# Package 'sitePath'

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Type Package

Title Phylogeny-based sequence clustering with site polymorphism

**Version** 1.23.0

**Description** Using site polymorphism is one of the ways to cluster DNA/protein sequences but it is possible for the sequences with the same polymorphism on a single site to be genetically distant. This package is aimed at clustering sequences using site polymorphism and their corresponding phylogenetic trees. By considering their location on the tree, only the structurally adjacent sequences will be clustered. However, the adjacent sequences may not necessarily have the same polymorphism. So a branch-and-bound like algorithm is used to minimize the entropy representing the purity of site polymorphism of each cluster.

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**Depends** R (>= 4.1)

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allSitesName

Retrieve position of all the sites

# Description

The function is a way to get position of the resulting sites from SNPsites, fixationSites and parallelSites. The numbering is consistent with what's being set by setSiteNumbering

# Usage

```
allSitesName(x, ...)
## S3 method for class 'SNPsites'
allSitesName(x, ...)
## S3 method for class 'sitesMinEntropy'
allSitesName(x, ...)
## S3 method for class 'fixationSites'
allSitesName(x, ...)
## S3 method for class 'parallelSites'
allSitesName(x, ...)
## S3 method for class 'paraFixSites'
allSitesName(x, type = c("paraFix", "fixation", "parallel"), ...)
```

#### **Arguments**

x The object containing the sites from analysis... Other argumentstype Return fixation or parallel sites

#### Value

An integer vector for sites position

```
data(zikv_tree)
msaPath <- system.file('extdata', 'ZIKV.fasta', package = 'sitePath')
tree <- addMSA(zikv_tree, msaPath = msaPath, msaFormat = 'fasta')
snp <- SNPsites(tree)
allSitesName(snp)</pre>
```

as.data.frame.fixationSites

```
as.data.frame.fixationSites
```

Convert results to Data Frame

#### **Description**

Convert return of functions in sitePath package to a data.frame so can be better worked with. The group name for each tip is the same as groupTips.

A fixationSites object will output the mutation name of the fixation and the cluster name before and after the mutation.

An SNPsites object will output the tip name with the SNP and its position.

An parallelSites object will output the tip name with the group name and mutation info.

#### Usage

```
## S3 method for class 'fixationSites'
as.data.frame(x, row.names = NULL, optional = FALSE, ...)
## S3 method for class 'SNPsites'
as.data.frame(x, row.names = NULL, optional = FALSE, ...)
## S3 method for class 'parallelSites'
as.data.frame(x, row.names = NULL, optional = FALSE, ...)
```

# Arguments

```
x The object to be converted to data.frame.row.names Unimplemented.optional Unimplemented.... Other arguments.
```

#### Value

A data. frame object.

```
data(zikv_tree_reduced)
data(zikv_align_reduced)
tree <- addMSA(zikv_tree_reduced, alignment = zikv_align_reduced)
fixations <- fixationSites(lineagePath(tree))
as.data.frame(fixations)</pre>
```

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extractSite	Extract tips for a single site	

#### **Description**

The functions in sitePath usually include the results on more than one site. The function extractSite can be used to extract the predicted result on a single site.

#### Usage

```
extractSite(x, site, ...)
## S3 method for class 'fixationSites'
extractSite(x, site, ...)
```

#### **Arguments**

A fixationSites or a parallelSites object. More type will be supported in the later version.
 A site included in the result.
 Other arguments

#### Value

The predicted result of a single site

#### **Examples**

```
data(zikv_tree_reduced)
data(zikv_align_reduced)
tree <- addMSA(zikv_tree_reduced, alignment = zikv_align_reduced)
mutations <- fixationSites(lineagePath(tree))
extractSite(mutations, 139)</pre>
```

extractTips

Extract grouped tips for a single site

#### **Description**

The result of fixationSites and sitePath contains all the possible sites with fixation mutation. The function extractTips retrieves the name of the tips involved in the fixation.

For lineagePath, the function extractTips groups all the tree tips according to the amino acid/nucleotide of the site.

For parallelSites and sitePara object, the function extractTips retrieve all the tips with parallel mutation.

