

Package ‘memes’

May 20, 2024

Type Package

Title motif matching, comparison, and de novo discovery using the MEME Suite

Version 1.13.0

Description A seamless interface to the MEME Suite family of tools for motif analysis. 'memes' provides data aware utilities for using GRanges objects as entrypoints to motif analysis, data structures for examining & editing motif lists, and novel data visualizations. 'memes' functions and data structures are amenable to both base R and tidyverse workflows.

License MIT + file LICENSE

Encoding UTF-8

LazyData true

Imports Biostrings, dplyr, cmdfun (>= 1.0.2), GenomicRanges, ggplot2, ggseqlogo, magrittr, matrixStats, methods, patchwork, processx, purrr, rlang, readr, stats, tools, tibble, tidyr, utils, usethis, universalmotif (>= 1.9.3), xml2

Suggests cowplot, BSgenome.Dmelanogaster.UCSC.dm3, BSgenome.Dmelanogaster.UCSC.dm6, forcats, testthat (>= 2.1.0), knitr, MotifDb, heatmap, PMCMRplus, plyranges (>= 1.9.1), rmarkdown, covr

biocViews DataImport, FunctionalGenomics, GeneRegulation, MotifAnnotation, MotifDiscovery, SequenceMatching, Software

RoxygenNote 7.1.1

SystemRequirements Meme Suite (v5.3.3 or above)
<<http://meme-suite.org/doc/download.html>>

VignetteBuilder knitr

URL <https://snystrom.github.io/memes/>,
<https://github.com/snystrom/memes>

BugReports <https://github.com/snystrom/memes/issues>

Depends R (>= 4.1)

Config/testthat/edition 3**git_url** <https://git.bioconductor.org/packages/memes>**git_branch** devel**git_last_commit** 10e69e6**git_last_commit_date** 2024-04-30**Repository** Bioconductor 3.20**Date/Publication** 2024-05-20**Author** Spencer Nystrom [aut, cre, cph]
(<https://orcid.org/0000-0003-1000-1579>>)**Maintainer** Spencer Nystrom <nystromdev@gmail.com>**Contents**

memes-package	3
add_sequence	3
ame_compare_heatmap_methods	4
check_meme_install	5
drop_best_match	6
example_ame	6
example_ame_large	7
example_chip_summits	7
example_dreme	8
example_dreme_by_binding	9
example_dreme_tomtom	9
example_fimo	9
example_peaks	10
example_rnaseq	10
example_tomtom	11
force_best_match	11
get_sequence	12
has_duplicate_motifs	13
importAme	13
importDremeXML	14
importFimo	15
importMeme	15
importStremeXML	16
importTomTomXML	17
meme_is_installed	18
nest_tomtom	19
plot_ame_heatmap	19
plot_sequence_heatmap	21
remove_duplicate_motifs	22
runAme.list	23
runDreme	26
runFimo	29

<i>memes-package</i>	3
runMeme	31
runStreme	34
runTomTom	36
update_best_match	38
view_tomtom_hits	39
write_fasta	40
Index	41

<i>memes-package</i>	<i>memes: motif matching, comparison, and de novo discovery using the MEME Suite</i>
----------------------	--

Description

A seamless interface to the MEME Suite family of tools for motif analysis. 'memes' provides data aware utilities for using GRanges objects as entrypoints to motif analysis, data structures for examining & editing motif lists, and novel data visualizations. 'memes' functions and data structures are amenable to both base R and tidyverse workflows.

Author(s)

Maintainer: Spencer Nystrom <nystromdev@gmail.com> ([ORCID](#)) [copyright holder]

See Also

Useful links:

- <https://snystrom.github.io/memes/>
- <https://github.com/snystrom/memes>
- Report bugs at <https://github.com/snystrom/memes/issues>

<i>add_sequence</i>	<i>Add nucleic acid sequence of regions to metadata column</i>
---------------------	--

Description

Add nucleic acid sequence of regions to metadata column

Usage

```
add_sequence(ranges, genome, name = "sequence")
```

Arguments

ranges	GRanges object
genome	BSgenome object or any other valid input to 'Biostrings::getSeq()' (Do 'showMethods(Biostrings::getSeq)' to show valid types)
name	name of metadata column to hold sequence information (default: "sequence"). Note, this will overwrite existing columns without warning if the name already exists.

Value

'ranges' with new metadata column named "sequence" (or another value passed to 'name') holding the DNA or RNA sequence from 'genome'

Examples

```
data(example_peaks, package = "memes")
dm.genome <- BSgenome.Dmelanogaster.UCSC.dm3::BSgenome.Dmelanogaster.UCSC.dm3
add_sequence(example_peaks, dm.genome)
```

ame_compare_heatmap_methods

Compare AME heatmap methods

Description

This helper function allows the user to visualize the distribution of their AME results data on different scales to help understand the implications of using different values in 'plot_ame_heatmap()'

Usage

```
ame_compare_heatmap_methods(ame, group, value = -log10(adj.pvalue))
```

Arguments

ame	ame results data.frame
group	optional name of group to split results by
value	value to compare to "normalize" method (default: -log10(adj.pvalue))

Value

a cowplot 2 panel plot comparing the distribution of 'value' to normalized rank values

Examples

```
data("example_ame", package = "memes")
ame_compare_heatmap_methods(example_ame$Decreasing)

ame_compare_heatmap_methods(dplyr::bind_rows(example_ame, .id = "type"), type)
```

check_meme_install	<i>Check user's MEME install</i>
--------------------	----------------------------------

Description

In order to use the run* family of functions, memes must detect a local install of the MEME Suite. MEME is installed in a directory named meme/bin/ which can be located anywhere on the filesystem, but is typically found in ~/meme/bin. If the MEME Suite is installed at ~/meme/bin, memes can autodetect the install. However, in the case that the MEME Suite is found at a nonstandard location, the user may specify the location of their meme/bin in three ways:

Usage

```
check_meme_install(meme_path = NULL)
```

Arguments

meme_path	path to "meme/bin" (if unset will search MEME_BIN environment variable or meme_bin option)
-----------	--

Details

1. provide the full path to meme/bin to the meme_path argument to each run* function.
2. set the meme_bin option using options(meme_bin = "path/to/meme/bin") once per R session.
3. set the MEME_BIN environment variable either in .Renviro or ~/.bashrc with the path to meme/bin

To aid the user in determining if memes can detect their meme/bin install, check_meme_install() will search the aforementioned locations for a valid meme/bin, returning green checks for each detected tool, or red X's for undetected tools. Alternatively, users can run meme_is_installed() to get a boolean value indicating whether their MEME Suite can be detected.

check_meme_install() searches using the following heirarchy. This heirarchy mimics how all run* functions search for meme/bin, thus the paths printed from check_meme_install() will indicate the paths used by each run* function. The heirarchy is as follows:

1. the meme_path function argument if set
2. the meme_bin option
3. the MEME_BIN environment variable
4. the default location at ~/meme/bin

Value

message indicating which MEME utilities are installed and their location on disk

Examples

```
check_meme_install()
```

drop_best_match	<i>Drop best match columns from tomtom results</i>
-----------------	--

Description

Convenience function for dropping all columns created by runTomTom prefixed by "best_match_" and the "best_db_name" column. Keeps the "tomtom" data.frame column. Can be useful if you want to unnest the 'tomtom' data without propagating these columns.

Usage

```
drop_best_match(res)
```

Arguments

res results of runTomTom

Value

'res' without the tomtom best_match_ columns

Examples

```
data("example_dreme_tomtom")
names(example_dreme_tomtom)
names(drop_best_match(example_dreme_tomtom))
```

example_ame	<i>Example runAme() output</i>
-------------	--------------------------------

Description

Result when running AME using 100bp window around 'example_chip_summits' for "Increasing" and "Decreasing" sites, using "Static" as background.

