

Package ‘sesame’

September 30, 2022

Type Package

Title Sensible Step-wise Analysis of DNA METHylation BeadChips

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

Version 1.15.10

Depends R (>= 4.1), sesameData

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RoxygenNote 7.2.0

Imports graphics, BiocParallel, utils, methods, stringr, readr, tibble, illuminaio, MASS, wheatmap (>= 0.2.0), GenomicRanges, IRanges, grid, preprocessCore, S4Vectors, ggplot2, BiocFileCache, GenomeInfoDb, stats, SummarizedExperiment, dplyr, reshape2

Suggests scales, knitr, DNACopy, e1071, randomForest, RPMM, rmarkdown, testthat, tidyr, BiocStyle, ggrepel, grDevices, pals

Encoding UTF-8

VignetteBuilder knitr

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

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

biocViews DNAMethylation, MethylationArray, Preprocessing, QualityControl

Collate 'sex.R' 'species.R' 'QC.R' 'GEO.R' 'SigDFMethods.R' 'sesame.R' 'age.R' 'background.R' 'cell_composition.R' 'channel_inference.R' 'cnv.R' 'ethnicity.R' 'deidentify.R' 'detection.R' 'dm.R' 'dye_bias.R' 'feature_selection.R' 'fileSet.R' 'mask.R' 'sesameAnno.R' 'open.R' 'strain.R' 'tissue.R' 'track.R' 'match_design.R' 'utils.R' 'vcf.R' 'visualize.R' 'visualizeHelper.R' 'zzz.R' 'KYCG.R' 'KYCG_plot.R' 'palgen.R'

git_url <https://git.bioconductor.org/packages/sesame>

git_branch master

git_last_commit abddfbf

git_last_commit_date 2022-09-22

Date/Publication 2022-09-30

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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

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

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

addMask *Add probes to mask*

Description

This function essentially merge existing probe masking with new probes to mask

Usage

```
addMask(sdf, probes)
```

Arguments

sdf a SigDF
probes a vector of probe IDs or a logical vector with TRUE representing masked probes

Value

a SigDF with added mask

Examples

```
sdf <- sesameDataGet('EPIC.1.SigDF')  
sum(sdf$mask)  
sum(addMask(sdf, c("cg14057072", "cg22344912"))$mask)
```

aggregateTestEnrichments
Aggregate test enrichment results

Description

Aggregate test enrichment results

Usage

```
aggregateTestEnrichments(result_list, column = "estimate", return_df = FALSE)
```

Arguments

result_list a list of results from testEnrichment
column the column name to aggregate (Default: estimate)
return_df whether to return a merged data frame

Value

a matrix for all results

Examples

```
## pick some big TFBS-overlapping CpG groups
cg_lists <- KYCG_getDBs("MM285.TFBS")
queries <- cg_lists[(sapply(cg_lists, length) > 40000)]
result_list <- lapply(queries, testEnrichment, "MM285.chromHMM")
mtx <- aggregateTestEnrichments(result_list)
```

assemble_plots	<i>assemble_plots</i>
----------------	-----------------------

Description

assemble plots

Usage

```
assemble_plots(
  betas,
  txns,
  probes,
  plt.txns,
  plt.mapLines,
  plt.cytoband,
  heat.height = NULL,
  show.probeNames = TRUE,
  show.samples.n = NULL,
  show.sampleNames = TRUE,
  sample.name.fontsize = 10,
  dmin = 0,
  dmax = 1
)
```

Arguments

betas	beta value
txns	transcripts GRanges
probes	probe GRanges
plt.txns	transcripts plot objects
plt.mapLines	map line plot objects
plt.cytoband	cytoband plot objects

heat.height heatmap height (auto inferred based on rows)
show.probeNames whether to show probe names
show.samples.n number of samples to show (default: all)
show.sampleNames whether to show sample names
sample.name.fontsize sample name font size
dmin data min
dmax data max

Value

a grid object

BetaValueToMValue *Convert beta-value to M-value*

Description

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

Usage

BetaValueToMValue(b)

Arguments

b 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(sdf, verbose = FALSE)
```

Arguments

sdf	a SigDF
verbose	print more messages

Value

GCT score (the higher, the more incomplete conversion)

Examples

```
sesameDataCache() # if not done yet  
sdf <- sesameDataGet('EPIC.1.SigDF')  
bisConversionControl(sdf)
```

calcEffectSize	<i>Compute effect size for different variables from prediction matrix</i>
----------------	---

Description

The effect size is defined by the maximum variation of a variable with all the other variables controlled constant.

Usage

```
calcEffectSize(pred)
```

Arguments

pred	predictions
------	-------------

Value

a data.frame of effect sizes. Columns are different variables. Rows are different probes.

Examples

```
data <- sesameDataGet('HM450.76.TCGA.matched')
res <- DMLpredict(data$betas[1:10,], ~type, meta=data$sampleInfo)
head(calcEffectSize(res))
```

checkLevels	<i>filter data matrix by factor completeness only works for discrete factors</i>
-------------	--

Description

filter data matrix by factor completeness only works for discrete factors

Usage

```
checkLevels(betas, fc)
```

Arguments

betas	matrix data
fc	factors, or characters

Value

a boolean vector whether there is non-NA value for each tested group for each probe

Examples

```
se0 <- sesameDataGet("MM285.10.SE.tissue")[1:100,]
se_ok <- checkLevels(SummarizedExperiment::assay(se0),
  SummarizedExperiment::colData(se0)$tissue)
sum(se_ok) # number of good probes
se1 <- se0[se_ok,]

sesameDataGet_resetEnv()
```

chipAddressToSignal *Lookup address in one sample*

Description

Lookup address and transform address to probe

Usage

```
chipAddressToSignal(dm, mft)
```

Arguments

dm	data frame in chip address, 2 columns: cy3/Grn and cy5/Red
mft	a data frame with columns Probe_ID, M, U and col

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 SigDF, 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 SigDF for the target sample and a set of normal SigDF 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(sdf, sdfs.normal = NULL, genome = NULL, verbose = FALSE)
```

Arguments

sdf	SigDF
sdfs.normal	a list of SigDFs for normalization, if not given, use the stored normal data from sesameData. However, we do recommend using a matched copy number normal dataset for normalization.
genome	hg38, mm10, ..., will infer if not given. For additional mapping, download the GRanges object from http://zwdzwd.github.io/InfiniumAnnotation and provide the following argument ..., genome = sesameAnno_buildManifestGRanges("downloaded_file"),... to this function.
verbose	print more messages

Value

an object of CNSegment

Examples

```
sesameDataCache()  
  
## sdf <- sesameDataGet('EPIC.1.SigDF')  
## sdfs.normal <- sesameDataGet('EPIC.5.SigDF.normal')  
## seg <- cnSegmentation(sdf, sdfs.normal)
```

```
compareDatabaseSetOverlap
```

calculates the pairwise overlap between given list of database sets using a distance metric.

Description

calculates the pairwise overlap between given list of database sets using a distance metric.

Usage

```
compareDatabaseSetOverlap(databases = NA, metric = "Jaccard")
```

Arguments

databases	List of vectors corresponding to the database sets of interest with associated meta data as an attribute to each element. Optional. (Default: NA)
metric	String representing the similarity metric to use. Optional. (Default: "Jaccard").

Value

An upper triangular matrix containing a metric (Jaccard) comparing the pairwise distances between database sets.

```
compareMouseStrainReference
```

Compare Strain SNPs with a reference panel

Description

Compare Strain SNPs with a reference panel

Usage

```
compareMouseStrainReference(
  betas = NULL,
  show_sample_names = FALSE,
  query_width = NULL
)
```

Arguments

betas	beta value vector or matrix (for multiple samples)
show_sample_names	whether to show sample name
query_width	optional argument for adjusting query width

Value

grid object that contrast the target sample with pre-built mouse strain reference

Examples

```
sesameDataCache() # if not done yet
compareMouseStrainReference()
```

compareMouseTissueReference

Compare mouse array data with mouse tissue references

Description

Compare mouse array data with mouse tissue references

Usage

```
compareMouseTissueReference(
  betas = NULL,
  ref = NULL,
  color = "blueYellow",
  query_width = 0.3
)
```

Arguments

betas	matrix of betas for the target sample This argument is optional. If not given, only the reference will be shown.
ref	the reference beta values in SummarizedExperiment. This argument is optional. If not given, the reference will be downloaded from the sesameData package.
color	either blueYellow or fullJet
query_width	the width of the query beta value matrix

Value

grid object that contrast the target sample with pre-built mouse tissue reference

Examples

```
cat("Deprecated, see compareReference")
```

compareReference	<i>Compare array data with references (e.g., tissue, cell types)</i>
------------------	--

Description

Compare array data with references (e.g., tissue, cell types)

Usage

```
compareReference(  
  ref,  
  betas = NULL,  
  stop.points = NULL,  
  query_width = 0.3,  
  show_sample_names = FALSE  
)
```

Arguments

ref	the reference beta values in SummarizedExperiment. One can download them from the sesameData package. See examples.
betas	matrix of betas for the target sample This argument is optional. If not given, only the reference will be shown.
stop.points	stop points for the color palette. Default to blue, yellow.
query_width	the width of the query beta value matrix
show_sample_names	whether to show sample names (default: FALSE)

Value

grid object that contrast the target sample with references.

