# Package 'jackalope'

November 28, 2023

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 <a href="https://www.illumina.com/">https://www.illumina.com/</a>

and Pacific Biosciences (PacBio) <a href="https://www.pacb.com/">https://www.pacb.com/</a>> 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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**Encoding UTF-8** 

LazyData true

**Depends** R (>= 2.10)

biocViews

Imports ape, R6, Rcpp (>= 0.12.11), zlibbioc

LinkingTo Rcpp, RcppArmadillo, RcppProgress, Rhtslib, zlibbioc

SystemRequirements GNU make

RoxygenNote 7.2.3

Suggests coala, knitr, markdown, rmarkdown, scrm, testthat

VignetteBuilder knitr

create\_genome

URL https://github.com/lucasnell/jackalope

 $\pmb{BugReports} \ \text{https://github.com/lucasnell/jackalope/issues}$ 

NeedsCompilation yes

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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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# 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 $\leq$ 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))
```

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,
```

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```
epsilon = 0.03,
n_threads = 1,
show_progress = FALSE
)
```

## Arguments

sub

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.

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  $\geq 0$  and  $\leq 1$ . 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.

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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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 ( $10^{-10}$  events per site per generation).

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

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

theta\_s Population mutation rate estimated using  $\theta_s$ .

pi\_s Population mutation rate estimated using  $\pi_s$ .

## 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.
 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$gc_prop(hap_ind, chrom_ind, start, end)
 Arguments:
 hap_ind Index for the focal haplotype.
 chrom_ind Index for the focal chromosome.
 start Point on the chromosome at which to start the calculation (inclusive).
 end Point on the chromosome at which to end the calculation (inclusive).
 Returns: A double in the range [0,1] representing the proportion of DNA sequence that is
 either G or C.
Method nt_prop(): View nucleotide content for part of one haplotype chromosome
 haplotypes$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. 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. 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

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haps\_functions

Organize higher-level information for creating 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 coalescent-simulator 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)
```

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# **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)
```

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#### **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.

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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

haps_vcf	Organize information to create haplotypes using a VCF file

## **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(obj,
    out_prefix,
    n_reads,
    read_length,
    paired,
    frag_mean = 400,
    frag_sd = 100,
    matepair = FALSE,
```

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```
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

del\_prob1

ins\_prob2

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 TRUE if matepair is 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 file for read 1. See "Sequencing profiles" section for more information 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 information 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.

Deletion probability for read 1. Defaults to 0.00011.

Insertion probability for read 2. Defaults to 0.00015.

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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<sup>32-1</sup>, 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 sequencing 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. Logical indicating whether to make separate files for each haplotype. This arsep\_files 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 > 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. read\_pool\_size The number of reads to store before writing to disk. Increasing this number should improve speed but take up more memory. Defaults to 1000. Logical for whether to show a progress bar. Defaults to FALSE. show\_progress Logical for whether to overwrite existing FASTQ file(s) of the same name, if overwrite

#### Value

Nothing is returned.

they exist.

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## 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.

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## **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
- 2. 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

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- 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.

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# 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 param for the chi^2 distribution. Defaults to c(0.01214, -5.12, 675, 48303.07331.4691051212330266).			
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 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 $\emptyset.2$ .		
ins_prob	Probability for insertions for reads with one pass. Defaults to 0.11.		

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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.

min\_read\_length

Minium read length for lognormal distribution. Defaults to 50.

lognorm\_read\_length

Vector containing the 3 parameters for lognormal read length distribution. Defaults to c(0.200110276521, -10075.4363813, 17922.611306).

custom\_read\_lengths

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

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.

sep\_files 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.

> 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.

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".

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.

read\_pool\_size The number of reads to store before writing to disk. Increasing this number should improve speed but take up more memory. Defaults to 100.

Logical for whether to show a progress bar. Defaults to FALSE.

Logical for whether to overwrite existing FASTQ file(s) of the same name, if they exist.

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n\_threads

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# 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)</pre>
```

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

R6 Class Representing a Reference 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$print()
Method ptr(): View pointer to underlying C++ object (this is not useful to end users).
 ref_genome$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$gc_prop(chrom_ind, start, end)
 Arguments:
 chrom_ind Index for the focal chromosome.
 start Point on the chromosome at which to start the calculation (inclusive).
 end Point on the chromosome at which to end the calculation (inclusive).
 Returns: A double in the range [0,1] representing the proportion of DNA sequence that is
 either G or C.
Method nt_prop(): View nucleotide content for part of one reference chromosome
 ref_genome$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$set_names(new_names)
 Arguments:
 new_names Vector of new names to use. This must be the same length as the number of current
     names.
 Returns: This R6 object, invisibly.
 Examples:
 ref <- create_genome(4, 10)</pre>
 ref$set_names(c("a", "b", "c", "d"))
Method clean_names(): Clean chromosome names, converting ":;=%,\\|/\"\'" to "_".
 Usage:
 ref_genome$clean_names()
 Returns: This R6 object, invisibly.
 Examples:
 ref <- create_genome(4, 10)</pre>
 ref$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)
 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)</pre>
 ref$add_chroms("TCAGTCAG")
Method rm_chroms(): Remove one or more chromosomes by name
```

```
Usage:
 ref_genome$rm_chroms(chrom_names)
 Arguments:
 chrom_names Vector of the name(s) of the chromosome(s) to remove.
 Returns: This R6 object, invisibly.
 Examples:
 ref <- create_genome(4, 10)</pre>
 ref$set_names(c("a", "b", "c", "d"))
 ref$rm_chroms("b")
Method merge_chroms(): Merge chromosomes into one.
 Usage:
 ref_genome$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)</pre>
 ref$merge_chroms(ref$chrom_names()[1:2])
 ref$merge_chroms(NULL)
Method filter_chroms(): Filter chromosomes by size or for a proportion of total bases.
 ref_genome$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.
 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)</pre>
 ref$filter_chroms(90, "size")
 ref$filter_chroms(0.4, "prop")
```

**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 29

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,
```

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

## **Arguments**

lambda	Substitution	rate for al	ll 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. Val-

ues 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). De-

faults 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  $\geq 0$ , and they are forced to sum to 1.

kappa The transition/transversion rate ratio.

alpha\_1 Substitution rate for  $T \leftarrow C$  transition.

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alpha\_2 Substitution rate for  $A \leftarrow G$  transition.

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 func-

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

32 write\_fasta

## **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 specifying the level of compression, from 1 to 9. If TRUE, a compression level of 6 is

used. Defaults to FALSE.

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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 specify-

ing 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

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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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