Usage

```
example_ame
```

Format

A list object of AME results data.frames

Increasing 'runAme()' Results object for Increasing sites vs Static sites

Decreasing 'runAme()' Results object for Decreasing sites vs Static sites

Examples

```
# Data can be combined into 1 large data.frame using:  
# where the "behavior" column will hold the "Increasing"/"Decreasing" information  
dplyr::bind_rows(example_ame, .id = "behavior")
```

example_ame_large	<i>runAme() output for example_chip_summits split by binding description</i>
-------------------	--

Description

AME was run for "ectopic", "entopic", and "orphan" sites using shuffled background.

Usage

```
example_ame_large
```

Format

a list of runAme() results data.frames

Details

see ‘vignette("integrative_analysis", package = "memes)” for details.

Examples

```
# Data can be combined into 1 large data.frame using:  
dplyr::bind_rows(example_ame_large, .id = "binding_type")
```

example_chip_summits	<i>Annotated Transcription Factor ChIP-seq summits</i>
----------------------	--

Description

ChIP-seq summit positions on *Drosophila melanogaster* chromosome 3 for the transcription factor E93 using a union set of peaks from third-instar larval wings ("Early") and 24 hour Pupal ("Late") wings.

Usage

```
example_chip_summits
```

Format

A GRanges object of ChIP summit position with 2 metadata columns

peak_binding_description Binding profiles between Early and Late E93 were compared. Peaks are annotated as whether they are bound in Early wings only ("ectopic"), both Early and Late wings ("entopic"), or only bound in Late wings ("orphan").

e93_sensitive_behavior change in chromatin accessibility in response to E93 binding: Increasing, Decreasing, or Static

Details

E93 is a transcription factor normally present only in Late wings. An experimental perturbation precociously expressed E93 during Early stages. Binding profiles between Early and Late E93 were compared. Peaks are annotated as whether they are bound in Early wings only ("ectopic"), both Early and Late wings ("entopic"), or only bound in Late wings ("orphan").

DNA elements can be made "open" or "closed" in response to binding of transcription factors like E93. Accessibility of E93 binding sites before and after E93 expression was measured using FAIRE-seq. ChIP peaks are annotated by how their accessibility changes in response to E93 binding. Peaks can become more open ("Increasing"), more closed ("Decreasing"), or unchanged in accessibility ("Static"). These experiments demonstrate a causal relationship between E93 binding and both opening and closing of DNA elements.

Source

<https://www.ncbi.nlm.nih.gov/geo/download/?acc=GSE141738&format=file&file=GSE141738>

example_dreme

Example runDreme() output

Description

Result when running dreme using 100bp window around example_chip_summits using "Decreasing" sites as foreground, and "Static" sites as background.

Usage

```
example_dreme
```

Format

```
a runDreme results data.frame
```

```
example_dreme_by_binding
```

runDreme() output for example_chip_summits split by binding description

Description

See vignette("integrative_analysis", package = "memes") for details

Usage

```
example_dreme_by_binding
```

Format

a runDreme results data.frame

```
example_dreme_tomtom
```

Example runDreme() output after passing to runTomTom()

Description

Result when running 'runTomTom(example_dreme)' using FlyFactorSurvey as database.

Usage

```
example_dreme_tomtom
```

Format

a runDreme results data.frame with runTomTom results columns

```
example_fimo
```

Example runFimo() output

Description

Run using 100bp windows around 'example_chip_summits', using E93 motif as database.

Usage

```
example_fimo
```

Format

A GRanges object of E93 motif positions within 100bp windows of 'example_chip_summits'

example_peaks	<i>Example ChIP-seq peaks</i>
---------------	-------------------------------

Description

10 ChIP-seq peaks from GSE141738

Usage

example_peaks

Format

An object of class GRanges of length 10.

Details

A small number of transcription factor ChIP-seq peaks as a GRanges object, taken from [GSE141738](https://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE141738)

Source

<https://www.ncbi.nlm.nih.gov/geo/download/?acc=GSE141738&format=file&file=GSE141738>

example_rnaseq	<i>RNAseq data from Early and Late Drosophila wings</i>
----------------	---

Description

These data are a subset of RNAseq counts from the full FPKM table in GSE97956. Includes counts for all Drosophila transcription factors and ~200 other random genes.

Usage

example_rnaseq

Format

A data.frame of RNAseq FPKM data

symbol The FlyBase gene symbol

time Developmental stage of RNAseq experiment

fpkm RNAseq count in Fragments per Kilobase Million (FPKM)

Source

"<https://www.ncbi.nlm.nih.gov/geo/download/?acc=GSE97956&format=file&file=GSE97956>"

example_tomtom	<i>Example runTomTom() output</i>
----------------	-----------------------------------

Description

Result when running 'runTomTom(example_dreme\$motif)' using FlyFactorSurvey as database

Usage

```
example_tomtom
```

Format

a data.frame

force_best_match	<i>Force best tomtom match by id</i>
------------------	--------------------------------------

Description

Although TomTom assigns a best match, this is not always the most biologically relevant match. In these cases, it is useful to "force" the best match to another lower ranked, but still significant TomTom match. This function allows users to select a new best match motif from the set of lower-ranked matches in the 'tomtom' list column. This function also reorders the 'tomtom' data.frame such that the forced match is the first row of the 'tomtom' entry.

Usage

```
force_best_match(res, matches)
```

Arguments

res	results from runTomTom
matches	named vector where name is the input motif name, and value is the match_name to use as the new best match

Value

'res' with new best_* columns and re-ranked tomtom data in the 'tomtom' list column for the updated entries.

See Also

[update_best_match()]

Examples

```

if (meme_is_installed()){
  motif <- universalmotif::create_motif("CCRAAAW", name = "example_motif")
  db <- system.file("extdata", "flyFactorSurvey_cleaned.meme", package = "memes")
  res <- runTomTom(motif, database = db)
  res$best_match_name
  res2 <- force_best_match(res, c("example_motif" = "Eip93F_SANGER_10"))
  res2$best_match_name
}

```

get_sequence

Get sequence from GRanges

Description

A light wrapper around `Biostrings::getSeq` to return named `DNASTringSets`, from input genomic coordinates.

Usage

```
get_sequence(regions, genome, score_column, ...)
```

Arguments

<code>regions</code>	<code>GRanges</code> , or <code>GRangesList</code> object. Will also accept a <code>data.frame</code> as long as it can be coerced to a <code>GRanges</code> object, or a string in the format: "chr:start-end" (NOTE: use 1-based closed intervals, not BED format 0-based half-open intervals).
<code>genome</code>	object of any valid type in <code>'showMethods(Biostrings::getSeq)'</code> . Commonly a <code>BSgenome</code> object, or fasta file. Used to look up sequences in regions.
<code>score_column</code>	optional name of column (in <code>mcols()</code> of <code>'regions'</code>) containing a fasta score that is added to the fasta header of each entry. Used when using <code>[runAme()]</code> in partitioning mode. (default: <code>'NULL'</code>)
<code>...</code>	additional arguments passed to <code>Biostrings::getSeq</code> .

Value

`'Biostrings::DNASTringSet'` object with names corresponding to genomic coordinates. If input is a list object, output will be a `'Biostrings::BStringSetList'` with list names corresponding to input list names.

