Package: jackalope (via r-universe)

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Type Package

Title A Swift, Versatile Phylogenomic and High-Throughput Sequencing Simulator

Version 1.1.5

Description Simply and efficiently simulates (i) variants from reference genomes and (ii) reads from both Illumina <https://www.illumina.com/> and Pacific Biosciences (PacBio) <https://www.pacb.com/> platforms. It can either read reference genomes from FASTA files or simulate new ones. Genomic variants can be simulated using summary statistics, phylogenies, Variant Call Format (VCF) files, and coalescent simulations-the latter of which can include selection, recombination, and demographic fluctuations. 'jackalope' can simulate single, paired-end, or mate-pair Illumina reads, as well as PacBio reads. These simulations include sequencing errors, mapping qualities, multiplexing, and optical/polymerase chain reaction (PCR) duplicates. Simulating Illumina sequencing is based on ART by Huang et al. (2012) <doi:10.1093/bioinformatics/btr708>. PacBio sequencing simulation is based on SimLoRD by Stöcker et al. (2016) <doi:10.1093/bioinformatics/btw286>. All outputs can be written to standard file formats.

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BugReports https://github.com/lucasnell/jackalope/issues

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Author Lucas A. Nell [cph, aut, cre] (<https://orcid.org/0000-0003-3209-0517>)

Maintainer Lucas A. Nell <lucnell@gmail.com>

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create_genome

Create a reference genome.

Description

Random chromosomes are generated to create a new ref_genome object. Note that this function will never generate empty chromosomes.

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create_haplotypes

Usage

```
create_genome(
  n_chroms,
  len_mean,
  len_sd = 0,
  pi_tcag = rep(0.25, 4),
  n_threads = 1
)
```

Arguments

n_chroms	Number of chromosomes.		
len_mean	Mean for the gamma distribution of chromosome sizes.		
len_sd	Standard deviation for the gamma distribution of chromosome sizes. If set to <= 0, all chromosomes will be the same length. Defaults to 0.		
pi_tcag	Vector of length 4 containing the nucleotide equilibrium frequencies for "T", "C", "A", and "G", respectively. Defaults to rep(0.25, 4).		
n_threads	Number of threads to use for parallel processing. This argument is ignored if OpenMP is not enabled. Defaults to 1.		

Value

A ref_genome object.

Examples

genome <- create_genome(10, 100e3, 100, pi_tcag = c(0.1, 0.2, 0.3, 0.4))</pre>

create_haplotypes Create haplotypes from a reference genome.

Description

Uses one of multiple methods to create variant haplotypes from a reference genome. See haps_functions for the methods available.

Usage

```
create_haplotypes(
  reference,
  haps_info,
  sub = NULL,
  ins = NULL,
  del = NULL,
  epsilon = 0.03,
```

```
n_threads = 1,
show_progress = FALSE
)
```

Arguments

reference	A ref_genome object from which to generate haplotypes. This argument is required.
haps_info	Output from one of the haps_functions. These functions organize higher-level information for use here. See haps_functions for brief descriptions and links to each method. If this argument is NULL, all arguments other than reference are ignored, and an empty haplotypes object with no haplotypes is returned. This is designed for use when you'd like to add mutations manually. If you create a blank haplotypes object, you can use its add_haps method to add haplotypes manually.
sub	Output from one of the sub_models functions that organizes information for the substitution models. See sub_models for more information on these models and their required parameters. This argument is ignored if you are using a VCF file to create haplotypes. Passing NULL to this argument results in no substitutions. Defaults to NULL.
ins	Output from the indels function that specifies rates of insertions by length. This argument is ignored if you are using a VCF file to create haplotypes. Passing NULL to this argument results in no insertions. Defaults to NULL.
del	Output from the indels function that specifies rates of deletions by length. This argument is ignored if you are using a VCF file to create haplotypes. Passing NULL to this argument results in no deletions. Defaults to NULL.
epsilon	Error control parameter for the "tau-leaping" approximation to the Doob–Gillespie algorithm, as used for the indel portion of the simulations. Smaller values result in a closer approximation. Larger values are less exact but faster. Values must be ≥ 0 and ≤ 1 . For more information on the approximation, see Cao et al. (2006) and Wieder et al. (2011), listed below. If epsilon is 0, then it reverts to the exact Doob–Gillespie algorithm. Defaults to 0.03 .
n_threads	Number of threads to use for parallel processing. This argument is ignored if OpenMP is not enabled. Threads are spread across chromosomes, so it doesn't make sense to supply more threads than chromosomes in the reference genome. Defaults to 1.
show_progress	Boolean for whether to show a progress bar during processing. Defaults to FALSE.

Value

A haplotypes object.

References

Cao, Y., D. T. Gillespie, and L. R. Petzold. 2006. Efficient step size selection for the tau-leaping simulation method. *The Journal of Chemical Physics* **124**(4): 044109.

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evo_rates

Doob, J. L. 1942. Topics in the theory of markoff chains. *Transactions of the American Mathematical Society* **52**(1): 37–64.

Gillespie, D. T. 1976. A general method for numerically simulating the stochastic time evolution of coupled chemical reactions. *Journal of Computational Physics* **22**(4): 403–434.

Wieder, N., R. H. Fink, and F. von Wegner. 2011. Exact and approximate stochastic simulation of intracellular calcium dynamics. *Journal of Biomedicine and Biotechnology* **2011**: 572492.

Examples

```
r <- create_genome(10, 1000)
v_phylo <- create_haplotypes(r, haps_phylo(ape::rcoal(5)), sub_JC69(0.1))
v_theta <- create_haplotypes(r, haps_theta(0.001, 5), sub_K80(0.1, 0.2))</pre>
```

evo_rates

Table of evolutionary rates.

Description

From Table 1 in Sung et al. (2016).

Usage

evo_rates

Format

A data frame with 15 rows and 8 variables:

domain Either Bacteria or Eukarya for what type of organism the species is.

species Species name.

- Ge Effective genome size using only coding DNA.
- Gc_Gnc Effective genome size using coding DNA and non-coding DNA that is under purifying selection.
- indels Rate of insertions and deletions $(10^{-10} \text{ events per site per generation})$.

subs Base-substitution mutation rate $(10^{-10} \text{ events per site per generation})$.

