

Package ‘AnaCoDa’

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Type Package

Title Analysis of Codon Data under Stationarity using a Bayesian Framework

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Description Is a collection of models to analyze genome scale codon data using a Bayesian framework. Provides visualization routines and checkpointing for model fittings. Currently published models to analyze gene data for selection on codon usage based on Ribosome Overhead Cost (ROC) are: ROC (Gilchrist et al. (2015) <doi:10.1093/gbe/evv087>), and ROC with phi (Wallace & Drummond (2013) <doi:10.1093/molbev/mst051>). In addition 'AnaCoDa' contains three currently unpublished models. The FONSE (First order approximation On NonSense Error) model analyzes gene data for selection on codon usage against of nonsense error rates. The PA (PAusing time) and PANSE (PAusing time + NonSense Error) models use ribosome footprinting data to analyze estimate ribosome pausing times with and without nonsense error rate from ribosome footprinting data.

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AAToCodon *Amino Acid to codon set*

Description

Converts one character amino acid code to the set of codon encoding that amino acid

Usage

```
AAToCodon(aa, focal = FALSE)
```

Arguments

aa	Amino acid in single character notation
focal	logical, Include the alphabetically first (focal) codon

Value

Returns the names of the codon encoding the give amino acid

See Also

[codonToAA](#)

acfCSP *Plots ACF for codon specific parameter traces*

Description

The function calculates and by defaults plots the acf and estimates the autocorrelation in the trace

Usage

```
acfCSP(parameter, csp = "Mutation", numMixtures = 1, samples = NULL,
lag.max = 40, plot = TRUE)
```

Arguments

parameter	object of class Parameter
csp	"Selection" or "Mutation", defaults to "Mutation"
numMixtures	indicates the number of CSP mixtures used
samples	number of samples at the end of the trace used to calculate the acf
lag.max	Maximum amount of lag to calculate acf. Default is $10 \cdot \log_{10}(N)$, where N is the number of observations.
plot	logical. If TRUE (default) a plot of the acf is created

See Also

[acfMCMC](#)

acfMCMC

Autocorrelation function for the likelihood or posterior trace

Description

The function calculates and by defaults plots the acf and estimates the autocorrelation in the trace.

Usage

```
acfMCMC(mcmc, type = "LogPosterior", samples = NULL, lag.max = 40,
        plot = TRUE)
```

Arguments

mcmc	object of class MCMC
type	"LogPosterior" or "LogLikelihood", defaults to "LogPosterior"
samples	number of samples at the end of the trace used to calculate the acf
lag.max	Maximum amount of lag to calculate acf. Default is $10 \cdot \log_{10}(N)$, where N is the number of observations.
plot	logical. If TRUE (default) a plot of the acf is created

See Also

[acfCSP](#)

`addObservedSynthesisRateSet`*Add gene observed synthesis rates*

Description

`addObservedSynthesisRateSet` returns the observed synthesis rates of the genes within the genome specified.

Usage

```
addObservedSynthesisRateSet(genome, observed.expression.file,  
                             match.expression.by.id = TRUE)
```

Arguments

`genome` A genome object initialized with `initializeGenomeObject` to add observed expression data.

`observed.expression.file`
 A string containing the location of a file containing empirical expression rates (optional).

`match.expression.by.id`
 If TRUE (default) observed expression values will be assigned by matching sequence identifier. If FALSE observed expression values will be assigned by order

Value

Returns the genome after adding the new gene expression values

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")  
expression_file <- system.file("extdata", "expression.csv", package = "AnaCoDa")  
## reading genome  
genome <- initializeGenomeObject(file = genome_file)  
  
## add expression values after the genome was initialized,  
## or adding an additional set of expression values  
genome <- addObservedSynthesisRateSet(genome = genome,  
                                     observed.expression.file = expression_file)
```

aminoAcids

Amino acids

Description

Returns a vector of all amino acids

Usage

```
aminoAcids()
```

Value

Returns a vector of all amino acids

See Also

[codons](#)

codons

Codons

Description

Returns a vector of all codons

Usage

```
codons()
```

Value

Returns a vector of all codons

See Also

[aminoAcids](#)

codonToAA	<i>translates codon to amino acid</i>
-----------	---------------------------------------

Description

Translates a given codon into the amino acid encoded by it.

Usage

```
codonToAA(codon)
```

Arguments

codon character, codon to translate

Value

Returns the amino acid encoded by the given codon as character

See Also

[AAToCodon](#)

convergence.test	<i>Convergence Test</i>
------------------	-------------------------

Description

Convergence Test

Usage

```
convergence.test(object, samples = 10, frac1 = 0.1, frac2 = 0.5,
  thin = 1, plot = FALSE, what = "Mutation", mixture = 1)
```

Arguments

object	an object of either class Trace or MCMC
samples	number of samples at the end of the trace used to determine convergence (< length of trace)
frac1	fraction to use from beginning of chain
frac2	fraction to use from end of chain
thin	the thinning interval between consecutive observations
plot	(logical) plot result instead of returning an object

what	Character describing which trace should be tested for convergence (only for Trace object). Valid options are Mutation, Selection, MixtureProbability, Sphi, Mphi, ExpectedPhi, or Expression
mixture	Integer determining for which mixture distribution the convergence test should be applied (only for trace object).

Value

geweke score object

Examples

