# Package 'phyloscannerR'

October 18, 2017
<b>Title</b> Phylogenetics between and within hosts at once, all along the genome
Version 1.3.0
<b>Description</b> An R package for the second half of phyloscanner (tree analysis).
<b>Depends</b> R (>= $3.4.2$ )
<b>Imports</b> ape, data.table, dplyr, dtplyr, ff, GGally, ggplot2, ggtree, grid, gridExtra, gtable, kimisc, network, pegas, phangorn, phytools, prodlim, RColorBrewer, reshape2, scales, sna
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Encoding UTF-8
LazyData true
RoxygenNote 6.0.1
draw.summary.statistics gather.summary.statistics   multipage.summary.statistics gather.summary.statistics   phyloscanner.analyse.tree gather.summary.statistics   phyloscanner.analyse.tree gather.summary.statistics   phyloscanner.analyse.tree gather.summary.statistics   phyloscanner.analyse.tree gather.summary.statistics   phyloscanner.analyse.tree gather.summary.statistics   simplified.transmission.summary gather.summary.statistics   transmission.summary gather.summary.sum
draw.summary.statistics  Graph summary statistics for a single host
Description Graph summary statistics for a single host Usage

draw.summary.statistics(phyloscanner.trees, sum.stats, host, verbose = F)

#### **Arguments**

phyloscanner.trees

A list of class phyloscanner. trees

sum.stats The output of a call to gather.summary.statistics.

host The host to obtain graphs for.

verbose Verbose output

gather.summary.statistics

Make a data.table of per-window host statistics

#### **Description**

This function performs a parsimony reconstruction and classification of pairwise host relationships.

## Usage

```
gather.summary.statistics(phyloscanner.trees,
hosts = all.hosts.from.trees(phyloscanner.trees),
tip.regex = "^(.*)_read_([0-9]+)_count_([0-9]+)$", verbose = F)
```

## **Arguments**

phyloscanner.trees

 $A\ list\ of\ class\ {\tt phyloscanner.trees}$ 

hosts A list of hosts to record statistics for. If not specified, every identifiable host in

phyloscanner.trees

tip.regex Regular expression identifying tips from the dataset. This expects up to three

capture groups, for host ID, read ID, and read count (in that order). If the latter two groups are missing then read information will not be used. The default matches input from the phyloscanner pipeline where the host ID is the BAM file

name.

verbose Produce verbose output

## Value

A data.table

```
multipage.summary.statistics
```

Draw summary statistics to file for many hosts as a multipage file

#### **Description**

Draw summary statistics to file for many hosts as a multipage file

#### Usage

```
multipage.summary.statistics(phyloscanner.trees, sum.stats,
hosts = all.hosts.from.trees(phyloscanner.trees), file.name,
height = 11.6929, width = 8.26772, verbose = F)
```

## **Arguments**

```
phyloscanner.trees
```

A list of class phyloscanner. trees

sum.stats The output of a call to gather.summary.statistics.

hosts A vector of hosts to obtain graphs for. By default, all hosts detected in phyloscanner. trees.

file.name Output file name (should have a .pdf file extension)

height The height of each page of the output file in inches (defaults to A4 size) width The width of each page of the output file in inches (defaults to A4 size)

verbose Verbose output

phyloscanner.analyse.tree

Perform a phyloscanner analysis on a single tree

## **Description**

This function performs a parsimony reconstruction and classification of pairwise host relationships.

## Usage

```
phyloscanner.analyse.tree(tree.file.name, splits.rule = c("s", "r", "f"),
    sankoff.k = 0, sankoff.unsampled.switch.threshold = 0,
    continuation.unsampled.proximity.cost = 1000, outgroup.name = NULL,
    multifurcation.threshold = 0, guess.multifurcation.threshold = F,
    user.blacklist.file.name = NULL, duplicate.file.name = NULL,
    recombination.file.name = NULL,
    tip.regex = "^(.*)_read_([0-9]+)_count_([0-9]+)$",
    file.name.regex = "^\\D*([0-9]+)_to_([0-9]+)\D*$",
    seed = sample(1:1e+07, 1), norm.ref.file.name = NULL,
    norm.standardise.gag.pol = F, norm.constants = NULL,
    parsimony.blacklist.k = 0, raw.blacklist.threshold = 0,
    ratio.blacklist.threshold = 0, do.dual.blacklisting = F,
    max.reads.per.host = Inf, blacklist.underrepresented = F, use.ff = F,
    prune.blacklist = F, read.counts.matter.on.zero.length.tips = T,
    verbose = F)
```

