

Package ‘sesame’

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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.

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Author Wanding Zhou [aut, cre] (ORCID:
 <<https://orcid.org/0000-0001-9126-1932>>),
 Wubin Ding [ctb],
 David Goldberg [ctb],
 Ethan Moyer [ctb],
 Bret Barnes [ctb],
 Timothy Triche [ctb],
 Hui Shen [aut]
Maintainer Wanding Zhou <zhouwanding@gmail.com>

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sesame-package *Analyze DNA methylation data*

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.

Value

package

Author(s)

Wanding Zhou <Wanding.Zhou@vai.org>, Hui Shen <Hui.Shen@vai.org> Timothy J Triche Jr <Tim.Triche@vai.org>

References

Zhou W, Triche TJ, Laird PW, Shen H (2018)

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)
```

assemble_plots *assemble plots*

Description

assemble plots

Usage

```
assemble_plots(  
  betas,  
  txns,  
  probes,  
  plt.txns,  
  plt.mapLines,  
  plt.cytoband,  
  heat.height = NULL,  
  mapLine.height = 0.2,  
  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) |
| mapLine.height | height of the map lines |
| 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

betasCollapseToPfx *Collapse betas by averagng probes with common probe ID prefix*

Description

Collapse betas by averagng probes with common probe ID prefix

Usage

```
betasCollapseToPfx(betas, BPPARAM = SerialParam())
```

Arguments

betas either a named numeric vector or a numeric matrix (row: probes, column: samples)

BPPARAM use MulticoreParam(n) for parallel processing

Value

either named numeric vector or a numeric matrix of collapsed beta value matrix

Examples

```
## input is a matrix
m <- matrix(seq(0,1,length.out=9), nrow=3)
rownames(m) <- c("cg00004963_TC21", "cg00004963_TC22", "cg00004747_TC21")
colnames(m) <- c("A","B","C")
betasCollapseToPfx(m)

## input is a vector
m <- setNames(seq(0,1,length.out=3),
  c("cg00004963_TC21", "cg00004963_TC22", "cg00004747_TC21"))
betasCollapseToPfx(m)
```

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, probeCoords)
```

Arguments

| | |
|---------------|-------------------|
| probe.signals | probe signals |
| bin.coords | bin coordinates |
| probeCoords | 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, extR = NULL, extA = NULL, verbose = FALSE)
```

Arguments

| | |
|---------|--|
| sdf | a SigDF |
| extR | a vector of probe IDs for Infinium-I probes that extend to converted A |
| extA | a vector of probe IDs for Infinium-I probes that extend to original A |
| 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)

## For more recent platforms like EPICv2, MSA:
## One need extR and extA of other arrays using the sesameAnno
## Not run:
mft = sesameAnno_buildManifestGRanges(sprintf(
  "%s/EPICv2/EPICv2.hg38.manifest.tsv.gz",
  "https://github.com/zhou-lab/InfiniumAnnotationV1/raw/main/Anno/"),
  columns="nextBase")
extR = names(mft)[!is.na(mft$nextBase) & mft$nextBase=="R"]
extA = names(mft)[!is.na(mft$nextBase) & mft$nextBase=="A"]

## End(Not run)

```

calcEffectSize
Compute effect size for different variables from prediction matrix

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 | <i>Lookup address in one sample</i> |
|---------------------|-------------------------------------|

Description

Lookup address and transform address to probe

Usage

```
chipAddressToSignal(dm, mft, min_beads = NULL)
```

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 |
| min_beads | minimum bead counts, otherwise masked |

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,
  genomeInfo = NULL,
  probeCoords = NULL,
  tilewidth = 50000,
  verbose = FALSE,
  return.probe.signals = 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. assembly |
| genomeInfo | the genomeInfo files. The default is retrieved from sesameData. Alternative genomeInfo files can be found at https://github.com/zhou-lab/GenomeInfo |
| probeCoords | the probe coordinates in the corresponding genome if NULL (default), then the default genome assembly is used. Default genome is given by, e.g., sesameData_check_genome(NULL, "EPIC") For additional mapping, download the GRanges object from http://zwdzwd.github.io/InfiniumAnnotation and provide |

the following argument ..., probeCoords = sesameAnno_buildManifestGRanges("downloaded_file"),...
to this function.

tilewidth tile width for smoothing
verbose print more messages
return.probe.signals
 return probe-level instead of bin-level signal

Value

an object of CNSegment

Examples