```
## check for convergence after a run:

genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")

genome <- initializeGenomeObject(file = genome_file)
sphi_init <- c(1,1)
numMixtures <- 2
geneAssignment <- sample(1:2, length(genome), replace = TRUE) # random assignment to mixtures
parameter <- initializeParameterObject(genome = genome, sphi = sphi_init,
                                       num.mixtures = numMixtures,
                                       gene.assignment = geneAssignment,
                                       mixture.definition = "allUnique")

samples <- 2500
thinning <- 50
adaptiveWidth <- 25
mcmc <- initializeMCMCObject(samples = samples, thinning = thinning,
                             adaptive.width=adaptiveWidth, est.expression=TRUE,
                             est.csp=TRUE, est.hyper=TRUE, est.mix = TRUE)

divergence.iteration <- 10
## Not run:
runMCMC(mcmc = mcmc, genome = genome, model = model,
        ncores = 4, divergence.iteration = divergence.iteration)
# check if posterior trace has converged
convergence.test(object = mcmc, samples = 500, plot = TRUE)

trace <- getTrace(parameter)
# check if Mutation trace has converged
convergence.test(mcmc, samples = 500, plot = TRUE, what = "Mutation")
# check if Sphi trace has converged
convergence.test(mcmc, samples = 500, plot = TRUE, what = "Sphi")
# check if ExpectedPhi trace has converged
convergence.test(mcmc, samples = 500, plot = TRUE, what = "ExpectedPhi")

## End(Not run)
```

getCAI	<i>Calculate the Codon Adaptation Index</i>
--------	---

Description

getCAI returns the Codon Adaptation Index for a genome based on a provided reference.

Usage

```
getCAI(referenceGenome, testGenome)
```

Arguments

referenceGenome

A genome object initialized with `initializeGenomeObject`. Serves as reference set to calculate the necessary codon weights.

testGenome

A genome object initialized with `initializeGenomeObject`. The genome for which the CAI is supposed to be calculated

Value

Returns a named vector with the CAI for each gene

Examples

```
genome_file1 <- system.file("extdata", "more_genes.fasta", package = "AnaCoDa")
genome_file2 <- system.file("extdata", "genome.fasta", package = "AnaCoDa")

## reading genome
referenceGenome <- initializeGenomeObject(file = genome_file1)
testGenome <- initializeGenomeObject(file = genome_file2)

cai <- getCAI(referenceGenome, testGenome)
```

getCAIweights	<i>Calculate the CAI codon weights for a reference genome</i>
---------------	---

Description

getCAIweights returns the weights for the Codon Adaptation Index based on a reference genome.

Usage

```
getCAIweights(referenceGenome)
```

Arguments

referenceGenome
A genome object initialized with `initializeGenomeObject`.

Value

Returns a named vector with the CAI weights for each codon

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")  
  
## reading genome  
referenceGenome <- initializeGenomeObject(file = genome_file)  
  
wi <- getCAIweights(referenceGenome)
```

getCodonCountsForAA *Get Codon Counts For Each Amino Acid*

Description

provides the codon counts for a given amino acid across all genes

Usage

```
getCodonCountsForAA(aa, genome)
```

Arguments

aa A one character representation of an amino acid.
genome A genome object from which the counts of each codon can be obtained.

Details

The returned matrix contains a row for each gene and a column for each synonymous codon of aa.

Value

Returns a matrix storing the codonCounts for the given amino acid.

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")

## reading genome
genome <- initializeGenomeObject(file = genome_file)
countsForA <- getCodonCountsForAA("A", genome)

counts <- lapply(X = c("A", "C"), FUN = getCodonCountsForAA, genome = genome)
```

getCSPEstimates	<i>Return Codon Specific Parameters (or write to csv) estimates as data.frame</i>
-----------------	---

Description

getCSPEstimates returns the codon specific parameter estimates for a given parameter and mixture or write it to a csv file.

Usage

```
getCSPEstimates(parameter, filename = NULL, CSP = "Mutation", mixture = 1,
  samples = 10)
```

Arguments

parameter	parameter an object created by initializeParameterObject.
filename	Posterior estimates will be written to file instead of returned if specified (format: csv).
CSP	which type of codon specific parameter should be returned (mutation (default) or selection)
mixture	estimates for which mixture should be returned
samples	The number of samples used for the posterior estimates.

Value

returns a data.frame with the posterior estimates of the models codon specific parameters or writes it directly to a csv file if filename is specified

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")

genome <- initializeGenomeObject(file = genome_file)
sphi_init <- c(1,1)
```

```

numMixtures <- 2
geneAssignment <- sample(1:2, length(genome), replace = TRUE) # random assignment to mixtures
parameter <- initializeParameterObject(genome = genome, sphi = sphi_init,
                                      num.mixtures = numMixtures,
                                      gene.assignment = geneAssignment,
                                      mixture.definition = "allUnique")

model <- initializeModelObject(parameter = parameter, model = "ROC")
samples <- 2500
thinning <- 50
adaptiveWidth <- 25
mcmc <- initializeMCMCObject(samples = samples, thinning = thinning,
                             adaptive.width=adaptiveWidth, est.expression=TRUE,
                             est.csp=TRUE, est.hyper=TRUE, est.mix = TRUE)

divergence.iteration <- 10
## Not run:
runMCMC(mcmc = mcmc, genome = genome, model = model,
        ncores = 4, divergence.iteration = divergence.iteration)

## return estimates for codon specific parameters
csp_mat <- getCSPEstimates(parameter, CSP="Mutation")

# write the result directly to the filesystem as a csv file. No values are returned
getCSPEstimates(parameter, , filename=file.path(tempdir(), "csp_out.csv"), CSP="Mutation")

## End(Not run)

```

getExpressionEstimates

Returns the estimated phi posterior for a gene

Description

Posterior estimates for the phi value of specified genes

Usage

```
getExpressionEstimates(parameter, gene.index, samples, quantiles = c(0.025,
0.975))
```

Arguments

parameter	on object created by initializeParameterObject.
gene.index	a integer or vector of integers representing the gene(s) of interesst.
samples	number of samples for the posterior estimate
quantiles	vector of quantiles, (default: c(0.025, 0.975))

Details

The returned vector is unnamed as gene ids are only stored in the genome object, but the `gene.index` vector can be used to match the assignment to the genome.

