mBPCR

March 24, 2012

computeMBPCR

Estimate the copy number profile

Description

Function to estimate the copy number profile with a piecewise constant function using mBPCR. Eventually, it is possible to estimate the profile with a smoothing curve using either the Bayesian Regression Curve with K_2 (BRC with K_2) or the Bayesian Regression Curve Averaging over k (BRCAk). It is also possible to choose the estimator of the variance of the levels rhoSquare (i.e. either $\hat{\rho}_1^2$ or $\hat{\rho}^2$) and by default $\hat{\rho}_1^2$ is used.

Usage

computeMBPCR(y, kMax=50, nu=NULL, rhoSquare=NULL, sigmaSquare=NULL, typeEstRho=

Arguments

У	array containing the log2ratio of the copy number data
kMax	maximum number of segments
nu	mean of the segment levels. If $\verb"nu=NULL"$, then the algorithm estimates it on the sample.
rhoSquare	variance of the segment levels. If ${\tt rhoSquare=NULL}$, then the algorithm estimates it on the sample.
sigmaSquare	variance of the noise. If ${\tt sigmaSquare=NULL}$, then the algorithm estimates it on the sample.
typeEstRho	choice of the estimator of rhoSquare. If typeEstRho=1, then the algorithm estimates rhoSquare with $\hat{\rho}_1^2$, while if typeEstRho=0, it estimates rhoSquare with $\hat{\rho}^2$.
regr	choice of the computation of the regression curve. If <code>regr=NULL</code> , then the regression curve is not computed, if <code>regr="BRC"</code> the Bayesian Regression Curve with K_2 is computed (BRC with K_2), if <code>regr="BRCAk"</code> the Bayesian Regression Curve Averaging over k is computed (BRCAk).

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Details

By default, the function estimates the copy number profile with mBPCR and estimating rhoSquare on the sample, using $\hat{\rho}_1^2$. It is also possible to use $\hat{\rho}^2$ as estimator of rhoSquare, by setting typeEstRho=0, or to directly set the value of the parameter.

The function gives also the possibility to estimate the profile with a Bayesian regression curve: if regr="BRC" the Bayesian Regression Curve with K_2 is computed (BRC with K_2), if regr="BRCAk" the Bayesian Regression Curve Averaging over k is computed (BRCAk).

Value

A list containing:

```
estK the estimated number of segments
estBoundaries
the estimated boundaries
estPC the estimated profile with mBPCR
regrCurve the estimated bayesian regression curve. It is returned only if regr!=NULL.
nu
rhoSquare
sigmaSquare
postProbT for each probe, the posterior probablity to be a breakpoint
```

References

```
Rancoita, P. M. V., Hutter, M., Bertoni, F., Kwee, I. (2009). Bayesian DNA copy number analysis. BMC Bioinformatics 10: 10. http://www.idsia.ch/~paola/mBPCR
```

See Also

```
estProfileWithMBPCR, plotEstProfile, writeEstProfile, estGlobParam
```

#p <- jekoChr11Array250Knsp\$PhysicalPosition[10600:11600]</pre>

Examples

```
##import the 250K NSP data of chromosome 11 of cell line JEKO-1
data(jekoChr11Array250Knsp)

##first example
## we select a part of chromosome 11
y <- jekoChr11Array250Knsp$log2ratio[6400:6900]
p <- jekoChr11Array250Knsp$PhysicalPosition[6400:6900]
##we estimate the profile using the global parameters estimated on the whole genome
##the profile is estimated with mBPCR and with the Bayesian Regression Curve
results <- computeMBPCR(y, nu=-3.012772e-10, rhoSquare=0.0479, sigmaSquare=0.0699, regr='
plot(p, y)
points(p, results$estPC, type='l', col='red')
points(p, results$regrCurve,type='l', col='green')

###second example
### we select a part of chromosome 11
#y <- jekoChr11Array250Knsp$log2ratio[10600:11600]</pre>
```

