rGADEM

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GADEM

Motif Analysis with rGADEM

Description

It is an R implementation of GADEM, a powerful computational tools for de novo motif discovery.

Usage

```
GADEM(Sequences, seed=1, genome=NULL, verbose=FALSE, numWordGroup=3, numTop3mer=20, numTop4mer=40, numTop5mer=60, numGeneration=5, populationSize=100, pValue=0.0002, eValue=0.0, extTrim=1, minSpaceWidth=0, maxSpaceWidth=10, useChIPscore=0, numEM=40, fEM=0.5, widthWt=80, fullScan=0, slideWinPWM=6, stopCriterion=1, numBackgSets=10, weightType=0, bFileName="NULL", Spwm="NULL", minSites =-1, maskR=0, nmotifs=25)
```

Arguments

Sequences Sequences from BED or FASTA file are converted into XString object view

when a seed is specified, the run results are deterministic

genome Specify the genome

verbose Print immediate results on screen [TRUE-yes (default), FALSE-no]. These re-

sults include the motif consensus sequence, number of sites (in sequences sub-

jected to EM optimization, see -fEM, above), and ln(E-value).

numWordGroup number of non-zero k-mer groups

numTop3mer Number of top-ranked trimers for spaced dyads (default: 20).

NumTop4mer Number of top-ranked tetramers for spaced dyads (default: 40).

Number of top-ranked pentamers for spaced dyads (default: 60).

numGeneration

Number of genetic algorithm (GA) generations (default: 5).

populationSize

GA population size (default: 100). Both default settings should work well for most datasets (ChIP-chip and ChIP-seq). The above two arguments are ignored in a seeded analysis, because spaced dyads and GA are no longer needed (num-Generation is set to 1 and populationSize is set to 10 internally, corresponding to the 10 maxp choices).

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pValue P-value cutoff for declaring BINDING SITES (default: 0.0002). Depending on

data size and the motif, you might want to assess more than one value. For ChIP-seq data (e.g., 10 thousand +/-200-bp max-center peak cores), p=0.0002 often seems appropriate. However, short motifs may require a less stringent setting.

eValue ln(E-value) cutoff for selecting MOTIFS (default: 0.0). If a seeded analysis fails

to identify the expected motif, run GADEM with -verbose 1 to show motif ln(E-value)s on screen, then rerun with a larger ln(E-value) cutoff. This can help in identifying short and/or low abundance motifs, for which the default E-value

threshold may be too low.

extTrim Base extension and trimming (1 -yes, 0 -no) (default: 1).

minSpaceWidth

Minimal number of unspecified nucleotides in spaced dyads (default: 0).

maxSpaceWidth

Maximal number of unspecified nucleotides in spaced dyads (default: 10). minSpaceWidth and maxSpaceWidth control the lengths of spaced dyads, and, with exTrim, control motif lengths. Longer motifs can be discovered by setting maxSpaceWidth

to larger values (e.g. 50).

useChIPscore Use top-scoring sequences for deriving PWMs. Sequence (quality) scores are

stored in sequence header (see documentation). 0 - no (default, randomly select

sequences), 1 - yes.

numEM Number of EM steps (default: 40). One might want to set it to a larger value

(e.g. 80) in a seeded run, because such runs are fast.

Fraction of sequences used in EM to obtain PWMs in an unseeded analysis

(default: 0.5). For unseeded motif discovery in a large dataset (e.g. >10 million nt), one might want to set -fEM to a smaller value (e.g., 0.3 or 0.4) to reduce run

time.

widthWt For -posWt 1 or 3, width of central sequence region with large EM weights for

PWM optimization (default: 50). This argument is ignored when weight Type is

0 (uniform prior) or 2 (Gaussian prior).

fullScan GADEM keeps two copies of the input sequences internally: one (D) for discov-

ering PWMs and one (S) for scanning for binding sites using the PWMs Once a motif is identified, its instances in set D are always masked by Ns. However, masking motif instances in set S is optional, and scanning unmasked sequences

allows sites of discovered motifs to overlap.

slideWinPWM sliding window for comparing pwm similarity (default: 6).

stopCriterion

Number of generations without new motifs before stopping analysis.

numBackgSets Number of sets of background sequences (default: 10). The background se-

quences are simulated using the [a,c,g,t] frequencies in the input sequences, with length matched between the two sets. The background sequences are used as the

random sequences for assessing motif enrichment in the input data.

weight Type Weight profile for positions on the sequence. 0 - no weight (uniform spatial

prior, default), 1 (gaussian prior) and 2 (triangle prior) - small or zero weights for the ends and large weights for the center (e.g. the center 50 bp). Consider using 1 or 2 if you expect strong central enrichment (as in ChIP-seq) and your

sequences are long(e.g. >200 bp).

bFileName Reading user-specified background models.