Examples

```
# using character string as coordinates
# using BSgenome object for genome
drosophila.genome <- BSgenome.Dmelanogaster.UCSC.dm6::BSgenome.Dmelanogaster.UCSC.dm6
get_sequence("chr2L:100-200", drosophila.genome)

# using GRanges object for coordinates
data(example_peaks, package = "memes")
get_sequence(example_peaks, drosophila.genome)
```

has_duplicate_motifs *Check for duplicated motif matrices*

Description

This function identifies whether any motif matrices in the input universalmotif list or universalmotif_df are identical to each other. Note: this operation is slow on large motif lists

Usage

```
has_duplicate_motifs(x)
```

Arguments

x a universalmotif list or universalmotif_df

Value

logical value indicating presence or absence of duplicated motif matrices

Examples

```
motif <- universalmotif::create_motif()
duplicated <- c(motif, motif)
has_duplicate_motifs(duplicated)
```

importAme *Parse AME output*

Description

This imports AME results using the "ame.tsv" output, and optionally the "sequences.tsv" output if run with "method = fisher". AME results differ based on the method used, thus this must be set on import or the column names will be incorrect.

Usage

```
importAme(path, method = "fisher", sequences = FALSE)
```

Arguments

`path` path to ame results file ("ame.tsv")

`method` ame run method used (one of: c("fisher", "ranksum", "dmhg3", "dmhg4", "pearson", "spearman")). Default: "fisher".

`sequences` FALSE or path to sequences file (only valid for method = "fisher")

Value

data.frame with method-specific results. See [AME results](http://meme-suite.org/doc/ame-output-format.html) webpage for more information. If `sequences` is set to a path to the sequences.tsv and `method = "fisher"`, the list-column 'sequences' will be appended to resulting data.frame.

See Also

[runAme()]

Examples

```
ame_tsv <- system.file("extdata", "ame.tsv", package = "memes", mustWork = TRUE)
importAme(ame_tsv)
```

importDremeXML

Import Dreme output from previous run

Description

Import Dreme output from previous run

Usage

```
importDremeXML(dreme_xml_path)
```

Arguments

`dreme_xml_path` path to dreme.xml file

Value

data.frame with statistics for each discovered motif. The 'motifs' column contains a universal motif object representation in PCM format of each DREME motif. If no motifs are discovered, returns NULL.

See Also

[runDreme()]

Examples

```
dreme_xml <- system.file("extdata", "dreme.xml", package = "memes")
importDremeXML(dreme_xml)
```

importFimo	<i>Import FIMO results</i>
------------	----------------------------

Description

Import FIMO results

Usage

```
importFimo(fimo_tsv)
```

Arguments

fimo_tsv path to fimo.tsv output file

Value

GenomicRanges object for each match position. Note unless coordinates are genomic positions, each 'seqnames' entry will be the fasta header, and start/end will be the position within that sequence of the match.

Examples

```
fimo_tsv <- system.file("extdata", "fimo.tsv", package = "memes")
importFimo(fimo_tsv)
```

importMeme	<i>Import MEME results</i>
------------	----------------------------

Description

This is a light wrapper around [universalmotif::read_meme()] that imports MEME results as universalmotif data.frame. If MEME is run with genomic coordinates in the fasta header, in "chr:start-end" format (base 1 indexed), the genomic coordinates of the motif match from input sequences can be parsed from the header.

Usage

```
importMeme(meme_txt, parse_genomic_coord = FALSE, combined_sites = FALSE)
```

Arguments

`meme_txt` path to "meme.txt" output

`parse_genomic_coord` whether to parse sequence headers into genomic coordinates for motif position information, only works if fasta files were written such that the sequence headers are in the form: "chr:start-end", or some variation of this form (delimiters can be any of: "[^[:alnum:]]+" (ie non-alphanumeric characters)), (default = FALSE).

`combined_sites` whether to add 'combined_sites' output which contains coordinates of each sequence, the motif sequence (if 'parse_genomic_coord = TRUE'), and the 'diagram' column row output from MEME indicating the relative locations of motifs in the sequence.

Value

MEME results in `universalmotif` data.frame format (see: `[as_universalmotif_dataframe()]`). 'sites_hits' is a nested data.frame column containing the position within each input sequence of matches to the identified motif.

See Also

`[runMeme()]` `[universalmotif::read_meme()]`

Examples

```
example_meme_txt <- system.file("extdata", "meme_full.txt", package = "universalmotif")
importMeme(example_meme_txt)
```

<code>importStremeXML</code>	<i>Import Streme output from previous run</i>
------------------------------	---

Description

Import Streme output from previous run

Usage

```
importStremeXML(streme_xml_path)
```

Arguments

`streme_xml_path`
path to streme.xml file

Value

data.frame with statistics for each discovered motif. The 'motifs' column contains a `universalmotif` object representation in PCM format of each DREME motif. If no motifs are discovered, returns NULL.

See Also

[runStreme()]

Examples

```
streme_xml <- system.file("extdata", "streme.xml", package = "memes")
importStremeXML(streme_xml)
```

```
importTomTomXML
```

Import tomtom data from previous run

Description

Import tomtom data from previous run

Usage

```
importTomTomXML(tomtom_xml_path)
```

Arguments

```
tomtom_xml_path
                path to tomtom.xml
```

Details

tomtom list column format the 'tomtom' list column contains data.frames with the following format:
 - name: name of query PWM - altname: alternate name of query PWM - match_name: name of matched PWM - match_altname: alt name of matched PWM - match_pval: p-value of match - match_eval: E-value of match - match_qval: q-value of match - match_offset: number of letters the query was offset from the target match - match_strand: whether the motif was found on input strand (+) or as reverse-complement (-) - db_name: database source of matched motif - match_motif: universalmotif object containing the PWM that was matched

Value

will return data.frame with input motifs & results for best match. 'tomtom' list column contains full tomtom data for each input motif. NOTE: if tomtom detects no matches for any input motif, currently will print a message & return NA values for 'tomtom', 'best_match_name', and 'best_match_motif'.

See Also

[runTomTom()]

Examples

```
tomtom_xml <- system.file("extdata", "tomtom.xml", package = "memes")
importTomTomXML(tomtom_xml)
```

meme_is_installed	Returns logical vector indicating valid MEME-Suite install status
-------------------	---

Description

Checks for a valid meme install using same heirarchy as `check_meme_install()`. Returns TRUE if all supported utilities are found in the meme install location, FALSE if not.

Usage

```
meme_is_installed(path = NULL)
```

Arguments

path	optional path to "meme/bin/". If unset, will follow the search heirarchy listed above.
------	--

Details

The search heirarchy is as follows:

1. the `meme_path` function argument if set
2. the `meme_bin` option
3. the `MEME_BIN` environment variable
4. the default location at `~/meme/bin`

Value

logical(1) indicating whether meme is installed with all supported utilities

See Also

[check_meme_install\(\)](#)

Examples

```
meme_is_installed()
```

nest_tomtom	<i>Nest TomTom results columns into a data.frame column named "tomtom"</i>
-------------	--

Description

This is a convenience function for re-nesting the ‘tomtom’ list column if the user unnests it. Additionally, it will update the best_match information based on the ranking of the resulting ‘tomtom’ data.frame. This avoids having out-of-date best_match information after manipulating the ‘tomtom’ entries.

Usage

```
nest_tomtom(data)
```

Arguments

data tomtom results data.frame after unnesting the ‘tomtom’ column

Details

****NOTE:**** that the resulting columns may not be in the same order, so operations like ‘identical()’ before & after a nest/reneest operation may fail even though the column values are unchanged.