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#### Usage

```
extractTips(x, ...)
## S3 method for class 'lineagePath'
extractTips(x, site, ...)
## S3 method for class 'sitesMinEntropy'
extractTips(x, site, ...)
## S3 method for class 'fixationSites'
extractTips(x, site, select = 1, ...)
## S3 method for class 'sitePath'
extractTips(x, select = 1, ...)
## S3 method for class 'parallelSites'
extractTips(x, site, ...)
## S3 method for class 'sitePara'
extractTips(x, ...)
```

#### **Arguments**

x A fixationSites or a sitePath object.

... Other arguments

site A site predicted to experience fixation.

select For a site, there theoretically might be more than one fixation on different lin-

eages. You may use this argument to extract for a specific fixation of a site. The

default is the first fixation of the site.

#### Value

Tree tips grouped as list

```
data(zikv_tree_reduced)
data(zikv_align_reduced)
tree <- addMSA(zikv_tree_reduced, alignment = zikv_align_reduced)
mutations <- fixationSites(lineagePath(tree))
extractTips(mutations, 139)</pre>
```

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fixationIndels

Fixation indels prediction

# Description

The fixation of insertions of deletions.

#### Usage

```
fixationIndels(x, ...)
## S3 method for class 'sitesMinEntropy'
fixationIndels(x, ...)
```

#### **Arguments**

x The return from sitesMinEntropy function.
... Other arguments.

# Value

A fixationIndels object.

#### **Examples**

```
data(zikv_tree_reduced)
data(zikv_align_reduced)
tree <- addMSA(zikv_tree_reduced, alignment = zikv_align_reduced)
fixationIndels(sitesMinEntropy(tree))</pre>
```

fixationPath

Accumulation of fixed mutation as a tree

# **Description**

The tips are clustered according to the fixation sites. The transition of fixation sites will be plotted as a phylogenetic tree. The length of each branch represents the number of fixation mutation between two clusters. The name of the tree tips indicate the number of sequences in the cluster.

# Usage

```
fixationPath(x, ...)
## S3 method for class 'sitesMinEntropy'
fixationPath(x, minEffectiveSize = NULL, ...)
## S3 method for class 'fixationSites'
fixationPath(x, minEffectiveSize = NULL, ...)
```

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#### **Arguments**

```
x The return from fixationSites function.
... Further arguments passed to or from other methods.
minEffectiveSize
```

The minimum size for a tip cluster.

#### Value

An fixationPath object

#### **Examples**

```
data(zikv_tree_reduced)
data(zikv_align_reduced)
tree <- addMSA(zikv_tree_reduced, alignment = zikv_align_reduced)
paths <- lineagePath(tree)
mutations <- fixationSites(paths)
fixationPath(mutations)</pre>
```

fixationSites

Fixation sites prediction

# **Description**

After finding the lineagePath of a phylogenetic tree, fixationSites uses the result to find those sites that show fixation on some, if not all, of the lineages. The number of tips before and after the fixation mutation is expected to be more than minEffectiveSize. Also, the fixation will be skipped if the amino acid/nucleotide is gap or ambiguous character. A lineage has to have at least one fixation mutation to be reported.

### Usage

```
fixationSites(paths, ...)

## S3 method for class 'lineagePath'
fixationSites(
  paths,
  minEffectiveSize = NULL,
  searchDepth = 1,
  method = c("compare", "insert", "delete"),
  ...
)

## S3 method for class 'sitesMinEntropy'
fixationSites(paths, ...)

## S3 method for class 'paraFixSites'
fixationSites(paths, ...)
```

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#### **Arguments**

paths A lineagePath object returned from lineagePath function.

... further arguments passed to or from other methods.

minEffectiveSize

The minimum number of tips in a group.

searchDepth The function uses heuristic search but the termination of the search cannot be

intrinsically decided. searchDepth is needed to tell the search when to stop.

method The strategy for predicting the fixation. The basic approach is entropy minimiza-

tion and can be achieved by adding or removing fixation point, or by comparing

the two.

#### Value

A fixationSites object.

#### See Also

```
as.data.frame.fixationSites
```

#### **Examples**

```
data(zikv_tree_reduced)
data(zikv_align_reduced)
tree <- addMSA(zikv_tree_reduced, alignment = zikv_align_reduced)
fixationSites(lineagePath(tree))</pre>
```

groupTips

The grouping of tree tips

# Description

The tips between divergent nodes or fixation mutations on the lineages are each gathered as group.