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```
###we estimate the profile using the global parameters estimated on the whole genome
###the profile is estimated with mBPCR and with the Bayesian Regression Curve Ak
#results <- computeMBPCR(y, nu=-3.012772e-10, rhoSquare=0.0479, sigmaSquare=0.0699, regr=
#plot(p,y)
#points(p, results$estPC, type='l', col='red')
#points(p, results$regrCurve, type='l', col='green')</pre>
```

estGlobParam

Estimate global parameters of copy number data

Description

Function to estimate the global parameters of copy number data: the mean and the variance of the segment levels (called nu and rhoSquare, respectively), the variance of the noise (sigmaSquare). It is possible to choose the estimator of rhoSquare (i.e. either $\hat{\rho}_1^2$ or $\hat{\rho}^2$) and by default $\hat{\rho}_1^2$ is used.

Usage

```
estGlobParam(y, nu=NULL, rhoSquare=NULL, sigmaSquare=NULL, typeEstRho=1)
```

Arguments

У	array containing the log2ratio of the copy number data
nu	mean of the segment levels. If $\verb"nu=NULL"$, then the algorithm estimates it on the sample.
rhoSquare	variance of the segment levels. If ${\tt rhoSquare=NULL}$, then the algorithm estimates it on the sample.
sigmaSquare	variance of the noise. If ${\tt sigmaSquare=NULL}$, then the algorithm estimates it on the sample.
typeEstRho	choice of the estimator of rhoSquare. If typeEstRho=1, then the algorithm estimates rhoSquare with $\hat{\rho}_1^2$, while if typeEstRho=0, it estimates rhoSquare with $\hat{\rho}^2$.

Value

A list containing:

nu rhoSquare sigmaSquare

References

Rancoita, P. M. V., Hutter, M., Bertoni, F., Kwee, I. (2009). Bayesian DNA copy number analysis. *BMC Bioinformatics* 10: 10. http://www.idsia.ch/~paola/mBPCR

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Examples

```
##import the 10K data of cell line REC
data(rec10k)
##estimation of all the global parameters (the variance of the segment is estimated with
estGlobParam(rec10k$log2ratio)
```

estProfileWithMBPCR

Estimate and print the copy number profile of some chromosomes of a sample

Description

Function to estimate the copy number profile with a piecewise constant function using mBPCR. Eventually, it is possible to estimate the profile with a smoothing curve, using either the Bayesian Regression Curve with K_2 (BRC with K_2) or the Bayesian Regression Curve Averaging over k (BRCAk). It is also possible to choose the estimator of the variance of the levels rhoSquare (i.e. either $\hat{\rho}_1^2$ or $\hat{\rho}^2$) and by default $\hat{\rho}_1^2$ is used.

Usage

```
estProfileWithMBPCR(snpName, chr, position, logratio, chrToBeAnalyzed, maxProbrestProfileWithMBPCR(snpName, chr, position, logratio, chr, p
```

Arguments

snpName array containing the name of each probe

chr array containing the name of the chromosome to which each of the probes be-

longs. The possible values of the elements of chr are: the integers from 1 to

22, 'X' and 'Y'.

position array containing the physical position of each probe

logratio array containing the log2ratio of the raw copy number data

chrToBeAnalyzed

array containing the name of the chromosomes that the user wants to analyze. The possible values of the chromosomes are: the integers from 1 to 22, 'X' and

'Y'.

maxProbeNumber

maximum number of probes that a chromosome (or arm of a chromosome) can have to be analyzed. The procedure of profile estimation needs the computation of an array of length (length(chromosome)+1)*(length(chromosome)+2)/2. To be sure to have set this parameter correctly, try to create the array A <-array(1, dim=(maxProbeNumber+1)*(maxProbeNumber+2)/2),

before starting with the estimation procedure.

rhoSquare variance of the segment levels. If rhoSquare=NULL, then the algorithm esti-

mates it on the sample.

kMax maximum number of segments

nu mean of the segment levels. If nu=NULL, then the algorithm estimates it on the

sample.

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 $\verb|sigmaSquare=NULL|, then the algorithm estimates it \\$

on the sample.

typeEstRho choice of the estimator of rhoSquare. If typeEstRho=1, then the algo-

rithm estimates rhoSquare with $\hat{\rho}_1^2$, while if typeEstRho=0, it estimates

rhoSquare with $\hat{\rho}^2$.

regr choice of the computation of the regression curve. If regr=NULL, then the

regression curve is not computed, if regr="BRC" the Bayesian Regression Curve is computed (BRC with K_2), if regr="BRCAk" the Bayesian Regres-

sion Curve Averaging over k is computed (BRCAk).

Details

By default, the function estimates the copy number profile with mBPCR and estimating rhoSquare on the sample, using $\hat{\rho}_1^2$. It is also possible to use $\hat{\rho}^2$ as estimator of rhoSquare, by setting typeEstRho=0, or to directly set the value of the parameter.

The function gives also the possibility to estimate the profile with a Bayesian regression curve: if regr="BRC" the Bayesian Regression Curve with K_2 is computed (BRC with K_2), if regr="BRCAk" the Bayesian Regression Curve Averaging over k is computed (BRCAk).

See function writeEstProfile, to have the results in nicer tables or to write them on files.

Value

A list containing:

estPC an array containing the estimated profile with mBPCR

estBoundaries

the list of estimated breakpoints for each of the analyzed chomosomes

postProbT the list of the posterior probablity to be a breakpoint for each estimated break-

point of the analyzed chomosomes

regrCurve an array containing the estimated bayesian regression curve

estPC and regrCurve have the same length of logratio, hence their components, corresponding to the not analyzed chromosomes, are equal to NA.

References

Rancoita, P. M. V., Hutter, M., Bertoni, F., Kwee, I. (2009). Bayesian DNA copy number analysis. *BMC Bioinformatics* 10: 10. http://www.idsia.ch/~paola/mBPCR

See Also

plotEstProfile, writeEstProfile, computeMBPCR

Examples