Value

the input data.frame with the match_* columns nested into a column named ‘tomtom’

Examples

```
if (meme_is_installed()){
  motif <- universalmotif::create_motif("CCRAAAW")
  db <- system.file("extdata/flyFactorSurvey_cleaned.meme", package = "memes")
  res <- runTomTom(motif, database = db)
  data <- tidyr::unnest(res, "tomtom")
  identical(nest_tomtom(data), res)
}
```

plot_ame_heatmap	<i>Plot AME heatmap clustered by similarity in detected motifs</i>
------------------	--

Description

Plot AME heatmap clustered by similarity in detected motifs

Usage

```
plot_ame_heatmap(
  ame,
  id = motif_id,
  group = NULL,
  value = -log10(adj.pvalue),
  group_name = NULL,
  scale_max = NA
)
```

Arguments

ame	ame results data.frame
id	column of motif ids to use (default: motif_id).
group	grouping column if comparing across multiple ame runs (optional, default: NULL).
value	value to display as heatmap intensity. Default: -log10(adj.pvalue). Takes function or column name as input. If set to "normalize", will use normalized rank within 'group' as the heatmap values. **If in doubt** , prefer the -log10(adj.pvalue) plot potentially in conjunction with adjusting 'scale_max'. (See "Normalized rank visualization" section below for more details on how to interpret these data)
group_name	when group = NULL, name to use for input regions. Ignored if group is set.
scale_max	max heatmap value to limit upper-value of scale. Useful if distribution of 'value's vary greatly between groups. Usually a better to tweak this option than to use value = "normalize". The cumulative distribution plot generated by 'ame_compare_heatmap_methods()' can be useful for selecting this value, try to pick a value which captures the largest fraction of hits across all groups while excluding outliers.

Details

Normalized rank visualization ****NOTE:**** The normalized rank visualization eliminates all real values related to statistical significance! Instead, this visualization represents the relative ranks of hits within an AME run, which already pass a significance threshold set during 'runAME()'. This means that even if several motifs have similar or even identical pvalues, their heatmap representation will be a different color value based on their ranked order in the results list. This also means that using the normalized rank visualization will be misleading if there are only a few AME hits; it is only worth using if the number of hits is very large (>100). Both visualizations can be useful and reveal different properties of the data to the user during data exploration. Use 'ame_compare_heatmap_methods()' to help assess differences in the two visualizations. ****If in doubt****, prefer the '-log10(adj.pvalue)' representation.

Common mistake: if 'value' is set to a string that is not "normalize", it will return: "Error: Discrete value supplied to continuous scale". To use a column by name, do not quote the column name.

Value

'ggplot' object

Examples

```
data("example_ame", package = "memes")

# Plot a single category heatmap
plot_ame_heatmap(example_ame$Decreasing)

# Plot a multi category heatmap
grouped_ame <- dplyr::bind_rows(example_ame, .id = "category")
plot_ame_heatmap(grouped_ame, group = category)
```

plot_sequence_heatmap *Visualize a heatmap of aligned sequences*

Description

Sometimes it is useful to visualize individual motif matches in aggregate to understand how sequence variability contributes to motif matches. This function creates a heatmap where each row represents a single sequence and each column represents a position. Cells are colored by the sequence at that position. Sequences are optionally aggregated into a sequence logo aligned in register with the heatmap to visualize how sequence variability contributes to motif makeup.

Usage

```
plot_sequence_heatmap(
  sequence,
  title = NULL,
  logo = TRUE,
  alph = c("DNA", "RNA", "AA"),
  title_hjust = 0,
  heights = c(1, 5),
  legend = "none"
)
```

Arguments

sequence	character vector of sequences, plot will be ranked in order of the sequences. Each sequence must be equal length. Alternately, sequence can be a named list in which case each plot will be titled by the names of the list entries.
title	title of the plot. Default: NULL. If sequence is a named list of sequences, title defaults to the list entry names. Set to NULL to override this behavior. To use a different title than the list entry name, pass a vector of names to ‘title’.
logo	whether to include a sequence logo above the heatmap
alph	alphabet colorscheme to use. One of: DNA, RNA, AA.
title_hjust	value from 0 to 1 determining the horizontal justification of the title. Default: 0.
heights	ratio of logo:heatmap heights. Given as: c(logo_height, heatmap_height). Values are not absolute. Ignored when logo = FALSE.
legend	passed to ggplot2::theme(legend.position). Default: "none". Values can be: "none", "left", "right", "top", "bottom", or coordinates in c(x,y) format.

Value

a ggplot object of the sequence heatmap ranked by the order of sequences

See Also

runFimo

Examples

```
data(example_fimo, package = "memes")
genome <- BSgenome.Dmelanogaster.UCSC.dm3::BSgenome.Dmelanogaster.UCSC.dm3
motifs <- add_sequence(example_fimo, genome)
plot_sequence_heatmap(motifs$sequence)

# Use on named list
sequences <- list("set 1" = motifs$sequence[1:100],
                 "set 2" = motifs$sequence[101:200])
plot_sequence_heatmap(sequences)

# Use different titles for list input
plot_sequence_heatmap(sequences, title = c("A", "B"))
```

remove_duplicate_motifs

Remove duplicated motif entries

Description

This function identifies motif matrices which are duplicated in a universalmotif list or universalmotif_df and removes them. This operation ignores motif metadata and operates by removing all entries whose motif matrices are identical. The first instance of a duplicated motif in the input list is the one returned.

Usage

```
remove_duplicate_motifs(x)
```

Arguments

x a universalmotif list or universalmotif_df

Value

A deduplicated list or universalmotif_df

Examples

```
motif <- universalmotif::create_motif()
duplicated <- c(motif, motif)
remove_duplicate_motifs(duplicated)
```

`runAme.list`*Motif enrichment using AME*

Description

AME identifies known motifs (provided by the user) that are enriched in your input sequences.

Usage

```
## S3 method for class 'list'
runAme(
  input,
  control = "shuffle",
  outdir = "auto",
  method = "fisher",
  database = NULL,
  meme_path = NULL,
  sequences = FALSE,
  silent = TRUE,
  ...
)

## S3 method for class 'BStringSetList'
runAme(
  input,
  control = "shuffle",
  outdir = "auto",
  method = "fisher",
  database = NULL,
  meme_path = NULL,
  sequences = FALSE,
  silent = TRUE,
  ...
)

## Default S3 method:
runAme(
  input,
  control = "shuffle",
  outdir = "auto",
  method = "fisher",
  database = NULL,
  meme_path = NULL,
  sequences = FALSE,
  silent = TRUE,
  ...
)
```

```

runAme(
  input,
  control = "shuffle",
  outdir = "auto",
  method = "fisher",
  database = NULL,
  meme_path = NULL,
  sequences = FALSE,
  silent = TRUE,
  ...
)

```

Arguments

input	path to fasta, or DNASTringset (optional: DNASTringSet object names contain fasta score, required for partitioning mode)
control	default: "shuffle", or set to DNASTringset or path to fasta file to use those sequences in discriminative mode. If input is a list of DNASTringSet objects, and control is set to a character vector of names in input, those regions will be used as background in discriminative mode and AME will skip running on any control entries (NOTE: if input contains an entry named "shuffle" and control is set to "shuffle", it will use the input entry, not the AME shuffle algorithm). If control is a Biostrings::BStringSetList (generated using get_sequence), all sequences in the list will be combined as the control set. Set to NA for partitioning based on input fasta score (see get_sequence() for assigning fasta score). If input sequences are not assigned fasta scores but AME is run in partitioning mode, the sequence order will be used as the score.
outdir	Path to output directory location to save data files. If set to "auto", will use location of input files if passing file paths, otherwise will write to a temporary directory. default: "auto"
method	default: fisher (allowed values: fisher, ranksum, pearson, spearman, 3dmhg, 4dmhg)
database	path to .meme format file, universalmotif list object, dreme results data.frame, or list() of multiple of these. If objects are assigned names in the list, that name will be used as the database id in the output. It is highly recommended you set a name if not using a file path so the database name will be informative, otherwise the position in the list will be used as the database id. For example, if the input is: list("motifs.meme", list_of_motifs), the database id's will be: "motifs.meme" and "2". If the input is list("motifs.meme", "customMotifs" = list_of_motifs), the database id's will be "motifs.meme" and "customMotifs".
meme_path	path to "meme/bin/" (default: NULL). Will use default search behavior as described in check_meme_install() if unset.
sequences	logical(1) add results from sequences.tsv to sequences list column to returned data.frame. Valid only if method = "fisher". See AME outputs webpage for more information (Default: FALSE).
silent	whether to suppress stdout (default: TRUE), useful for debugging.