Value

returns a vector with the mixture assignment of each gene corresponding to `gene.index` in the same order as the genome.

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")

genome <- initializeGenomeObject(file = genome_file)
sphi_init <- c(1,1)
numMixtures <- 2
geneAssignment <- sample(1:2, length(genome), replace = TRUE) # random assignment to mixtures
parameter <- initializeParameterObject(genome = genome, sphi = sphi_init,
                                       num.mixtures = numMixtures,
                                       gene.assignment = geneAssignment,
                                       mixture.definition = "allUnique")

model <- initializeModelObject(parameter = parameter, model = "ROC")
samples <- 2500
thinning <- 50
adaptiveWidth <- 25
mcmc <- initializeMCMCObject(samples = samples, thinning = thinning,
                            adaptive.width=adaptiveWidth, est.expression=TRUE,
                            est.csp=TRUE, est.hyper=TRUE, est.mix = TRUE)

divergence.iteration <- 10
## Not run:
runMCMC(mcmc = mcmc, genome = genome, model = model,
        ncores = 4, divergence.iteration = divergence.iteration)

# get the estimated expression values for all genes based on the mixture
# they are assigned to at each step
estimatedExpression <- getExpressionEstimates(parameter, 1:length(genome), 1000)

## End(Not run)
```

```
getMixtureAssignmentEstimate
```

Returns mixture assignment estimates for each gene

Description

Posterior estimates for the mixture assignment of specified genes

Usage

```
getMixtureAssignmentEstimate(parameter, gene.index, samples)
```

Arguments

parameter	on object created by initializeParameterObject
gene.index	a integer or vector of integers representing the gene(s) of interesst.
samples	number of samples for the posterior estimate

Details

The returned vector is unnamed as gene ids are only stored in the genome object, but the gene.index vector can be used to match the assignment to the genome.

Value

returns a vector with the mixture assignment of each gene corresponding to gene.index in the same order as the genome.

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")

genome <- initializeGenomeObject(file = genome_file)
sphi_init <- c(1,1)
numMixtures <- 2
geneAssignment <- sample(1:2, length(genome), replace = TRUE) # random assignment to mixtures
parameter <- initializeParameterObject(genome = genome, sphi = sphi_init,
                                       num.mixtures = numMixtures,
                                       gene.assignment = geneAssignment,
                                       mixture.definition = "allUnique")

model <- initializeModelObject(parameter = parameter, model = "ROC")
samples <- 2500
thinning <- 50
adaptiveWidth <- 25
mcmc <- initializeMCMCObject(samples = samples, thinning = thinning, adaptive.width=adaptiveWidth,
                             est.expression=TRUE, est.csp=TRUE, est.hyper=TRUE, est.mix = TRUE)
divergence.iteration <- 10
## Not run:
runMCMC(mcmc = mcmc, genome = genome, model = model,
        ncores = 4, divergence.iteration = divergence.iteration)

# get the mixture assignment for all genes
mixAssign <- getMixtureAssignmentEstimate(parameter = parameter,
                                         gene.index = 1:length(genome), samples = 1000)

# get the mixture assignment for a subsample
mixAssign <- getMixtureAssignmentEstimate(parameter = parameter,
                                         gene.index = 5:100, samples = 1000)

# or
```

```
mixAssign <- getMixtureAssignmentEstimate(parameter = parameter,  
                                           gene.index = c(10, 30:50, 3, 90), samples = 1000)  
  
## End(Not run)
```

getNames	<i>Gene Names of Genome</i>
----------	-----------------------------

Description

returns the identifiers of the genes within the genome specified.

Usage

```
getNames(genome, simulated = FALSE)
```

Arguments

genome	A genome object initialized with initializeGenomeObject .
simulated	A logical value denoting if the gene names to be listed are simulated or not. The default value is FALSE.

Value

gene.names Returns the names of the genes as a vector of strings.

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")  
  
## reading genome  
genome <- initializeGenomeObject(file = genome_file)  
  
## return all gene ids for the genome  
geneIDs <- getNames(genome, FALSE)
```

`getNc`*Calculate the Effective Number of Codons*

Description

`getNc` returns the Effective Number of Codons for a genome.

Usage

```
getNc(genome)
```

Arguments

`genome` A genome object initialized with `initializeGenomeObject`.

Value

Returns a named vector with the Effective Number of Codons for each gene

Examples

```
genome_file <- system.file("extdata", "more_genes.fasta", package = "AnaCoDa")
## reading genome
genome <- initializeGenomeObject(file = genome_file)

nc <- getNc(genome)
```

`getNcAA`*Calculate the Effective Number of Codons for each Amino Acid*

Description

`getNcAA` returns the Effective Number of Codons for each Amino Acid.

Usage

```
getNcAA(genome)
```

Arguments

`genome` A genome object initialized with `initializeGenomeObject`.

Value

Returns an object of type `data.frame` with the Effective Number of Codons for each amino acid in each gene.

Examples

```
genome_file <- system.file("extdata", "more_genes.fasta", package = "AnaCoDa")
## reading genome
genome <- initializeGenomeObject(file = genome_file)

nc <- getNcAA(genome)
```

```
getObservedSynthesisRateSet
      Get gene observed synthesis rates
```

Description

`getObservedSynthesisRateSet` returns the observed synthesis rates of the genes within the genome specified.

Usage