Ne Effective population size ($\times 10^6$).

theta_s Population mutation rate estimated using θ_s .

pi_s Population mutation rate estimated using π_s .

Source

doi:10.1534/g3.116.030890

References

Sung, W., M. S. Ackerman, M. M. Dillon, T. G. Platt, C. Fuqua, V. S. Cooper, and M. Lynch. 2016. Evolution of the insertion-deletion mutation rate across the tree of life. *G3: Genes* | *Genomes* | *Genetics* 6:2583–2591.

haplotypes

An R6 Class Representing Haploid Variants

Description

Interactive wrapper for a pointer to a C++ object that stores information about variant haplotypes from a single reference genome.

Details

This class should NEVER be created using haplotypes\$new. Only use create_haplotypes. Because this class wraps a pointer to a C++ object, there are no fields to manipulate directly. All manipulations are done through this class's methods.

Connections to ref_genome objects

Regarding the ref_genome object you use to create a haplotypes object, you should note the following:

- This point is the most important. Both the ref_genome and haplotypes objects use the same underlying C++ object to store reference genome information. Thus, if you make any changes to the ref_genome object, those changes will also show up in the haplotypes object. For example, if you make a haplotypes object named V based on an existing ref_genome object named R, then you merge chromosomes in R, V will now have merged chromosomes. If you've already started adding mutations to V, then all the indexes used to store those mutations will be inaccurate. So when you do anything with V later, your R session will crash or have errors. The lesson here is that you shouldn't edit the reference genome after using it to create haplotypes.
- If a ref_genome object is used to create a haplotypes object, deleting the ref_genome object won't cause issues with the haplotypes object. However, the haplotypes class doesn't provide methods to edit chromosomes, so only remove the ref_genome object when you're done editing the reference genome.

Methods

Public methods:

- haplotypes\$new()
- haplotypes\$print()
- haplotypes\$ptr()
- haplotypes\$n_chroms()
- haplotypes\$n_haps()

- haplotypes\$sizes()
- haplotypes\$chrom_names()
- haplotypes\$hap_names()
- haplotypes\$chrom()
- haplotypes\$gc_prop()
- haplotypes\$nt_prop()
- haplotypes\$set_names()
- haplotypes\$add_haps()
- haplotypes\$dup_haps()
- haplotypes\$rm_haps()
- haplotypes\$add_sub()
- haplotypes\$add_ins()
- haplotypes\$add_del()

Method new(): Do NOT use this; only use create_haplotypes to make new haplotypes.

Usage:

haplotypes\$new(genomes_ptr, reference_ptr)

Arguments:

- genomes_ptr An externalptr object pointing to a C++ object that stores the information about the haplotypes.
- reference_ptr An externalptr object pointing to a C++ object that stores the information about the reference genome.

Method print(): Print a haplotypes object.

Usage:

```
haplotypes$print()
```

Method ptr(): View pointer to underlying C++ object (this is not useful to end users).

Usage:

haplotypes\$ptr()

Returns: An externalptr object.

Method n_chroms(): View number of chromosomes.

Usage:

haplotypes\$n_chroms()

Returns: Integer number of chromosomes.

Method n_haps(): View number of haplotypes.

Usage:

haplotypes\$n_haps()

Returns: Integer number of haplotypes.

Method sizes(): View chromosome sizes for one haplotype.

Usage: haplotypes\$sizes(hap_ind) Arguments:

hap_ind Index for the focal haplotype.

Returns: Integer vector of chromosome sizes for focal haplotype.

Method chrom_names(): View chromosome names.

Usage:

haplotypes\$chrom_names()

Returns: Character vector of chromosome names.

Method hap_names(): View haplotype names.

Usage:

haplotypes\$hap_names()

Returns: Character vector of haplotype names.

Method chrom(): View one haplotype chromosome.

Usage:

haplotypes\$chrom(hap_ind, chrom_ind)

Arguments:

hap_ind Index for the focal haplotype.

chrom_ind Index for the focal chromosome.

Returns: A single string representing the chosen haplotype chromosome's DNA sequence.

Method gc_prop(): View GC proportion for part of one haplotype chromosome.

Usage:

haplotypes\$gc_prop(hap_ind, chrom_ind, start, end)

Arguments:

hap_ind Index for the focal haplotype.

chrom_ind Index for the focal chromosome.

start Point on the chromosome at which to start the calculation (inclusive).

end Point on the chromosome at which to end the calculation (inclusive).

Returns: A double in the range [0,1] representing the proportion of DNA sequence that is either G or C.

Method nt_prop(): View nucleotide content for part of one haplotype chromosome

Usage:

haplotypes\$nt_prop(nt, hap_ind, chrom_ind, start, end)

Arguments:

nt Which nucleotide to calculate the proportion that the DNA sequence is made of. Must be one of T, C, A, G, or N.

hap_ind Index for the focal haplotype.

chrom_ind Index for the focal chromosome.

start Point on the chromosome at which to start the calculation (inclusive).

end Point on the chromosome at which to end the calculation (inclusive).

Returns: A double in the range [0,1] representing the proportion of DNA sequence that is nt.

Method set_names(): Change haplotype names.

Usage:

haplotypes\$set_names(new_names)

Arguments:

new_names Vector of new names to use. This must be the same length as the number of current names.

Returns: This R6 object, invisibly.

Method add_haps(): Add one or more blank, named haplotypes

Usage: haplotypes\$add_haps(new_names)

Arguments:

new_names Vector of name(s) for the new haplotype(s).

Returns: This R6 object, invisibly.

Method dup_haps(): Duplicate one or more haplotypes by name.

Usage:

haplotypes\$dup_haps(hap_names, new_names = NULL)

Arguments:

hap_names Vector of existing haplotype name(s) that you want to duplicate.

new_names Optional vector specifying the names of the duplicates. If NULL, their names are auto-generated. Defaults to NULL.

Returns: This R6 object, invisibly.

Method rm_haps(): Remove one or more haplotypes by name.

Usage:

haplotypes\$rm_haps(hap_names)

Arguments:

hap_names Vector of existing haplotype name(s) that you want to remove.

Returns: This R6 object, invisibly.

Method add_sub(): Manually add a substitution.