... additional arguments passed to AME (see AME Flag table below)

Details

AME can be run using several modes:

1. Discriminative mode: to discover motifs enriched relative to shuffled input, or a set of provided control sequences
2. Partitioning mode: in which AME uses some biological signal to identify the association between the signal and motif presence.

To read more about how AME works, see the [AME Tutorial](#)

Additional AME arguments

memes allows passing any valid flag to it's target programs via ... For additional details for all valid AME arguments, see the [AME Manual](#) webpage. For convenience, a table of valid parameters, and brief descriptions of their function are provided below:

AME Flag	allowed values	default	description
kmer	integer(1)	2	kmer frequency to preserve when shuffling
seed	integer(1)	1	seed for random number generator when
scoring	"avg", "max", "sum", "totalhits"	"avg"	Method for scoring a sequence for match
hit_lo_fraction	numeric(1)	0.25	fraction of hit log odds score to exceed to
evaluate_report_threshold	numeric(1)	10	E-value threshold for reporting a motif as
fasta_threshold	numeric(1)	0.001	AME will classify sequences with FASTA
fix_partition	numeric(1)	NULL	AME evaluates only the partition of the f
poslist	"pwm", "fasta"	"fasta"	When using partitioning mode (control
log_fscores	logical(1)	FALSE	Convert FASTA scores into log-space (onl
log_pwmcores	logical(1)	FALSE	Convert PWM scores into log-space (onl
lingreg_switchxy	logical(1)	FALSE	Make the x-points FASTA scores and y-p
xalph	file path	NULL (1)	alphabet file to use if input motifs are in
bfile	"motif", "motif-file", "uniform", path to file	NULL	source of 0-order background model. If "
motif_pseudo	numeric(1)	0.1	Add this pseudocount when converting
inc	character(1)	NULL	use only motifs with names matching thi
exc	character(1)	NULL	exclude motifs with names matching thi

Value

a data.frame of AME enrichment results. If run using a BStringsSetList input, returns a list of data.frames.

Citation

If you use runAme() in your analysis, please cite:

Robert McLeay and Timothy L. Bailey, "Motif Enrichment Analysis: A unified framework and method evaluation", BMC Bioinformatics, 11:165, 2010, doi:10.1186/1471-2105-11-165. [full text](#)

Licensing:

The MEME Suite is free for non-profit use, but for-profit users should purchase a license. See the [MEME Suite Copyright Page](#) for details.

Examples

```

if (meme_is_installed()) {
# Create random named sequences as input for example
seqs <- universalmotif::create_sequences(rng.seed = 123)
names(seqs) <- seq_along(seqs)

# An example path to a motif database file in .meme format
motif_file <- system.file("extdata", "flyFactorSurvey_cleaned.meme", package = "memes")

runAme(seqs, database = motif_file)

# Dreme results dataset for example
dreme_xml <- system.file("extdata", "dreme.xml", package = "memes")
dreme_results <- importDremeXML(dreme_xml)

# database can be set to multiple values like so:
runAme(seqs, database = list(motif_file, "my_dreme_motifs" = dreme_results))
}

```

runDreme

Denovo motif discovery of target regions using DREME

Description

DREME discovers short, ungapped, *de-novo* motifs that are relatively enriched relative to a control set of sequences. DREME can be run to discover motifs relative to a shuffled set of input sequences, or against a separately provided set of "control" sequences.

Usage

```
runDreme(input, control, outdir = "auto", meme_path = NULL, silent = TRUE, ...)
```

Arguments

input	regions to scan for motifs. Can be any of: <ul style="list-style-type: none"> • path to fasta file • DNASTringSet object (can be generated from GRanges using <code>get_sequence()</code>) • List of DNASTringSet objects (generated from <code>get_sequence()</code>) • <i>NOTE</i>: if using StringSet inputs, each entry must be named (set with <code>names()</code>). • <i>NOTE</i>: If you want to retain the raw dreme output files, you must use a path to fasta file as input, or specify an "outdir"
control	regions to use as background for motif search. Can be any of: <ul style="list-style-type: none"> • path to fasta file • DNASTringSet object (can be generated from GRanges using <code>get_sequence()</code>) • A Biostrings::BStringSetList (generated using <code>get_sequence()</code>), in which case all sequences in the list will be combined as the control set.

	<ul style="list-style-type: none"> • if input is a list of DNAStrngSet objects, a character vector of names in input will use those sequences as background. runDreme will not scan those regions as input. • "shuffle" to use dreme's built-in dinucleotide shuffle feature (NOTE: if input is a list object with an entry named "shuffle", the list entry will be used instead). Optionally can also pass seed = <any number> to . . . to use as the random seed during shuffling. If no seed is passed, dreme will use 1 as the random seed, so results will be reproducible if rerunning. NOTE: beware system-specific differences. As of v5, dreme will compile using the default python installation on a system (either python2.7 or python3). The random number generator changed between python2.7 and python3, so results will not be reproducible between systems using python2.7 vs 3.
outdir	path to output directory of dreme files, or "auto" to autogenerate path. Default: location of input fasta in dir named "\<input>\vs\<control>". If input is DNAStringset, will be temporary path. This means that if you want to save the raw output files, you must use fasta files as input or use an informative (and unique) outdir name. memes will not check if it overwrites files in a directory. Directories will be recursively created if needed.
meme_path	optional, path to "meme/bin/" on your local machine. runDreme will search 3 places in order for meme if this flag is unset: <ol style="list-style-type: none"> 1. the option "meme_bin" (set with options(meme_bin = "path/to/meme/bin")) 2. the environment variable "MEME_PATH" (set in .Renviro) 3. "~/meme/bin/" as the default location <ul style="list-style-type: none"> • If the user sets meme_path in the function call, this value will always be used
silent	whether to suppress printing dreme stdout as a message when finishing with no errors. Can be useful for troubleshooting in situations where no motifs are discovered, but command completes successfully. (default: TRUE)
. . .	dreme flags can be passed as R function arguments to use non-default behavior. For a full list of valid arguments, run your local install of dreme -h, or visit the dreme documentation website . See list below for aliases of common flags. To set flags with no values (ex. -dna), pass the argument as a boolean value (ex. dna = TRUE).

Details

Properly setting the control parameter is key to discovering biologically relevant motifs. Often, using control = "shuffle" will produce a suboptimal set of motifs; however, some discriminative analysis designs don't have proper "control" regions other than to shuffle.

As of MEME version 5.2.0, DREME is deprecated. Consider [runStreme\(\)](#) instead.

In addition to allowing any valid flag of dreme to be passed to . . . , we provide a few user-friendly aliases for common flags which are more readable (see list below). For example, e = 1 will use a max evalue cutoff of 1. This is equivalent to setting evalue = 1. For additional details about each DREME flag, see the [DREME Manual Webpage](#).