Examples

```
sesameDataCache() # if not done yet  
compareReference(sesameDataGet("MM285.tissueSignature"))  
sesameDataGet_resetEnv()
```

controls *get the controls attributes*

Description

get the controls attributes

Usage

```
controls(sdf, verbose = FALSE)
```

Arguments

sdf a SigDF
verbose print more messages

Value

the controls data frame

Examples

```
sesameDataCache() # if not done yet  
sdf <- sesameDataGet('EPIC.1.SigDF')  
head(controls(sdf))
```

createDBNetwork *createGeneNetwork creates database network using the Jaccard index.*

Description

createGeneNetwork creates database network using the Jaccard index.

Usage

```
createDBNetwork(databases)
```

Arguments

databases Vector of probes corresponding to a single database set of interest.

Value

ggplot lollipop plot

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", genome = "hg38")
```

Arguments

betas	a named numeric vector
output	output file name
platform	HM450, EPIC etc.
genome	hg38, mm10, ..., will infer if not given. For additional mapping, download the GRanges object from http://zwdzwd.github.io/InfiniumAnnotation and provide the following argument ..., genome = sesameAnno_buildManifestGRanges("downloaded_file"),... to this function.

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
```

dataFrame2sesameQC	<i>Convert data frame to sesameQC object</i>
--------------------	--

Description

The function convert a data frame back to a list of sesameQC objects

Usage

```
dataFrame2sesameQC(df)
```

Arguments

df a publicQC data frame

Value

a list sesameQC objects

Examples

```
df <- sesameDataGet("MM285.publicQC")
qcs <- dataFrame2sesameQC(df[1:2,])
```

dbStats	<i>dbStats builds dataset for a given betas matrix composed of engineered features from the given database sets</i>
---------	---

Description

dbStats builds dataset for a given betas matrix composed of engineered features from the given database sets

Usage

```
dbStats(betas, databases, fun = mean, na.rm = TRUE, n_min = NULL, f_min = 0.1)
```

Arguments

betas matrix of beta values where probes are on the rows and samples are on the columns

databases List of vectors corresponding to probe locations for which the features will be extracted

fun aggregation function, default to mean

na.rm whether to remove NA

n_min min number of non-NA for aggregation function to apply, overrides f_min

f_min min fraction of non-NA for aggregation function to apply

Value

matrix with samples on the rows and database set on the columns

Examples

```
library(SummarizedExperiment)
se <- sesameDataGet('MM285.467.SE.tissue20Kprobes')
head(dbStats(assay(se), "MM285.probeType")[,1:3])
sesameDataGet_resetEnv()
```

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)
```

detectionIB	<i>Mask detection by intermediate beta values</i>
-------------	---

Description

Mask detection by intermediate beta values

Usage

```
detectionIB(
  sdf,
  return.pval = FALSE,
  pval.threshold = 0.05,
  capMU = 3000,
  window = 100
)
```

Arguments

sdf	a SigDF
return.pval	whether to return p-values, instead of a SigDF
pval.threshold	minimum p-value to mask
capMU	the maximum M+U to search for intermediate betas
window	window size for smoothing and beta fraction calc.

Value

a SigDF with mask added

Examples

```
sdf <- sesameDataGet("EPIC.1.SigDF")
sum(sdf$mask)
sum(detectionIB(sdf)$mask)
```

detectionPnegEcdf	<i>Detection P-value based on ECDF of negative control</i>
-------------------	--

Description

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

Usage

```
detectionPnegEcdf(sdf, return.pval = FALSE, pval.threshold = 0.05)
```

Arguments

sdf	a SigDF
return.pval	whether to return p-values, instead of a masked SigDF
pval.threshold	minimum p-value to mask

Value

a SigDF, or a p-value vector if return.pval is TRUE

Examples

```
sdf <- sesameDataGet("EPIC.1.SigDF")
sum(sdf$mask)
sum(detectionPnegEcdf(sdf)$mask)
```

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'))
sesameDataGet_resetEnv()
```

dmContrasts	<i>List all contrasts of a DMLSummary</i>
-------------	---

Description

List all contrasts of a DMLSummary

Usage

```
dmContrasts(smry)
```

Arguments

smry a DMLSummary object

Value

a character vector of contrasts

Examples

```
data <- sesameDataGet('HM450.76.TCGA.matched')
smry <- DML(data$betas[1:10,], ~type, meta=data$sampleInfo)
dmContrasts(smry)

sesameDataGet_resetEnv()
```

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 (meta) and formula for testing. The function outputs a list of coefficient tables for each factor tested.

Usage

```
DML(betas, fm, meta = NULL, BPPARAM = SerialParam())
```

Arguments

betas	beta values, matrix or SummarizedExperiment rows are probes and columns are samples.
fm	formula
meta	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. When the betas argument is a SummarizedExperiment object, this is ignored. colData(betas) will be used instead.
BPPARAM	number of cores for parallel processing, default to SerialParam() Use Multi-coreParam(mc.cores) for parallel processing. For Windows, try DoparParam or SnowParam.

Value

a list of test summaries, summary.lm objects

Examples

```
sesameDataCache() # in case not done yet
data <- sesameDataGet('HM450.76.TCGA.matched')
smry <- DML(data$betas[1:1000,], ~type, meta=data$sampleInfo)

sesameDataGet_resetEnv()
```

DMLpredict

Predict new data from DML

Description

This function is also important for investigating factor interactions.

Usage

```
DMLpredict(betas, fm, pred = NULL, meta = NULL, BPPARAM = SerialParam())
```

Arguments

betas	beta values, matrix or SummarizedExperiment rows are probes and columns are samples.
fm	formula
pred	new data for prediction, useful for studying effect size. This argument is a data.frame to specify new data. If the argument is NULL, all combinations of all contrasts will be used as input. It might not work if there is a continuous variable input. One may need to explicitly provide the input in a data frame.

meta	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. When the betas argument is a SummarizedExperiment object, this is ignored. colData(betas) will be used instead.
BPPARAM	number of cores for parallel processing, default to SerialParam() Use Multi-coreParam(mc.cores) for parallel processing. For Windows, try DoparParam or SnowParam.

Value

a SummarizedExperiment of predictions. The colData describes the input of the prediction.

Examples

```
data <- sesameDataGet('HM450.76.TCGA.matched')

## use all contrasts as new input
res <- DMLpredict(data$betas[1:10,], ~type, meta=data$sampleInfo)

## specify new input
res <- DMLpredict(data$betas[1:10,], ~type, meta=data$sampleInfo,
  pred = data.frame(type=c("Normal", "Tumour")))

## note that the prediction needs to be a factor of the same
## level structure as the original training data.
pred = data.frame(type=factor(c("Normal"), levels=c("Normal", "Tumour")))
res <- DMLpredict(data$betas[1:10,], ~type,
  meta=data$sampleInfo, pred = pred)
```

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,
  smry,
  contrast,
  platform = NULL,
  genome = NULL,
```



```

    dist.cutoff = NULL,
    seg.per.locus = 0.5
  )

```

Arguments

betas	beta values for distance calculation
smry	DML
contrast	the pair-wise comparison or contrast check colnames(attr(smry, "model.matrix")) if uncertain
platform	EPIC, HM450, MM285, ...
genome	hg38, mm10, ..., will infer if not given. For additional mapping, download the GRanges object from http://zwdzwd.github.io/InfiniumAnnotation and provide the following argument ..., genome = sesameAnno_buildManifestGRanges("downloaded_file"),... to this function.
dist.cutoff	cutoff of beta value differences for two neighboring CGs to be considered the same DMR (by default it's determined using the quantile function on seg.per.locus)
seg.per.locus	number of segments per locus higher value leads to more segments

Value

coefficient table with segment ID and segment P-value each row is a locus, multiple loci may share a segment ID if they are merged to the same segment. Records are ordered by Seg_Est.

Examples

```

sesameDataCache() # in case not done yet

data <- sesameDataGet('HM450.76.TCGA.matched')
smry <- DML(data$betas[1:1000,], ~type, meta=data$sampleInfo)
colnames(attr(smry, "model.matrix")) # pick a contrast from here
## showing on a small set of 100 CGs
merged_segs <- DMR(data$betas[1:100,], smry, "typeTumour")

sesameDataGet_resetEnv()

```

dyeBiasCorr

Correct dye bias in by linear scaling.

Description

The function takes a SigDF 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 SigDF with dye bias corrected.

Usage

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

Arguments

```
sdf          a SigDF
ref          reference signal level
```

Value

a normalized SigDF

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
sdf.db <- dyeBiasCorr(sdf)
```

dyeBiasCorrMostBalanced

Correct dye bias using most balanced sample as the reference

Description

The function chose the reference signal level from a list of SigDF. 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 SigDFs with dye bias corrected.

Usage

```
dyeBiasCorrMostBalanced(sdfs)
```

Arguments

```
sdfs          a list of normalized SigDFs
```

Value

a list of normalized SigDFs

Examples

```
sesameDataCache() # if not done yet
sdfs <- sesameDataGet('HM450.10.SigDF')[1:2]
sdfs.db <- dyeBiasCorrMostBalanced(sdfs)
```

dyeBiasL	<i>Correct dye bias in by linear scaling.</i>
----------	---

Description

The function takes a SigDF 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 SigDF with dye bias corrected.