#### **Arguments**

tree.file.name The name of the tree file (Newick or NEXUS format).

splits.rule

The rules by which the sets of hosts are split into groups in order to ensure that all groups can be members of connected subgraphs without causing conflicts. Options: s=Sankoff with optional within-host diversity penalty (slow, rigorous, recommended), r=Romero-Severson (quick, less rigorous with >2 hosts), f=Sankoff with continuation costs (experimental).

sankoff.k

For splits.rule = s or f only. The *k* parameter in the Sankoff reconstruction, representing the within-host diversity penalty.

sankoff.unsampled.switch.threshold

For splits.rule = s only. Threshold at which a lineage reconstructed as infecting a host will transition to the unsampled state, if it would be equally parsimonious to remain in that host.

continuation.unsampled.proximity.cost

For splits.rule = f only. The branch length at which an node is reconstructed as unsampled if all its neighbouring nodes are a greater distance away. The default is 1000, intended to be effectively infinite, such a node will never normally receive the unsampled state.

outgroup.name

The name of the tip in the phylogeny/phylogenies to be used as outgroup (if unspecified, trees will be assumed to be already rooted). This should be sufficiently distant to any sequence obtained from a host that it can be assumed that the MRCA of the entire tree was not a lineage present in any sampled individual.

multifurcation.threshold

If specified, branches shorter than this in the input tree will be collapsed to form multifurcating internal nodes. This is recommended; many phylogenetics packages output binary trees with short or zero-length branches indicating multifurcations.

guess.multifurcation.threshold

Whether to guess the multifurcation threshold from the branch lengths of the trees and the width of the genomic window (if that information is available). It is recommended that trees are examined by eye to check that they do appear to have multifurcations if using this option.

user.blacklist.file.name

The path of a text file containing the user-specified list of tips to be blacklisted duplicate.file.name

The path of a .csv file specifying which tree tips are from duplicate reads. Normally this is produced by phyloscanner\_make\_trees.py.

recombination.file.name

The path for file containing the results of the phyloscanner\_make\_trees.py recombination metric analysis.

tip.regex

Regular expression identifying tips from the dataset. This expects up to three capture groups, for host ID, read ID, and read count (in that order). If the latter two groups are missing then read information will not be used. The default matches input from the phyloscanner pipeline where the host ID is the BAM file name.

file.name.regex

Regular expression identifying window coordinates. Two capture groups: start and end; if the latter is missing then the first group is a single numerical identifier for the window. The default matches input from the phyloscanner pipeline.

seed

Random number seed; used by the downsampling process, and also ties in some parsimony reconstructions can be broken randomly.

#### norm.ref.file.name

Name of a file giving a normalisation constant for every genome position. Cannot be used simultaneously with norm. constants. If neither is given then no normalisation will be performed.

#### norm.standardise.gag.pol

Use only if norm.ref.file.name is given. An HIV-specific option: if true, the normalising constants are standardised so that the average on gag+pol equals 1. Otherwise they are standardised so the average on the whole genome equals 1.

norm. constants Either the path of a CSV file listing the file name for each tree (column 1) and the respective normalisation constant (column 2) or a single numerical normalisation constant to be applied to every tree. Cannot be used simultaneously with norm.ref.file.name. If neither is given then no normalisation will be performed.

#### parsimony.blacklist.k

The k parameter of the single-host Sankhoff parsimony reconstruction used to identify probable contaminants. A value of 0 is equivalent to not performing parsimony blacklisting.

#### raw.blacklist.threshold

Used to specify a read count to be used as a raw threshold for duplicate or parsimony blacklisting. Use with parsimony.blacklist.korduplicate.file.regex or both. Parsimony blacklisting will blacklist any subgraph with a read count strictly less than this threshold. Duplicate blacklisting will black list any duplicate read with a count strictly less than this threshold. The default value of 0 means nothing is blacklisted.