```
getObservedSynthesisRateSet(genome, simulated = FALSE)
```

Arguments

<code>genome</code>	A genome object initialized with <code>initializeGenomeObject</code> .
<code>simulated</code>	A logical value denoting if the synthesis rates to be listed are simulated or not. The default value is <code>FALSE</code> .

Value

Returns a `data.frame` with the observed expression values in genome

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")
expression_file <- system.file("extdata", "expression.csv", package = "AnaCoDa")
## reading genome
genome <- initializeGenomeObject(file = genome_file)

## return expression values as a data.frame with gene ids in the first column.
expressionValues <- getObservedSynthesisRateSet(genome = genome)
```

```
getSelectionCoefficients
```

Calculate Selection coefficients

Description

getSelectionCoefficients calculates the selection coefficient of each codon in each gene.

Usage

```
getSelectionCoefficients(genome, parameter, samples = 100)
```

Arguments

genome	A genome object initialized with <code>initializeGenomeObject</code> to add observed expression data.
parameter	an object created by <code>initializeParameterObject</code> .
samples	The number of samples used for the posterior estimates.

Value

A matrix with selection coefficients.

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")

genome <- initializeGenomeObject(file = genome_file)
sphi_init <- 1
numMixtures <- 1
geneAssignment <- rep(1, length(genome))
parameter <- initializeParameterObject(genome = genome, sphi = sphi_init,
                                       num.mixtures = numMixtures,
                                       gene.assignment = geneAssignment,
                                       mixture.definition = "allUnique")

model <- initializeModelObject(parameter = parameter, model = "ROC")
samples <- 2500
thinning <- 50
adaptiveWidth <- 25
mcmc <- initializeMCMCObject(samples = samples, thinning = thinning,
                             adaptive.width=adaptiveWidth, est.expression=TRUE,
                             est.csp=TRUE, est.hyper=TRUE, est.mix = TRUE)

divergence.iteration <- 10
## Not run:
runMCMC(mcmc = mcmc, genome = genome, model = model,
        ncores = 4, divergence.iteration = divergence.iteration)

## return estimates for selection coefficients s for each codon in each gene
selection.coefficients <- getSelectionCoefficients(genome = genome,
```

```
parameter = parameter, samples = 1000)

## End(Not run)
```

getTrace	<i>extracts an object of traces from a parameter object.</i>
----------	--

Description

extracts an object of traces from a parameter object.

Usage

```
getTrace(parameter)
```

Arguments

parameter A Parameter object that corresponds to one of the model types.

Value

trace Returns an object of type Trace extracted from the given parameter object

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")

genome <- initializeGenomeObject(file = genome_file)
sphi_init <- c(1,1)
numMixtures <- 2
geneAssignment <- sample(1:2, length(genome), replace = TRUE) # random assignment to mixtures
parameter <- initializeParameterObject(genome = genome, sphi = sphi_init,
                                       num.mixtures = numMixtures,
                                       gene.assignment = geneAssignment,
                                       mixture.definition = "allUnique")

trace <- getTrace(parameter) # empty trace object since no MCMC was performed
```

```
initializeCovarianceMatrices
```

Initialize Covariance Matrices

Description

Initialize Covariance Matrices

Usage

```
initializeCovarianceMatrices(parameter, genome, numMixtures, geneAssignment,
  init.csp.variance = 0.0025)
```

Arguments

parameter	A Parameter object that corresponds to one of the model types. Valid values are "ROC", "PA", and "FONSE".
genome	An object of type Genome necessary for the initialization of the Parameter object.
numMixtures	The number of mixture elements for the underlying mixture distribution (numMixtures > 0).
geneAssignment	A vector holding the initial mixture assignment for each gene. The vector length has to equal the number of genes in the genome. Valid values for the vector range from 1 to numMixtures. It is possible but not advised to leave a mixture element empty.
init.csp.variance	initial proposal variance for codon specific parameter, default is 0.0025.

Value

parameter Returns the Parameter argument, now modified with initialized mutation, selection, and covariance matrices.

```
initializeGenomeObject
```

Genome Initialization

Description

initializeGenomeObject initializes the Rcpp Genome object

Usage

```
initializeGenomeObject(file, genome = NULL, observed.expression.file = NULL,
  fasta = TRUE, match.expression.by.id = TRUE, append = FALSE)
```

Arguments

file	A file of coding sequences in fasta or RFPData format
genome	A genome object can be passed in to concatenate the input file to it (optional).
observed.expression.file	A string containing the location of a file containing empirical expression rates (optional).
fasta	A boolean value which decides whether to initialize with a fasta file or an RFP-Data file. (TRUE for fasta, FALSE for RFPData)
match.expression.by.id	If TRUE (default), observed expression values will be assigned by matching sequence identifier. If FALSE, observed expression values will be assigned by order.
append	If TRUE (FALSE is default), function will read in additional genome data to append to an existing genome. If FALSE, genome data is cleared before reading in data (no preexisting data).

Value

This function returns the initialized Genome object.