```
sesameDataCache()

## Not run:
sdfs <- sesameDataGet('EPICv2.8.SigDF')
sdf <- sdfs[["K562_206909630040_R01C01"]]
seg <- cnSegmentation(sdf)
seg <- cnSegmentation(sdf, return.probe.signals=TRUE)
visualizeSegments(seg)

## End(Not run)
```

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 | <i>get the controls attributes</i> |
|----------|------------------------------------|

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))
```

| | |
|----------------|-------------------------|
| convertProbeID | <i>Convert Probe ID</i> |
|----------------|-------------------------|

Description

Convert Probe ID

Usage

```
convertProbeID(
  x,
  target_platform,
  source_platform = NULL,
  mapping = NULL,
  target_uniq = TRUE,
  include_new = FALSE,
  include_old = FALSE,
  return_mapping = FALSE
)
```


Arguments

| | |
|-----------------|---|
| x | source probe IDs |
| target_platform | the platform to take the data to |
| source_platform | optional source platform |
| mapping | a liftOver mapping file. Typically this file contains empirical evidence whether a probe mapping is reliable. If given, probe ID-based mapping will be skipped. This is to perform more stringent probe ID mapping. |
| target_uniq | whether the target Probe ID should be kept unique. |
| include_new | if true, include mapping of added probes |
| include_old | if true, include mapping of deleted probes |
| return_mapping | return mapping table, instead of the target IDs. |

Value

mapped probe IDs, or mapping table if return_mapping = T

| | |
|-----------------|---|
| 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

| | |
|------------|---|
| 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)
```

| | |
|-------------------|--|
| 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

*Test differential methylation on each locus***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. The row order of the data frame must match the column order of the beta value matrix. |
| 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,
  probe.coords = NULL,
  dist.cutoff = NULL,
  seg.per.locus = 0.5
)
```

Arguments

| | |
|----------------------------|---|
| <code>betas</code> | beta values for distance calculation |
| <code>smry</code> | DML |
| <code>contrast</code> | the pair-wise comparison or contrast check <code>colnames(attr(smry, "model.matrix"))</code> if uncertain |
| <code>platform</code> | EPIC, HM450, MM285, ... |
| <code>probe.coords</code> | <code>GRanges</code> object that defines CG coordinates if <code>NULL</code> (default), then the default genome assembly is used. Default genome is given by, e.g., <code>sesameData_check_genome(NULL, "EPIC")</code> For additional mapping, download the <code>GRanges</code> object from http://zwdzwd.github.io/InfiniumAnnotation and provide the following argument ..., <code>probe.coords = sesameAnno_buildManifestGRanges("downloaded_file"),...</code> to this function. |
| <code>dist.cutoff</code> | 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 <code>seg.per.locus</code>) |
| <code>seg.per.locus</code> | 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[1:1000,], smry, "typeTumour", platform="HM450")

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`*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

```
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

Dye bias correction by matching green and red to mid point

Description

This function compares the Type-I Red probes and Type-I Grn probes and generates and mapping to correct signal of the two channels to the middle. The function takes one single 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)
```

ELBAR*ELiminate BAcground-dominated Reading (ELBAR)*

Description

ELiminate BAcground-dominated Reading (ELBAR)

Usage

```
ELBAR(  
  sdf,  
  return.pval = FALSE,  
  pval.threshold = 0.05,  
  margin = 0.05,  
  capMU = 3000,  
  delta.beta = 0.2,  
  n.windows = 500  
)
```

Arguments

| | |
|----------------|--|
| sdf | a SigDF |
| return.pval | whether to return p-values, instead of a SigDF |
| pval.threshold | minimum p-value to mask |
| margin | the percentile margin to define envelope, the smaller the value the more aggressive the masking. |
| capMU | the maximum M+U to search for intermediate betas |
| delta.beta | maximum beta value change from sheer background-dominated readings |
| n.windows | number of windows for smoothing |