Usage

```
dyeBiasL(sdf, ref = NULL)
```

Arguments

sdf	a SigDF
ref	reference signal level

Value

a normalized SigDF

Examples

```
sesameDataCache() # if not done yet  
sdf <- sesameDataGet('EPIC.1.SigDF')  
sdf.db <- dyeBiasL(sdf)
```

dyeBiasNL	<i>Dye bias correction by matching green and red to mid point</i>
-----------	---

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 SigDF and returns a SigDF with dye bias corrected.

Usage

```
dyeBiasNL(sdf, mask = TRUE, verbose = FALSE)
```

```
dyeBiasCorrTypeINorm(sdf, mask = TRUE, verbose = FALSE)
```

Arguments

sdf	a SigDF
mask	include masked probes in Infinium-I probes. No big difference is noted in practice. More probes are generally better.
verbose	print more messages

Value

a SigDF after dye bias correction.

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
sdf.db <- dyeBiasNL(sdf)
sdf <- sesameDataGet('EPIC.1.SigDF')
sdf <- dyeBiasCorrTypeINorm(sdf)
```

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	<i>Estimate leukocyte fraction using a two-component model</i>
-------------------	--

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)  
sesameDataGet_resetEnv()
```

`formatVCF`*Convert SNP from Infinium array to VCF file*

Description

Convert SNP from Infinium array to VCF file

Usage

```
formatVCF(  
  sdf,  
  vcf = NULL,  
  genome = "hg19",  
  annoS = NULL,  
  annoI = NULL,  
  verbose = FALSE  
)
```

Arguments

<code>sdf</code>	SigDF
<code>vcf</code>	output VCF file path, if NULL output to console
<code>genome</code>	genome
<code>annoS</code>	SNP variant annotation, download if not given
<code>annoI</code>	Infinium-I variant annotation, download if not given hg19 and hg38 in human
<code>verbose</code>	print more messages

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

```
sesameDataCacheAll() # if not done yet  
sdf <- sesameDataGet('EPIC.1.SigDF')  
  
## output to console  
head(formatVCF(sdf))
```

getAFs	<i>Get allele frequency</i>
--------	-----------------------------

Description

Get allele frequency

Usage

```
getAFs(sdf, ...)
```

Arguments

sdf	SigDF
...	additional options to getBetas

Value

allele frequency

Examples

```
sesameDataCache() # if not done yet  
sdf <- sesameDataGet('EPIC.1.SigDF')  
af <- getAFs(sdf)
```

getAFTTypeIbySumAlleles	<i>Get allele frequency treating type I by summing alleles</i>
-------------------------	--

Description

Takes a SigDF 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 SigDF, then a numeric(0) is returned.

Usage

```
getAFTTypeIbySumAlleles(sdf, known.ccs.only = TRUE)
```

Arguments

sdf	SigDF
known.ccs.only	consider only known CCS probes

Value

beta values

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
af <- getAFTYPEIbySumAlleles(sdf)
```

getBetas

Get beta Values

Description

sum.typeI is used for rescuing beta values on Color-Channel-Switching CCS probes. The function takes a SigDF 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(sdf, mask = TRUE, sum.TypeI = FALSE, collapseToPfx = FALSE)
```

Arguments

sdf	SigDF
mask	whether to use mask
sum.TypeI	whether to sum type I channels
collapseToPfx	remove replicate to prefix (e.g., cg number) and remove the suffix

Value

a numeric vector, beta values

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
betas <- getBetas(sdf)
```

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

Description

requires GenomicRanges, IRanges

Usage

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

Arguments

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

Value

bin.coords

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')
sesameDataGet_resetEnv()
```

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

Description

The function takes a SigDF 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(sdf, verbose = FALSE)
```

Arguments

sdf	a SigDF
verbose	print more messages

Value

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

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
getSexInfo(sdf)
```

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(sdf, verbose = FALSE)
```

Arguments

sdf a SigDF
 verbose print more messages

Details

s better be background subtracted and dyebias corrected for best accuracy
 Please note: the betas should come from SigDF *without* channel inference.

Value

string of ethnicity

Examples

```
sdf <- sesameDataGet('EPIC.1.SigDF')
inferEthnicity(sdf)
```

`inferInfiniumIChannel` *Infer and reset color channel for Type-I probes instead of using what is specified in manifest. The results are stored to `sdf@extra$IGG` and `sdf@extra$IRR` slot.*

Description

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

Usage

```
inferInfiniumIChannel(
  sdf,
  switch_failed = FALSE,
  verbose = FALSE,
  summary = FALSE
)
```

Arguments

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

Value

a SigDF, or numerics if summary == TRUE

Examples

```
sdf <- sesameDataGet('EPIC.1.SigDF')
inferInfiniumIChannel(sdf)
```

inferSex

Infer Sex

Description

Infer Sex

Usage

```
inferSex(x, platform = NULL, verbose = FALSE)
```

Arguments

x	either a raw SigDF or a beta value vector named by probe ID SigDF is preferred over beta values.
platform	Only MM285, EPIC and HM450 are supported.
verbose	print more messages

Value

'F' or 'M' We established our sex calling based on the CpGs hypermethylated in inactive X (XiH), CpGs hypomethylated in inactive X (XiL) and signal intensity ratio of Y-chromosome over autosomes. Currently human inference uses a random forest and mouse inference uses a support vector machine.

The function checks the sample quality. If the sample is of poor quality the inference return NA.

Note many factors such as Dnmt genotype, XXY male (Klinefelter's), 45,X female (Turner's) can confuse the model sometimes. This function works on a single sample.

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
inferSex(sdf)
```

inferSexKaryotypes *Infer Sex Karyotype*

Description

The function takes a SigDF 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(sdf)
```

Arguments

sdf a SigDF

Value

Karyotype string, with XCI

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
inferSexKaryotypes(sdf)
```

inferSpecies *Infer Species*

Description

We infer species based on probes pvalues and alignment score. AUC was calculated for each specie, y_true is 1 or 0 for pval < threshold.pos or pval > threshold.neg, respectively,

Usage

```
inferSpecies(
  sdf,
  topN = 1000,
  threshold.pos = 0.01,
  threshold.neg = 0.1,
  return.auc = FALSE,
  return.species = FALSE,
  verbose = FALSE
)
```

Arguments

sdf	a SigDF
topN	Top n positive and negative probes used to infer species. increase this number can sometimes improve accuracy (DEFAULT: 1000)
threshold.pos	pvalue < threshold.pos are considered positive (default: 0.01).
threshold.neg	pvalue > threshold.neg are considered negative (default: 0.2).
return.auc	return AUC calculated, override return.species
return.species	return a string to represent species
verbose	print more messages

Value

a SigDF

Examples

```
sdf <- sesameDataGet("MM285.1.SigDF")
sdf <- inferSpecies(sdf)

## all available species
all_species <- names(sesameDataGet(sprintf(
  "%s.addressSpecies", sdfPlatform(sdf))))$species)
```

inferStrain	<i>Infer strain information for mouse array</i>
-------------	---

Description

Infer strain information for mouse array

Usage

```
inferStrain(
  sdf,
  return.strain = FALSE,
  return.probability = FALSE,
  return.pval = FALSE,
  min_frac_dt = 0.2,
  verbose = FALSE
)
```

Arguments

sdf	SigDF
return.strain	return strain name
return.probability	return probability vector for all strains
return.pval	return p-value
min_frac_dt	minimum fraction of detected signal (DEFAULT: 0.2) otherwise, we give up strain inference and return NA.
verbose	print more messages

Value

a list of best guess, p-value of the best guess and the probabilities of all strains

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('MM285.1.SigDF')
inferStrain(sdf, return.strain = TRUE)
sdf.strain <- inferStrain(sdf)
```

inferTissue	<i>inferTissue infers the tissue of a single sample (as identified through the branchIDs in the row data of the reference) by reporting independent composition through cell type deconvolution.</i>
-------------	--

Description

inferTissue infers the tissue of a single sample (as identified through the branchIDs in the row data of the reference) by reporting independent composition through cell type deconvolution.

Usage

```
inferTissue(
  betas,
  reference = NULL,
  platform = NULL,
  abs_delta_beta_min = 0.3,
  auc_min = 0.99,
  coverage_min = 0.8,
  topN = 15
)
```

Arguments

betas	Named vector with probes and their corresponding beta value measurement
reference	Summarized Experiment with either hypomethylated or hypermethylated probe selection (row data), sample selection (column data), meta data, and the betas (assay)
platform	String representing the array type of the betas and reference
abs_delta_beta_min	Numerical value indicating the absolute minimum required delta beta for the probe selection criteria
auc_min	Numeric value corresponding to the minimum AUC value required for a probe to be considered
coverage_min	Numeric value corresponding to the minimum coverage requirement for a probe to be considered. Coverage is defined here as the proportion of samples without an NA value at a given probe.
topN	number of probes to at most use for each branch

Value

inferred tissue as a string

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet("MM285.1.SigDF")
inferTissue(getBetas(dyeBiasNL(noob(sdf))))

sesameDataGet_resetEnv()
```

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 sdfPlatform(sdf)
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'))
```

KYCG_annoProbes	<i>Annotate Probe IDs using KYCG databases</i>
-----------------	--

Description

see sesameData_annoProbes if you'd like to annotate by genomic coordinates (in GRanges)

Usage