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")
genes_file <- system.file("extdata", "more_genes.fasta", package = "AnaCoDa")
expression_file <- system.file("extdata", "expression.csv", package = "AnaCoDa")

## reading genome
genome <- initializeGenomeObject(file = genome_file)

## reading genome and observed expression data
genome <- initializeGenomeObject(file = genome_file, observed.expression.file = expression_file)

## add additional genes to existing genome
genome <- initializeGenomeObject(file = genome_file)
genome <- initializeGenomeObject(file = genes_file, genome = genome, append = TRUE)
```

initializeMCMCObject *Initialize MCMC*

Description

initializeMCMCObject initializes a MCMC object to perform a model fitting for a parameter and model object.

Usage

```
initializeMCMCObject(samples, thinning = 1, adaptive.width = 100,
  est.expression = TRUE, est.csp = TRUE, est.hyper = TRUE,
  est.mix = TRUE)
```

Arguments

<code>samples</code>	Number of samples to be produced when running the MCMC algorithm. No default value.
<code>thinning</code>	The thinning interval between consecutive observations. If set to 1, every step will be saved as a sample. Default value is 1.
<code>adaptive.width</code>	Number that determines how often the acceptance/rejection window should be altered. Default value is 100 samples.
<code>est.expression</code>	Boolean that tells whether or not synthesis rate values should be estimated in the MCMC algorithm run. Default value is TRUE.
<code>est.csp</code>	Boolean that tells whether or not codon specific values should be estimated in the MCMC algorithm run. Default value is TRUE.
<code>est.hyper</code>	Boolean that tells whether or not hyper parameters should be estimated in the MCMC algorithm run. Default value is TRUE.
<code>est.mix</code>	Boolean that tells whether or not the genes' mixture element should be estimated in the MCMC algorithm run. Default value is TRUE.

Details

`initializeMCMCObject` sets up the MCMC object (monte carlo markov chain) and returns the object so a model fitting can be done. It is important to note that `est.expression` and `est.hyper` will affect one another negatively if their values differ.

Value

`mcmc` Returns an initialized MCMC object.

Examples

```
## initializing an object of type mcmc

samples <- 2500
thinning <- 50
adaptiveWidth <- 25

## estimate all parameter types
mcmc <- initializeMCMCObject(samples = samples, thinning = thinning, adaptive.width=adaptiveWidth,
  est.expression=TRUE, est.csp=TRUE, est.hyper=TRUE, est.mix = TRUE)

## do not estimate expression values, initial conditions will remain constant
mcmc <- initializeMCMCObject(samples = samples, thinning = thinning, adaptive.width=adaptiveWidth,
  est.expression=FALSE, est.csp=TRUE, est.hyper=TRUE, est.mix = TRUE)
```

```
## do not estimate hyper parameters, initial conditions will remain constant
mcmc <- initializeMCMCObject(samples = samples, thinning = thinning, adaptive.width=adaptiveWidth,
                             est.expression=TRUE, est.csp=TRUE, est.hyper=FALSE, est.mix = TRUE)
```

initializeModelObject *Model Initialization*

Description

initializes the model object.

Usage

```
initializeModelObject(parameter, model = "ROC", with.phi = FALSE,
                      fix.observation.noise = FALSE, rfp.count.column = 1)
```

Arguments

parameter	An object created with initializeParameterObject.
model	A string containing the model to run (ROC, FONSE, or PA), has to match parameter object.
with.phi	(ROC only) A boolean that determines whether or not to include empirical phi values (expression rates) for the calculations.
fix.observation.noise	(ROC only) Allows to fix the noise in the observed expression dataset to the initial condition. The initial condition for the observed expression noise can be set in the parameter object.
rfp.count.column	(PA and PANSE only) A number representing the RFP count column to use.

Details

initializeModelObject initializes a model. The type of model is determined based on the string passed to the model argument. The Parameter object has to match the model that is initialized. E.g. to initialize a ROC model, it is required that a ROC parameter object is passed to the function.

Value

This function returns the model object created.

Examples

```

#initializing a model object

genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")
expression_file <- system.file("extdata", "expression.csv", package = "AnaCoDa")

genome <- initializeGenomeObject(file = genome_file,
                                observed.expression.file = expression_file)

sphi_init <- c(1,1)
numMixtures <- 2
geneAssignment <- sample(1:2, length(genome), replace = TRUE) # random assignment to mixtures
parameter <- initializeParameterObject(genome = genome, sphi = sphi_init,
                                       num.mixtures = numMixtures,
                                       gene.assignment = geneAssignment,
                                       mixture.definition = "allUnique")

# initializing a model object assuming we have observed expression (phi)
# values stored in the genome object.
initializeModelObject(parameter = parameter, model = "ROC", with.phi = TRUE)

# initializing a model object ignoring observed expression (phi)
# values stored in the genome object.
initializeModelObject(parameter = parameter, model = "ROC", with.phi = FALSE)

```

```

initializeParameterObject
      Initialize Parameter

```

Description

initializeParameterObject initializes a new parameter object or reconstructs one from a restart file

Usage