Value

a SigDF with mask added

Examples

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

| | |
|-------------------|--|
| 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 | <i>Convert SNP from Infinium array to VCF file</i> |
|-----------|--|

Description

Convert SNP from Infinium array to VCF file

Usage

```
formatVCF(sdf, anno, vcf = NULL, genome = "hg38", verbose = FALSE)
```

Arguments

| | |
|---------|--|
| sdf | SigDF |
| anno | SNP variant annotation, available at https://github.com/zhou-lab/InfiniumAnnotationV1/tree/main/Anno/EPIC.hg38.snp.tsv.gz |
| vcf | output VCF file path, if NULL output to console |
| genome | genome |
| verbose | 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 output vcf is not sorted.

Examples

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

## Not run:
## download anno from
## http://zwdzwd.github.io/InfiniumAnnotation
## output to console
anno = read_tsv(sesameAnno_download("EPICv2.hg38.snp.tsv.gz"))
head(formatVCF(sdf, anno))

## End(Not run)
```

getAFs

Get allele frequency

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)
```

getAFTypeIbySumAlleles

Get allele frequency treating type I by summing alleles

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 an numeric(0) is returned.

Usage

```
getAFTypeIbySumAlleles(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,
  collapseMethod = c("mean", "minPval")
)
```

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 |
| collapseMethod | mean or minPval |

Value

a numeric vector, beta values

Examples

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

getBinCoordinates *Get bin coordinates*

Description

requires GenomicRanges, IRanges

Usage

```
getBinCoordinates(seqLength, gapInfo, tilewidth = 50000, probeCoords)
```

Arguments

| | |
|-------------|-------------------------------|
| seqLength | chromosome information object |
| gapInfo | chromosome gap information |
| tilewidth | tile width for smoothing |
| probeCoords | probe coordinates |

Value

bin.coords

getMask *get probe masking by mask names*

Description

get probe masking by mask names

Usage

```
getMask(platform = "EPICv2", mask_names = "recommended")
```

Arguments

| | |
|------------|--|
| platform | EPICv2, EPIC, HM450, HM27, ... |
| mask_names | mask names (see listAvailableMasks) by default: "recommended" see recommendedMaskNames() for detail. |

Value

a vector of probe ID

Examples

```
length(getMask("MSA", "recommended"))
length(getMask("EPICv2", "recommended"))
length(getMask("EPICv2", c("recommended", "M_SNPcommon_1pt")))
length(getMask("EPICv2", "M_mapping"))
length(getMask("EPIC"))
length(getMask("HM450"))
length(getMask("MM285"))
```

| | |
|-----------|-------------------------------|
| 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()
```

| | |
|-------------|--|
| imputeBetas | <i>Impute of missing data of specific platform</i> |
|-------------|--|

Description

Impute of missing data of specific platform

Usage

```
imputeBetas(  
  betas,  
  platform = NULL,  
  BPPARAM = SerialParam(),  
  celltype = NULL,  
  sd_max = 999  
)
```

Arguments

| | |
|----------|--|
| betas | named vector of beta values |
| platform | platform |
| BPPARAM | use MulticoreParam(n) for parallel processing |
| celltype | celltype/tissue context of imputation, if not given, will use nearest neighbor to determine. |
| sd_max | maximum standard deviation in imputation confidence |

Value

imputed data, vector or matrix

Examples

```
betas = openSesame(sesameDataGet("EPIC.1.SigDF"))  
sum(is.na(betas))  
betas2 = imputeBetas(betas, "EPIC")  
sum(is.na(betas2))
```

`imputeBetasByGenomicNeighbors`*Impute missing data based on genomic neighbors.*

Description

Impute missing data based on genomic neighbors.

Usage