#### ratio.blacklist.threshold

Used to specify a read count ratio (between 0 and 1) to be used as a threshold for duplicate or parsimony blacklisting. Use with parsimony.blacklist.k or duplicate.file.regex or both. Parsimony blacklisting will blacklist a subgraph if the ratio of its read count to the total read count from the same host is strictly less than this threshold. Duplcate blacklisting will blacklist a duplicate read if the ratio of its count to the count of the duplicate (from another host) is strictly less than this threshold.

#### do.dual.blacklisting

Blacklist all reads from the minor subgraphs for all hosts established as dual by parsimony blacklisting (which must have been done for this to do anything).

#### max.reads.per.host

Used to turn on downsampling. If given, reads will be blacklisted such that read counts (or tip counts if no read counts are identified) from each host are equal (although see blacklist.underrepresented.

#### blacklist.underrepresented

If TRUE and max.reads.per.host is given, blacklist hosts from trees where their total tip count does not reach the maximum.

use.ff Use the ff package to store parsimony reconstruction matrices. Use if you run out of memory.

#### prune.blacklist

If TRUE, all blacklisted and reference tips (except the outgroup) are pruned away before starting parsimony-based reconstruction.

```
read.counts.matter.on.zero.length.tips

If TRUE, read counts on tips will be taken into account in parsimony reconstructions at the parents of zero-length terminal branches. Not applicable for the Romero-Severson-like reconstruction method.
```

verbose Give verbose output.

#### Value

A list of class phyloscanner. trees with a single item of class phyloscanner. tree.

```
phyloscanner.analyse.trees

Perform a phyloscanner analysis on a set of trees
```

### **Description**

This function performs a parsimony reconstruction and classification of pairwise host relationships.

## Usage

```
phyloscanner.analyse.trees(tree.directory,
  tree.file.regex = "^RAxML_bestTree.InWindow_([0-9]+_to_[0-9]+)\\.tree$",
  splits.rule = c("s", "r", "f"), sankoff.k = 0,
  sankoff.unsampled.switch.threshold = 0,
 continuation.unsampled.proximity.cost = 1000, outgroup.name = NULL,
 multifurcation.threshold = 0, guess.multifurcation.threshold = F,
 user.blacklist.directory = NULL, user.blacklist.file.regex = NULL,
 duplicate.file.directory = NULL,
 duplicate.file.regex = "^DuplicateReadCountsProcessed_InWindow_([0-9]+_to_[0-9]+).csv$",
 recombination.file.directory = NULL,
 recombination.file.regex = "^RecombinantReads_InWindow_([0-9]+_to_[0-9]+).csv$",
 tip.regex = "^(.*)_read_([0-9]+)_count_([0-9]+)$",
 file.name.regex = "^\D*([0-9]+)_to_([0-9]+)\D*$",
 seed = sample(1:1e+07, 1), norm.ref.file.name = NULL,
 norm.standardise.gag.pol = F, norm.constants = NULL,
 parsimony.blacklist.k = 0, raw.blacklist.threshold = 0,
 ratio.blacklist.threshold = 0, do.dual.blacklisting = F,
 max.reads.per.host = Inf, blacklist.underrepresented = F, use.ff = F,
 prune.blacklist = F, read.counts.matter.on.zero.length.tips = T,
 verbose = F)
```

#### **Arguments**

```
tree.directory The directory containing all input trees. tree.file.regex
```

A regular expression identifying every file in tree.directory that is to be included in the analysis. The first capture group, if present, gives a unique string identifying each tree. If this is NULL then phyloscanner will attempt to open every file in tree.directory.

splits.rule

The rules by which the sets of hosts are split into groups in order to ensure that all groups can be members of connected subgraphs without causing conflicts. Options: s=Sankoff with optional within-host diversity penalty (slow, rigorous, recommended), r=Romero-Severson (quick, less rigorous with >2 hosts), f=Sankoff with continuation costs (experimental).