```
##import the 10K data of cell line REC
data(rec10k)
##estimation of the profile of chromosome 5
results <- estProfileWithMBPCR(rec10k$SNPname, rec10k$Chromosome, rec10k$PhysicalPosition
##plot the estimated profile of chromosome 5
y <- rec10k$log2ratio[rec10k$Chromosome == 5]
p <- rec10k$PhysicalPosition[rec10k$Chromosome == 5]
plot(p, y)</pre>
```

```
points(p, results$estPC[rec10k$Chromosome == 5], type='l', col='red')
###for the estimation of the profile of all chromosomes
#results <- estProfileWithMBPCR(rec10k$SNPname, rec10k$Chromosome, rec10k$PhysicalPosition</pre>
```

estProfileWithMBPCRforOligoSnpSet

Estimate and print the copy number profile of some chromosomes of samples in an oligoSnpSet object

Description

Function to estimate the copy number profile with a piecewise constant function using mBPCR. Eventually, it is possible to estimate the profile with a smoothing curve, using either the Bayesian Regression Curve with K_2 (BRC with K_2) or the Bayesian Regression Curve Averaging over k (BRCAk). It is also possible to choose the estimator of the variance of the levels rhoSquare (i.e. either $\hat{\rho}_1^2$ or $\hat{\rho}_2^2$) and by default $\hat{\rho}_1^2$ is used.

Usage

estProfileWithMBPCRforOligoSnpSet(sampleData, sampleToBeAnalyzed, chrToBeAnalyrnosquare=NULL, kMax=50, nu=NULL, sigmaSquare=NULL, typeEs

Arguments

sampleData

object of type oligoSnpSet. The following fields must not be empty: assayData(sampleData) \$ (it contains the raw copy number values), featureNames (featureData(sampleData)) (it contains the names of the SNPs), featureData(sampleData) \$chromosome (it contains the names of the chromosomes to which each of the SNPs belongs), featureData(sampleData) \$position (it contains the physical positions of the SNPs).

sampleToBeAnalyzed

vector containing the number of the columns corresponding to the samples the user wants to analyze.

chrToBeAnalyzed

array containing the name of the chromosomes that the user wants to analyze. The possible values of the chromosomes are: the integers from 1 to 22, 'X' and 'Y'.

maxProbeNumber

maximum number of probes that a chromosome (or arm of a chromosome) can have to be analyzed. The procedure of profile estimation needs the computation of an array of length (length(chromosome)+1)*(length(chromosome)+2)/2. To be sure to have set this parameter correctly, try to create the array A <-array(1, dim=(maxProbeNumber+1)*(maxProbeNumber+2)/2), before starting with the estimation procedure.

ifLogRatio

denotes if the data are either the log2ratio of raw copy number data or raw copy number data. By default, they are considered as log2ratio data, otherwise (ifLogRatio=0) they are transformed in log2ratio data.

rhoSquare variance of the segment levels. If rhoSquare=NULL, then the algorithm esti-

mates it on the sample.

kMax maximum number of segments

nu mean of the segment levels. If nu=NULL, then the algorithm estimates it on the

sample.

sigmaSquare variance of the noise. If sigmaSquare=NULL, then the algorithm estimates it

on the sample.

typeEstRho choice of the estimator of rhoSquare. If typeEstRho=1, then the algo-

rithm estimates rhoSquare with $\hat{\rho}_1^2$, while if typeEstRho=0, it estimates

rhoSquare with $\hat{
ho}^2$.

regr choice of the computation of the regression curve. If regr=NULL, then the