```

initializeParameterObject(genome = NULL, sphi = NULL, num.mixtures = 1,
                          gene.assignment = NULL, initial.expression.values = NULL, model = "ROC",
                          split.serine = TRUE, mixture.definition = "allUnique",
                          mixture.definition.matrix = NULL, init.with.restart.file = NULL,
                          mutation.prior.sd = 0.35, init.csp.variance = 0.0025,
                          init.sepsilon = 0.1, init.w.obs.phi = FALSE)

```

Arguments

genome An object of type Genome necessary for the initialization of the Parameter object. The default value is NULL.

sphi	Initial values for sphi. Expected is a vector of length numMixtures. The default value is NULL.
num.mixtures	The number of mixtures elements for the underlying mixture distribution (numMixtures > 0). The default value is 1.
gene.assignment	A vector holding the initial mixture assignment for each gene. The vector length has to equal the number of genes in the genome. Valid values for the vector range from 1 to numMixtures. It is possible but not advised to leave a mixture element empty. The default Value is NULL.
initial.expression.values	(Optional) A vector with initial phi values. The length of the vector has to equal the number of genes in the Genome object. The default value is NULL.
model	Specifies the model used. Valid options are "ROC", "PA", "PANSE", or "FONSE". The default model is "ROC". ROC is described in Gilchrist et al. 2015. PA, PANSE and FONSE are currently unpublished.
split.serine	Whether serine should be considered as one or two amino acids when running the model. TRUE and FALSE are the only valid values. The default value for split.serine is TRUE.
mixture.definition	A string describing how each mixture should be treated with respect to mutation and selection. Valid values consist of "allUnique", "mutationShared", and "selectionShared". The default value for mixture.definition is "allUnique". See details for more information.
mixture.definition.matrix	A matrix representation of how the mutation and selection categories correspond to the mixtures. The default value for mixture.definition.matrix is NULL. If provided, the model will use the matrix to initialize the mutation and selection categories instead of the definition listed directly above. See details for more information.
init.with.restart.file	File name containing information to reinitialize a previous Parameter object. If given, all other arguments will be ignored. The default value for init.with.restart.file is NULL.
mutation.prior.sd	Controlling the standard deviation of the normal prior on the mutation parameters
init.csp.variance	specifies the initial proposal width for codon specific parameter (default is 0.0025). The proposal width adapts during the runtime to reach a target acceptance rate of ~0.25
init.sepsilon	specifies the initial value for sepsilon. default is 0.1
init.w.obs.phi	TRUE: initialize phi values with observed phi values (data from RNAseq, mass spectrometry, ribosome footprinting) Default is FALSE. If multiple observed phi values exist for a gene, the geometric mean of these values is used as initial phi. When using this function, one should remove any genes with missing phi values, as these genes will not have an initial phi value.

Details

`initializeParameterObject` checks the values of the arguments given to insure the values are valid.

The mixture definition and mixture definition matrix describes how the mutation and selection categories are set up with respect to the number of mixtures. For example, if `mixture.definition = "allUnique"` and `numMixtures = 3`, a matrix representation would be `matrix(c(1,2,3,1,2,3), ncol=2)` where each row represents a mixture, the first column represents the mutation category, and the second column represents the selection category. Another example would be `mixture.definition = "selectionShared"` and `numMixtures = 4` (`matrix(c(1,2,3,4,1,1,1,1), ncol=2)`). In this case, the selection category is the same for every mixture. If a matrix is given, and it is valid, then the mutation/selection relationship will be defined by the given matrix and the keyword will be ignored. A matrix should only be given in cases where the keywords would not create the desired matrix.

Value

parameter Returns an initialized Parameter object.

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")
restart_file <- system.file("extdata", "restart_file.rst", package = "AnaCoDa")

genome <- initializeGenomeObject(file = genome_file)

## initialize a new parameter object
sphi_init <- 1
numMixtures <- 1
geneAssignment <- rep(1, length(genome))
parameter <- initializeParameterObject(genome = genome, sphi = sphi_init,
                                       num.mixtures = numMixtures,
                                       gene.assignment = geneAssignment,
                                       mixture.definition = "allUnique")

## re-initialize a parameter object from a restart file. Useful for checkpointing
parameter <- initializeParameterObject(init.with.restart.file = restart_file)

## initialize a parameter object with a custom mixture definition matrix
def.matrix <- matrix(c(1,1,1,2), ncol=2)
geneAssignment <- sample(1:2, length(genome), replace = TRUE) # random assignment to mixtures
parameter <- initializeParameterObject(genome = genome, sphi = c(0.5, 2), num.mixtures = 2,
                                       gene.assignment = geneAssignment,
                                       mixture.definition.matrix = def.matrix)
```

Description

length gives the length of a genome

Usage

```
## S3 method for class 'Rcpp_Genome'  
length(x)
```

Arguments

x A genome object initialized with `initializeGenomeObject`.

Value

returns the number of genes in a genome

Examples

```
genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")  
  
## reading genome  
genome <- initializeGenomeObject(file = genome_file)  
length(genome) # 10
```

loadMCMCObject	<i>Load MCMC Object</i>
----------------	-------------------------

Description

loadMCMCObject creates a new MCMC object and fills it with the information in the file given.

Usage

```
loadMCMCObject(files)
```

Arguments

files The filenames where the data will be stored.

Details

This MCMC object is not intended to be used to do another model fitting, only to graph the stored results.

Value

This function has no return value.

Examples

```
## loading mcmc objects from the filesystem
## Not run:
# load one mcmc object
mcmc <- loadMCMCObject(files = "mcmc.Rda")

# load and combine multiple mcmc objects. Useful when using checkpointing
mcmc <- loadMCMCObject(files = c("mcmc1.Rda", "mcmc2.Rda"))

## End(Not run)
```

loadParameterObject *Load Parameter Object*

Description

loadParameterObject will load a parameter object from the filesystem

Usage

```
loadParameterObject(files)
```

Arguments

files A list of parameter filenames to be loaded. If multiple files are given, the parameter objects will be concatenated in the order provided

Details

The function loads one or multiple files. In the case of multiple file, e.g. due to the use of checkpointing, the files will be concatenated to one parameter object. See [writeParameterObject](#) for the writing of parameter objects

Value

Returns an initialized Parameter object.

Examples

```
## Not run:
# load a single parameter object
parameter <- loadParameterObject("parameter.Rda")

# load and concatenate multiple parameter object
parameter <- loadParameterObject(c("parameter1.Rda", "parameter2.Rda"))

## End(Not run)
```

plot.Rcpp_FONSEModel *Plot Model Object*

Description

Plots traces from the model object such as synthesis rates for each gene. Will work regardless of whether or not expression/synthesis rate levels are being estimated. If you wish to plot observed/empirical values, these values MUST be set using the initial.expression.values parameter found in initializeParameterObject. Otherwise, the expression values plotted will just be SCUO values estimated upon initialization of the Parameter object.