 ${\sf sankoff.k}$ 

For splits.rule = s or f only. The *k* parameter in the Sankoff reconstruction, representing the within-host diversity penalty.

sankoff.unsampled.switch.threshold

For splits.rule = s only. Threshold at which a lineage reconstructed as infecting a host will transition to the unsampled state, if it would be equally parsimonious to remain in that host.

continuation.unsampled.proximity.cost

For splits.rule = f only. The branch length at which an node is reconstructed as unsampled if all its neighbouring nodes are a greater distance away. The default is 1000, intended to be effectively infinite, such a node will never normally receive the unsampled state.

outgroup.name

The name of the tip in the phylogeny/phylogenies to be used as outgroup (if unspecified, trees will be assumed to be already rooted). This should be sufficiently distant to any sequence obtained from a host that it can be assumed that the MRCA of the entire tree was not a lineage present in any sampled individual.

multifurcation.threshold

If specified, branches shorter than this in the input tree will be collapsed to form multifurcating internal nodes. This is recommended; many phylogenetics packages output binary trees with short or zero-length branches indicating multifurcations.

guess.multifurcation.threshold

Whether to guess the multifurcation threshold from the branch lengths of the trees and the width of the genomic window (if that information is available). It is recommended that trees are examined by eye to check that they do appear to have multifurcations if using this option.

user.blacklist.directory

An optional path for a folder containing pre-existing blacklist files. These tips are specified by the user to be excluded from the analysis.

user.blacklist.file.regex

A regular expression identifying every file in user.blacklist.directory that contains a blacklist. If a capture group is specified then its contents will uniquely identify the tree it belongs to, which must matches the IDs found by user.blacklist.regex. If these IDs cannot be identified then matching will be attempted using genome window coordinates.

duplicate.file.directory

An optional path for a folder containing information on duplicate reads, to be used for duplicate blacklisting. Normally this is produced by phyloscanner\_make\_trees.py.

duplicate.file.regex

A regular expression identifying every file in duplicate.file.directory that contains a duplicates file. If a capture group is specified then its contents will uniquely identify the tree it belongs to, which must matches the IDs found by duplicate.file.regex. If these IDs cannot be identified then matching will be attempted using genome window coordinates.

recombination.file.directory

An optional path for a folder containing results of the phyloscanner\_make\_trees.py recombination metric analysis.

recombination.file.regex

A regular expression identifying every file in recombination. file.directory that contains a recombination file. If a capture group is specified then its contents will uniquely identify the tree it belongs to, which must matches the IDs found by recombintion.file.regex. If these IDs cannot be identified then matching will be attempted using genome window coordinates.

tip.regex

Regular expression identifying tips from the dataset. This expects up to three capture groups, for host ID, read ID, and read count (in that order). If the latter two groups are missing then read information will not be used. The default matches input from the phyloscanner pipeline where the host ID is the BAM file name.

file.name.regex

Regular expression identifying window coordinates. Two capture groups: start and end; if the latter is missing then the first group is a single numerical identifier for the window. The default matches input from the phyloscanner pipeline.

seed

Random number seed; used by the downsampling process, and also ties in some parsimony reconstructions can be broken randomly.

norm.ref.file.name

Name of a file giving a normalisation constant for every genome position. Cannot be used simultaneously with norm. constants. If neither is given then no normalisation will be performed.

norm.standardise.gag.pol

Use only if norm.ref.file.name is given. An HIV-specific option: if true, the normalising constants are standardised so that the average on gag+pol equals 1. Otherwise they are standardised so the average on the whole genome equals 1.

norm.constants Either the path of a CSV file listing the file name for each tree (column 1) and the respective normalisation constant (column 2) or a single numerical normalisation constant to be applied to every tree. Cannot be used simultaneously with norm.ref.file.name. If neither is given then no normalisation will be performed.

parsimony.blacklist.k

The k parameter of the single-host Sankhoff parsimony reconstruction used to identify probable contaminants. A value of 0 is equivalent to not performing parsimony blacklisting.

raw.blacklist.threshold

Used to specify a read count to be used as a raw threshold for duplicate or parsimony blacklisting. Use with parsimony.blacklist.korduplicate.file.regex or both. Parsimony blacklisting will blacklist any subgraph with a read count strictly less than this threshold. Duplicate blacklisting will black list any duplicate read with a count strictly less than this threshold. The default value of 0 means nothing is blacklisted.