List of values which can be passed to . . . : **NOTE:** values must be referred to using their name in the "memes alias" column, not the "DREME Flag" column.

memes alias	DREME Flag	description	default
nmotifs	m	max number of motifs to discover	NULL
sec	t	max number of seconds to run	NULL
evaluate	e	max E-value cutoff	0.05
seed	s	random seed if using "shuffle" as control	1
ngen	g	nuber of REs to generalize	100
mink	mink	minimum motif width to search	3
maxk	maxk	maximum motif width to search	7
k	k	set mink and maxk to this value	NULL
norc	norc	search only the input strand for sequences	FALSE
dna	dna	use DNA alphabet	TRUE
rna	rna	use RNA alphabet	FALSE
protein	protein	use protein alphabet (NOT RECCOMENDED)	FALSE

Value

universalmotif_df with statistics for each discovered motif. The motif column contains a universalmotif object representation in PCM format of each DREME motif. If no motifs are discovered, returns NULL. The column values are as follows:

- rank = ranked order of discovered motif
- name = primary name of motif
- altname = alternative name of motif
- seq = consensus sequence of the motif
- length = length of discovered motif
- nsites = number of times the motif is found in input sequences
- positive_hits = number of sequences in input containing at least 1 of the motif
- negative_hits = number of sequences in control containing at least 1 of the motif
- pval = p-value
- eval = E-value
- unerasd_eval = Unerased E-Value
- positive_total = number of sequences in input
- negative_total = number of sequences in control
- pos_frac = fraction of positive sequences with a hit
- neg_frac = fraction of negative sequences with a hit
- motif = a universalmotif object of the discovered motif

Citation

If you use runDreme() in your analysis, please cite:

Timothy L. Bailey, "DREME: Motif discovery in transcription factor ChIP-seq data", *Bioinformatics*, 27(12):1653-1659, 2011. [full text](#)

Licensing:

The MEME Suite is free for non-profit use, but for-profit users should purchase a license. See the [MEME Suite Copyright Page](#) for details.

Examples

```

if (meme_is_installed()) {
# Create random named sequences as input for example
seqs <- universalmotif::create_sequences(rng.seed = 123)
names(seqs) <- seq_along(seqs)

# Runs dreame with default settings, shuffles input as background
runDreame(seqs, "shuffle")

# Runs searching for max 2 motifs, e-value cutoff = 0.1, explicitly using the DNA alphabet
runDreame(seqs, "shuffle", nmotifs = 2, e = 0.1, dna = TRUE)
}

```

runFimo

*Find instances of motifs using FIMO***Description**

FIMO scans input sequences to identify the positions of matches to each input motif. FIMO has no sequence length or motif number restrictions.

Usage

```

runFimo(
  sequences,
  motifs,
  bfile = "motif",
  outdir = "auto",
  parse_genomic_coord = TRUE,
  skip_matched_sequence = FALSE,
  max_strand = TRUE,
  text = TRUE,
  meme_path = NULL,
  silent = TRUE,
  ...
)

```

Arguments

sequences	path to fasta file, or stringset input.
motifs	path to .meme format file, or universalmotif/universalmotif list input.
bfile	path to background file, or special values: "motif" to use 0-order frequencies contained in the motif, or "uniform" to use a uniform letter distribution. (default: "motif")
outdir	output directory location. Only used if text = FALSE. Default: "auto" to auto-generate directory name. Note: if not using a fasta path as input, this will be a temporary location unless explicitly set.

parse_genomic_coord	logical(1) whether to parse genomic position from fasta headers. Fasta headers must be UCSC format positions (ie "chr:start-end"), but base 1 indexed (GRanges format). If names of fasta entries are genomic coordinates and parse_genomic_coord == TRUE, results will contain genomic coordinates of motif matches, otherwise FIMO will return relative coordinates (i.e. positions from 1 to length of the fasta entry).
skip_matched_sequence	logical(1) whether or not to include the DNA sequence of the match. Default: FALSE. Note: jobs will complete faster if set to TRUE. add_sequence() can be used to lookup the sequence after data import if parse_genomic_coord is TRUE, so setting this flag is not strictly needed.
max_strand	if match is found on both strands, only report strand with best match (default: TRUE).
text	logical(1) (default: TRUE). No output files will be created on the filesystem. The results are unsorted and no q-values are computed. This setting allows fast searches on very large inputs. When set to FALSE FIMO will discard 50% of the lower significance matches if >100,000 matches are detected. text = FALSE will also incur a performance penalty because it must first read a file to disk, then read it into memory. For these reasons, I suggest keeping text = TRUE.
meme_path	path to meme/bin/ (optional). Default: NULL, searches "MEME_PATH" environment variable or "meme_path" option for path to "meme/bin/".
silent	logical(1) whether to suppress stdout/stderr printing to console (default: TRUE). If the command is failing or giving unexpected output, setting silent = FALSE can aid troubleshooting.
...	additional commandline arguments to fimo. See the FIMO Flag table below.

Details

Additional arguments passed to See: [Fimo web manual](#) for a complete description of FIMO flags.

FIMO Flag	allowed values	default	description
alpha	numeric(1)	1	alpha for calculating position-specific priors. Rep
bfile	"motif", "motif-file", "uniform", file path,	"motif"	If "motif" or "motif-file", use 0-order letter frequ
max_stored_scores	integer(1)	NULL	maximum number of scores to be stored for comp
motif_pseudo	numeric(1)	0.1	pseudocount added to motif matrix
no_qvalue	logical(1)	FALSE	only needed when text = FALSE, do not compute
norc	logical(1)	FALSE	Do not score reverse complement strand
prior_dist	file path	NULL	file containing binned distribution of priors
psp	file path	NULL	file containing position specific priors. Requires p
qv_thresh	logical(1)	FALSE	use q-values for the output threshold
thresh	numeric(1)	1e-4	output threshold for returning a match, only matc

Licensing:

The MEME Suite is free for non-profit use, but for-profit users should purchase a license. See the [MEME Suite Copyright Page](#) for details.

Value

GRanges object containing positions of each match. Note: if `parse_genomic_coords = FALSE`, each `seqnames` entry will be the full fasta header, and `start/end` will be the relative position within that sequence of the match. The GRanges object has the following additional `mcols`: * `motif_id` = primary name of matched motif * `motif_alt_id` = alternate name of matched motif * `score` = score of match (higher score is a better match) * `pvalue` = pvalue of the match * `qvalue` = qvalue of the match * `matched_sequence` = sequence that matches the motif

Citation

If you use `runFimo()` in your analysis, please cite:

Charles E. Grant, Timothy L. Bailey, and William Stafford Noble, "FIMO: Scanning for occurrences of a given motif", *Bioinformatics*, 27(7):1017-1018, 2011. [full text](#)

Examples

```
if (meme_is_installed()){
  # Generate some example input sequences
  seq <- universalmotif::create_sequences()
  # sequences must have values in their fasta headers
  names(seq) <- seq_along(seq)
  # Create random example motif to search for
  motif <- universalmotif::create_motif()

  # Search for motif in sequences
  # parse_genomic_coord set to FALSE since fasta headers aren't in "chr:start-end" format.
  runFimo(seq, motif, parse_genomic_coord = FALSE)
}
```

runMeme

Identify motifs with MEME

Description

MEME performs *de-novo* discovery of ungapped motifs present in the input sequences. It can be used in both discriminative and non-discriminative modes.