regression curve is not computed, if regr="BRC" the Bayesian Regression Curve is computed (BRC with K_2), if regr="BRCAk" the Bayesian Regres-

sion Curve Averaging over k is computed (BRCAk).

Details

By default, the function estimates the copy number profile with mBPCR and estimating rhoSquare on the sample, using $\hat{\rho}_1^2$. It is also possible to use $\hat{\rho}^2$ as estimator of rhoSquare, by setting typeEstRho=0, or to directly set the value of the parameter.

The function gives also the possibility to estimate the profile with a Bayesian regression curve: if regr="BRC" the Bayesian Regression Curve with K_2 is computed (BRC with K_2), if regr="BRCAk" the Bayesian Regression Curve Averaging over k is computed (BRCAk).

Value

A list containing:

estPC an oligoSnpSet equal to sampleData apart from the field assayData (estPC) \$copyNumber,

which contains the estimated profile with mBPCR

regrCurve an oligoSnpSet equal to sampleData apart from the field assayData (regrCurve) \$copyNumbe

which contains the estimated bayesian regression curve. This object is returned

only if regr!=NULL.

The matrices assayData (estPC) \$copyNumber and assayData (regrCurve) \$copyNumber have the same dimension of assayData (sampleData) \$copyNumber, hence their elements, corresponding to the not analyzed chromosomes and samples, are equal to NA.

References

Rancoita, P. M. V., Hutter, M., Bertoni, F., Kwee, I. (2009). Bayesian DNA copy number analysis. *BMC Bioinformatics* 10: 10. http://www.idsia.ch/~paola/mBPCR

See Also

estProfileWithMBPCR, computeMBPCR

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Examples

```
###import an example of oligoSnpSet data
#data(sample.snpset)
##estimation of chromosome 1 in sample 3
#r <- estProfileWithMBPCRforOligoSnpSet(sample.snpset, sampleToBeAnalyzed=3, chrToBeAnaly
##plot of the estimated chromosomes
#cc <- r$estPC
#cc1 <- cc[chromosome(cc) == "1",3]
#graph.par <- plotSnp(cc1)
#graph.par$ylim <- c(-0.23, 0.1)
#graph.par$cytoband.ycoords <- c(-0.22, -0.18)
#print(graph.par)</pre>
```

importCNData

Import the copy number data

Description

Function to import the raw copy number data from a tab delimited file.

Usage

```
importCNData(path, NRowSkip, ifLogRatio=1)
```

Arguments

path	path of the tab delimited file containing the copy number data. The file must contain a table, where in the first column there are the names of the probes (snpName), in the second one, the chromosome to which each probe belongs (the possible values of the chromosomes are: the integers from 1 to 22, 'X' and 'Y'), in the third one, the phisical positions of the probes and in the forth one, the copy number data.
NRowSkip	number of row to skip in the file, before the table. The names of the columns are to be skipped.
ifLogRatio	denotes if the data are either the log2ratio of raw copy number data or raw copy number data. By default, they are considered as log2ratio data, otherwise (ifLogRatio=0) they are transformed in log2ratio data.

Value

A list containing:

an array containing the names of the probes

chr an array containing the name of the chromosome to which each probe belongs

position an array containing the physical position of each probe

logratio an array containing the log2ratio of the raw copy number data

Examples