Usage:

haplotypes\$add_sub(hap_ind, chrom_ind, pos, nt)

Arguments:

hap_ind Index for the focal haplotype.

chrom_ind Index for the focal chromosome.

pos Position at which to add the mutation.

nt Single character representing the nucleotide to change the current one to.

Returns: This R6 object, invisibly.

Method add_ins(): Manually add an insertion.

Usage:

haplotypes\$add_ins(hap_ind, chrom_ind, pos, nts)

Arguments:

hap_ind Index for the focal haplotype.

chrom_ind Index for the focal chromosome.

pos Position at which to add the mutation.

nts String representing the nucleotide(s) that will be inserted after the designated position.

Returns: This R6 object, invisibly.

\item{`add_del(hap_ind, chrom_ind, pos, n_nts)`}{Manually add a deletion
 for a given haplotype (`hap_ind`), chromosome (`chrom_ind`), and position (`pos`).
 The designated number of nucleotides to delete (`n_nts`) will be deleted
 starting at `pos`, unless `pos` is near the chromosome end and doesn't have
 `n_nts` nucleotides to remove; it simply stops at the chromosome end in
 this case.}

Method add_del(): Manually add a deletion.

Usage:

haplotypes\$add_del(hap_ind, chrom_ind, pos, n_nts)

Arguments:

hap_ind Index for the focal haplotype.

chrom_ind Index for the focal chromosome.

pos Position at which to add the mutation.

n_nts Single integer specifying the number of nucleotides to delete. These will be deleted starting at pos. If pos is near the chromosome end and doesn't have n_nts nucleotides to remove, it simply removes nucleotides from pos to the chromosome end.

Returns: This R6 object, invisibly.

See Also

create_haplotypes

Description

The following functions organize information that gets passed to create_haplotypes to generate haplotypes from a reference genome. Each function represents a method of generation and starts with "haps_". The first three are phylogenomic methods, and all functions but haps_vcf will use molecular evolution information when passed to create_haplotypes.

Details

- haps_theta Uses an estimate for theta, the population-scaled mutation rate, and a desired number of haplotypes.
- haps_phylo Uses phylogenetic tree(s) from phylo object(s) or NEWICK file(s), one tree per chromosome or one for all chromosomes.
- haps_gtrees Uses gene trees, either in the form of an object from the scrm or coala package or a file containing output in the style of the ms program.
- haps_ssites Uses matrices of segregating sites, either in the form of scrm or coala coalescentsimulator object(s), or a ms-style output file.
- haps_vcf Uses a haplotype call format (VCF) file that directly specifies haplotypes.

See Also

create_haplotypes

haps_gtrees

Organize information to create haplotypes using gene trees

Description

This function organizes higher-level information for creating haplotypes from gene trees output from coalescent simulations. Note that all gene trees must be rooted and binary.

Usage

```
haps_gtrees(obj = NULL, fn = NULL)
write_gtrees(gtrees, out_prefix)
```

Arguments

obj	Object containing gene trees. This can be one of the following: (1) A single list with a trees field inside. This field must contain a set of gene trees for each chromosome. (2) A list of lists, each sub-list containing a trees field of length 1. The top-level list must be of the same length as the number of chromosomes. Defaults to NULL.
fn	A single string specifying the name of the file containing the ms-style coalescent output with gene trees. Defaults to NULL.
gtrees	A haps_gtrees_info object output from haps_gtrees.
out_prefix	Prefix for the output file of gene trees. The extension will be .trees.

Details

Using the obj argument is designed after the trees fields in the output from the scrm and coala packages. (These packages are not required to be installed when installing jackalope.) To get gene trees, make sure to add + sumstat_trees() to the coalmodel for coala, or make sure that "-T" is present in args for scrm. If using either of these packages, I encourage you to cite them. For citation information, see output from citation("scrm") or citation("coala").

If using an output file from a command-line program like ms/msms, add the -T option.

Value

A haps_gtrees_info object containing information used in create_haplotypes to create variant haplotypes. This class is just a wrapper around a list of NEWICK tree strings, one for each gene tree, which you can view (but not change) using the object's trees() method.

Functions

• write_gtrees(): Write gene trees to ms-style output file.

haps_phylo

Organize information to create haplotypes using phylogenetic tree(s)

Description

This function organizes higher-level information for creating haplotypes from phylogenetic tree(s) output as phylo or multiPhylo objects (both from the ape package) or NEWICK files. Note that all phylogenetic trees must be rooted and binary. If using this function, I encourage you to cite ape. For citation information, see output from citation("ape").

Usage

haps_phylo(obj = NULL, fn = NULL)

haps_ssites

Arguments

obj	Object containing phylogenetic tree(s). This can be (1) a single phylo object that represents all chromosomes in the genome or (2) a list or multiPhylo object containing a phylo object for each reference chromosome. In the latter case, phylogenies will be assigned to chromosomes in the order provided. Defaults to NULL.
fn	One or more string(s), each of which specifies the file name of a NEWICK file containing a phylogeny. If one name is provided, that phylogeny will be used for all chromosomes. If more than one is provided, there must be a phylogeny for each reference genome chromosome, and phylogenies will be assigned to chromosomes in the order provided. Defaults to NULL.

Details

See ?ape::write.tree for writing phylogenies to an output file.

Value

A haps_phylo_info object containing information used in create_haplotypes to create variant haplotypes. This class is just a wrapper around a list containing phylogenetic tree information for each reference chromosome, which you can view (but not change) using the object's phylo() method.

haps_ssites	Organize information to create haplotypes using segregating sites ma-
	trices

Description

This function organizes higher-level information for creating haplotypes from matrices of segregating sites output from coalescent simulations.

Usage

haps_ssites(obj = NULL, fn = NULL)

Arguments

```
obj
```

Object containing segregating sites information. This can be one of the following: (1) A single list with a seg_sites field inside. This field must contain a matrix for segregating sites for each chromosome. The matrix itself should contain the haplotype information, coded using 0s and 1s: 0s indicate the ancestral state and 1s indicate mutant. The matrix column names should indicate the positions of the polymorphisms on the chromosome. If positions are in the range (0, 1), they're assumed to come from an infinite- sites model and are relative positions. If positions are integers in the range $[0, chromosome \ length - 1]$ or $[1, chromosome \ length]$, they're assumed to come from an finite-sites model and are absolute positions. Defaults to NULL. fn A single string specifying the name of the file containing the ms-style coalescent output with segregating site info. Defaults to NULL.