```
KYCG_annoProbes(  
  query,  
  databases,  
  db_names = NULL,  
  platform = NULL,  
  sep = ",",  
  indicator = FALSE,  
  silent = FALSE  
)
```

Arguments

query	probe IDs in a character vector
databases	character or actual database (i.e. list of probe IDs)
db_names	specific database (default to all databases)
platform	EPIC, MM285 etc. will infer from probe IDs if not given
sep	delimiter used in paste
indicator	return the indicator matrix instead of a concatenated annotation (in the case of have multiple annotations)
silent	suppress message

Value

named annotation vector, or indicator matrix

Examples

```
query <- names(sesameData_getManifestGRanges("MM285"))  
anno <- KYCG_annoProbes(query, "designGroup", silent = TRUE)
```

KYCG_buildGeneDBs *build gene-probe association database*

Description

build gene-probe association database

Usage

```
KYCG_buildGeneDBs(  
  query = NULL,  
  platform = NULL,  
  genome = NULL,  
  max_distance = 10000,  
  silent = FALSE  
)
```

Arguments

query	the query probe list. If NULL, use all the probes on the platform
platform	HM450, EPIC, MM285, Mammal40, will infer from query if not given
genome	hg38, mm10, ..., will infer if not given. For additional mapping, download the GRanges object from http://zwdzwd.github.io/InfiniumAnnotation and provide the following argument ..., genome = sesameAnno_buildManifestGRanges("downloaded_file"),... to this function.
max_distance	probe-gene distance for association
silent	suppress messages

Value

gene databases

Examples

```
query <- c("cg04707299", "cg13380562", "cg00480749")  
dbs <- KYCG_buildGeneDBs(query, platform = "EPIC")  
testEnrichment(query, dbs, platform = "EPIC")
```

`KYCG_getDBs`*Get databases by full or partial names of the database group(s)*

Description

Get databases by full or partial names of the database group(s)

Usage

```
KYCG_getDBs(  
  group_nms,  
  db_names = NULL,  
  platform = NULL,  
  summary = FALSE,  
  allow_multi = FALSE,  
  type = NULL,  
  silent = FALSE  
)
```

Arguments

<code>group_nms</code>	database group names
<code>db_names</code>	name of the database, fetch only the given databases
<code>platform</code>	EPIC, HM450, MM285, ... If given, will restrict to that platform.
<code>summary</code>	return a summary of database instead of db itself
<code>allow_multi</code>	allow multiple groups to be returned for
<code>type</code>	numerical, categorical, default: all
<code>silent</code>	no messages each query.

Value

a list of databases, return NULL if no database is found

Examples

```
dbs <- KYCG_getDBs("MM285.chromHMM")  
dbs <- KYCG_getDBs(c("MM285.chromHMM", "MM285.probeType"))
```

KYCG_listDBGroups *List database group names*

Description

List database group names

Usage

```
KYCG_listDBGroups(filter = NULL, type = NULL)
```

Arguments

filter	keywords for filtering
type	categorical, numerical (default: all)

Value

a list of db group names

Examples

```
head(KYCG_listDBGroups("chromHMM"))
```

KYCG_loadDBs *Load database groups*

Description

Load database groups

Usage

```
KYCG_loadDBs(in_paths)
```

Arguments

in_paths	folder that contains all databases
----------	------------------------------------

Value

a list of db group names

Examples

```
## download regulatory annotations from
## http://zwdzwd.github.io/InfiniumAnnotation
## unzip the file
if (FALSE) {
  dbs <- KYCG_loadDBs(path_to_unzipped_folder)
}
```

KYCG_plotBar

Bar plot to show most enriched CG groups from testEnrichment

Description

The input data frame should have an "estimate" and a "FDR" columns.

Usage

```
KYCG_plotBar(df, n_min = 10, n_max = 30, max_fdr = 0.05, min_cap = -5)
```

Arguments

df	KYCG result data frame
n_min	minimum number of databases to report
n_max	maximum number of databases to report
max_fdr	maximum FDR
min_cap	the minimum log ₂ (OR), value below this are capped

Details

Top CG groups are determined by estimate (descending order).

Value

grid plot object

Examples

```
KYCG_plotBar(data.frame(
  estimate=runif(10,0,10), FDR=runif(10,0,1), nD=10,
  overlap=as.integer(runif(10,0,30)), group="g", dbname=seq_len(10)))
```

`KYCG_plotDot`*Dot plot to show most enriched CG groups from testEnrichment*

Description

The input data frame should have an "estimate" and a "FDR" columns.

Usage

```
KYCG_plotDot(df, n_min = 10, n_max = 30, max_fdr = 0.05)
```

Arguments

<code>df</code>	KYCG result data frame
<code>n_min</code>	minimum number of databases to report
<code>n_max</code>	maximum number of databases to report
<code>max_fdr</code>	maximum FDR

Details

Top CG groups are determined by estimate (descending order).

Value

grid plot object

Examples

```
KYCG_plotDot(data.frame(
  estimate=runif(10,0,10), FDR=runif(10,0,1), nD=runif(10,10,20),
  overlap=as.integer(runif(10,0,30)), group="g", dbname=seq_len(10)))
```

`KYCG_plotEnrichAll`*plot enrichment test result*

Description

plot enrichment test result

Usage

```
KYCG_plotEnrichAll(df, fdr_max = 25, n_label = 15, min_estimate = 0)
```

Arguments

df	test enrichment result data frame
fdr_max	maximum fdr for capping
n_label	number of database to label
min_estimate	minimum estimate

Value

grid object

Examples

```
query <- KYCG_getDBs("MM285.designGroup")[[ "PGCMeth" ]]
res <- testEnrichment(query)
KYCG_plotEnrichAll(res)
```

KYCG_plotLollipop *creates a lollipop plot of log(estimate) given data with fields estimate.*

Description

creates a lollipop plot of log(estimate) given data with fields estimate.

Usage

```
KYCG_plotLollipop(df, label_column = "dbname", n = 20)
```

Arguments

df	DataFrame where each row is a database name with its estimate.
label_column	column in df to be used as the label (default: dbname)
n	Integer representing the number of top enrichments to report. Optional. (Default: 10)

Value

ggplot lollipop plot

Examples

```
KYCG_plotLollipop(data.frame(
  estimate=runif(10,0,10), FDR=runif(10,0,1), nD=runif(10,10,20),
  overlap=as.integer(runif(10,0,30)), group="g",
  dbname=as.character(seq_len(10))))
```

KYCG_plotManhattan	<i>KYCG_plotManhattan makes a manhattan plot to summarize EWAS results</i>
--------------------	--

Description

KYCG_plotManhattan makes a manhattan plot to summarize EWAS results

Usage

```
KYCG_plotManhattan(
  vals,
  platform = NULL,
  genome = NULL,
  title = NULL,
  label_min = 100,
  col = c("wheat1", "sienna3"),
  ylabel = "Value"
)
```

Arguments

vals	named vector of values (P,Q etc), vector name is Probe ID.
platform	String corresponding to the type of platform to use for retrieving GRanges coordinates of probes. Either MM285, EPIC, HM450, or HM27. If it is not provided, it will be inferred from the query set probeIDs (Default: NA).
genome	hg38, mm10, ..., will infer if not given. For additional mapping, download the GRanges object from http://zwdzwd.github.io/InfiniumAnnotation and provide the following argument ..., genome = sesameAnno_buildManifestGRanges("downloaded_file"),... to this function.
title	title for plot
label_min	Threshold above which data points will be labelled with Probe ID
col	color
ylabel	y-axis label

Value

a ggplot object

Examples

```
## see vignette for examples
sesameDataGet_resetEnv()
```

KYCG_plotMeta	<i>Plot meta gene or other meta genomic features</i>
---------------	--

Description

Plot meta gene or other meta genomic features

Usage

```
KYCG_plotMeta(betas, platform = NULL)
```

Arguments

betas	a named numeric vector or a matrix (row: probes; column: samples)
platform	if not given and x is a SigDF, will be inferred the meta features

Value

a grid plot object

Examples

```
sdf <- sesameDataGet("EPIC.1.SigDF")  
KYCG_plotMeta(getBetas(sdf))
```

KYCG_plotMetaEnrichment	<i>Plot meta gene or other meta genomic features</i>
-------------------------	--

Description

Plot meta gene or other meta genomic features

Usage

```
KYCG_plotMetaEnrichment(result_list)
```

Arguments

result_list	one or a list of testEnrichment
-------------	---------------------------------

Value

a grid plot object

Examples

```
cg_lists <- KYCG_getDBs("MM285.TFBS")
queries <- cg_lists[(sapply(cg_lists, length) > 40000)]
result_list <- lapply(queries, testEnrichment,
  "MM285.metagene", silent=TRUE)

KYCG_plotMetaEnrichment(result_list)
```

KYCG_plotPointRange *Plot point range for a list of enrichment testing results against the same set of databases*

Description

Plot point range for a list of enrichment testing results against the same set of databases

Usage

```
KYCG_plotPointRange(result_list)
```

Arguments

result_list a list of testEnrichment resultsx

Value

grid plot object

Examples

```
## pick some big TFBS-overlapping CpG groups
cg_lists <- KYCG_getDBs("MM285.TFBS")
queries <- cg_lists[(sapply(cg_lists, length) > 40000)]
result_list <- lapply(queries, testEnrichment, "MM285.chromHMM")
KYCG_plotPointRange(result_list)
```

`KYCG_plotSetEnrichment`*Plot Set Enrichment*

Description

Plot Set Enrichment

Usage`KYCG_plotSetEnrichment(result, n_sample = 1000, n_presence = 200)`**Arguments**

<code>result</code>	result object as returned from an element of the list of <code>testEnrichmentSEA(..., prepPlot=TRUE)</code>
<code>n_sample</code>	number of CpGs to sample
<code>n_presence</code>	number of overlap to sample for the plot

Value

grid object for plot

Examples