```
###import the 10K data of cell line REC
path <- system.file("extdata", "rec10k.txt", package = "mBPCR")
rec10k <- importCNData(path, NRowSkip=1)
plot(rec10k$position[rec10k$chr == 3], rec10k$logratio[rec10k$chr == 3])</pre>
```

```
jekoChr11Array250Knsp
```

Affymetrix GeneChip Mapping 250K NSP Array data of JEKO-1 cell line (chr. 11)

Description

Affymetrix GeneChip Mapping 250K NSP Array data of JEKO-1 cell line.

Usage

```
data(jekoChr11Array250Knsp)
```

Format

A data frame containing four variables: first is SNP name ('SNPname'), second is probe chromosome ('Chromosome'), third is probe position ('PhysicalPosition') and fourth is probe raw log2ratio ('log2ratio').

Source

Poretti, G. Rancoita, P.M.V. Kwee, I. Bertoni, F., unpublished

logAdd

Overflow-safe computation of the logarithm of a sum

Description

Function to compute the logarithm of a sum of small numbers, avoiding overflow.

Usage

```
logAdd(x)
```

Arguments

Х

array or matrix containing the logarithm of the terms of the sum. If x is a matrix, the function return the results by column.

Value

If x is an array, the function returns $log(sum_i(e^x[i]))$, otherwise it returns an array containing the results by column.

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Examples

```
x <- \log(c(0.0001, 0.0003, 0.000006))

y <- \log Add(x)

##verification that the computation is correct

z <- sum(c(0.0001, 0.0003, 0.000006))

z

exp(y)
```

plotEstProfile

Plot the estimated profile of copy number data

Description

Function to plot the estimated profiles of copy number data.

Usage

Arguments

sampleName name of the sample, if the user wants to put it in the title of the graph

chr array containing the name of the chromosome to which each probe belongs. The

possible values of the elements of chr are: the integers from 1 to 22, 'X' and

Υ'.

position array containing the physical position of each probe

logratio array containing the log2ratio of the raw copy number data

chrToBePlotted

array containing the name of the estimated chromosomes, that the user wants to plot. The possible values of the chromosomes are: the integers from 1 to 22, 'X'

and 'Y'.

estPC array containing the estimated copy number profile as a piecewise constant func-

tion. If estPC=NULL, only the estimated Bayesian regression curve is plotted.

 ${\tt maxProbeNumber}$

maximum number of probes that a chromosome (or arm of a chromosome) can have to be analyzed. The procedure of profile estimation needs the computation of an array of length (length(chromosome)+1)*(length(chromosome)+2)/2. To be sure to have set this parameter correctly, try to create the array A <-array(1, dim=(maxProbeNumber+1)*(maxProbeNumber+2)/2),

before starting with the estimation procedure.

legendPosition

string containing the position of the legend in the plot. The possible values are

the same used in the function plot.

regrCurve array containing the estimated regression curve. If regrCurve=NULL, then the estimated Bayesian regression curve is not plotted. If regrCurve!=NULL

and also estPC! =NULL both estimated profiles are plotted on the same graph.

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regr

choice of the computation of the regression curve. If regr=NULL, then the regression curve was not computed (then the estimated Bayesian regression curve is not plotted), if regr="BRC" the Bayesian Regression Curve was computed (mBRC with K_2), if regr="BRCAk" the Bayesian Regression Curve Averaging over k was computed (BRCAk).

Details

The function plots the estimated profiles of the chromosomes of chrToBePlotted, separately.

Examples

```
##import the 10K data of cell line REC
data(rec10k)
##estimation of chromosomes 3 and 5
results <- estProfileWithMBPCR(rec10k$SNPname, rec10k$Chromosome, rec10k$PhysicalPosition
##plot the corresponding estimated profiles
plotEstProfile(sampleName='rec10k', rec10k$Chromosome, rec10k$PhysicalPosition, rec10k$le</pre>
```

rec10k

Affymetrix GeneChip Mapping 10K Array data of REC-1 cell line

Description

Affymetrix GeneChip Mapping 10K Array data of REC-1 cell line taken from the reference below.

Usage