```
imputeBetasByGenomicNeighbors(  
  betas,  
  platform = NULL,  
  BPPARAM = SerialParam(),  
  max_neighbors = 3,  
  max_dist = 10000  
)
```

Arguments

| | |
|----------------------------|--|
| <code>betas</code> | named vector of beta values |
| <code>platform</code> | platform |
| <code>BPPARAM</code> | use <code>MulticoreParam(n)</code> for parallel processing |
| <code>max_neighbors</code> | maximum neighbors to use for dense regions |
| <code>max_dist</code> | maximum distance to count as neighbor |

Value

imputed data, vector or matrix

Examples

```
betas = openSesame(sesameDataGet("EPICv2.8.SigDF")[[1]])  
sum(is.na(betas))  
betas2 = imputeBetasByGenomicNeighbors(betas, "EPICv2")  
sum(is.na(betas2))
```

```
imputeBetasMatrixByMean
```

Impute Missing Values with Mean This function replaces missing values (NA) in a matrix, default is row means.

Description

Impute Missing Values with Mean This function replaces missing values (NA) in a matrix, default is row means.

Usage

```
imputeBetasMatrixByMean(mx, axis = 1)
```

Arguments

| | |
|------|--|
| mx | A matrix |
| axis | A single integer. Use 1 to impute column means (default), and 2 to impute row means. |

Value

A matrix with missing values imputed.

Examples

```
mx <- cbind(c(1, 2, NA, 4), c(NA, 2, 3, 4))
imputeBetasMatrixByMean(mx, axis = 1)
imputeBetasMatrixByMean(mx, axis = 2)
```

```
inferEthnicity
```

Infer Ethnicity

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,
  mask_failed = FALSE,
  verbose = FALSE,
  summary = FALSE
)
```

Arguments

| | |
|----------------------------|--|
| <code>sdf</code> | a SigDF |
| <code>switch_failed</code> | whether to switch failed probes (default to FALSE) |
| <code>mask_failed</code> | whether to mask failed probes (default to FALSE) |
| <code>verbose</code> | whether to print correction summary |
| <code>summary</code> | return summarized numbers only. |

Value

a SigDF, or numerics if `summary == TRUE`

Examples

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

inferSex

Infer sex.

Description

We established our sex calling based on the CpGs hypermethylated in inactive X (XiH), CpGs hypomethylated in inactive X (XiL).

Usage

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

Arguments

| | |
|----------|----------------------------------|
| betas | DNA methylation beta |
| platform | EPICv2, EPIC, HM450, MM285, etc. |

Details

Note genotype abnormalities 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.

Value

Inferred sex of sample

Examples

```
## EPICv2 input
betas = openSesame(sesameDataGet("EPICv2.8.SigDF")[[1]])
inferSex(betas)

## Not run:
## MM285 input
betas = openSesame(sesameDataGet("MM285.1.SigDF"))
inferSex(betas)

## EPIC input
betas = openSesame(sesameDataGet('EPIC.1.SigDF'))
inferSex(betas)

## HM450 input
betas = openSesame(sesameDataGet("HM450.10.SigDF")[[1]])
```

```
inferSex(betas)

## End(Not run)
```

| | |
|--------------|----------------------|
| inferSpecies | <i>Infer Species</i> |
|--------------|----------------------|

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

*Infer strain information for mouse array***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'))
```

| | |
|----------|--|
| liftOver | <i>liftOver, see mLiftOver (renamed)</i> |
|----------|--|

Description

liftOver, see mLiftOver (renamed)

Usage

```
liftOver(...)
```

Arguments

... see mLiftOver

Value

imputed data, vector, matrix, SigDF(s)

| | |
|--------------------|--|
| listAvailableMasks | <i>list existing quality masks for a SigDF</i> |
|--------------------|--|

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("EPICv2")
```

| | |
|------------|--|
| mapFileSet | <i>Deposit data of one sample to a fileSet (and hence to file)</i> |
|------------|--|

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 bimodal.