Details

For what the seg_sites field should look like in a list, see output from the scrm or coala package. (These packages are not required to be installed when installing jackalope.) If using either of these packages, I encourage you to cite them. For citation information, see output from citation("scrm") or citation("coala").

Value

A haps_ssites_info object containing information used in create_haplotypes to create variant haplotypes. This class is just a wrapper around a list of matrices of segregating site info, which you can view (but not change) using the object's mats() method.

haps_theta	Organize information to create haplotypes using theta parameter
------------	---

Description

This function organizes higher-level information for creating haplotypes from the population-scaled mutation rate and a desired number of haplotypes.

Usage

```
haps_theta(theta, n_haps)
```

Arguments

theta	Population-scaled mutation rate.		
n_haps	Number of desired haplotypes.		

Value

A haps_theta_info object containing information used in create_haplotypes to create variant haplotypes. This class is just a wrapper around a list containing the phylogenetic tree and theta parameter, which you can view (but not change) using the object's phylo() and theta() methods, respectively.

haps_vcf

Description

This function organizes higher-level information for creating haplotypes from Variant Call Format (VCF) files.

Usage

haps_vcf(fn, print_names = FALSE)

Arguments

fn	A single string specifying the name of the VCF file
print_names	Logical for whether to print all unique chromosome names from the VCF file when VCF chromosome names don't match those from the reference genome. This printing doesn't happen until this object is passed to create_haplotypes. This can be useful for troubleshooting. Defaults to FALSE.

Value

A haps_vcf_info object containing information used in create_haplotypes to create variant haplotypes. This class is just a wrapper around a list containing the arguments to this function, which you can view (but not change) using the object's fn() and print_names() methods.

illumina

Create and write Illumina reads to FASTQ file(s).

Description

From either a reference genome or set of variant haplotypes, create Illumina reads from error profiles and write them to FASTQ output file(s). I encourage you to cite the reference below in addition to jackalope if you use this function.

Usage

illumina

```
seq_sys = NULL,
profile1 = NULL,
profile2 = NULL,
ins_prob1 = 0.00009,
del_prob1 = 0.00011,
ins_prob2 = 0.00015,
del_prob2 = 0.00023,
frag_len_min = NULL,
frag_len_max = NULL,
haplotype_probs = NULL,
barcodes = NULL,
prob_dup = 0.02,
sep_files = FALSE,
compress = FALSE,
comp_method = "bgzip",
n_{threads} = 1L,
read_pool_size = 1000L,
show_progress = FALSE,
overwrite = FALSE)
```

Arguments

obj	Sequencing object of class ref_genome or haplotypes.
out_prefix	Prefix for the output file(s), including entire path except for the file extension.
n_reads	Number of reads you want to create.
read_length	Length of reads.
paired	Logical for whether to use paired-end reads. This argument is changed to \ensuremath{TRUE} if matepair is \ensuremath{TRUE} .
frag_mean	Mean of the Gamma distribution that generates fragment sizes. Defaults to 400.
frag_sd	Standard deviation of the Gamma distribution that generates fragment sizes. Defaults to 100.
matepair	Logical for whether to simulate mate-pair reads. Defaults to FALSE.
seq_sys	Full or abbreviated name of sequencing system to use. See "Sequencing systems" section for options. See "Sequencing profiles" section for more information on how this argument, profile1, and profile2 are used to specify profiles. Defaults to NULL.
profile1	Custom profile for read 1. See "Sequencing profiles" section for more in- formation on how this argument, profile2, and seq_sys are used to specify profiles. Defaults to NULL.
profile2	Custom profile file for read 2. See "Sequencing profiles" section for more in- formation on how this argument, profile1, and seq_sys are used to specify profiles. Defaults to NULL.
ins_prob1	Insertion probability for read 1. Defaults to 0.00009.
del_prob1	Deletion probability for read 1. Defaults to 0.00011.
ins_prob2	Insertion probability for read 2. Defaults to 0.00015.

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illumina

del_prob2	Deletion probability for read 2. Defaults to 0.00023.
frag_len_min	Minimum fragment size. A NULL value results in the read length. Defaults to NULL.
frag_len_max	Maximum fragment size. A NULL value results in 2^32-1 , the maximum allowed value. Defaults to NULL
haplotype_probs	
	Relative probability of sampling each haplotype. This is ignored if sequencing a reference genome. NULL results in all having the same probability. Defaults to NULL.
barcodes	Character vector of barcodes for each haplotype, or a single barcode if sequenc- ing a reference genome. NULL results in no barcodes. Defaults to NULL.
prob_dup	A single number indicating the probability of duplicates. Defaults to 0.02.
sep_files	Logical indicating whether to make separate files for each haplotype. This ar- gument is coerced to FALSE if the obj argument is not a haplotypes object. Defaults to FALSE.
compress	Logical specifying whether or not to compress output file, or an integer specifying the level of compression, from 1 to 9. If TRUE, a compression level of 6 is used. Defaults to FALSE.
comp_method	Character specifying which type of compression to use if any is desired. Options include "gzip" and "bgzip". This is ignored if compress is FALSE, and it throws an error if it's set to "gzip" when n_threads > 1 (since I don't have a method to do gzip compression in parallel). Defaults to "bgzip".
n_threads	The number of threads to use in processing. If compress is TRUE or > \emptyset (indicating compressed output), setting n_threads to 2 or more makes this function first create an uncompressed file/files using n_threads threads, then compress that/those file/files also using n_threads threads. There is no speed increase if you try to use multiple threads to create compressed output on the fly, so that option is not included. If you want to be conservative with disk space (by not having an uncompressed file present even temporarily), set n_threads to 1. Threads are NOT spread across chromosomes or haplotypes, so you don't need to think about these when choosing this argument's value. However, all threads write to the same file/files, so there are diminishing returns for providing many threads. This argument is ignored if the package was not compiled with OpenMP. Defaults to 1.
<pre>read_pool_size</pre>	The number of reads to store before writing to disk. Increasing this number should improve speed but take up more memory. Defaults to 1000.
show_progress	Logical for whether to show a progress bar. Defaults to FALSE.
overwrite	Logical for whether to overwrite existing FASTQ file(s) of the same name, if they exist.