Usage

```
runMeme(
  input,
  control = NA,
  outdir = "auto",
  alph = "dna",
  parse_genomic_coord = TRUE,
  combined_sites = FALSE,
  silent = TRUE,
```

```

    meme_path = NULL,
    ...
)

## S3 method for class 'list'
runMeme(
  input,
  control = NA,
  outdir = "auto",
  alph = "dna",
  parse_genomic_coord = TRUE,
  combined_sites = FALSE,
  silent = TRUE,
  meme_path = NULL,
  ...
)

## S3 method for class 'BStringSetList'
runMeme(
  input,
  control = NA,
  outdir = "auto",
  alph = "dna",
  parse_genomic_coord = TRUE,
  combined_sites = FALSE,
  silent = TRUE,
  meme_path = NULL,
  ...
)

## Default S3 method:
runMeme(
  input,
  control = NA,
  outdir = "auto",
  alph = "dna",
  parse_genomic_coord = TRUE,
  combined_sites = FALSE,
  silent = TRUE,
  meme_path = NULL,
  ...
)

```

Arguments

input	path to fasta, Biostrings::BStringSet list, or list of Biostrings::BStringSet (can generate using <code>get_sequence()</code>)
control	any data type as in input, or a character vector of names(input) to use those

	regions as control sequences. Using sequences as background requires an alternative objective function. Users must pass a non-default value of <code>objfun</code> to ... if using a non-NA control set (default: NA)
<code>outdir</code>	(default: "auto") Directory where output data will be stored.
<code>alph</code>	one of <code>c("dna", "rna", "protein")</code> or path to alphabet file (default: "dna").
<code>parse_genomic_coord</code>	<code>logical(1)</code> whether to parse genomic coordinates from fasta headers. Requires headers are in the form: "chr:start-end", or will result in an error. Automatically set to FALSE if <code>alph = "protein"</code> . This setting only needs to be changed if using a custom-built fasta file without genomic coordinates in the header.
<code>combined_sites</code>	<code>logical(1)</code> whether to return combined sites information (coerces output to list) (default: FALSE)
<code>silent</code>	Whether to suppress printing stdout to terminal (default: TRUE)
<code>meme_path</code>	path to "meme/bin/". If unset, will use default search behavior: <ol style="list-style-type: none"> 1. <code>meme_path</code> setting in <code>options()</code> 2. <code>MEME_PATH</code> setting in <code>.Renvirom</code> or <code>.bashrc</code>
<code>...</code>	additional arguments passed to MEME (see below)

Details

Note that MEME can take a long time to run. The more input sequences used, the wider the motifs searched for, and the more motifs MEME is asked to discover will drastically affect runtime. For this reason, MEME usually performs best on a few (<50) short (100-200 bp) sequences, although this is not a requirement. Additional details on how data size affects runtime can be found [here](#).

MEME works best when specifically tuned to the analysis question. The default settings are unlikely to be ideal. It has several complex arguments [documented here](#), which `runMeme()` accepts as R function arguments (see details below).

If discovering motifs within ChIP-seq, ATAC-seq, or similar peaks, MEME may perform best if using sequences flanking the summit (the site of maximum signal) of each peak rather than the center. ChIP-seq or similar data can also benefit from setting `revcomp = TRUE`, `minw = 5`, `maxw = 20`. For more tips on using MEME to analyze ChIP-seq data, see the following [tips page](#).

Additional arguments:

`runMeme()` accepts all valid arguments to `meme` as arguments passed to `...`. For flags without values, pass them as `flag = TRUE`. The `dna`, `rna`, and `protein` flags should instead be passed to the `alph` argument of `runMeme()`. The arguments passed to MEME often have many interactions with each other, for a detailed description of each argument see [MEME Commandline Documentation](#).

Value

MEME results in `universalmotif_df` format (see: `universalmotif::to_df()`). `sites_hits` is a nested data.frame column containing the position within each input sequence of matches to the identified motif.

Citation

If you use runMeme() in your analysis, please cite:

Timothy L. Bailey and Charles Elkan, "Fitting a mixture model by expectation maximization to discover motifs in biopolymers", Proceedings of the Second International Conference on Intelligent Systems for Molecular Biology, pp. 28-36, AAAI Press, Menlo Park, California, 1994. [pdf](#)

Licensing

The MEME Suite is free for non-profit use, but for-profit users should purchase a license. See the [MEME Suite Copyright Page](#) for details.

Examples

```
if (meme_is_installed()) {  
  seqs <- universalmotif::create_sequences("CCRAAAW", seqnum = 4)  
  names(seqs) <- 1:length(seqs)  
  runMeme(seqs, parse_genomic_coord = FALSE)  
}
```

runStreme

Denovo motif discovery of target regions using STREME

Description

STREME discovers short, ungapped, *de-novo* motifs that are enriched or relatively enriched relative to a control set of sequences. STREME can be run to discover motifs relative to a shuffled set of input sequences, against a separately provided set of "control" sequences, or to determine whether motifs are centrally enriched within input sequences.

Usage

```
runStreme(  
  input,  
  control,  
  outdir = "auto",  
  objfun = "de",  
  alph = "dna",  
  meme_path = NULL,  
  silent = TRUE,  
  ...  
)
```

Arguments

input	regions to scan for motifs. If using 'objfun = "cd"' to test for centrally enriched motifs, be sure to include sufficient flanking sequence (e.g. +/- 500bp) for an accurate estimate. Can be any of: - path to fasta file - DNASTringSet object (can be generated from GRanges using 'get_sequence()') - List of DNASTringSet objects (generated from 'get_sequence()') - *NOTE:* if using StringSet inputs, each entry must be named (set with 'names()'). - *NOTE:* If you want to retain the raw streme output files, you must use a path to fasta file as input, or specify an "outdir"
control	regions to use as background for motif search. These should have a similar length distribution as the input sequences. Can be any of: - path to fasta file - DNASTringSet object (can be generated from GRanges using get_sequence) - A Biostrings::BStringSetList (generated using 'get_sequence'), in which case all sequences in the list will be combined as the control set. - if 'input' is a list of DNASTringSet objects, a character vector of names in 'input' will use those sequences as background. runstreme will not scan those regions as input. - "shuffle" to use streme's built-in dinucleotide shuffle feature (NOTE: if 'input' is a list object with an entry named "shuffle", the list entry will be used instead). Optionally can also pass 'seed = <any number>' to '...' to use as the random seed during shuffling. If no seed is passed, streme will use 0 as the random seed, so results will be reproducible if rerunning.
outdir	path to output directory of streme files, or "auto" to autogenerate path. Default: location of input fasta in dir named "\<input>_vs_\<control>". If input is DNASTringset, will be temporary path. This means that if you want to save the raw output files, you must use fasta files as input or use an informative (and unique) outdir name. memes will **not check** if it overwrites files in a directory. Directories will be recursively created if needed. (default: "auto")
objfun	one of c("de", "cd"). Default: "de" for differential enrichment. "cd" for central distance (control must be set to NA for "cd").
alph	one of c("dna", "rna", "protein") or a path to a MEME format alph file. (default: "dna")
meme_path	path to "meme/bin"
silent	Whether to suppress printing stdout & stderr to console (default: TRUE). Warnings are always printed regardless of this setting.
...	pass any commandline options as R function arguments. For a complete list of STREME options, see [the STREME manual](https://meme-suite.org/meme/doc/streme.html).

Details

Properly setting the 'control' parameter is key to discovering biologically relevant motifs. Often, using 'control = "shuffle"' will produce a suboptimal set of motifs; however, some discriminative analysis designs don't have proper "control" regions other than to shuffle.

If you have fewer than 50 sequences, consider using [runMeme()] instead.

Citation

If you use 'runStreme()' in your analysis, please cite:

Timothy L. Bailey, "STREME: Accurate and versatile sequence motif discovery", Bioinformatics, 2021. <https://doi.org/10.1093/bioinformatics/btab203>

Licensing The MEME Suite is free for non-profit use, but for-profit users should purchase a license. See the [MEME Suite Copyright Page](<http://meme-suite.org/doc/copyright.html>) for details.

Value

a 'universalmotif_df' of STREME Motifs

See Also

'?universalmotif::tidy-motifs'

runTomTom	<i>Run TomTom on target motifs</i>
-----------	------------------------------------

Description

TomTom compares input motifs to a database of known, user-provided motifs to identify matches.