```
query <- KYCG_getDBs("KYCG.MM285.designGroup")[[ "VMR" ]]  
db <- KYCG_getDBs("MM285.seqContextN", "distToTSS")  
res <- testEnrichmentSEA(query, db, prepPlot = TRUE)  
KYCG_plotSetEnrichment(res[[1]])
```

<code>KYCG_plotVolcano</code>	<i>creates a volcano plot of $-\log_2(p.value)$ and $\log(estimate)$ given data with fields estimate and p.value.</i>
-------------------------------	---

Descriptioncreates a volcano plot of $-\log_2(p.value)$ and $\log(estimate)$ given data with fields estimate and p.value.**Usage**`KYCG_plotVolcano(data, label_column = "dbname", alpha = 0.05)`

Arguments

data	DataFrame where each field is a database name with two fields for the estimate and p.value.
label_column	column in df to be used as the label (default: dbname)
alpha	Float representing the cut-off alpha value for the plot. Optional. (Default: 0.05)

Value

ggplot volcano plot

Examples

```
KYCG_plotVolcano(data.frame(
  estimate=runif(10,0,10), FDR=runif(10,0,1), nD=runif(10,10,20),
  overlap=as.integer(runif(10,0,30)), group="g", dbname=seq_len(10)))
```

KYCG_plotWaterfall *create a waterfall plot of log(estimate) given test enrichment*

Description

create a waterfall plot of log(estimate) given test enrichment

Usage

```
KYCG_plotWaterfall(df, n_label = 10, label_column = "dbname")
```

Arguments

df	data frame where each row is a database with test enrichment result
n_label	number of datapoints to label
label_column	column in df to be used as the label (default: dbname)

Value

grid

Examples

```
library(SummarizedExperiment)
df <- rowData(sesameDataGet('MM285.tissueSignature'))
query <- df$Probe_ID[df$branch == "fetal_brain" & df$type == "Hypo"]
results <- testEnrichment(query, "TFBS")
KYCG_plotWaterfall(results)
```

listAvailableMasks *list existing quality masks for a SigDF*

Description

list existing quality masks for a SigDF

Usage

```
listAvailableMasks(platform, verbose = FALSE)
```

Arguments

platform	EPIC, MM285, HM450 etc
verbose	print more messages

Value

a tibble of masks

Examples

```
listAvailableMasks("EPIC")
```

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')
```

mapToMammal40	<i>Map the SDF (from overlap array platforms) Replicates are merged by picking the best detection</i>
---------------	---

Description

Map the SDF (from overlap array platforms) Replicates are merged by picking the best detection

Usage

```
mapToMammal40(sdf)
```

Arguments

sdf a SigDF object

Value

a named numeric vector for beta values

Examples

```
sdf <- sesameDataGet("Mammal40.1.SigDF")
betas <- mapToMammal40(sdf[1:10,])
```

matchDesign	<i>normalize Infinium I probe betas to Infinium II</i>
-------------	--

Description

This is designed to counter tail inflation in Infinium I probes.

Usage

```
matchDesign(sdf, min_dbeta = 0.3)
```

Arguments

sdf	SigDF
min_dbeta	the default algorithm perform 2-state quantile-normalization of the unmethylated and methylated modes separately. However, when the two modes are too close, we fall back to a one-mode normalization. The threshold defines the maximum inter-mode distance.

Value

SigDF

Examples

```
library(RPMM)
sdf <- sesameDataGet("MM285.1.SigDF")
sesameQC_plotBetaByDesign(sdf)
sesameQC_plotBetaByDesign(matchDesign(sdf))
```

meanIntensity	<i>Whole-dataset-wide Mean Intensity</i>
---------------	--

Description

The function takes one single SigDF 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(sdf, mask = TRUE)
```

Arguments

sdf	a SigDF
mask	whether to mask probes using mask column

Details

Note: mean in this case is more informative than median because methylation level is mostly bi-modal.

Value

mean of all intensities

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
meanIntensity(sdf)
```

medianTotalIntensity *Whole-dataset-wide Median Total Intensity (M+U)*

Description

The function takes one single SigDF and computes median intensity of M+U for each probe. This function outputs a single numeric for the median.

Usage

```
medianTotalIntensity(sdf, mask = TRUE)
```

Arguments

sdf	a SigDF
mask	whether to mask probes using mask column

Value

median of all intensities

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
medianTotalIntensity(sdf)
```

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))
```

negControls	<i>get negative control signal</i>
-------------	------------------------------------

Description

get negative control signal

Usage

```
negControls(sdf)
```

Arguments

sdf a SigDF

Value

a data frame of negative control signals

noMasked	<i>remove masked probes from SigDF</i>
----------	--

Description

remove masked probes from SigDF

Usage

```
noMasked(sdf)
```

Arguments

sdf	input SigDF object
-----	--------------------

Value

a SigDF object without masked probes

Examples

```
sesameDataCache()  
sdf <- sesameDataGet("EPIC.1.SigDF")  
sdf <- p00BAH(sdf)  
  
sdf_noMasked <- noMasked(sdf)
```

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

Description

The function takes a SigDF and returns a modified SigDF 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. For species-specific processing, one should call inferSpecies on SigDF first. Multi-mapping probes are excluded.

Usage

```
noob(sdf, combine.neg = TRUE, offset = 15)
```

Arguments

sdf	a SigDF
combine.neg	whether to combine negative control probe.
offset	offset

Details

When `combine.neg = TRUE`, background will be parameterized by both negative control and out-of-band probes.

Value

a new SigDF with noob background correction

Examples

```
sdf <- sesameDataGet('EPIC.1.SigDF')
sdf.nb <- noob(sdf)
```

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

Description

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

Usage

```
normControls(sdf, average = FALSE, verbose = FALSE)
```

Arguments

sdf	a SigDF
average	whether to average
verbose	print more messages

Value

a data frame of normalization control signals

 openSesame

The openSesame pipeline

Description

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

Usage

```
openSesame(
  x,
  prep = "QCDPB",
  prep_args = NULL,
  manifest = NULL,
  func = getBetas,
  BPPARAM = SerialParam(),
  platform = "",
  ...
)
```

Arguments

x	SigDF(s), IDAT prefix(es)
prep	preprocessing code, see ?prepSesame
prep_args	optional preprocessing argument list, see ?prepSesame
manifest	optional dynamic manifest
func	either getBetas or getAFs, if NULL, then return SigDF list
BPPARAM	get parallel with MulticoreParam(n)
platform	optional platform string
...	parameters to getBetas

Details

If the input is an IDAT prefix or a SigDF, the output is the beta value numerics.

Value

a numeric vector for processed beta values

Examples

```
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'))
```

palgen *Generate some additional color palettes*

Description

Generate some additional color palettes

Usage

```
palgen(pal, n = 150, space = "Lab")
```

Arguments

pal	a string for adhoc pals
n	the number of colors for interpolation
space	rgb or Lab

Value

a palette-generating function

Examples

```
library(pals)
pal.bands(palgen("whiteturbo"))
```

parseGEOsignalMU	<i>Convert signal M and U to SigDF</i>
------------------	--

Description

This overcomes the issue of missing IDAT files. However, out-of-band signals will be missing or faked (sampled from a normal distribution).

Usage

```
parseGEOsignalMU(
  sigM,
  sigU,
  Probe_IDs,
  oob.mean = 500,
  oob.sd = 300,
  platform = NULL
)
```

Arguments

sigM	methyated signal, a numeric vector
sigU	unmethyated signal, a numeric vector
Probe_IDs	probe ID vector
oob.mean	assumed mean for out-of-band signals
oob.sd	assumed standard deviation for out-of-band signals
platform	platform code, will infer if not given

Value

SigDF

Examples

```
sigM <- c(11436, 6068, 2864)
sigU <- c(1476, 804, 393)
probes <- c("cg07881041", "cg23229610", "cg03513874")
sdf <- parseGEOsignalMU(sigM, sigU, probes, platform = "EPIC")
```

pOOBAH

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

Description

aka pOOBAH (p-val's by Out-Of-Band Array Hybridization)

Usage

```
pOOBAH(  
  sdf,  
  return.pval = FALSE,  
  combine.neg = TRUE,  
  pval.threshold = 0.05,  
  verbose = FALSE  
)
```

Arguments

sdf	a SigDF
return.pval	whether to return p-values, instead of a masked SigDF
combine.neg	whether to combine negative control probes with the out-of-band probes in simulating the signal background
pval.threshold	minimum p-value to mask
verbose	print more messages

Details

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

Value

a SigDF, or a p-value vector if return.pval is TRUE

Examples

```
sdf <- sesameDataGet("EPIC.1.SigDF")  
sum(sdf$mask)  
sum(pOOBAH(sdf)$mask)
```

predictAge	<i>Predict age using linear models</i>
------------	--

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 different models.

Usage

```
predictAge(betas, model, na_fallback = TRUE, min_nonna = 10)
```

Arguments

betas	a probeID-named vector of beta values
model	a model object from sesameDataGet. should contain param, intercept, response2age. default to the Horvath353 model.
na_fallback	use fall back values if na
min_nonna	the minimum number of non-NA values.