Value

Nothing is returned.

Sequencing profiles

This section outlines how to use the seq_sys, profile1, and profile2 arguments. If all arguments are NULL (their defaults), a sequencing system is chosen based on the read length. If, however, one or more arguments has been provided, then how they're provided should depend on whether you want single- or paired-end reads.

For single-end reads

- profile2 should be NULL.
- Only seq_sys or profile1 should be provided, not both.

For paired-end reads

- If providing seq_sys, don't provide either profile1 or profile2.
- If providing profile1, you must also provide profile2 (they can be the same if you want) and you cannot provide seq_sys.

Sequencing systems

Sequencing system options are the following, where, for each system, "name" is the full name, "abbrev" is the abbreviated name, "max_len" indicates the maximum allowed read length, and "paired" indicates whether paired-end sequencing is allowed.

name	abbrev	max_len	paired
Genome Analyzer I	GA1	44	Yes
Genome Analyzer II	GA2	75	Yes
HiSeq 1000	HS10	100	Yes
HiSeq 2000	HS20	100	Yes
HiSeq 2500	HS25	150	Yes
HiSeqX v2.5 PCR free	HSXn	150	Yes
HiSeqX v2.5 TruSeq	HSXt	150	Yes
MiniSeq TruSeq	MinS	50	No
MiSeq v1	MSv1	250	Yes
MiSeq v3	MSv3	250	Yes
NextSeq 500 v2	NS50	75	Yes

ID lines

The ID lines for FASTQ files are formatted as such:

@<genome name>-<chromosome name>-<starting position>-<strand>[/<read#>]

where the part in [] is only for paired-end Illumina reads, and where genome name is always REF for reference genomes (as opposed to haplotypes).

References

Huang, W., L. Li, J. R. Myers, and G. T. Marth. 2012. ART: a next-generation sequencing read simulator. *Bioinformatics* 28:593–594.

indels

Examples

indels

Insertions and deletions (indels) specification

Description

Construct necessary information for insertions and deletions (indels) that will be used in create_haplotypes.

Usage

```
indels(rate, max_length = 10, a = NULL, rel_rates = NULL)
```

Arguments

rate	Single number specifying the overall indel rate among all lengths.
max_length	Maximum length of indels. Defaults to 10.
a	Extra parameter necessary for generating rates from a Lavalette distribution. See Details for more info. Defaults to NULL.
rel_rates	A numeric vector of relative rates for each indel length from 1 to the maximum length. If provided, all arguments other than rate are ignored. Defaults to NULL.

Details

All indels require the rate parameter, which specifies the overall indels rate among all lengths. The rate parameter is ultimately combined with a vector of relative rates among the different lengths of indels from 1 to the maximum possible length. There are three different ways to specify/generate relative-rate values.

- 1. Assume that rates are proportional to exp(-L) for indel length L from 1 to the maximum length (Albers et al. 2011). This method will be used if the following arguments are provided:
 - rate
 - max_length
- Generate relative rates from a Lavalette distribution (Fletcher and Yang 2009), where the rate for length L is proportional to {L * max_length / (max_length L + 1)}^(-a). This method will be used if the following arguments are provided:
 - rate
 - max_length
 - a

- 3. Directly specify values by providing a numeric vector of relative rates for each insertion/deletion length from 1 to the maximum length. This method will be used if the following arguments are provided:
 - rate
 - rel_rates

Value

An indel_info object, which is an R6 class that wraps the info needed for the create_haplotypes function. It does not allow the user to directly manipulate the info inside, as that should be done using this function. You can use the rates() method to view the indel rates by size.

References

Albers, C. A., G. Lunter, D. G. MacArthur, G. McVean, W. H. Ouwehand, and R. Durbin. 2011. Dindel: accurate indel calls from short-read data. Genome Research 21:961–973.

Fletcher, W., and Z. Yang. 2009. INDELible: a flexible simulator of biological sequence evolution. Molecular Biology and Evolution 26:1879–1888.

Examples

```
# relative rates are proportional to `exp(-L)` for indel
# length `L` from 1 to 5:
indel_rates1 <- indels(0.1, max_length = 5)
# relative rates are proportional to Lavalette distribution
# for length from 1 to 10:
indel_rates2 <- indels(0.2, max_length = 10, a = 1.1)
# relative rates are all the same for lengths from 1 to 100:
```

indel_rates3 <- indels(0.2, rel_rates = rep(1, 100))</pre>

pacbio

Create and write PacBio reads to FASTQ file(s).

Description

From either a reference genome or set of variant haplotypes, create PacBio reads and write them to FASTQ output file(s). I encourage you to cite the reference below in addition to jackalope if you use this function.

pacbio

Usage

```
pacbio(obj,
       out_prefix,
       n_reads,
       chi2_params_s = c(0.01214, -5.12, 675, 48303.0732881,
                         1.4691051212330266),
       chi2_params_n = c(0.00189237136, 2.53944970, 5500),
       max_passes = 40,
       sqrt_params = c(0.5, 0.2247),
       norm_params = c(0, 0.2),
       prob_thresh = 0.2,
       ins_prob = 0.11,
       del_prob = 0.04,
       sub_prob = 0.01,
       min_read_length = 50,
       lognorm_read_length = c(0.200110276521, -10075.4363813,
                               17922.611306),
       custom_read_lengths = NULL,
       prob_dup = 0.0,
       haplotype_probs = NULL,
       sep_files = FALSE,
       compress = FALSE,
       comp_method = "bgzip",
       n_{threads} = 1L,
       read_pool_size = 100L,
       show_progress = FALSE,
       overwrite = FALSE)
```