ratio.blacklist.threshold

Used to specify a read count ratio (between 0 and 1) to be used as a threshold for duplicate or parsimony blacklisting. Use with parsimony.blacklist.k or duplicate.file.regex or both. Parsimony blacklisting will blacklist a subgraph if the ratio of its read count to the total read count from the same host is strictly less than this threshold. Duplcate blacklisting will blacklist a duplicate read if the ratio of its count to the count of the duplicate (from another host) is strictly less than this threshold.

do.dual.blacklisting

Blacklist all reads from the minor subgraphs for all hosts established as dual by parsimony blacklisting (which must have been done for this to do anything).

max.reads.per.host

Used to turn on downsampling. If given, reads will be blacklisted such that read counts (or tip counts if no read counts are identified) from each host are equal (although see blacklist.underrepresented.

blacklist.underrepresented

If TRUE and max.reads.per.host is given, blacklist hosts from trees where their total tip count does not reach the maximum.

use.ff Use the ff package to store parsimony reconstruction matrices. Use if you run out of memory.

prune.blacklist

If TRUE, all blacklisted and reference tips (except the outgroup) are pruned away before starting parsimony-based reconstruction.

read.counts.matter.on.zero.length.tips

If TRUE, read counts on tips will be taken into account in parsimony reconstructions at the parents of zero-length terminal branches. Not applicable for the Romero-Severson-like reconstruction method.

verbose Give verbose output.

#### Value

A list of class phyloscanner. trees.

simplified.transmission.summary

Simplfy and visually display the pairwise host relationships across all trees

#### **Description**

Simplfy and visually display the pairwise host relationships across all trees

## Usage

```
simplified.transmission.summary(phyloscanner.trees, transmission.summary,
    arrow.threshold, plot = F)
```

## **Arguments**

phyloscanner.trees

A list of class phyloscanner. trees

arrow.threshold

The proportion of trees in which a pair of hosts need to show a direction of transmission for that direction to be indicated as an arrow. If both directions meet this threshold, the arrow is in the direction with the larger proportion of trees.

plot

If TRUE, the returned list has an item called simp.diagram, a ggplot object plotting the simplified relationship diagram.

 $trans.summary \quad The \ output \ of \ transmission.summary; a \ data.table.$ 

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transmission.summary Summarise the pairwise host relationships across all trees

## **Description**

Summarise the pairwise host relationships across all trees

## Usage

```
transmission.summary(phyloscanner.trees, win.threshold = 0,
  dist.threshold = Inf, allow.mt = T, verbose = F)
```

## **Arguments**

phyloscanner.trees

A list of class phyloscanner. trees

win.threshold The proportion of windows that a pair of hosts need to be related (adjacent and

within dist. threshold of each other) in order for them to appear in the sum-

mary.

dist.threshold The patristic distance within which the subgraphs from two hosts need to be in

order for them to be declared related (default is infinity, so adjacent hosts are

always related).

allow.mt If FALSE, directionality is only inferred between pairs of hosts where a single

clade from one host is nested in one from the other; this is more conservative.

verbose Give verbose output

#### Value

A data.table, every line of which counts the number of pairwise relationships of a particular type between a pair of hosts

write.annotated.tree Write the phylogeny with reconstructed host annotations to file

## **Description**

Write the phylogeny with reconstructed host annotations to file

## Usage

```
write.annotated.tree(phyloscanner.tree, file.name, format = c("pdf", "nexus"),
   pdf.scale.bar.width = 0.01, pdf.w = 50, pdf.hm = 0.15)
```

write.annotated.tree

## Arguments

phyloscanner.tree

 $A\ list\ of\ class\ phyloscanner.\ tree\ (usually\ an\ item\ in\ a\ list\ of\ class\ phyloscanner.\ trees)$ 

file.name The name of the output file

format The format - PDF or NEXUS - in which to write the output.

pdf.scale.bar.width

The width, in substitutions per site, of the scale bar in PDF output

pdf.w The width of the output PDF file, in inches

pdf.hm The height, in inches per tip, of the output PDF file

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