#### Usage

```
groupTips(tree, ...)
## S3 method for class 'phyMSAmatched'
groupTips(
   tree,
   similarity = NULL,
   simMatrix = NULL,
   forbidTrivial = TRUE,
   tipnames = TRUE,
   ...
)
```

h3n2\_align

```
## S3 method for class 'lineagePath'
groupTips(tree, tipnames = TRUE, ...)
## S3 method for class 'sitesMinEntropy'
groupTips(tree, tipnames = TRUE, ...)
## S3 method for class 'fixationSites'
groupTips(tree, tipnames = TRUE, ...)
## S3 method for class 'fixationPath'
groupTips(tree, tipnames = TRUE, ...)
```

#### Arguments

tree The return from addMSA, lineagePath, sitesMinEntropy or other functions.

... Other arguments.

similarity This decides how minor SNPs are to remove. If provided as fraction between

0 and 1, then the minimum number of SNP will be total tips times similarly. If provided as integer greater than 1, the minimum number will be similarly.

The default similarly is 0.05 for lineagePath.

simMatrix Deprecated and will not have effect.
forbidTrivial Does not allow trivial trimming.
tipnames If return tips as integer or tip names.

#### Value

groupTips returns grouping of tips.

#### **Examples**

```
data(zikv_tree)
data(zikv_align)
tree <- addMSA(zikv_tree, alignment = zikv_align)
groupTips(tree)</pre>
```

h3n2\_align

Multiple sequence alignment of H3N2's HA protein

#### **Description**

The raw protein sequences were downloaded from NCBI database and aligned using MAFFT (https://mafft.cbrc.jp/alignment/software/).

h3n2\_align\_reduced is a truncated version of h3n2\_align

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#### Usage

```
data(h3n2_align)
data(h3n2_align_reduced)
```

#### **Format**

an alignment object an alignment object

h3n2\_tree

Phylogenetic tree of H3N2's HA protein

# Description

Tree was built from h3n2\_align using RAxML (http://www.exelixis-lab.org/) with default settings.

h3n2\_tree\_reduced is a truncated version of h3n2\_tree

# Usage

```
data(h3n2_tree)
data(h3n2_tree_reduced)
```

#### **Format**

a phylo object a phylo object

lineagePath

Resolving lineage paths using SNP

# **Description**

lineagePath finds the lineages of a phylogenetic tree providing the corresponding sequence alignment. This is done by finding 'major SNPs' which usually accumulate along the evolutionary pathways.

sneakPeek is intended to plot 'similarity' (actually the least percentage of 'major SNP') as a threshold against number of output lineagePath. This plot is intended to give user a rough view about how many lineages they could expect from the 'similarity' threshold in the function lineagePath. The number of lineagePath is preferably not be too many or too few. The result excludes where the number of lineagePath is greater than number of tips divided by 20 or user-defined maxPath. The zero lineagePath result will also be excluded.

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When used on the return of sneakPeek, a lineagePath with the closest similarity will be retrieved from the returned value.

similarity has no effect when using on paraFixSites object

#### Usage

```
lineagePath(tree, similarity, ...)
## S3 method for class 'phylo'
lineagePath(
  tree,
  similarity = NULL,
  alignment = NULL,
  seqType = c("AA", "DNA", "RNA"),
  reference = NULL,
  gapChar = "-",
 minSkipSize = NULL,
)
## S3 method for class 'treedata'
lineagePath(tree, ...)
## S3 method for class 'phyMSAmatched'
lineagePath(
  tree,
  similarity = NULL,
  simMatrix = NULL,
  forbidTrivial = TRUE,
)
sneakPeek(tree, step = 9, maxPath = NULL, minPath = 0, makePlot = TRUE)
## S3 method for class 'sneakPeekedPaths'
lineagePath(tree, similarity, ...)
## S3 method for class 'paraFixSites'
lineagePath(tree, similarity = NULL, ...)
```

# Arguments

The return from addMSA or sneakPeek function.

The parameter for identifying phylogenetic pathway using SNP. If provided as fraction between 0 and 1, then the minimum number of SNP will be total tips times Nmin. If provided as integer greater than 1, the minimum number will be Nmin.

... Other arguments.