Arguments

obj	Sequencing object of class ref_genome or haplotypes.
out_prefix	Prefix for the output file(s), including entire path except for the file extension.
n_reads	Number of reads you want to create.
chi2_params_s	Vector containing the 5 parameters for the curve determining the scale parameter for the chi^2 distribution. Defaults to c(0.01214, -5.12, 675, 48303.0732881, 1.4691051212330266).
chi2_params_n	Vector containing the 3 parameters for the function determining the n parameter for the chi^2 distribution. Defaults to $c(0.00189237136, 2.53944970, 5500)$.
max_passes	Maximal number of passes for one molecule. Defaults to 40.
sqrt_params	Vector containing the 2 parameters for the square root function for the quality increase. Defaults to $c(0.5, 0.2247)$.
norm_params	Vector containing the 2 parameters for normal distributed noise added to quality increase square root function Defaults to $c(0, 0.2)$.
prob_thresh	Upper bound for the modified total error probability. Defaults to 0.2.
ins_prob	Probability for insertions for reads with one pass. Defaults to 0.11.

del_prob	Probability for deletions for reads with one pass. Defaults to 0.04.
sub_prob	Probability for substitutions for reads with one pass. Defaults to 0.01.
<pre>min_read_length</pre>	
	Minium read length for lognormal distribution. Defaults to 50.
lognorm_read_le	Vector containing the 3 parameters for lognormal read length distribution. De- faults to c(0.200110276521, -10075.4363813, 17922.611306).
custom_read_len	gths
	Sample read lengths from a vector or column in a matrix; if a matrix, the second column specifies the sampling weights. If NULL, it samples read lengths from the lognormal distribution using parameters in lognorm_read_length. Defaults to NULL.
prob_dup	A single number indicating the probability of duplicates. Defaults to 0.0.
haplotype_probs	
	Relative probability of sampling each haplotype. This is ignored if sequencing a reference genome. NULL results in all having the same probability. Defaults to NULL.
<pre>sep_files</pre>	Logical indicating whether to make separate files for each haplotype. This argument is coerced to FALSE if the obj argument is not a haplotypes object. Defaults to FALSE.
compress	Logical specifying whether or not to compress output file, or an integer specifying the level of compression, from 1 to 9. If TRUE, a compression level of 6 is used. Defaults to FALSE.
comp_method	Character specifying which type of compression to use if any is desired. Options include "gzip" and "bgzip". This is ignored if compress is FALSE, and it throws an error if it's set to "gzip" when n_threads > 1 (since I don't have a method to do gzip compression in parallel). Defaults to "bgzip".
n_threads	The number of threads to use in processing. If compress is TRUE or > 0 (indicating compressed output), setting n_threads to 2 or more makes this function first create an uncompressed file/files using n_threads threads, then compress that/those file/files also using n_threads threads. There is no speed increase if you try to use multiple threads to create compressed output on the fly, so that option is not included. If you want to be conservative with disk space (by not having an uncompressed file present even temporarily), set n_threads to 1. Threads are NOT spread across chromosomes or haplotypes, so you don't need to think about these when choosing this argument's value. However, all threads write to the same file/files, so there are diminishing returns for providing many threads. This argument is ignored if the package was not compiled with OpenMP. Defaults to 1.
<pre>read_pool_size</pre>	The number of reads to store before writing to disk. Increasing this number should improve speed but take up more memory. Defaults to 100.
show_progress	Logical for whether to show a progress bar. Defaults to FALSE.
overwrite	Logical for whether to overwrite existing FASTQ file(s) of the same name, if they exist.

read_fasta

Value

Nothing is returned.

ID lines

The ID lines for FASTQ files are formatted as such: @<genome name>-<chromosome name>-<starting position>-<strand>

where genome name is always REF for reference genomes (as opposed to haplotypes).

References

Stöcker, B. K., J. Köster, and S. Rahmann. 2016. SimLoRD: simulation of long read data. *Bioinformatics* **32**:2704–2706.

Examples

```
rg <- create_genome(10, 100e3, 100)
dir <- tempdir(TRUE)
pacbio(rg, paste0(dir, "/pacbio_reads"), n_reads = 100)
```

read_fasta Read a fasta file.

Description

Accepts uncompressed and gzipped fasta files.

Usage

```
read_fasta(fasta_files, fai_files = NULL, cut_names = FALSE)
```

Arguments

fasta_files	File name(s) of the fasta file(s).
fai_files	File name(s) of the fasta index file(s). Providing this argument speeds up the reading process significantly. If this argument is provided, it must be the same length as the fasta_files argument. Defaults to NULL, which indicates the fasta file(s) is/are not indexed.
cut_names	Boolean for whether to cut chromosome names at the first space. This argument is ignored if fai_file is not NULL. Defaults to FALSE.

Value

A ref_genome object.

ref_genome

Description

Interactive wrapper for a pointer to a C++ object that stores reference genome information.

Details

This class should NEVER be created using ref_genome\$new. Only use read_fasta or create_genome. Because this class wraps a pointer to a C++ object, there are no fields to manipulate directly. All manipulations are done through this class's methods.

Methods

Public methods:

- ref_genome\$new()
- ref_genome\$print()
- ref_genome\$ptr()
- ref_genome\$n_chroms()
- ref_genome\$sizes()
- ref_genome\$chrom_names()
- ref_genome\$chrom()
- ref_genome\$gc_prop()
- ref_genome\$nt_prop()
- ref_genome\$set_names()
- ref_genome\$clean_names()
- ref_genome\$add_chroms()
- ref_genome\$rm_chroms()
- ref_genome\$merge_chroms()
- ref_genome\$filter_chroms()
- ref_genome\$replace_Ns()

Method new(): Do NOT use this; only use read_fasta or create_genome to make a new ref_genome.

Usage:

ref_genome\$new(genome_ptr)

Arguments:

genome_ptr An externalptr object pointing to a C++ object that stores the information about the reference genome.

Method print(): Print a ref_genome object.

Usage:

ref_genome\$print()

Method ptr(): View pointer to underlying C++ object (this is not useful to end users).

Usage:

ref_genome\$ptr()

Returns: An externalptr object.

Method n_chroms(): View number of chromosomes.

Usage: ref_genome\$n_chroms()

Returns: Integer number of chromosomes.

Method sizes(): View chromosome sizes.

Usage: ref_genome\$sizes() Returns: Integer vector of chromosome sizes.

Method chrom_names(): View chromosome names.

Usage:
ref_genome\$chrom_names()

Returns: Character vector of chromosome names.

Method chrom(): View one reference chromosome.

