName,
  betas,
  platform = c("EPIC", "HM450"),
  upstream = 2000,
  dwnstream = 2000,
  refversion = c("hg38", "hg19"),
  ...
)
```

Arguments

geneName	gene name
betas	beta value matrix (row: probes, column: samples)
platform	HM450 or EPIC (default)
upstream	distance to extend upstream
dwnstream	distance to extend downstream
refversion	hg19 or hg38 (default)
...	additional options, see visualizeRegion

Value

None

Examples

```
betas <- sesameDataGet('HM450.76.TCGA.matched')$betas
visualizeGene('ADA', betas, 'HM450')
```

visualizeProbes

Visualize Region that Contains the Specified Probes

Description

Visualize the beta value in heatmaps for the genomic region containing specified probes. The function works only if specified probes can be spanned by a single genomic region. The region can cover more probes than specified. Hence the plotting heatmap may encompass more probes. The function takes as input a string vector of probe IDs (cg/ch/rs-numbers). if draw is FALSE, the function returns the subset beta value matrix otherwise it returns the grid graphics object.

Usage

```
visualizeProbes(
  probeNames,
  betas,
  platform = c("EPIC", "HM450"),
  refversion = c("hg38", "hg19"),
  upstream = 1000,
  dstream = 1000,
  ...
)
```

Arguments

probeNames	probe names
betas	beta value matrix (row: probes, column: samples)
platform	HM450 or EPIC (default)
refversion	hg19 or hg38 (default)
upstream	distance to extend upstream
dstream	distance to extend downstream
...	additional options, see visualizeRegion

Value

None

Examples

```
betas <- sesameDataGet('HM450.76.TCGA.matched')$betas
visualizeProbes(c('cg22316575', 'cg16084772', 'cg20622019'), betas, 'HM450')
```

visualizeRegion	<i>Visualize Region</i>
-----------------	-------------------------

Description

The function takes a genomic coordinate (chromosome, start and end) and a beta value matrix (probes on the row and samples on the column). It plots the beta values as a heatmap for all probes falling into the genomic region. If 'draw=TRUE' the function returns the plotted grid graphics object. Otherwise, the selected beta value matrix is returned. 'cluster.samples=TRUE/FALSE' controls whether hierarchical clustering is applied to the subset beta value matrix.

Usage

```
visualizeRegion(
  chr,
  plt.beg,
  plt.end,
  betas,
  platform = c("EPIC", "HM450"),
  refversion = c("hg38", "hg19"),
  sample.name.fontsize = 10,
  heat.height = NULL,
  draw = TRUE,
  show.sampleNames = TRUE,
  show.samples.n = NULL,
  show.probeNames = TRUE,
  cluster.samples = FALSE,
  nprobes.max = 1000,
  na.rm = FALSE,
  dmin = 0,
  dmax = 1
)
```

Arguments

chr	chromosome
plt.beg	begin of the region
plt.end	end of the region
betas	beta value matrix (row: probes, column: samples)
platform	EPIC or HM450
refversion	hg38 or hg19
sample.name.fontsize	sample name font size
heat.height	heatmap height (auto inferred based on rows)
draw	draw figure or return betas
show.sampleNames	whether to show sample names
show.samples.n	number of samples to show (default: all)

show.probeNames	whether to show probe names
cluster.samples	whether to cluster samples
nprobes.max	maximum number of probes to plot
na.rm	remove probes with all NA.
dmin	data min
dmax	data max

Value

graphics or a matrix containing the captured beta values

Examples

```
betas <- sesameDataGet('HM450.76.TCGA.matched')$betas
visualizeRegion('chr20', 44648623, 44652152, betas, 'HM450')
```

visualizeSegments	<i>Visualize segments</i>
-------------------	---------------------------

Description

The function takes a CNSegment object obtained from cnSegmentation and plot the bin signals and segments (as horizontal lines).

Usage

```
visualizeSegments(seg, to.plot = NULL)
```

Arguments

seg	a CNSegment object
to.plot	chromosome to plot (by default plot all chromosomes)

Details

require ggplot2, scales

Value

plot graphics

Examples

```
seg <- sesameDataGet('EPIC.1.LNCaP')$seg
visualizeSegments(seg)
```

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