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An alignment object. This commonly can be from sequence parsing function in the seqinr package. Sequence names in the alignment should include all tip.label in the tree  seqType The type of the sequence in the alignment file. The default is "AA" for amino acid. The other options are "DNA" and "RNA".  reference Name of reference for site numbering. The name has to be one of the sequences' name. The default uses the intrinsic alignment numbering  gapChar The character to indicate gap. The numbering will skip the gapChar for the reference sequence.  minSkipSize The minimum number of tips to have gap or ambiguous amino acid/nucleotide for a site to be ignored in other analysis. This will not affect the numbering. The default is 0.8.  simMatrix Deprecated and will not have effect.  forbidTrivial Does not allow trivial trimming.  step the 'similarity' window for calculating and plotting. To better see the impact of threshold on path number. The default is 10.  maxPath maximum number of path to return show in the plot. The number of path in the raw tree can be far greater than trimmed tree. To better see the impact of threshold on path number. This is preferably specified. The default is one 20th of tree tip number.  minPath minimum number of path to return show in the plot. To better see the impact of threshold on path number. This default is 1.		
acid. The other options are "DNA" and "RNA".  reference Name of reference for site numbering. The name has to be one of the sequences' name. The default uses the intrinsic alignment numbering  gapChar The character to indicate gap. The numbering will skip the gapChar for the reference sequence.  minSkipSize The minimum number of tips to have gap or ambiguous amino acid/nucleotide for a site to be ignored in other analysis. This will not affect the numbering. The default is 0.8.  simMatrix Deprecated and will not have effect.  forbidTrivial Does not allow trivial trimming.  step the 'similarity' window for calculating and plotting. To better see the impact of threshold on path number. The default is 10.  maxPath maximum number of path to return show in the plot. The number of path in the raw tree can be far greater than trimmed tree. To better see the impact of threshold on path number. This is preferably specified. The default is one 20th of tree tip number.  minPath minimum number of path to return show in the plot. To better see the impact of threshold on path number. The default is 1.	alignment	in the seqinr package. Sequence names in the alignment should include all
mame. The default uses the intrinsic alignment numbering  gapChar The character to indicate gap. The numbering will skip the gapChar for the reference sequence.  minSkipSize The minimum number of tips to have gap or ambiguous amino acid/nucleotide for a site to be ignored in other analysis. This will not affect the numbering. The default is 0.8.  simMatrix Deprecated and will not have effect.  forbidTrivial Does not allow trivial trimming.  step the 'similarity' window for calculating and plotting. To better see the impact of threshold on path number. The default is 10.  maxPath maximum number of path to return show in the plot. The number of path in the raw tree can be far greater than trimmed tree. To better see the impact of threshold on path number. This is preferably specified. The default is one 20th of tree tip number.  minPath minimum number of path to return show in the plot. To better see the impact of threshold on path number. The default is 1.	seqType	7.2
reference sequence.  The minimum number of tips to have gap or ambiguous amino acid/nucleotide for a site to be ignored in other analysis. This will not affect the numbering. The default is 0.8.  SimMatrix Deprecated and will not have effect.  forbidTrivial Does not allow trivial trimming.  step the 'similarity' window for calculating and plotting. To better see the impact of threshold on path number. The default is 10.  maxPath maximum number of path to return show in the plot. The number of path in the raw tree can be far greater than trimmed tree. To better see the impact of threshold on path number. This is preferably specified. The default is one 20th of tree tip number.  minPath minimum number of path to return show in the plot. To better see the impact of threshold on path number. The default is 1.	reference	
for a site to be ignored in other analysis. This will not affect the numbering. The default is 0.8.  simMatrix Deprecated and will not have effect.  forbidTrivial Does not allow trivial trimming.  step the 'similarity' window for calculating and plotting. To better see the impact of threshold on path number. The default is 10.  maxPath maximum number of path to return show in the plot. The number of path in the raw tree can be far greater than trimmed tree. To better see the impact of threshold on path number. This is preferably specified. The default is one 20th of tree tip number.  minPath minimum number of path to return show in the plot. To better see the impact of threshold on path number. The default is 1.	gapChar	
forbidTrivial Does not allow trivial trimming.  step the 'similarity' window for calculating and plotting. To better see the impact of threshold on path number. The default is 10.  maxPath maximum number of path to return show in the plot. The number of path in the raw tree can be far greater than trimmed tree. To better see the impact of threshold on path number. This is preferably specified. The default is one 20th of tree tip number.  minPath minimum number of path to return show in the plot. To better see the impact of threshold on path number. The default is 1.	minSkipSize	for a site to be ignored in other analysis. This will not affect the numbering. The
the 'similarity' window for calculating and plotting. To better see the impact of threshold on path number. The default is 10.  maxPath  maximum number of path to return show in the plot. The number of path in the raw tree can be far greater than trimmed tree. To better see the impact of threshold on path number. This is preferably specified. The default is one 20th of tree tip number.  minPath  minimum number of path to return show in the plot. To better see the impact of threshold on path number. The default is 1.	simMatrix	Deprecated and will not have effect.
maxPath maximum number of path to return show in the plot. The number of path in the raw tree can be far greater than trimmed tree. To better see the impact of threshold on path number. This is preferably specified. The default is one 20th of tree tip number.  minPath minimum number of path to return show in the plot. To better see the impact of threshold on path number. The default is 1.	forbidTrivial	Does not allow trivial trimming.
the raw tree can be far greater than trimmed tree. To better see the impact of threshold on path number. This is preferably specified. The default is one 20th of tree tip number.  minPath  minimum number of path to return show in the plot. To better see the impact of threshold on path number. The default is 1.	step	• • • • • • • • • • • • • • • • • • • •
threshold on path number. The default is 1.	maxPath	the raw tree can be far greater than trimmed tree. To better see the impact of threshold on path number. This is preferably specified. The default is one 20th
makeDlet Whether make a plot when return	minPath	minimum number of path to return show in the plot. To better see the impact of
makePlot Whether make a plot when return.		threshold on path number. The default is 1.