Details

You can get the models such as the Horvath aging model (Horvath 2013 Genome Biology) from sesameDataGet. The function outputs a single numeric of age in years.

Here are some built-in age models: Anno/HM450/Clock_Horvath353.rds Anno/HM450/Clock_Hannum.rds Anno/HM450/Clock_SkinBlood.rds Anno/EPIC/Clock_PhenoAge.rds Anno/MM285/Clock_Zhou347.rds see vignette [inferences.html#Age__Epigenetic_Clock](#) for details

Value

age in the unit specified in the model (usually in year, but sometimes can be month, like in the mouse clocks).

Examples

```
betas <- sesameDataGet('HM450.1.TCGA.PAAD')$betas
## model <- sesameAnno_get("Anno/HM450/Clock_Horvath353.rds")
## predictAge(betas, model)
```

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

```
cat("Deprecated. See predictAge")
```

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

```
cat("Deprecated. See predictAge")
```

```
predictMouseAgeInMonth
```

Mouse age predictor

Description

The function takes a named numeric vector of beta values. The name attribute contains the probe ID. The function looks for overlapping probes and estimate age using an aging model built from 321 MM285 probes. The function outputs a single numeric of age in months. The clock is most accurate with the sesame preprocessing.

Usage

```
predictMouseAgeInMonth(betas, na_fallback = TRUE)
```

Arguments

betas	a probeID-named vector of beta values
na_fallback	use the fallback default for NAs.

Value

age in month

Examples

```
cat("Deprecated. See predictAge")
```

```
prefixMask
```

Mask SigDF by probe ID prefix

Description

Mask SigDF by probe ID prefix

Usage

```
prefixMask(sdf, prefixes = NULL, invert = FALSE)
```

Arguments

sdf	SigDF
prefixes	prefix characters
invert	use the complement set

Value

SigDF

Examples

```
sdf <- resetMask(sesameDataGet("MM285.1.SigDF"))
sum(prefixMask(sdf, c("ct1", "rs"))$mask)
sum(prefixMask(sdf, c("ct1"))$mask)
sum(prefixMask(sdf, c("ct1", "rs", "ch"))$mask)
```

prefixMaskButC	<i>Mask all but C probes in SigDF</i>
----------------	---------------------------------------

Description

Mask all but C probes in SigDF

Usage

```
prefixMaskButC(sdf)
```

Arguments

sdf	SigDF
-----	-------

Value

SigDF

Examples

```
sdf <- resetMask(sesameDataGet("MM285.1.SigDF"))
sum(prefixMaskButC(sdf)$mask)
```

prefixMaskButCG	<i>Mask all but CG probes in SigDF</i>
-----------------	--

Description

Mask all but CG probes in SigDF

Usage

```
prefixMaskButCG(sdf)
```

Arguments

sdf	SigDF
-----	-------

Value

SigDF

Examples

```
sdf <- resetMask(sesameDataGet("MM285.1.SigDF"))
sum(prefixMaskButCG(sdf)$mask)
```

prepSesame	<i>Apply a chain of sesame preprocessing functions in an arbitrary order</i>
------------	--

Description

Notes on the order of operation: 1. qualityMask and inferSpecies should go before noob and pOOBAH, otherwise the background is too high because of Multi, uk and other probes 2. dyeBias correction needs to happen early 3. channel inference before dyebias 4. noob should happen last, pOOBAH before noob because noob modifies oob

Usage

```
prepSesame(sdf, prep = "QCDPB", prep_args = NULL)
```

Arguments

sdf	SigDF
prep	code that indicates preprocessing functions and their execution order (functions on the left is executed first).
prep_args	optional argument list to individual functions, e.g., prepSesame(sdf, prep_args=list(Q=list(mask_names = "design_issue"))) sets qualityMask(sdf, mask_names = "design_issue")

Value

SigDF

Examples

```
sdf <- sesameDataGet("MM285.1.SigDF")
sdf1 <- prepSesame(sdf, "QCDPB")
```

```
prepSesameList      List supported prepSesame functions
```

Description

List supported prepSesame functions

Usage

```
prepSesameList()
```

Value

a data frame with code, func, description

Examples

```
prepSesameList()
```

```
print.DMLSummary    Print DMLSummary object
```

Description

Print DMLSummary object

Usage

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

Arguments

```
x          a DMLSummary object
...        extra parameter for print
```

Value

print DMLSummary result on screen

Examples

```
sesameDataCache() # in case not done yet
data <- sesameDataGet('HM450.76.TCGA.matched')
## test the first 10
smry <- DML(data$betas[1:10,], ~type, meta=data$sampleInfo)
smry

sesameDataGet_resetEnv()
```

print.fileSet	<i>Print a fileSet</i>
---------------	------------------------

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
```

probeID_designType	<i>Extract the probe type field from probe ID This only works with the new probe ID system. See https://github.com/zhou-lab/InfiniumAnnotation for illustration</i>
--------------------	--

Description

Extract the probe type field from probe ID This only works with the new probe ID system. See <https://github.com/zhou-lab/InfiniumAnnotation> for illustration

Usage

```
probeID_designType(Probe_ID)
```

Arguments

Probe_ID	Probe ID
----------	----------

Value

a vector of '1' and '2' suggesting Infinium-I and Infinium-II

Examples

```
probeID_designType("cg36609548_TC21")
```

probeSuccessRate	<i>Whole-dataset-wide Probe Success Rate</i>
------------------	--

Description

This function calculates the probe success rate using pOOBAH detection p-values. Probes that has a detection p-value higher than a specific threshold are considered failed probes.

Usage

```
probeSuccessRate(sdf, mask = TRUE, max_pval = 0.05)
```

Arguments

sdf	a SigDF
mask	whether or not we count the masked probes in SigDF
max_pval	the maximum p-value to consider detection success

Value

a fraction number as probe success rate

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
probeSuccessRate(sdf)
```

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

Description

Currently quality masking only supports three platforms see also `listAvailableMasks(sdfPlatform(sdf))`

Usage

```
qualityMask(sdf, mask_names = "recommended", prefixes = NULL, verbose = FALSE)
```

Arguments

sdf	a SigDF object
mask_names	mask names, default to "recommended", can be a vector of multiple masks, e.g., <code>c("design_issue", "multi")</code> , NULL to skip
prefixes	mask by probe ID prefixes, e.g., cg
verbose	print more messages

Value

a filtered SigDF

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
sum(sdf$mask)
sum(qualityMask(sdf)$mask)
sum(qualityMask(sdf, mask_names = NULL, prefixes = "rs")$mask)

## list available masks, the mask_name column
listAvailableMasks(sdfPlatform(sdf))
```

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 SigDF.

Usage

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

Arguments

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

Value

a SigDF

Examples

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

reIdentify

Re-identify IDATs by restoring scrambled SNP intensities

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)
```

resetMask

Reset Masking

Description

Reset Masking

Usage

```
resetMask(sdf, verbose = FALSE)
```

Arguments

sdf	a SigDF
verbose	print more messages

Value

a new SigDF with mask reset to all FALSE

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
sum(sdf$mask)
sdf <- addMask(sdf, c("cg14057072", "cg22344912"))
sum(sdf$mask)
sum(resetMask(sdf)$mask)
```

scrub	<i>SCRUB background correction</i>
-------	------------------------------------

Description

This function takes a SigDF and returns a modified SigDF with background subtracted. scrub subtracts residual background using background median

Usage

```
scrub(sdf)
```

Arguments

sdf a SigDF

Details

This function is meant to be used after noob.

Value

a new SigDF with noob background correction

Examples

```
sdf <- sesameDataGet('EPIC.1.SigDF')
sdf.nb <- noob(sdf)
sdf.nb.scrub <- scrub(sdf.nb)
```

scrubSoft	<i>SCRUB background correction</i>
-----------	------------------------------------

Description

This function takes a SigDF and returns a modified SigDF with background subtracted. scrubSoft subtracts residual background using a noob-like procedure.

Usage

```
scrubSoft(sdf)
```

Arguments

sdf a SigDF

Details

This function is meant to be used after noob.

Value

a new SigDF with noob background correction

Examples

```
sdf <- sesameDataGet('EPIC.1.SigDF')
sdf.nb <- noob(sdf)
sdf.nb.scrubSoft <- scrubSoft(sdf.nb)
```

SDFcollapseToPfx	<i>collapse to probe prefix</i>
------------------	---------------------------------

Description

collapse to probe prefix

Usage

```
SDFcollapseToPfx(sdf)
```

Arguments

sdf	a SigDF object
-----	----------------

Value

a data frame with updated Probe_ID

sdfPlatform	<i>Convenience function to output platform attribute of SigDF</i>
-------------	---

Description

Convenience function to output platform attribute of SigDF

Usage

```
sdfPlatform(sdf, verbose = FALSE)
```

Arguments

sdf	a SigDF object
verbose	print more messages

Value

the platform string for the SigDF object

Examples

```
sesameDataCache()
sdf <- sesameDataGet('EPIC.1.SigDF')
sdfPlatform(sdf)
```

sdf_read_table	<i>read a table file to SigDF</i>
----------------	-----------------------------------

Description

read a table file to SigDF

Usage