Usage

```
## S3 method for class 'Rcpp_FONSEModel'
plot(x, genome, samples = 100, mixture = 1,
     simulated = FALSE, ...)
```

Arguments

x	An Rcpp model object initialized with initializeModelObject.
genome	An Rcpp genome object initialized with initializeGenomeObject.
samples	The number of samples in the trace
mixture	The mixture for which to graph values.
simulated	A boolean value that determines whether to use the simulated genome.
...	Optional, additional arguments. For this function, a possible title for the plot in the form of a list if set with "main".

Value

This function has no return value.

plot.Rcpp_FONSEParameter
Plot Parameter

Description

plot graphs the mutation or selection parameter for a ROC or FONSE parameter object for each mixture element.

Usage

```
## S3 method for class 'Rcpp_FONSEParameter'
plot(x, what = "Mutation", samples = 100,
     mixture.name = NULL, with.ci = TRUE, ...)
```

Arguments

<code>x</code>	A parameter object
<code>what</code>	Which aspect of the parameter to plot. Default value is "Mutation".
<code>samples</code>	Number of samples to plot using the posterior mean. Default value is 100.
<code>mixture.name</code>	a vector with names/descriptions of the mixture distributions in the parameter object
<code>with.ci</code>	Plot with or without confidence intervals. Default value is TRUE
<code>...</code>	Arguments to be passed to methods, such as graphical parameters.

Details

Graphs are based off the last # samples for the posterior mean.

Value

This function has no return value.

`plot.Rcpp_MCMCArgorithm`

Plot MCMC algorithm

Description

This function will plot the logLikelihood trace, and if the Hmisc package is installed, it will plot a subplot of the logLikelihood trace with the first few samples removed.

Usage

```
## S3 method for class 'Rcpp_MCMCArgorithm'
plot(x, what = "LogPosterior",
     zoom.window = NULL, ...)
```

Arguments

<code>x</code>	An Rcpp_MCMC object initialized with <code>initializeMCMCObject</code> .
<code>what</code>	character defining if <code>log(Posterior)</code> (Default) or <code>log(Likelihood)</code> options are: <code>LogPosterior</code> or <code>logLikelihood</code>
<code>zoom.window</code>	A vector describing the start and end of the zoom window.
<code>...</code>	Arguments to be passed to methods, such as graphical parameters.

Value

This function has no return value.

plot.Rcpp_ROCModel *Plot Model Object*

Description

Plots traces from the model object such as synthesis rates for each gene. Will work regardless of whether or not expression/synthesis rate levels are being estimated. If you wish to plot observed/empirical values, these values **MUST** be set using the `initial.expression.values` parameter found in `initializeParameterObject`. Otherwise, the expression values plotted will just be SCUO values estimated upon initialization of the `Parameter` object.

Usage

```
## S3 method for class 'Rcpp_ROCModel'
plot(x, genome = NULL, samples = 100, mixture = 1,
     simulated = FALSE, ...)
```

Arguments

<code>x</code>	An Rcpp model object initialized with <code>initializeModelObject</code> .
<code>genome</code>	An Rcpp genome object initialized with <code>initializeGenomeObject.#</code>
<code>samples</code>	The number of samples in the trace
<code>mixture</code>	The mixture for which to graph values.
<code>simulated</code>	A boolean value that determines whether to use the simulated genome.
<code>...</code>	Optional, additional arguments. For this function, a possible title for the plot in the form of a list if set with "main".

Value

This function has no return value.

plot.Rcpp_ROCParameter *Plot Parameter*

Description

plot graphs the mutation or selection parameter for a ROC or FONSE parameter object for each mixture element.

Usage

```
## S3 method for class 'Rcpp_ROCParameter'
plot(x, what = "Mutation", samples = 100,
     mixture.name = NULL, with.ci = TRUE, ...)
```

Arguments

x	A parameter object
what	Which aspect of the parameter to plot. Default value is "Mutation".
samples	Number of samples to plot using the posterior mean. Default value is 100.
mixture.name	a vector with names/descriptions of the mixture distributions in the parameter object
with.ci	Plot with or without confidence intervals. Default value is TRUE
...	Arguments to be passed to methods, such as graphical parameters.

Details

Graphs are based off the last # samples for the posterior mean.

Value

This function has no return value.

plot.Rcpp_Trace	<i>Plot Trace Object</i>
-----------------	--------------------------

Description

Plots different traces, specified with the what parameter.

Usage

```
## S3 method for class 'Rcpp_Trace'
plot(x, what = c("Mutation", "Selection",
  "MixtureProbability", "Sphi", "Mphi", "Aphi", "Sepsilon", "ExpectedPhi",
  "Expression"), geneIndex = 1, mixture = 1, ...)
```

Arguments

x	An Rcpp trace object initialized with initializeTraceObject.
what	A string containing one of the following to graph: Mutation, Selection, Alpha, LambdaPrime, Mean, MixtureProbability, Sphi, Mphi, Aphi, Spesilon, ExpectedPhi, Expression.
geneIndex	When plotting expression, the index of the gene to be plotted.
mixture	The mixture for which to plot values.
...	Optional, additional arguments. For this function, may be a logical value determining if the trace is ROC-based or not.

Value

This function has no return value.

plotCodonSpecificParameters
Plot Codon Specific Parameter

Description

Plots a codon-specific set of traces, specified with the type parameter.