#### Value

Lineage path represent by node number.

sneakPeek return the similarity threhold against number of lineagePath. There will be a simple dot plot between threshold and path number if makePlot is TRUE.

```
data('zikv_tree')
data('zikv_align')
tree <- addMSA(zikv_tree, alignment = zikv_align)
lineagePath(tree)
sneakPeek(tree, step = 3)
x <- sneakPeek(tree, step = 3)
lineagePath(x, similarity = 0.05)</pre>
```

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paraFixSites

The fixation sites with mutation on parallel lineage

#### **Description**

The operation between the results of fixationSites and parallelSites.

# Usage

```
paraFixSites(x, ...)
## S3 method for class 'phylo'
paraFixSites(
  Х,
  alignment = NULL,
  seqType = c("AA", "DNA", "RNA"),
 Nmin = NULL,
  reference = NULL,
  gapChar = "-",
 minSkipSize = NULL,
)
## S3 method for class 'treedata'
paraFixSites(x, ...)
## S3 method for class 'lineagePath'
paraFixSites(
  Х,
 minEffectiveSize = NULL,
  searchDepth = 1,
 method = c("compare", "insert", "delete"),
)
## S3 method for class 'sitesMinEntropy'
paraFixSites(
  Х,
  category = c("intersect", "union", "parallelOnly", "fixationOnly"),
 minSNP = NULL,
 mutMode = c("all", "exact", "pre", "post"),
)
```

# **Arguments** ×

A lineagePath object returned from lineagePath function.

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... further arguments passed to or from other methods.

alignment An alignment object. This commonly can be from sequence parsing function

in the seqinr package. Sequence names in the alignment should include all

tip.label in the tree

seqType The type of the sequence in the alignment file. The default is "AA" for amino

acid. The other options are "DNA" and "RNA".

Nmin The parameter for identifying phylogenetic pathway using SNP. If provided as

fraction between 0 and 1, then the minimum number of SNP will be total tips times Nmin. If provided as integer greater than 1, the minimum number will be

Nmin.

reference Name of reference for site numbering. The name has to be one of the sequences'

name. The default uses the intrinsic alignment numbering

gapChar The character to indicate gap. The numbering will skip the gapChar for the

reference sequence.

minSkipSize The minimum number of tips to have gap or ambiguous amino acid/nucleotide

for a site to be ignored in other analysis. This will not affect the numbering. The

default is 0.8.

minEffectiveSize

The minimum number of tips in a group.