Value

mean of all intensities

Examples

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

| | |
|----------------------|--|
| medianTotalIntensity | <i>Whole-dataset-wide Median Total Intensity (M+U)</i> |
|----------------------|--|

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)
```

| | |
|-----------|--|
| mLiftOver | <i>Lift over beta values or SigDFs to another Infinium platform This function wraps ID conversion and provide optional imputation functionality.</i> |
|-----------|--|

Description

Lift over beta values or SigDFs to another Infinium platform This function wraps ID conversion and provide optional imputation functionality.

Usage

```
mLiftOver(
  x,
  target_platform,
  source_platform = NULL,
  BPPARAM = SerialParam(),
  mapping = NULL,
  impute = FALSE,
  sd_max = 999,
  celltype = "Blood",
  ...
)
```

Arguments

x either named beta value (vector or matrix), probe IDs or SigDF(s) if input is a matrix, probe IDs should be in the row names if input is a numeric vector, probe IDs should be in the vector names. If input is a character vector, the input will be considered probe IDs.

target_platform the platform to take the data to

| | |
|-----------------|---|
| source_platform | optional information of the source data platform (when there might be ambiguity). |
| BPPARAM | use MulticoreParam(n) for parallel processing |
| mapping | a liftOver mapping file. Typically this file contains empirical evidence whether a probe mapping is reliable. If given, probe ID-based mapping will be skipped. This is to perform more stringent probe ID mapping. |
| impute | whether to impute or not, default is FALSE |
| sd_max | the maximum standard deviation for filtering low confidence imputation. |
| celltype | the cell type / tissue context of imputation, if not given, will use nearest neighbor to find out. |
| ... | extra arguments, see ?convertProbeID |

Value

imputed data, vector, matrix, SigDF(s)

Examples

```
## Not run:
sesameDataCache()

## lift SigDF

sdf = sesameDataGet("EPICv2.8.SigDF")[["GM12878_206909630042_R08C01"]]
dim(mLiftOver(sdf, "EPICv2"))
dim(mLiftOver(sdf, "EPIC"))
dim(mLiftOver(sdf, "HM450"))

sdfs = sesameDataGet("EPICv2.8.SigDF")[1:2]
sdfs_hm450 = mLiftOver(sdfs, "HM450")
## parallel processing
sdfs_hm450 = mLiftOver(sdfs, "HM450", BPPARAM=BiocParallel::MulticoreParam(2))

sdf = sesameDataGet("EPIC.5.SigDF.normal")[["1"]]
dim(mLiftOver(sdf, "EPICv2"))
dim(mLiftOver(sdf, "EPIC"))
dim(mLiftOver(sdf, "HM450"))

sdf = sesameDataGet("HM450.10.SigDF")[["1"]]
dim(mLiftOver(sdf, "EPICv2"))
dim(mLiftOver(sdf, "EPIC"))
dim(mLiftOver(sdf, "HM450"))

## lift beta values

betas = openSesame(sesameDataGet("EPICv2.8.SigDF")[["1"]])
betas_hm450 = mLiftOver(betas, "HM450", impute=TRUE)
length(betas_hm450)
sum(is.na(betas_hm450))
```



```

betas_hm450 <- mLiftOver(betas, "HM450", impute=FALSE)
length(betas_hm450)
sum(is.na(betas_hm450))
betas_epic1 <- mLiftOver(betas, "EPIC", impute=TRUE)
length(betas_epic1)
sum(is.na(betas_epic1))
betas_epic1 <- mLiftOver(betas, "EPIC", impute=FALSE)
length(betas_epic1)
sum(is.na(betas_epic1))

betas_matrix = openSesame(sesameDataGet("EPICv2.8.SigDF")[1:4])
dim(betas_matrix)
betas_matrix_hm450 = mLiftOver(betas_matrix, "HM450", impute=T)
dim(betas_matrix_hm450)
## parallel processing
betas_matrix_hm450 = mLiftOver(betas_matrix, "HM450", impute=T,
BPPARAM=BiocParallel::MulticoreParam(4))