Usage

```
runTomTom(
  input,
  database = NULL,
  outdir = "auto",
  thresh = 10,
  min_overlap = 5,
  dist = "ed",
  evaluate = TRUE,
  silent = TRUE,
  meme_path = NULL,
  ...
)
```

Arguments

input	path to .meme format file of motifs, a list of universalmotifs, or a universalmotif data.frame object (such as the output of runDreme())
database	path to .meme format file to use as reference database (or list of universalmotifs). NOTE: p-value estimates are inaccurate when the database has fewer than 50 entries.
outdir	directory to store tomtom results (will be overwritten if exists). Default: location of input fasta file, or temporary location if using universalmotif input.

thresh	report matches less than or equal to this value. If <code>eval</code> = TRUE (default), set an e-value threshold (default = 10). If <code>eval</code> = FALSE, set a value between 0-1 (default = 0.5).
min_overlap	only report matches that overlap by this value or more, unless input motif is shorter, in which case the shorter length is used as the minimum value
dist	distance metric. Valid arguments: <code>allr</code> <code>ed</code> <code>kullback</code> <code>pearson</code> <code>sandelin</code> <code>blic1</code> <code>blic5</code> <code>llr1</code> <code>llr5</code> . Default: <code>ed</code> (euclidean distance).
eval	whether to use E-value as significance threshold (default: TRUE). If <code>eval</code> = FALSE, uses <i>q-value</i> instead.
silent	suppress printing stderr to console (default: TRUE).
meme_path	path to "meme/bin/" (optional). If unset, will check R environment variable "MEME_DB (set in .Renviron), or option "meme_db" (set with <code>option(meme_db = "path/to/meme/bin")</code>)
...	additional flags passed to tomtom using <code>cmdfun</code> formatting (see table below for details)

Details

runTomTom will rank matches by significance and return a best match motif for each input (whose properties are stored in the `best_match_*` columns) as well as a ranked list of all possible matches stored in the `tomtom list` column.

Additional arguments

runTomTom() can accept all valid tomtom arguments passed to ... as described in the [tomtom commandline reference](#). For convenience, below is a table of valid arguments, their default values, and their description.

TomTom Flag	allowed values	default	description
<code>bfile</code>	file path	NULL	path to background model for converting frequency matrix to log-odds score
<code>motif_pseudo</code>	numeric	0.1	pseudocount to add to motifs
<code>xalph</code>	logical	FALSE	convert alphabet of target database to alphabet of query database
<code>norc</code>	logical	FALSE	Do not score reverse complements of motifs
<code>incomplete_scores</code>	logical	FALSE	Compute scores using only aligned columns
<code>thresh</code>	numeric	0.5	only report matches with significance values \leq this value. Unless <code>eval</code> =
<code>internal</code>	logical	FALSE	forces the shorter motif to be completely contained in the longer motif
<code>min_overlap</code>	integer	1	only report matches that overlap by this number of positions or more. If quer
<code>time</code>	integer	NULL	Maximum runtime in CPU seconds (default: no limit)

Value

data.frame of match results. Contains `best_match_motif` column of `universalmotif` objects with the matched PWM from the database, a series of `best_match_*` columns describing the TomTom results of the match, and a `tomtom list` column storing the ranked list of possible matches to each motif. If a `universalmotif` data.frame is used as input, these columns are appended to the data.frame. If no matches are returned, `tomtom` and `best_match_motif` columns will be set to NA and a message indicating this will print.

Citation

If you use `runTomTom()` in your analysis, please cite:

Shobhit Gupta, JA Stamatoyannopolous, Timothy Bailey and William Stafford Noble, "Quantifying similarity between motifs", *Genome Biology*, 8(2):R24, 2007. [full text](#)

Licensing:

The MEME Suite is free for non-profit use, but for-profit users should purchase a license. See the [MEME Suite Copyright Page](#) for details.

Examples

```
if (meme_is_installed()) {  
  motif <- universalmotif::create_motif("CCRAAAW")  
  database <- system.file("extdata", "flyFactorSurvey_cleaned.meme", package = "memes")  
  
  runTomTom(motif, database)  
}
```

update_best_match	<i>Update best match info by ranking of tomtom data</i>
-------------------	---

Description

This function updates the `best_match` columns based on the rankings on the `tomtom` list data. By re-ordering the entries of a `'tomtom'` object, the `best_match` columns can be updated to reflect the new rankings using `[update_best_match()]`, where the first row of the `'tomtom'` `data.frame` is selected as the best match.

Usage

```
update_best_match(res)
```

Arguments

`res` results from `runTomTom`

Value

'res' with updated `best_*` columns

See Also

`[force_best_match()]`

Examples

```

data("example_dreme_tomtom")
# best match is "CG2052_SANGER_2.5"
example_dreme_tomtom$best_match_name[1]
# reorder the `tomtom` data.frame
example_dreme_tomtom$tomtom[[1]] <- dplyr::arrange(example_dreme_tomtom$tomtom[[1]],
                                                  dplyr::desc(match_eval))
# update_best_match will use the new order of rows, taking the top row as the new best match
new_res <- update_best_match(example_dreme_tomtom)
# best match is now altered:
new_res$best_match_name[1]

```

view_tomtom_hits	<i>Compare top tomtom hits to original motif</i>
------------------	--

Description

Although TomTom does a good job of matching unknown motifs to known motifs, sometimes the top hit is not the correct assignment. It can be useful to manually inspect the hits. This function provides a quick utility to compare matches.

Usage

```
view_tomtom_hits(results, top_n = "all")
```

Arguments

results	results data.frame from [runTomTom()]
top_n	number of matched motifs to return in plot (default: "all")

Details

This is intended to be a function used interactively and may not always be the best tool for creating publication-quality figures. Results with matches return ggseqlogo outputs which can be further manipulated using [ggplot2::theme()] calls, but results containing no matches are static plots.

Value

plot of input motif vs the top n number of tomtom matched motifs. If no match found, will plot "No Match". Note: the "No Match" plots are not amenable to ggplot theme() manipulations, while all others are.

Examples

```

results <- importTomTomXML(system.file("extdata", "tomtom.xml", package = "memes"))
# show top 3 hits
view_tomtom_hits(results, top_n = 3)

```

write_fasta	<i>Write fasta file from stringset</i>
-------------	--

Description

Write fasta file from stringset

Usage

```
write_fasta(seq, path = tempfile(fileext = ".fa"))
```

Arguments

seq	a 'Biostrings::XStringSet'
path	path of fasta file to write (default: temporary file)

Value

path to created fasta file

Examples

```
seq <- universalmotif::create_sequences()
write_fasta(seq)
```


Index

- * **datasets**
 - example_ame, 6
 - example_ame_large, 7
 - example_chip_summits, 7
 - example_dreme, 8
 - example_dreme_by_binding, 9
 - example_dreme_tomtom, 9
 - example_fimo, 9
 - example_peaks, 10
 - example_rnaseq, 10
 - example_tomtom, 11
- * **import**
 - importAme, 13
- * **internal**
 - memes-package, 3
- add_sequence, 3
- ame_compare_heatmap_methods, 4
- check_meme_install, 5
- check_meme_install(), 18
- drop_best_match, 6
- example_ame, 6
- example_ame_large, 7
- example_chip_summits, 7
- example_dreme, 8
- example_dreme_by_binding, 9
- example_dreme_tomtom, 9
- example_fimo, 9
- example_peaks, 10
- example_rnaseq, 10
- example_tomtom, 11
- force_best_match, 11
- get_sequence, 12
- has_duplicate_motifs, 13
- importAme, 13
- importDremeXML, 14
- importFimo, 15
- importMeme, 15
- importStremeXML, 16
- importTomTomXML, 17
- meme_is_installed, 18
- memes (memes-package), 3
- memes-package, 3
- nest_tomtom, 19
- plot_ame_heatmap, 19
- plot_sequence_heatmap, 21
- remove_duplicate_motifs, 22
- runAme (runAme.list), 23
- runAme.list, 23
- runDreme, 26
- runFimo, 29
- runMeme, 31
- runMeme(), 33
- runStreme, 34
- runStreme(), 27
- runTomTom, 36
- universalmotif::to_df(), 33
- update_best_match, 38
- view_tomtom_hits, 39
- write_fasta, 40