```
data(rec10k)
```

Format

A data frame containing five variables: first is SNP name ('SNPname'), second is probe chromosome ('Chromosome'), third is probe position ('PhysicalPosition'), fourth is probe raw log2ratio ('log2ratio') and fifth are is probe genotype ('call').

Source

Rinaldi et al. (2006), Genomic and expression profiling identifies the B-cell associated tyrosine kinase Syk as a possible therapeutic target in mantle cell lymphoma, *British Journal of Haematology*, 132, 303-316

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writeEstProfile

Write the estimated profile of copy number data

Description

Function to write nicely the results of the copy number profile estimation. The function either writes the tables directly on a tab delimited file or returns the corresponding tables.

Usage

Arguments

path	path of the folder where the user wants to write the results of the estimation (it must end with '\' in windows, or '//' in linux). If path=", they will be written in the working directory. If path=NULL, the tables will not be written on a file, but only returned by the function.	
sampleName	name of the sample. If the name of the sample if provided, it is used to named the files.	
snpName	array containing the name of each probe	
chr	array containing the name of the chromosome to which each probe belongs. The possible values of the elements of chr are: the integers from 1 to 22, 'X' and 'Y'.	
position	array containing the physical position of each probe	
logratio	array containing the log2ratio of the raw copy number data	
chrToBeWritt	en	
	array containing the name of the estimated chromosomes, of which the user wants to write the results. The possible values of the chromosomes are: the integers from 1 to 22, 'X' and 'Y'.	
estPC	array containing the estimated copy number profile as a piecewise constant function	
estBoundaries		
	list containing the vectors of the estimated breakpoints, for each of the chromo-	
	somes mentioned in chrToBeWritten. If estBoundaries=NULL, then this information is not written.	
postProbT	list containing the vectors of the posterior probabilities to be a breakpoint of the estimated breakpoints, for each of the chromosomes mentioned in chrToBeWritten. If postProbT=NULL, then this information is not written in the file containing the estimated breakpoints.	
regrCurve	array containing the estimated regression curve. If regrCurve=NULL, then the file containing this information is not written.	
regr	choice of the computation of the regression curve. If $regr=NULL$, then the regression curve was not computed (then the file containing this information is not written), if $regr="BRC"$ the Bayesian Regression Curve with K_2 was computed (BRC with K_2), if $regr="BRCAk"$ the Bayesian Regression Curve Averaging over k was computed (BRCAk).	

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Details

The function writes or returns at maximum three tables:

-one containing the estimated profile with mBPCR (the columns are: 'SNPname', 'chromosome', 'position', 'rawLog2ratio', 'mBPCRestimate')

-one containing a summary about the estimated profile with mBPCR (the columns are: 'SNPname(start)', 'SNPname(end)', 'chromosome', 'position(start)', 'position(end)', 'nProbes', 'mBPCRestimate' and, eventually, 'breakpointPostProb'). This table is not created if estBoundaries=NULL.

-one containing the estimated profile with a regression curve (the columns are: 'SNPname', 'chromosome', 'position', 'rawLog2ratio' and the name of the regression curve used). This table is not created if regrCurve=NULL.

Examples

```
##import the 10K data of cell line REC
data(rec10k)
##estimation of chromosome 5
results <- estProfileWithMBPCR(rec10k$SNPname, rec10k$Chromosome, rec10k$PhysicalPosition
\#\#write the estimated profile of chromosome 5 in a file in the working directory
writeEstProfile(path='', sampleName='rec10k', rec10k$SNPname, rec10k$Chromosome, rec10k$E
#### the same result can be obtained in the following way, by using the function computeN
##estimation of the global parameters
#param <- estGlobParam(rec10k$log2ratio)</pre>
##estimation of chromosome 5
#results <- computeMBPCR(rec10k$log2ratio[rec10k$Chromosome == 5], nu=param$nu, rhoSquare
##write the estimated profile of chromosome 5 in a file in the working directory
#estPC <- array(dim=length(rec10k$SNPname))</pre>
#estBoundaries <- list(dim=1)</pre>
#postProbT <- list(dim=1)</pre>
#estPC[rec10k$Chromosome == 5] <- results$estPC</pre>
#estBoundaries[[1]] <- results$estBoundaries</pre>
#postProbT[[1]] <- c(results$postProbT[results$estBoundaries[-results$estK]],1)</pre>
#writeEstProfile(path='', sampleName='rec10k', rec10k$SNPname, rec10k$Chromosome, rec10k$
```

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