## use empirical evidence in mLiftOver
mapping = sesameDataGet("liftOver.EPICv2ToEPIC")
betas_matrix = openSesame(sesameDataGet("EPICv2.8.SigDF")[1:4])
dim(mLiftOver(betas_matrix, "EPIC", mapping = mapping))
## compare to without using empirical evidence
dim(mLiftOver(betas_matrix, "EPIC"))

betas <- c("cg04707299"=0.2, "cg13380562"=0.9, "cg00000103"=0.1)
head(mLiftOver(betas, "HM450", impute=TRUE))

betas <- c("cg00004963_TC21"=0, "cg00004963_TC22"=0.5, "cg00004747_TC21"=1.0)
betas_hm450 <- mLiftOver(betas, "HM450", impute=TRUE)
head(na.omit(mLiftOver(betas, "HM450", impute=FALSE)))

## lift probe IDs

cg_epic2 = names(sesameData_getManifestGRanges("EPICv2"))
head(mLiftOver(cg_epic2, "HM450"))

cg_epic2 = grep("cg", names(sesameData_getManifestGRanges("EPICv2")), value=T)
head(mLiftOver(cg_epic2, "HM450"))

cg_hm450 = grep("cg", names(sesameData_getManifestGRanges("HM450")), value=T)
head(mLiftOver(cg_hm450, "EPICv2"))

rs_epic2 = grep("rs", names(sesameData_getManifestGRanges("EPICv2")), value=T)
head(mLiftOver(rs_epic2, "HM450", source_platform="EPICv2"))

probes_epic2 = names(sesameData_getManifestGRanges("EPICv2"))
head(mLiftOver(probes_epic2, "EPIC"))
head(mLiftOver(probes_epic2, "EPIC", target_uniq = TRUE))
head(mLiftOver(probes_epic2, "EPIC", include_new = FALSE))
head(mLiftOver(probes_epic2, "EPIC", include_old = FALSE))
head(mLiftOver(probes_epic2, "EPIC", return_mapping=TRUE))

```

```
## End(Not run)
```

| | |
|-------------------|--------------------------------------|
| 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 = "",
  min_beads = 1,
  ...
)
```

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 |
| min_beads | minimum bead number, probes with R or G smaller than this threshold will be masked. If NULL, no filtering based on bead count will be applied. Default to 1. |
| ... | parameters to getBetas |

Details

Please use mask=FALSE to turn off masking.

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

```
in_dir <- system.file("extdata", "", package = "sesameData")
betas <- openSesame(in_dir)
## or
IDATprefixes <- searchIDATprefixes(in_dir)
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 | <i>Generate some additional color palettes</i> |
|--------|--|

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 | methylated signal, a numeric vector |
| sigU | unmethylated 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 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 = FALSE, 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
## Not run:
## download age models from
## https://github.com/zhou-lab/InfiniumAnnotationV1/tree/main/Anno
## e.g., Anno/HM450/Clock_Horvath353.rds
predictAge(betas, model)

## End(Not run)
```

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("ctl", "rs"))$mask)
sum(prefixMask(sdf, c("ctl"))$mask)
sum(prefixMask(sdf, c("ctl", "rs", "ch"))$mask)
```

prefixMaskButC *Mask all but C probes in SigDF*

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 | <i>List supported prepSesame functions</i> |
|----------------|--|

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 *Print a fileSet*

Description

Print a fileSet

Usage

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

Arguments

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

Value

string representation

Examples

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

| | |
|--------------------|--|
| 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", verbose = TRUE)
```

Arguments

| | |
|------------|--|
| sdf | a SigDF object |
| mask_names | a vector of masking groups, see listAvailableMasks use "recommended" for recommended masking. One can also combine "recommended" with other masking groups by specifying a vector, e.g., c("recommended", "M_mapping") |
| verbose | be verbose |

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)$mask)

## list available masks, the dbname column
listAvailableMasks(sdfPlatform(sdf))
listAvailableMasks("EPICv2")
```

| | |
|-------------|--|
| 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,  
  manifest = NULL,  
  platform = "",  
  min_beads = NULL,  
  controls = NULL,  
  verbose = FALSE  
)
```