Usage

```
plotCodonSpecificParameters(trace, mixture, type = "Mutation",
  main = "Mutation Parameter Traces", ROC = TRUE)
```

Arguments

trace	An Rcpp trace object initialized with initializeTraceObject.
mixture	The mixture for which to plot values.
type	A string containing one of the following to graph: Mutation, Selection, Alpha, LambdaPrime, Mean
main	The title of the plot.
ROC	A logical value determining if the Parameter was ROC or not.

Value

This function has no return value.

runMCMC *Run MCMC*

Description

runMCMC will run a monte carlo markov chain algorithm for the given mcmc, genome, and model objects to perform a model fitting.

Usage

```
runMCMC(mcmc, genome, model, ncores = 1, divergence.iteration = 0)
```

Arguments

mcmc	MCMC object that will run the model fitting algorithm.
genome	Genome that the model fitting will run on. Should be the same genome associated with the parameter and model objects.
model	Model to run the fitting on. Should be associated with the given genome.
ncores	Number of cores to perform the model fitting with. Default value is 1.
divergence.iteration	Number of steps that the initial conditions can diverge from the original conditions given. Default value is 0.

Details

runMCMC will run for the number of samples times the number thinning given when the mcmc object is initialized. Updates are provided every 100 steps, and the state of the chain is saved every thinning steps.

Value

This function has no return value.

Examples

```
#fitting a model to a genome using the runMCMC function

genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")

genome <- initializeGenomeObject(file = genome_file)
sphi_init <- c(1,1)
numMixtures <- 2
geneAssignment <- sample(1:2, length(genome), replace = TRUE) # random assignment to mixtures
parameter <- initializeParameterObject(genome = genome, sphi = sphi_init,
                                       num.mixtures = numMixtures,
                                       gene.assignment = geneAssignment,
                                       mixture.definition = "allUnique")

model <- initializeModelObject(parameter = parameter, model = "ROC")
samples <- 2500
thinning <- 50
adaptiveWidth <- 25
mcmc <- initializeMCMCObject(samples = samples, thinning = thinning,
                             adaptive.width=adaptiveWidth, est.expression=TRUE,
                             est.csp=TRUE, est.hyper=TRUE, est.mix = TRUE)

divergence.iteration <- 10
## Not run:
runMCMC(mcmc = mcmc, genome = genome, model = model,
        ncores = 4, divergence.iteration = divergence.iteration)

## End(Not run)
```

setRestartSettings *Set Restart Settings*

Description

setRestartSettings sets the needed information (what the file is called, how often the file should be written) to write information to restart the MCMC algorithm from a given point.

Usage

```
setRestartSettings(mcmc, filename, samples, write.multiple = TRUE)
```

Arguments

mcmc	MCMC object that will run the model fitting algorithm.
filename	Filename for the restart files to be written.
samples	Number of samples that should occur before a file is written.
write.multiple	Boolean that determines if multiple restart files are written. Default value is TRUE.

Details

setRestartSettings writes a restart file every set amount of samples that occur. Also, if write.multiple is true, instead of overwriting the previous restart file, the sample number is prepended onto the file name and multiple rerstart files are generated for a run.

Value

This function has no return value.

Examples

```
## set restart settings for checkpointing

samples <- 2500
thinning <- 50
adaptiveWidth <- 25

## estimate all parameter types
mcmc <- initializeMCMCObject(samples = samples, thinning = thinning,
                             adaptive.width=adaptiveWidth, est.expression=TRUE,
                             est.csp=TRUE, est.hyper=TRUE, est.mix = TRUE)

# prompts the mcmc to write a restart file every 100 samples during the run.
setRestartSettings(mcmc = mcmc, filename = "test_restart", samples = 100)

# prompts the mcmc to write a restart file every 100 samples during the run,
# but will overwrite it each time.
setRestartSettings(mcmc = mcmc, filename = "test_restart", samples = 100,
                  write.multiple = FALSE)
```

summary.Rcpp_Genome *Summary of Genome*

Description

summary summarizes the description of a genome, such as number of genes and average gene length.


```

samples <- 2500
thinning <- 50
adaptiveWidth <- 25
mcmc <- initializeMCMCObject(samples = samples, thinning = thinning,
                             adaptive.width=adaptiveWidth, est.expression=TRUE,
                             est.csp=TRUE, est.hyper=TRUE, est.mix = TRUE)

divergence.iteration <- 10
## Not run:
runMCMC(mcmc = mcmc, genome = genome, model = model,
        ncores = 4, divergence.iteration = divergence.iteration)
writeMCMCObject(mcmc = mcmc, file = file.path(tempdir(), "file.Rda"))

## End(Not run)

```

writeParameterObject *Write Parameter Object to a File*

Description

writeParameterObject will write the parameter object as binary to the filesystem

Usage

```
writeParameterObject(parameter, file)
```

Arguments

parameter	parameter on object created by initializeParameterObject.
file	A filename that where the data will be stored.

Details

As Rcpp object are not serializable with the default R save function, therefore this custom save function is provided (see [loadParameterObject](#)).

Value

This function has no return value.

Examples

```

## Not run:

genome_file <- system.file("extdata", "genome.fasta", package = "AnaCoDa")

genome <- initializeGenomeObject(file = genome_file)
sphi_init <- c(1,1)
numMixtures <- 2

```

```
geneAssignment <- sample(1:2, length(genome), replace = TRUE) # random assignment to mixtures
parameter <- initializeParameterObject(genome = genome, phi = phi_init,
                                       num.mixtures = numMixtures,
                                       gene.assignment = geneAssignment,
                                       mixture.definition = "allUnique")

## writing an empty parameter object as the runMCMC routine was not called yet
writeParameterObject(parameter = parameter, file = file.path(tempdir(), "file.Rda"))

## End(Not run)
```

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