Usage: ref_genome\$chrom(chrom_ind) Arguments:

chrom_ind Index for the focal chromosome.

Returns: A single string representing the chosen chromosome's DNA sequence.

Method gc_prop(): View GC proportion for part of one reference chromosome.

Usage: ref_genome\$gc_prop(chrom_ind, start, end)

Arguments:

chrom_ind Index for the focal chromosome.

start Point on the chromosome at which to start the calculation (inclusive).

end Point on the chromosome at which to end the calculation (inclusive).

Returns: A double in the range [0,1] representing the proportion of DNA sequence that is either G or C.

Method nt_prop(): View nucleotide content for part of one reference chromosome

Usage:

ref_genome\$nt_prop(nt, chrom_ind, start, end)

Arguments:

nt Which nucleotide to calculate the proportion that the DNA sequence is made of. Must be one of T, C, A, G, or N.

chrom_ind Index for the focal chromosome.

start Point on the chromosome at which to start the calculation (inclusive).

end Point on the chromosome at which to end the calculation (inclusive).

Returns: A double in the range [0,1] representing the proportion of DNA sequence that is nt.

Method set_names(): Change chromosome names.

Usage: ref_genome\$set_names(new_names)

Arguments:

new_names Vector of new names to use. This must be the same length as the number of current names.

Returns: This R6 object, invisibly.

Examples:

ref <- create_genome(4, 10)
ref\$set_names(c("a", "b", "c", "d"))</pre>

Method clean_names(): Clean chromosome names, converting ":;=%,\\|/\"\'" to "_".

Usage: ref_genome\$clean_names() Returns: This R6 object, invisibly. Examples: ref <- create_genome(4, 10) ref\$set_names(c("a:", "b|", "c;", "d'"))

```
ref$clean_names()
```

Method add_chroms(): Add one or more chromosomes.

Usage: ref_genome\$add_chroms(new_chroms, new_names = NULL)

Arguments:

new_chroms Character vector of DNA strings representing new chromosomes.

new_names Optional character vector of names for the new chromosomes. It should be the same length as new_chroms. If NULL, new names will be automatically generated. Defaults to NULL.

Returns: This R6 object, invisibly.

Examples:

ref <- create_genome(4, 10)
ref\$add_chroms("TCAGTCAG")</pre>

Method rm_chroms(): Remove one or more chromosomes by name

ref_genome

Usage:

ref_genome\$rm_chroms(chrom_names)

Arguments:

chrom_names Vector of the name(s) of the chromosome(s) to remove.

Returns: This R6 object, invisibly.

Examples:

```
ref <- create_genome(4, 10)
ref$set_names(c("a", "b", "c", "d"))
ref$rm_chroms("b")</pre>
```

Method merge_chroms(): Merge chromosomes into one.

Usage:

ref_genome\$merge_chroms(chrom_names)

Arguments:

chrom_names Vector of the names of the chromosomes to merge into one. Duplicates are not allowed, and chromosomes are merged in the order they're provided. If this is NULL, then all chromosomes are merged after first shuffling their order.

Returns: This R6 object, invisibly.

Examples:

```
ref <- create_genome(4, 10)
ref$merge_chroms(ref$chrom_names()[1:2])
ref$merge_chroms(NULL)</pre>
```

Method filter_chroms(): Filter chromosomes by size or for a proportion of total bases.

Usage:

ref_genome\$filter_chroms(threshold, method)

Arguments:

- threshold Number used as a threshold. If method == "size", then this is the minimum length
 of a chromosome that will remain after filtering. If method == "prop", chromosomes are
 first size-sorted, then the largest N chromosomes are retained that allow at least threshold * sum(<all chromosome
 base pairs remaining after filtering.</pre>
- method String indicating which filter method to use: chromosome size (method = "size") or proportion of total bases (method = "prop").

Returns: This R6 object, invisibly.

Examples:

```
ref <- create_genome(4, 100, 50)
ref$filter_chroms(90, "size")
ref$filter_chroms(0.4, "prop")</pre>
```

Method replace_Ns(): Replace Ns in the reference genome.

Usage:

```
ref_genome$replace_Ns(pi_tcag, n_threads = 1, show_progress = FALSE)
```

Arguments:

- pi_tcag Numeric vector (length 4) indicating the sampling weights for T, C, A, and G, respectively, for generating new nucleotides with which to replace the Ns.
- n_threads Optional integer specifying the threads to use. Ignored if the package wasn't compiled with OpenMP. Defaults to 1.

show_progress Optional logical indicating whether to show a progress bar. Defaults to FALSE.

Returns: This R6 object, invisibly.

See Also

read_fasta create_genome

Examples

```
## -----
## Method `ref_genome$set_names`
## ------
ref <- create_genome(4, 10)</pre>
ref$set_names(c("a", "b", "c", "d"))
## -----
## Method `ref_genome$clean_names`
## -----
ref <- create_genome(4, 10)</pre>
ref$set_names(c("a:", "b|", "c;", "d'"))
ref$clean_names()
## ------
## Method `ref_genome$add_chroms`
## -----
ref <- create_genome(4, 10)</pre>
ref$add_chroms("TCAGTCAG")
## ------
## Method `ref_genome$rm_chroms`
## -----
ref <- create_genome(4, 10)</pre>
ref$set_names(c("a", "b", "c", "d"))
ref$rm_chroms("b")
```

sub_models

```
## Method `ref_genome$merge_chroms`
## ------
ref <- create_genome(4, 10)
ref$merge_chroms(ref$chrom_names()[1:2])
ref$merge_chroms(NULL)
## ------
## Method `ref_genome$filter_chroms`
## ------
ref <- create_genome(4, 100, 50)
ref$filter_chroms(90, "size")
ref$filter_chroms(0.4, "prop")</pre>
```

```
sub_models
```

Construct necessary information for substitution models.

Description

For a more detailed explanation, see vignette("sub-models").