Arguments

| | |
|--------------------------|--|
| <code>prefix.path</code> | sample prefix without <code>_Grn.idat</code> and <code>_Red.idat</code> |
| <code>manifest</code> | optional design manifest file |
| <code>platform</code> | EPIC, HM450 and HM27 etc. |
| <code>min_beads</code> | minimum bead number, probes with R or G smaller than this threshold will be masked. If <code>NULL</code> , no filtering based on bead count will be applied. |
| <code>controls</code> | optional control probe manifest file |
| <code>verbose</code> | be verbose? (<code>FALSE</code>) |

Value

a `SigDF`

Examples

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

recommendedMaskNames *Recommended mask names for each Infinium platform*

Description

The returned name is the db name used in KYCG.mask

Usage

```
recommendedMaskNames()
```

Value

a named list of mask names

Examples

```
recommendedMaskNames()[["EPICv2"]]
recommendedMaskNames()[["EPIC"]]
```

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 | <i>Reset Masking</i> |
|-----------|----------------------|

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 *SCRUB background correction*

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 *SCRUB background correction*

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_attachManifest | <i>Annotate a data.frame using manifest</i> |
|---------------------------|---|

Description

Annotate a data.frame using manifest

Usage

```
sesameAnno_attachManifest(
  df,
  probe_id = "Probe_ID",
  platform = NULL,
  genome = NULL
)
```

Arguments

| | |
|----------|---|
| df | input data frame with Probe_ID as a column |
| probe_id | the Probe_ID column name, default to "Probe_ID" or rownames |
| platform | which array platform, guess from probe ID if not given |
| genome | the genome build, use default if not given |

Value

a new data.frame with manifest attached

Examples

```
## Not run:
df <- data.frame(Probe_ID = c("cg00101675_BC21", "cg00116289_BC21"))
sesameAnno_attachManifest(df)

## End(Not run)
```

```
sesameAnno_buildAddressFile
```

Build sesame ordering address file from tsv

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

Examples

```
## Not run:
tsv = sesameAnno_download("HM450.hg38.manifest.tsv.gz")
addr <- sesameAnno_buildAddressFile(tsv)

## End(Not run)
```

sesameAnno_buildManifestGRanges
Build manifest GRanges from tsv

Description

manifest tsv files can be downloaded from <http://zwdzwd.github.io/InfiniumAnnotation>

Usage

```
sesameAnno_buildManifestGRanges(  
  tsv,  
  genome = NULL,  
  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

Examples

```
## Not run:
tsv = sesameAnno_download("HM450.hg38.manifest.tsv.gz")
gr <- sesameAnno_buildManifestGRanges(tsv)
## direct access
gr <- sesameAnno_buildManifestGRanges("HM450.hg38.manifest")

## End(Not run)
```

sesameAnno_download *Download SeSAmE annotation files*

Description

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

Usage

```
sesameAnno_download(url, destfile = tempfile(basename(url)))
```

Arguments

| | |
|----------|---|
| url | url or title of the annotation file |
| destfile | download to this file, a temp file if unspecified |

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

the path to downloaded file

Examples

```
## Not run:  
## avoid testing as this function uses external host  
sesameAnno_download("Test/3999492009_R01C01_Grn.idat")  
sesameAnno_download("EPIC.hg38.manifest.tsv.gz")  
sesameAnno_download("EPIC.hg38.snp.tsv.gz")  
  
## End(Not run)
```

sesameAnno_readManifestTSV
Read manifest file to a tsv format

Description

Read manifest file to a tsv format

Usage

```
sesameAnno_readManifestTSV(tsv_fn)
```

Arguments

tsv_fn tsv file path

Value

a manifest as a tibble

Examples

```
## Not run:  
tsv = sesameAnno_download("HM450.hg38.manifest.tsv.gz")  
mft <- sesameAnno_readManifestTSV(tsv)  
## direct access  
mft <- sesameAnno_readManifestTSV("HM450.hg38.manifest")  
  
## End(Not run)
```

sesameQC-class *An S4 class to hold QC statistics*

Description

An S4 class to hold QC statistics

Value

sesameQC object

Slots

stat a list to store qc stats

| | |
|--------------|---|
| sesameQCtoDF | <i>Convert a list of sesameQC to data frame</i> |
|--------------|---|

Description

Convert a list of sesameQC to data frame

Usage

