methylumi

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extractBarcodeAndPosition

Extract the Barcode and Position Information from Sentrix ID

Description

The sentrix IDs from an illumina sentrix array contain positional information that might be useful. This function simply extracts that information from the ID itself.

Usage

```
extractBarcodeAndPosition(sentrixids)
```

Arguments

sentrixids A character vector of sentrix IDs that look like: 1632405013_R001_C001

Value

A data.frame with three columns:

sentrix	numeric, the sentrix ID
row	numeric, the sentrix row
column	numeric, the sentrix column

Author(s)

Sean Davis <sdavis2@mail.nih.gov>

See Also

methylumiR

```
extractBarcodeAndPosition(c('12341234_R001_C001'))
```

MethyLumi-accessors

methylumi accessors

Description

These functions serve as getters and setters for information in methylumi classes.

Usage

```
betas(object)
pvals(object)
methylated(object)
unmethylated(object)
getHistory(object)
QCdata(object)
```

Arguments

object an object of class MethyLumi or a subclass

Details

See the methods definitions in MethyLumiSet and MethyLumiQC for details.

Author(s)

Sean Davis <sdavis2@mail.nih.gov>

See Also

normalizeMethyLumiSet, MethyLumiSet, MethyLumiQC, eSet

getAssayDataNameSubstitutions

Return a data.frame of AssayData name substitutions.

Description

The Illumina methylation platforms use two distinct platforms, the "goldengate" platform and the "infinium" platform. Each of these uses different file formats as well as different assay techologies. To make the downstream data handling more straightforward and uniform between the two different systems, a simple mapping from the column names in the output files from the Illumina software is used to convert things from Red/Green or Cy5/Cy3 to unmethylated/methylated. This function simply returns that mapping.

Usage

getAssayDataNameSubstitutions()

MethyLumi-class

Details

A file in the extdata directory called "substitutions.txt" contains two columns. The function loads this file and uses the first column as a match against column names in the data file (with the "sample part" removed). If matched, the second column gives the replacement.

Value

A data.frame with two columns, regex and replacement.

Author(s)

Sean Davis <seandavi@gmail.com>

Examples

getAssayDataNameSubstitutions()

MethyLumi-class The base class for storing Illumina Methylation data

Description

This class inherits from eSet from the Biobase package and is used as a base class for the other two methylumi classes, MethyLumiSet and MethyLumiQC.

Objects from the Class

The MethyLumi class is a virtual class and is not meant to be instantiated. Instead, one should instantiate a MethyLumiSet or a MethyLumiQC object.

Slots

assayData: Object of class "AssayData"

phenoData: Object of class "AnnotatedDataFrame"

- featureData: Object of class "AnnotatedDataFrame" that will hold the annotation columns from the Beadstudio output, if they are available.
- experimentData: Object of class "MIAME"
- annotation: Object of class "character"; note that this slot is not currently used, but may be used in the future to store the character name of the annotation package, if available.

.___classVersion__: Object of class "Versions"

Extends

Class "eSet", directly. Class "VersionedBiobase", by class "eSet", distance 2. Class "Versioned", by class "eSet", distance 3.

Methods

- pvals<- signature(object = "MethyLumi", value = "matrix"): Set the assay-Data slot of the same name and stores the P-values from BeadStudio
- pvals signature(object = "MethyLumi"): Get the assayData slot of the same name
- betas<- signature(object = "MethyLumi", value = "matrix"): Set the assay-Data slot of the same name and represents the methylation values for the samples, analogous to exprs() in gene expression data.
- betas signature (object = "MethyLumi"): Get the assayData slot of the same name
- methylated<- signature(object = "MethyLumi", value = "matrix"): Set the assayData slot that represents the Methylated single-channel signal
- methylated signature(object = "MethyLumi"): Get the assayData slot that represents
 the Methylated single-channel signal
- unmethylated<- signature(object = "MethyLumi", value = "matrix"): Set the
 assayData slot that represents the Unmethylated single-channel signal</pre>
- unmethylated signature(object = "MethyLumi"): Get the assayData slot that represents the Unmethylated single-channel signal
- controlTypes signature(object = "MethyLumi": Find the unique control type beeds in the QCdata slot.
- **qcplot** signature (object = "MethyLumi", what, ...): Plot of QC data. This plot can be useful for diagnosing the problems associated with specific samples or arrays. The value for "what" is one of the control types (which can be found by using controlTypes() on the object.

Author(s)

Sean Davis <sdavis2@mail.nih.gov>

See Also

methylumiR, MethyLumiSet, MethyLumiQC, eSet

```
## The class structure
showClass("MethyLumi")
## read in some data
## Read in sample information
samps <- read.table(system.file("extdata/samples.txt",</pre>
                                package = "methylumi"), sep="\t", header=TRUE)
## Perform the actual data reading
## This is an example of reading data from a
## Sentrix Array format file (actually two files,
## one for data and one for QC probes)
mldat <- methylumiR(system.file('extdata/exampledata.samples.txt',</pre>
        package='methylumi'),
      qcfile=system.file('extdata/exampledata.controls.txt',
        package="methylumi"),
      sampleDescriptions=samps)
mldat
```

MethyLumiQC-class

```
## Get history information
getHistory(mldat)
## Get QC data, which is another eSet-derived object
QCdata(mldat)
```

methylumi-package Handle Illumina methylation data

Description

This package contains a class structure for handling methylation data from Illumina as well as utility functions for loading the data from files generated by Illumina. Normalization that attempts to correct for dye bias is also included.

