

Package ‘normalize450K’

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Type Package

Title Preprocessing of Illumina Infinium 450K data

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Description Precise measurements are important for epigenome-wide studies investigating DNA methylation in whole blood samples, where effect sizes are expected to be small in magnitude. The 450K platform is often affected by batch effects and proper preprocessing is recommended. This package provides functions to read and normalize 450K '.idat' files. The normalization corrects for dye bias and biases related to signal intensity and methylation of probes using local regression. No adjustment for probe type bias is performed to avoid the trade-off of precision for accuracy of beta-values.

Depends R (>= 3.3), Biobase, illuminaio

Suggests minfiData, data.table

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biocViews Normalization, DNAMethylation, Microarray, TwoChannel, Preprocessing, MethylationArray

NeedsCompilation no

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read_and_normalize450K

Normalization of 450K data by LOESS method

Description

Read 450K '.idat' files and compute raw or normalized beta-values.

Usage

```
read450K(idat_files)
normalize450K(intensities)
dont_normalize450K(intensities)
```

Arguments

idat_files	a character vector containing the paths to the .idat files stripped from the '_Grn.idat' suffix with one entry for each sample (.idat files for green and red intensities have to be in the same folder).
intensities	List object containing raw signal intensities. Result of calling read450K.

Details

Function read450K reads .idat files and returns a list object containing raw signal intensities. dont_normalize450K returns an ExpressionSet containing beta-values without normalization. normalize450K performs dye bias correction using the extension controls probes followed by normalization by local regression (Heiss and Brenner, 2015) and returns an ExpressionSet containing beta-values, too.

Value

For read450K a list containing the methylated, unmethylated and control signal intensities. For dont_normalize450K and normalize450K an ExpressionSet containing beta-values, rows corresponding to CpG sites (named) and columns to samples (in the same order as 'idat_files').

Note

A benchmark comparing the performance of this method with other normalization approaches is provided in the vignette.

Author(s)

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References

Heiss, Jonathan A., and Hermann Brenner. *Between-array normalization for 450K data*. *Frontiers in genetics* 6 (2015). doi:[10.3389/fgene.2015.00092](https://doi.org/10.3389/fgene.2015.00092)

Examples

```
library(minfiData) ## this package includes some .idat files
library(data.table)

path <- system.file("extdata",package="minfiData")
samples = fread(file.path(path, 'SampleSheet.csv'),integer64='character')

samples[,file:=file.path(path,Sentrix_ID,paste0(Sentrix_ID,'_',Sentrix_Position))]
## samples$file is a character vector containing the location of the
## .idat files, but without the suffixes "_Red.idat" or "_Grn.idat"

raw = read450K(samples$file)
none = dont_normalize450K(raw) ## no normalization
norm = normalize450K(raw)
```

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