```
sesameQCtoDF(qcs, cols = c("frac_dt_cg", "RGdistort", "RGratio"))
```

Arguments

| | |
|------|------------------------------------|
| qcs | sesameQCs |
| cols | QC columns, use NULL to report all |

Value

a data frame

Examples

```
sdf <- sesameDataGet("EPIC.1.SigDF")
qcs <- sesameQC_calcStats(sdf, "detection")
sesameQCtoDF(qcs)
```

| | |
|--------------------|--------------------------------|
| 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 *Get stat numbers from an sesameQC object*

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),  
  pal = "whiteturbo",  
  ...  
)
```

Arguments

| | |
|---------------------------|---|
| <code>sdf</code> | a SigDF |
| <code>mask</code> | whether to remove probes that are masked |
| <code>use_max</code> | to use max(M,U) or M+U |
| <code>intens.range</code> | plot range of signal intensity |
| <code>pal</code> | color palette, whiteturbo, whiteblack, whitejet |
| <code>...</code> | 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 message text for deprecated function

Examples

```
cat("Deprecated. see https://github.com/zwdzwd/sesamize")
```

| | |
|---------|--|
| setMask | <i>Set mask to only the probes specified</i> |
|---------|--|

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, MU = FALSE)
```

Arguments

sdf a SigDF
mask whether to apply mask
MU add a column for M+U

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()
```

| | |
|------------------|------------------------------|
| totalIntensities | <i>M+U Intensities Array</i> |
|------------------|------------------------------|

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

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

Value

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

| | |
|-------------|--|
| updateSigDF | <i>Set color and mask using strain/species-specific manifest</i> |
|-------------|--|

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('Mammal40.1.SigDF')  
sdf_mouse <- updateSigDF(sdf, species="mus_musculus")
```

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

Description

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

Usage

```
visualizeGene(
  gene_name,
  betas,
  platform = NULL,
  genome = NULL,
  upstream = 2000,
  dwstream = 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 |
| dwstream | 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

Visualize Region that Contains the Specified Probes

Description

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

Usage

```
visualizeProbes(  
  probeNames,  
  betas,  
  platform = 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(
  chr,
  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

| | |
|------------------------------|---|
| <code>chr</code> | chromosome |
| <code>beg</code> | begin of the region |
| <code>end</code> | end of the region |
| <code>betas</code> | beta value matrix (row: probes, column: samples) |
| <code>platform</code> | EPIC, HM450, or MM285 |
| <code>genome</code> | 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 ..., <code>genome = sesameAnno_buildManifestGRanges("downloaded_file"),...</code> to this function. |
| <code>draw</code> | draw figure or return betas |
| <code>cluster.samples</code> | whether to cluster samples |
| <code>na.rm</code> | remove probes with all NA. |
| <code>nprobes.max</code> | maximum number of probes to plot |
| <code>txn.types</code> | default to <code>protein_coding</code> , use NULL for all |
| <code>txn.font.size</code> | transcript name font size |
| <code>...</code> | additional options, see <code>assemble_plots</code> |

Value

graphics or a matrix containing the captured beta values

Examples

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

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

Description

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

Usage

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

Arguments

| | |
|----------------|--|
| seg | a CNSegment object |
| to.plot | chromosome to plot (by default plot all chromosomes) |
| genes.to.label | gene(s) to label |

Details

require ggplot2, scales

Value

plot graphics

Examples

```
sesameDataCache()
## Not run:
sdfs <- sesameDataGet('EPICv2.8.SigDF')
sdf <- sdfs[["K562_206909630040_R01C01"]]
seg <- cnSegmentation(sdf)
seg <- cnSegmentation(sdf, return.probe.signals=TRUE)
visualizeSegments(seg)
visualizeSegments(seg, to.plot=c("chr9","chr22"))
visualizeSegments(seg, genes.to.label=c("ABL1","BCR"))

## End(Not run)

sesameDataGet_resetEnv()
```

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