Important data classes include: MethyLumiSet and MethyLumiQC, both of which are subsets of the MethyLumi class, which is a subset of the eSet class.

A worked example of the use of the package can be found by typing: openVignette().

A full listing of the available documentation can be obtained by typing help.start() and selecting methylumi from the Packages link or by typing library (help="methylumi").

Details

```
Package: methylumi
Type: Package
License: GPL
```

Author(s)

Sean Davis <sdavis2@mail.nih.gov>

References

http://watson.nci.nih.gov/~sdavis/software/R

See Also

Biobase

MethyLumiQC-class Class to hold Illumina Methylation QC data

Description

This class inherits from the MethyLumi class (and therefore, from eSet in Biobase) and is designed to hold QC data from Illumina Beadstudio output. These data can be potentially useful when determining the cause for quality problems.

Objects from the Class

```
Objects can be created by calls of the form new ("MethyLumiQC", assayData, phenoData, featureData, experimentData, annotation, betas).
```

Slots

assayData: Object of class "AssayData"

phenoData: Object of class "AnnotatedDataFrame"

featureData: Object of class "AnnotatedDataFrame" containing the annotation columns from the Illumina Beadstudio output. In particular, the names of the probes describe the types of control probes.

experimentData: Object of class "MIAME"

annotation: Object of class "character", not currently used

.___classVersion__: Object of class "Versions"

Extends

Class "MethyLumi", directly. Class "QCDataOrNULL", directly. Class "eSet", by class "MethyLumi", distance 2. Class "VersionedBiobase", by class "MethyLumi", distance 3. Class "Versioned", by class "MethyLumi", distance 4.

Methods

```
qcplot signature(object = "MethyLumiQC", what, ...): QC plots of various con-
troltypes
```

initialize signature(.Object = "MethyLumiQC")

Author(s)

Sean Davis <sdavis2@mail.nih.gov>

See Also

methylumiR, MethyLumiSet, MethyLumiQC, eSet

```
showClass("MethyLumiQC")
```

methylumiR

Description

This function is useful for loading Illumina methylation data into a MethyLumiSet object. Sample information can be supplied and will then be incorporated into the resulting phenoData slot.

Usage

```
methylumiR(filename,qcfile=NULL,sampleDescriptions = NULL,...)
```

Arguments

filename	A filename of the excel-like file from BeadStudio	
qcfile	A filename of the excel-like file from BeadStudio	
sampleDescriptions		
	A data frame that contains at least one column, SampleID (case insensitive).	
	This column MUST match the part of the column headers before the . ${\tt Avg_Beta},$	
	etc. Also, if a column called SampleLabel (case insensitive), it is used for sam-	
	ple labels, IF the sampleLabel column contains unique identifiers	
	Passed into read.delim()	

Details

Used to construct a MethyLumiSet object

Value

A MethyLumiSet object

Author(s)

Sean Davis <sdavis2@mail.nih.gov>

See Also

MethyLumiSet-class, MethyLumiQC-class

Examples

mldat

MethyLumiSet-class Class to hold Illumina Methylation data

Description

This class inherits from the MethyLumi class (and therefore, from eSet in Biobase) and is designed to hold both the intensities and the calculated betas, as well as pvalues if present.

Objects from the Class

Objects can be created by calls of the form new ("MethyLumiSet", assayData, phenoData, featureData, experimentData, annotation, betas). An object of this type is the main storage class for methylation data from Illumina. Subsetting, etc., works as normal (rows represent genes, columns represent samples). There is also a rudimentary history tracking system that is modeled after that from the lumi package.

Slots

QC: Object of class "QCDataOrNULL", containing either the MethyLumiQC object or NULL

- history: Object of class "data.frame", containing a running history of transforms to the data contained herein
- assayData: Object of class AssayData
- phenoData: Object of class AnnotatedDataFrame
- featureData: Object of class AnnotatedDataFrame, containing the annotation columns from the Illumina Beadstudio output
- experimentData: Object of class MIAME

annotation: Object of class "character", not currently used

.___classVersion__: Object of class "Versions"

Extends

Class "MethyLumi", directly. Class "eSet", by class "MethyLumi", distance 2. Class "VersionedBiobase", by class "MethyLumi", distance 3. Class "Versioned", by class "MethyLumi", distance 4.