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Spwm	File name for the seed PWM, when a seeded approach is used. can be used as the starting PWM for the EM algorithm. This will help find an expected motif and is much faster than unseeded de novo discovery. Also, when a seed PWM is specified, the run results are deterministic, so only a single run is needed (repeat runs with the same settings will give identical results). In contrast, unseeded runs are stochastic, and we recommend comparing results from several repeat runs.
minSites	Minimal number of sites required for a motif to be reported (default: num-Seq/20)
maskR	Mask low-complexity sequences or repeats; 'aaaaaaaaa', 'tttttttt', 'cacacaca', 'tgtgtgtg', 'tatatatat', 'ggaggaggagga', 'gaggaggaggag', 'agaagaagaaga', 'ctcctcctcctc', 'tcctcctcctcc', 'tc or 'cagcagcagcagcag' (default: 0-no masking, 1-masking)

nmotifs Number of motifs sought (default: 25)

Author(s)

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Examples

```
library(BSgenome.Hsapiens.UCSC.hg18)
pwd<-"" #INPUT FILES- BedFiles, FASTA, etc.
path<- system.file("extdata", "Test_100.bed", package="rGADEM")
BedFile<-paste(pwd, path, sep="")
BED<-read.table(BedFile, header=FALSE, sep="\t")
BED<-data.frame(chr=as.factor(BED[,1]), start=as.numeric(BED[,2]), end=as.numeric(BED[,3]))
#Create RD files
rgBED<-IRanges(start=BED[,2], end=BED[,3])
Sequences<-RangedData(rgBED, space=BED[,1])</pre>
gadem<-GADEM(Sequences, verbose=1, genome=Hsapiens)
```

align-class Class "align"

Description

This object contains the individual motifs identified but also the location (seqID and position) of the sites in the original sequence data. It also included the spaced dyad from which the motifs is derived, PWM score p-value cuttoff for the run.

Objects from the Class

Objects can be created by calls of the form new ("align", ...).

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Slots

```
seq :Motif identified .
  chr :Chromosome identified.
start :Sequence start.
end :Sequence end.
strand :Strand position.
seqID :Sequence identification.
pos :Position identification.
pval :p-Value for each identification.
fastaHeader :Fasta accession.

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```

See Also

```
gadem, motif, parameters
```

Examples

```
showClass("align")
```

gadem-class

Class "gadem"

Description

This object contains all gadem output information.

Objects from the Class

Objects can be created by calls of the form new ("gadem", ...).

Slots

```
motifList List of input PWM. parameters List of rGADEM parameters.
```

Methods

```
[ signature(x = "gadem"): subset gadem object.
[[ ] signature(x = "gadem"): subset gadem object.
nMotifs signature(x = "gadem"): Number of motifs identified
names signature(x = "gadem"): Assign motifs names.
dim signature(x = "gadem"): Number of sequences identified for each motifs.
consensus signature(x = "gadem"): Sequence of consensus motifs.
```

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```
nOccurrences signature (x = "gadem"): View of PWMs. 

startPos signature (x = "gadem"): Start position for each sequences. 

endPos signature (x = "gadem"): End position for each sequences. 

getPWM signature (x = "gadem"): End position for each sequences.
```

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See Also

```
motif, align, parameters
```

Examples

```
showClass("gadem")
```

motif-class

Class "motif"

Description

This object contains PWM, motif consensus, motif length and all aligned sequences for a specific motif

Objects from the Class

Objects can be created by calls of the form new("motif_gadem", ...).

Slots

```
pwm: PWM results.consensus: Sequences consensus.alignList: List of sequences alignment.name: Name of sequences.
```

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See Also

```
gadem, align, parameters
```

Examples

```
showClass("gadem")
```

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```
parameters-class Class "parameters"
```

Description

This object contains contains parameters of GADEM analysis

Objects from the Class

Objects can be created by calls of the form new("motif_gadem", ...).

Slots

```
numWordGroup: Number of non-zero k-mer groups.
numTop3mer: Number of top-ranked trimers for spaced dyads (default: 20).
verbose: Print immediate results on screen [1-yes (default), 0-no].
numTop4mer: Number of top-ranked tetramers for spaced dyads (default: 40).
numTop5mer: Number of top-ranked pentamers for spaced dyads (default: 60).
numGeneration: Number of genetic algorithm (GA) generations (default: 5).
populationSize :GA population size (default: 100).
pValue: P-value cutoff for declaring BINDING SITES (default: 0.0002).
eValue :ln(E-value) cutoff for selecting MOTIFS (default: 0.0).
extTrim: Base extension and trimming (1 -yes, 0 -no) (default: 1).
minSpaceWidth: Minimal number of unspecified nucleotides in spaced dyads (default: 0).
maxSpaceWidth: Maximal number of unspecified nucleotides in spaced dyads (default: 10).
useChIPscore: Use top-scoring sequences for deriving PWMs.
numEM: Number of EM steps (default: 40).
fEM: Fraction of sequences used in EM to obtain PWMs in an unseeded analysis (default: 0.5).
widthWt: For -posWt 1 or 3, width of central sequence region with large EM weights for PWM
     optimization (default: 50).
fullScan :GADEM keeps two copies of the input sequences internally.
slideWinPWM: Sliding window for comparing pwm similarity (default : 6).
stopCriterion
numBackgSets: Number of sets of background sequences (default: 10).
weight Type: Weight profile for positions on the sequence.
bFileName: Reading user-specified background models.
Spwm: File name for the seed PWM, when a seeded approach is used.
nSequences: Number of input sequences.
maskR : Mask low-complexity sequences or repeats.
nmotifs: Maximal number of motifs sought.
```

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See Also

```
gadem, align, motif
```

Examples

```
showClass("parameters")
```

readPWMfile

Read Transfac File

Description

This function is use to read standard Transfac type file.

Usage

```
readPWMfile(file)
```

Arguments

file

Transfac file's name.

Details

This function is designed to read standard Transfac type file. For more information about the format, please refere to http://mcast.sdsc.edu/doc/transfac-format.html

Value

A list of matrix.

Author(s)

```
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```

Examples

```
#####Database and Scores#####
path <- system.file("extdata","jaspar2009.txt",package="rGADEM")
jaspar <- readPWMfile(path)</pre>
```

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