```
sdf_read_table(fname, platform = NULL, verbose = FALSE, ...)
```

Arguments

fname	file name
platform	array platform (will infer if not given)
verbose	print more information
...	additional argument to read.table

Value

read table file to SigDF

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
fname <- sprintf("%s/sigdf.txt", tempdir())
sdf_write_table(sdf, file=fname)
sdf2 <- sdf_read_table(fname)
```

sdf_write_table	<i>write SigDF to table file</i>
-----------------	----------------------------------

Description

write SigDF to table file

Usage

```
sdf_write_table(sdf, ...)
```

Arguments

sdf	the SigDF to output
...	additional argument to write.table

Value

write SigDF to table file

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
sdf_write_table(sdf, file=sprintf("%s/sigdf.txt", tempdir()))
```

searchIDATprefixes	<i>Identify IDATs from a directory</i>
--------------------	--

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

sesameAnno_buildAddressFile	<i>Build sesame ordering address file from tsv</i>
-----------------------------	--

Description

Build sesame ordering address file from tsv

Usage

```
sesameAnno_buildAddressFile(tsv)
```


Arguments

tsv a platform name, a file path or a tibble/data.frame manifest file

Value

a list of ordering and controls

addr <- sesameAnno_buildAddressFile("HM450") ## or ## download tsv directly from ## <http://zwdzwd.github.io/Infinium>
or ## addr <- sesameAnno_buildAddressFile("downloaded_file")

sesameAnno_buildManifestGRanges

Build manifest GRanges from tsv

Description

Build manifest GRanges from tsv

Usage

```
sesameAnno_buildManifestGRanges(tsv, genome, decoy = FALSE, columns = NULL)
```

Arguments

tsv a file path, a platform (e.g., EPIC), or a tibble/data.frame object

genome a genome string, e.g., hg38, mm10

decoy consider decoy sequence in chromosome order

columns the columns to include in the GRanges

Value

GRanges

genome <- sesameAnno_buildManifestGRanges("HM450", "hg38") ## or ## download tsv directly from ## <http://zwdzwd.github.io/InfiniumAnnotation> or ## genome <- sesameAnno_buildManifestGRanges("downloaded_file", "hg38")

sesameAnno_download *Download additional annotation files*

Description

see also <http://zwdzwd.github.io/InfiniumAnnotation>

Usage

```
sesameAnno_download(title, dest_dir, version = 1)
```

Arguments

title	title of the annotation file
dest_dir	download to this directory
version	version number

Details

This function acts similarly as `sesameAnno_get` except that it directly download files without invoking `BiocFileCache`. This is needed in some situation because `BiocFileCache` may change the file name and downstream program may depend on the correct file names. It also lets you download files in a cleaner way without routing through `BiocFileCache`

Value

annotation file

Examples

```
## avoid testing as this function uses external host
if (FALSE) {
  sesameAnno_download("Test/3999492009_R01C01_Grn.idat", tempdir())
}
```

sesameAnno_get *download Infinium manifest from the associated Github repository*

Description

Since most of the annotation is not essential to sesame functioning, `sesameData` package no longer host the full manifest. This is the command to use to retrieve the full manifest and other annotation from the following Github host:

Usage

```
sesameAnno_get(title, return_path = FALSE, version = 1)
```

Arguments

title	the title of the resource
return_path	return cached file path
version	release version, default is the latest

Details

<https://github.com/zhou-lab/InfiniumAnnotationV1>

Please check the repo itself for what is available. See also <http://zwdzwd.github.io/InfiniumAnnotation>

Unless `return_path = TRUE`, This function calls import function depending on the resource name suffix. If the url ends with `.rds`, it will use `readRDS`. If the url ends with `.tsv.gz` it will use `read_tsv`. For all other cases, the function will return the cached file name.

This function replaces `sesameAnno_getManifestDF`.

Value

tibble

Examples

```
## avoid testing since it depends on external host
if (FALSE) {
  mapping <- sesameAnno_get("Mammal40/hg38.tsv.gz")
  annoI <- sesameAnno_get("Anno/EPIC/EPIC.hg19.typeI_overlap_b151.rds")
  mft <- sesameAnno_get("Anno/MM285/MM285.mm10.manifest.tsv.gz")
}
```

sesameData_getAnno	<i>retrieve additional annotation files</i>
--------------------	---

Description

retrieve additional annotation files

Usage

```
sesameData_getAnno(title, version = 1, dest_dir = NULL)
```

Arguments

title	title of the annotation file
version	version number
dest_dir	if not NULL, download to this directory

Value

annotation file

Examples

```
cat("Deprecated!")
```

sesameQC-class	<i>An S4 class to hold QC statistics</i>
----------------	--

Description

An S4 class to hold QC statistics

Slots

stat a list to store qc stats

sesameQC_calcStats	<i>Calculate QC statistics</i>
--------------------	--------------------------------

Description

It is a function to call one or multiple sesameQC_calcStats functions

Usage

```
sesameQC_calcStats(sdf, funs = NULL)
```

Arguments

sdf	a SigDF object
funs	a sesameQC_calcStats_* function or a list of them default to all functions. One can also use a string such as "detection" or c("detection", "intensity") to reduce typing

Details

currently supporting: detection, intensity, numProbes, channel, dyeBias, betas

Value

a sesameQC object

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
sesameQC_calcStats(sdf)
sesameQC_calcStats(sdf, "detection")
sesameQC_calcStats(sdf, c("detection", "channel"))
## retrieve stats as a list
sesameQC_getStats(sesameQC_calcStats(sdf, "detection"))
## or as data frames
as.data.frame(sesameQC_calcStats(sdf, "detection"))
```

sesameQC_getStats	<i>Get stat numbers from an sesameQC object</i>
-------------------	---

Description

Get stat numbers from an sesameQC object

Usage

```
sesameQC_getStats(qc, stat_names = NULL, drop = TRUE)
```

Arguments

qc	a sesameQC object
stat_names	which stat(s) to retrieve, default to all.
drop	whether to drop to a string when stats_names has only one element.

Value

a list of named stats to be retrieved

Examples

```
sdf <- sesameDataGet("EPIC.1.SigDF")
qc <- sesameQC_calcStats(sdf, "detection")
sesameQC_getStats(qc, "frac_dt")
```

sesameQC_plotBar *Bar plots for sesameQC*

Description

By default, it plots median_beta_cg, median_beta_ch, RGratio, RGdistort, frac_dt

Usage

```
sesameQC_plotBar(qcs, keys = NULL)
```

Arguments

qcs	a list of SigDFs
keys	optional, other key to plot, instead of the default keys can be found in the parenthesis of the print output of each sesameQC output.

Value

a bar plot comparing different QC metrics

Examples

```
sesameDataCache() # if not done yet
sdfs <- sesameDataGet("EPIC.5.SigDF.normal")[1:2]
sesameQC_plotBar(lapply(sdfs, sesameQC_calcStats, "detection"))
```

sesameQC_plotBetaByDesign
Plot betas distinguishing different Infinium chemistries

Description

Plot betas distinguishing different Infinium chemistries

Usage

```
sesameQC_plotBetaByDesign(
  sdf,
  prep = NULL,
  legend_pos = "top",
  mar = c(3, 3, 1, 1),
  main = "",
  ...
)
```

Arguments

sdf	SigDF
prep	prep codes to step through
legend_pos	legend position (default: top)
mar	margin of layout when showing steps of prep
main	main title in plots
...	additional options to plot

Value

create a density plot

Examples

```
sdf <- sesameDataGet("EPIC.1.SigDF")
sesameQC_plotBetaByDesign(sdf, prep="DB")
```

sesameQC_plotHeatSNPs *Plot SNP heatmap*

Description

Plot SNP heatmap

Usage

```
sesameQC_plotHeatSNPs(sdfs, cluster = TRUE, filter.nonvariant = TRUE)
```

Arguments

sdfs	beta value matrix, row: probes; column: samples
cluster	show clustered heatmap
filter.nonvariant	whether to filter nonvariant (range < 0.3)

Value

a grid graphics object

Examples

```
sdfs <- sesameDataGet("EPIC.5.SigDF.normal")[1:2]
plt <- sesameQC_plotHeatSNPs(sdfs, filter.nonvariant = FALSE)
```

sesameQC_plotIntensVsBetas

Plot Total Signal Intensities vs Beta Values This plot is helpful in revealing the extent of signal background and dye bias.

Description

Plot Total Signal Intensities vs Beta Values This plot is helpful in revealing the extent of signal background and dye bias.