Methods

```
[ signature(x = "MethyLumiSet"): subsetting, genes as rows, samples as columns
betas<- signature(object = "MethyLumiSet", value = "matrix"): Set the as-
sayData slot of the same name
betas signature(object = "MethyLumiSet"): Get the assayData slot of the same name
boxplot signature(x = "MethyLumiSet"): boxplot of all sample betas
combine signature(x = "MethyLumiSet", y = "MethyLumiSet")
corplot signature(x = "MethyLumiSet")
exprs<- signature(object = "MethyLumiSet", value = "matrix"): alias for "<-
betas"
exprs signature(object = "MethyLumiSet"): alias for "betas"
```

MethyLumiSet-class

- getHistory signature(object = "MethyLumiSet"): returns a data.frame containing the
 history for this object
- **hist** signature (x = "MethyLumiSet"): histogram of the betas for the data
- initialize signature(.Object = "MethyLumiSet")
- pairs signature(x = "MethyLumiSet"): pairs plot of the betas for the object. Note that
 pairs plots of more than a few samples are not very useful.
- **plotSampleIntensities** signature (x = "MethyLumiSet"): The intensities as output by the Beadstudio software often show a considerable amount of dye bias. This method shows a graphical example of this dye bias. In short, for each of the Cy3 and Cy5 channels, a cutoff in beta is used to calculate which Cy3 and Cy5 values should be plotted at high-methylation and low-methylation status. Any offset between Cy3 and Cy5 when plotted in this way likely represents dye bias and will lead to biases in the estimate of beta.
- QCdata<- signature(object = "MethyLumiSet", value = "MethyLumiQC"):assign QC data to the QC slot
- **QCdata** signature (object = "MethyLumiSet"): retrieve the QC data.
- show signature(object = "MethyLumiSet")
- methylated<- signature(object = "MethyLumiSet", value = "matrix"): Set
 the assayData slot associated with methylated intensity</pre>
- methylated signature(object = "MethyLumiSet"): Get the assayData slot associated
 with methylated intensity
- unmethylated<- signature(object = "MethyLumiSet", value = "matrix"): Set
 the assayData slot associated with unmethylated intensity</pre>
- unmethylated signature(object = "MethyLumiSet"): Get the assayData slot associated with unmethylated intensity
- controlTypes signature(object = "MethyLumiSet"): determine the character vector
 of control types from the QCdata information

Author(s)

Sean Davis <sdavis2@mail.nih.gov>

See Also

methylumiR, normalizeMethyLumiSet, MethyLumiSet, MethyLumiQC, eSet

Examples

showClass("MethyLumiSet")

mldat

Description

This is an example MethyLumiSet object.

Usage

data(mldat)

Examples

data(mldat)

normalizeMethyLumiSet

Normalize a MethyLumiSet, accounting for dye bias

Description

The Illumina methylation platform uses two colors, one to represent the unmethylated state and the other to represent the methylated state. This function corrects that dye bias and recalculates the betas based on the corrected intensities.

Usage

```
normalizeMethyLumiSet(x, beta.cuts = c(0.2, 0.8), mapfun = c("atan", "ratio"))
```

Arguments

Х	A MethyLumiSet object
beta.cuts	Two numeric values with the first less than the second and between 0 and 1, representing the beta cutoffs that will be used when determining the median intensities to which to correct. See details below.
mapfun	Either "atan" or "ratio". See details below.

Details

The Illumina methylation platform uses two colors, one to represent the unmethylated state and the other to represent the methylated state. This function corrects that dye bias and recalculates the betas based on the corrected intensities.

As a first step, the medians for each of Cy3 and Cy5 are calculated at high and low betas, representing the (nearly) fully methylated state and the (nearly) fully unmethylated states. Values of Cy3 and Cy5 that are negative are set to zero for this process. Then, the Cy5 medians are adjusted to match those of the Cy3 channel, thereby correcting the dye bias.

To map the new intensities back to betas, one of two map functions can be used. The default is the atan(Cy3/Cy5). The ratio maps using the function (Cy3/Cy3+Cy5). The differences should be very small, but we feel that the atan map function is probably the mathematically appropriate way of doing this.

Value

A new "MethyLumiSet" that contains the corrected betas and the adjusted intensities.

Author(s)

Sean Davis <sdavis2@mail.nih.gov>

Examples

```
plotSampleIntensities
```

Plot the sample intensities.

Description

The Illumina methylation platforms all show a significant dye bias. The plotSampleIntensities method shows the density plots for the two channels allowing direct visualization of the effect.

Usage

plotSampleIntensities(x, beta.cuts, s)

Arguments

Х	an object of class MethyLumi or a subclass
beta.cuts	cutoffs for low and high beta values
S	sample number to plot

```
data(mldat)
plotSampleIntensities(mldat, s=1)
```

qcplot

Description

The qcplot function simply generates a plot of the control probe information for a given controlType.

Usage

```
qcplot(object, controltype, ...)
controlTypes(object, ...)
```

Arguments

object	An object of class MethyLumiSet or MethyLumiQC
controltype	A single character value representing the bead type to plot from the quality con- trol data. The available types are accessible via the controlTypes method.
	passed to plot function

Details

The descriptions of the various control types can be obtained from the Illumina methylation user's guides.

Author(s)

Sean Davis <sdavis2@mail.nih.gov>

See Also

MethyLumiSet, MethyLumiQC

```
data(mldat)
controlTypes(mldat)
qcplot(mldat,controlTypes(mldat)[3])
```

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