Usage

```
sub_JC69(lambda, mu = 1, gamma_shape = NULL, gamma_k = 5, invariant = 0)
sub_K80(alpha, beta, mu = 1, gamma_shape = NULL, gamma_k = 5, invariant = 0)
sub_F81(pi_tcag, mu = 1, gamma_shape = NULL, gamma_k = 5, invariant = 0)
sub_HKY85(
 pi_tcag,
 alpha,
 beta,
 mu = 1,
 gamma_shape = NULL,
  gamma_k = 5,
  invariant = 0
)
sub_F84(
  pi_tcag,
 beta,
  kappa,
 mu = 1,
  gamma_shape = NULL,
```

```
gamma_k = 5,
 invariant = 0
)
sub_TN93(
 pi_tcag,
 alpha_1,
 alpha_2,
 beta,
 mu = 1,
 gamma_shape = NULL,
 gamma_k = 5,
 invariant = 0
)
sub_GTR(
 pi_tcag,
  abcdef,
 mu = 1,
 gamma_shape = NULL,
 gamma_k = 5,
 invariant = 0
)
```

sub_UNREST(Q, mu = 1, gamma_shape = NULL, gamma_k = 5, invariant = 0)

Arguments

lambda	Substitution rate for all possible substitutions.
mu	Total rate of substitutions. Defaults to 1, which makes branch lengths in units of substitutions per site. Passing NULL results in no scaling.
gamma_shape	Numeric shape parameter for discrete Gamma distribution used for among-site variability. Values must be greater than zero. If this parameter is NULL, among-site variability is not included. Defaults to NULL.
gamma_k	The number of categories to split the discrete Gamma distribution into. Values must be an integer in the range [2,255]. This argument is ignored if gamma_shape is NA. Defaults to 5.
invariant	Proportion of sites that are invariant. Values must be in the range $[0,1)$. Defaults to 0.
alpha	Substitution rate for transitions.
beta	Substitution rate for transversions.
pi_tcag	Vector of length 4 indicating the equilibrium distributions of T, C, A, and G respectively. Values must be ≥ 0 , and they are forced to sum to 1.
kappa	The transition/transversion rate ratio.
alpha_1	Substitution rate for T <-> C transition.
alpha_2	Substitution rate for A <-> G transition.

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abcdef	A vector of length 6 that contains the off-diagonal elements for the substitution rate matrix. See vignette("sub-models") for how the values are ordered in the matrix.
Q	Matrix of substitution rates for "T", "C", "A", and "G", respectively. Item Q[i,j] is the rate of substitution from nucleotide i to nucleotide j. Do not include indel rates here! Values on the diagonal are calculated inside the function so are ignored.

Value

A sub_info object, which is an R6 class that wraps the info needed for the create_haplotypes function. It does not allow the user to directly manipulate the info inside, as that should be done using the sub_models functions. You can use the following methods from the class to view information:

- Q() View a list of substitution rate matrices, one for each Gamma category.
- pi_tcag() View the equilibrium nucleotide frequencies.

gammas() View the discrete Gamma-class values.

invariant() View the proportion of invariant sites.

model() View the substitution model.

- U() View list of the U matrices (one matrix per Gamma category) used for calculating transitionprobability matrices. This is empty for UNREST models.
- Ui () View list of the U⁻¹ matrices (one matrix per Gamma category) used for calculating transitionprobability matrices. This is empty for UNREST models.
- L() View list of the lambda vectors (one vector per Gamma category) used for calculating transitionprobability matrices. This is empty for UNREST models.

Functions

- sub_JC69(): JC69 model.
- sub_K80(): K80 model.
- sub_F81(): F81 model.
- sub_HKY85(): HKY85 model.
- sub_F84(): F84 model.
- sub_TN93(): TN93 model.
- sub_GTR(): GTR model.
- sub_UNREST(): UNREST model.

See Also

create_haplotypes

Examples

```
write_fasta
```

Write a ref_genome or haplotypes object to a FASTA file.

Description

This file produces 1 FASTA file for a ref_genome object and one file for each haplotype in a haplotypes object.

Usage

```
write_fasta(
   obj,
   out_prefix,
   compress = FALSE,
   comp_method = "bgzip",
   text_width = 80,
   show_progress = FALSE,
   n_threads = 1,
   overwrite = FALSE
)
```

Arguments

obj	A ref_genome or haplotypes object.
out_prefix	Prefix for the output file.
compress	Logical specifying whether or not to compress output file, or an integer specify- ing the level of compression, from 1 to 9. If TRUE, a compression level of 6 is used. Defaults to FALSE.

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write_vcf

comp_method	Character specifying which type of compression to use if any is desired. Options include "gzip" and "bgzip". This is ignored if compress is FALSE. Defaults to "bgzip".
text_width	The number of characters per line in the output fasta file. Defaults to 80.
show_progress	Logical for whether to show a progress bar. Defaults to FALSE.
n_threads	Number of threads to use if writing from a haplotypes object. Threads are split among haplotypes, so it's not useful to provide more threads than haplotypes. This argument is ignored if obj is a ref_genome object, or if OpenMP is not enabled. Defaults to 1.
overwrite	Logical for whether to overwrite existing file(s) of the same name, if they exist. Defaults to FALSE.

Value

NULL

write_vcf

Write haplotype info from a haplotypes object to a VCF file.

Description

Compression in this function always uses "bgzip" for compatibility with "tabix".

Usage

```
write_vcf(
    haps,
    out_prefix,
    compress = FALSE,
    sample_matrix = NULL,
    show_progress = FALSE,
    overwrite = FALSE
)
```

Arguments

haps	A haplotypes object.
out_prefix	Prefix for the output file.
compress	Logical specifying whether or not to compress output file, or an integer specifying the level of compression, from 1 to 9. If TRUE, a compression level of 6 is used. Defaults to FALSE.
sample_matrix	Matrix to specify how haplotypes are grouped into samples if samples are not haploid. There should be one row for each sample, and each row should contain indices or names for the haplotypes present in that sample. Indices/names for haplotypes cannot be repeated. Instead of repeating indices here, you should

	use the dup_haps method of the haplotypes class to duplicate the necessary haplotype(s). The number of columns indicates the ploidy level: 2 columns for diploid, 3 for triploid, 4 for tetraploid, and so on; there is no limit to the ploidy level. If this argument is NULL, it's assumed that each haplotype is its own separate sample. Defaults to NULL.
show_progress	Logical for whether to show a progress bar. Defaults to FALSE.
overwrite	Logical for whether to overwrite existing file(s) of the same name, if they exist. Defaults to FALSE.

Value

NULL

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