Usage

```
sesameQC_plotIntensVsBetas(  
  sdf,  
  mask = TRUE,  
  use_max = FALSE,  
  intens.range = c(5, 15),  
  ...  
)
```

Arguments

sdf	a SigDF
mask	whether to remove probes that are masked
use_max	to use max(M,U) or M+U
intens.range	plot range of signal intensity
...	additional arguments to smoothScatter

Value

create a total signal intensity vs beta value plot

Examples

```
sesameDataCache() # if not done yet  
sdf <- sesameDataGet('EPIC.1.SigDF')  
sesameQC_plotIntensVsBetas(sdf)
```

sesameQC_plotRedGrnQQ *Plot red-green QQ-Plot using Infinium-I Probes*

Description

Plot red-green QQ-Plot using Infinium-I Probes

Usage

```
sesameQC_plotRedGrnQQ(sdf, main = "R-G QQ Plot", ...)
```

Arguments

sdf	a SigDF
main	plot title
...	additional options to qqplot

Value

create a qqplot

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
sesameQC_plotRedGrnQQ(sdf)
```

sesameQC_rankStats *This function compares the input sample with public data. Only overlapping metrics will be compared.*

Description

This function compares the input sample with public data. Only overlapping metrics will be compared.

Usage

```
sesameQC_rankStats(qc, publicQC = NULL, platform = "EPIC")
```

Arguments

qc	a sesameQC object
publicQC	public QC statistics, filtered from e.g.: EPIC.publicQC, MM285.publicQC and Mammal40.publicQC
platform	EPIC, MM285 or Mammal40, used when publicQC is not given

Value

a sesameQC

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
sesameQC_rankStats(sesameQC_calcStats(sdf, "intensity"))
```

setMask

Set mask to only the probes specified

Description

Set mask to only the probes specified

Usage

```
setMask(sdf, probes)
```

Arguments

sdf	a SigDF
probes	a vector of probe IDs or a logical vector with TRUE representing masked probes

Value

a SigDF with added mask

Examples

```
sdf <- sesameDataGet('EPIC.1.SigDF')
sum(sdf$mask)
sum(setMask(sdf, "cg14959801")$mask)
sum(setMask(sdf, c("cg14057072", "cg22344912"))$mask)
```

SigDF	<i>SigDF validation from a plain data frame</i>
-------	---

Description

SigDF validation from a plain data frame

Usage

```
SigDF(df, platform = "EPIC", ctl = NULL)
```

Arguments

df	a data.frame with Probe_ID, MG, MR, UG, UR, col and mask
platform	a string to specify the array platform
ctl	optional control probe data frame

Value

a SigDF object

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
```

signalMU	<i>report M and U for regular probes</i>
----------	--

Description

report M and U for regular probes

Usage

```
signalMU(sdf, mask = TRUE)
```

Arguments

sdf	a SigDF
mask	whether to apply mask

Value

a data frame of M and U columns

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
head(signalMU(sdf))
```

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')
```

summaryExtractTest	<i>Extract slope information from DMLSummary</i>
--------------------	--

Description

Extract slope information from DMLSummary

Usage

```
summaryExtractTest(smry)
```

Arguments

smry DMLSummary from DML command

Value

a table of slope and p-value

Examples

```
sesameDataCache() # in case not done yet
data <- sesameDataGet('HM450.76.TCGA.matched')
smry <- DML(data$betas[1:10,], ~type, meta=data$sampleInfo)
slopes <- summaryExtractTest(smry)

sesameDataGet_resetEnv()
```

testEnrichment	<i>testEnrichment tests for the enrichment of set of probes (query set) in a number of features (database sets).</i>
----------------	--

Description

testEnrichment tests for the enrichment of set of probes (query set) in a number of features (database sets).

Usage

```
testEnrichment(
  query,
  databases = NULL,
  universe = NULL,
  alternative = "greater",
  platform = NULL,
  silent = FALSE
)
```

Arguments

query	Vector of probes of interest (e.g., significant probes)
databases	List of vectors corresponding to the database sets of interest with associated meta data as an attribute to each element. Optional. (Default: NA)
universe	Vector of probes in the universe set containing all of the probes to be considered in the test. If it is not provided, it will be inferred from the provided platform. (Default: NA).
alternative	"two.sided", "greater", or "less"
platform	String corresponding to the type of platform to use. Either MM285, EPIC, HM450, or HM27. If it is not provided, it will be inferred from the query set probeIDs (Default: NA).
silent	output message? (Default: FALSE)

Value

One list containing features corresponding the test estimate, p-value, and type of test.

Examples

```
library(SummarizedExperiment)
df <- rowData(sesameDataGet('MM285.tissueSignature'))
query <- df$Probe_ID[df$branch == "B_cell"]
res <- testEnrichment(query, "chromHMM")
sesameDataGet_resetEnv()
```

testEnrichmentFisher *testEnrichmentFisher uses Fisher's exact test to estimate the association between two categorical variables.*

Description

Estimates log2 Odds ratio

Usage

```
testEnrichmentFisher(query, database, universe, alternative = "greater")
```

Arguments

query	Vector of probes of interest (e.g., significant probes)
database	Vectors corresponding to the database set of interest with associated meta data as an attribute to each element.
universe	Vector of probes in the universe set containing all of
alternative	greater or two.sided (default: greater) the probes to be considered in the test. (Default: NULL)

Value

A DataFrame with the estimate/statistic, p-value, and name of test for the given results.

testEnrichmentSEA	<i>uses the GSEA-like test to estimate the association of a categorical variable against a continuous variable.</i>
-------------------	---

Description

estimate represent enrichment score and negative estimate indicate a test for depletion

Usage

```
testEnrichmentSEA(
  query,
  databases,
  platform = NULL,
  silent = FALSE,
  precise = FALSE,
  prepPlot = FALSE
)
```

Arguments

query	query, if numerical, expect categorical database, if categorical expect numerical database
databases	database, numerical or categorical, but needs to be different from query
platform	EPIC, MM285, ..., infer if not given
silent	suppress message (default: FALSE)
precise	whether to compute precise p-value (up to numerical limit) of interest.
prepPlot	return the raw enrichment scores and presence vectors for plotting

Value

A DataFrame with the estimate/statistic, p-value, and name of test for the given results.

Examples

```
query <- KYCG_getDBs("KYCG.MM285.designGroup")[[ "TSS" ]]
res <- testEnrichmentSEA(query, "MM285.seqContextN")
```

testEnrichmentSpearman

testEnrichmentSpearman uses the Spearman statistical test to estimate the association between two continuous variables.

Description

testEnrichmentSpearman uses the Spearman statistical test to estimate the association between two continuous variables.

Usage

```
testEnrichmentSpearman(query, database)
```

Arguments

query	Vector of probes of interest (e.g., significant probes)
database	List of vectors corresponding to the database set of interest with associated meta data as an attribute to each element.

Value

A DataFrame with the estimate/statistic, p-value, and name of test for the given results.

totalIntensities *M+U Intensities Array*

Description

The function takes one single SigDF 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(sdf, mask = FALSE)
```

Arguments

sdf	a SigDF
mask	whether to mask probes using mask column

Value

a vector of M+U signal for each probe

Examples

```
sesameDataCache() # if not done yet
sdf <- sesameDataGet('EPIC.1.SigDF')
intensities <- totalIntensities(sdf)
```

`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

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

updateSigDF *Set color and mask using strain/species-specific manifest*

Description

also sets attr("species")

Usage

```
updateSigDF(sdf, species = NULL, strain = NULL, addr = NULL, verbose = FALSE)
```

Arguments

sdf	a SigDF
species	the species the sample is considered to be
strain	the strain the sample is considered to be
addr	species-specific address species, optional
verbose	print more messages

Value

a SigDF with updated color channel and mask

Examples

```
sdf <- sesameDataGet('Mamma140.1.SigDF')  
sdf_mouse <- updateSigDF(sdf, species="mus_musculus")
```

visualizeGene *Visualize Gene*

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(  
  gene_name,  
  betas,  
  platform = NULL,  
  genome = NULL,  
  upstream = 2000,  
  dwestream = 2000,  
  ...  
)
```

Arguments

gene_name	gene name
betas	beta value matrix (row: probes, column: samples)
platform	HM450, EPIC, or MM285 (default)
genome	hg19, hg38, or mm10 (default)
upstream	distance to extend upstream
dwestream	distance to extend downstream
...	additional options, see visualizeRegion, assemble_plots

Value

None

Examples

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

visualizeProbes	<i>Visualize Region that Contains the Specified Probes</i>
-----------------	--

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 = NULL,
  genome = NULL,
  upstream = 1000,
  dwestream = 1000,
  ...
)
```

Arguments

probeNames	probe names
betas	beta value matrix (row: probes, column: samples)
platform	HM450, EPIC or MM285 (default)
genome	hg19, hg38 or mm10 (default)
upstream	distance to extend upstream
dwestream	distance to extend downstream
...	additional options, see visualizeRegion and assemble_plots

Value

None

Examples

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

visualizeRegion

Visualize Region

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(
  chrm,
  beg,
  end,
  betas,
  platform = NULL,
  genome = NULL,
  draw = TRUE,
  cluster.samples = FALSE,
  na.rm = FALSE,
  nprobes.max = 1000,
  txn.types = "protein_coding",
  txn.font.size = 6,
  ...
)
```

Arguments

chrm	chromosome
beg	begin of the region
end	end of the region
betas	beta value matrix (row: probes, column: samples)
platform	EPIC, HM450, or MM285
genome	hg38, mm10, ..., will infer if not given. For additional mapping, download the GRanges object from http://zwdzwd.github.io/InfiniumAnnotation and provide the following argument ..., genome = sesameAnno_buildManifestGRanges("downloaded_file"),... to this function.
draw	draw figure or return betas
cluster.samples	whether to cluster samples
na.rm	remove probes with all NA.
nprobes.max	maximum number of probes to plot
txn.types	default to protein_coding, use NULL for all
txn.font.size	transcript name font size
...	additional options, see assemble_plots

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 *Visualize segments*

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

```
sesameDataCache()  
## sdf <- sesameDataGet('EPIC.1.SigDF')  
## sdf.normal <- sesameDataGet('EPIC.5.SigDF.normal')  
## seg <- cnSegmentation(sdf, sdf.normal)  
## visualizeSegments(seg)  
  
sesameDataGet_resetEnv()
```

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