searchDepth The function uses heuristic search but the termination of the search cannot be

intrinsically decided. searchDepth is needed to tell the search when to stop.

method The strategy for predicting the fixation. The basic approach is entropy minimiza-

tion and can be achieved by adding or removing fixation point, or by comparing

the two.

category Could be parallelOnly, fixationOnly, intersect or union.

minSNP The minimum number of mutations to be qualified as parallel on at least two

lineages. The default is 1.

mutMode The strategy for finding parallel site. The default all is to consider any mutation

regardless of the amino acid/nucleotide before and after mutation; Or exact to force mutation to be the same; Or pre/post to select the site having amino

acid/nucleotide before/after mutation.

#### Value

A paraFixSites object.

```
data(zikv_tree_reduced)
data(zikv_align_reduced)
paraFixSites(zikv_tree_reduced, alignment = zikv_align_reduced)
```

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parallelSites

Mutation across multiple phylogenetic lineages

#### **Description**

A site may have mutated on parallel lineages. Mutation can occur on the same site across the phylogenetic lineages solved by lineagePath. The site will be considered mutated in parallel if the mutation occurs on the non-overlap part of more than two lineages. The amino acid/nucleotide before and after the mutation can be allowed different on different lineages or only the exact same mutations are considered.

#### Usage

```
parallelSites(x, ...)

## S3 method for class 'lineagePath'
parallelSites(
    x,
    minSNP = NULL,
    mutMode = c("all", "exact", "pre", "post"),
    ...
)

## S3 method for class 'sitesMinEntropy'
parallelSites(
    x,
    minSNP = NULL,
    mutMode = c("all", "exact", "pre", "post"),
    ...
)

## S3 method for class 'paraFixSites'
parallelSites(x, ...)
```

#### **Arguments**

minSNP

x A lineagePath or a sitesMinEntropy object.

... The arguments in sitesMinEntropy.

The minimum number of mutations to be qualified as parallel on at least two

lineages. The default is 1.

mutMode The strategy for finding parallel site. The default all is to consider any mutation

regardless of the amino acid/nucleotide before and after mutation; Or exact to force mutation to be the same; Or pre/post to select the site having amino

acid/nucleotide before/after mutation.

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#### Value

A parallelSites object

#### **Examples**

```
data(zikv_tree_reduced)
data(zikv_align_reduced)
tree <- addMSA(zikv_tree_reduced, alignment = zikv_align_reduced)
paths <- lineagePath(tree)
x <- sitesMinEntropy(paths)
parallelSites(x)</pre>
```

phyMSAmatched

Add matching sequence alignment to the tree

#### **Description**

addMSA wraps read.alignment function in seqinr package and helps match names in tree and sequence alignment. Either provide the file path to an alignment file and its format or an alignment object from the return of read.alignment function. If both the file path and alignment object are given, the function will use the sequence in the alignment file.

#### Usage

```
addMSA(tree, ...)

## S3 method for class 'phylo'
addMSA(
    tree,
    msaPath = "",
    msaFormat = c("fasta", "clustal", "phylip", "mase", "msf"),
    alignment = NULL,
    seqType = c("AA", "DNA", "RNA"),
    ...
)

## S3 method for class 'treedata'
addMSA(tree, ...)
```

#### **Arguments**

A phylo object. This commonly can be from tree parsing function in ape or ggtree. All the tip.label should be found in the sequence alignment. The tree is supposed to be fully resolved (bifurcated) and will be resolved by multi2di if is.binary gives FALSE.

Other arguments.

msaPath The file path to the multiple sequence alignment file.

msa⊦ormat	The format of the multiple sequence alignment file. The internal uses the read. alignment
	from seqinr package to parse the sequence alignment. The default is "fasta" and
	it also accepts "clustal", "phylip", "mase", "msf".
alignment	An alignment object. This commonly can be from sequence parsing function
	in the seqinr package. Sequence names in the alignment should include all

tip.label in the tree

seqType The type of the sequence in the alignment file. The default is "AA" for amino

acid. The other options are "DNA" and "RNA".

#### Value

Since 1.5.12, the function returns a phyMSAmatched object to avoid S3 methods used on phylo (better encapsulation).

#### See Also

```
read.alignment
```

#### **Examples**

```
data(zikv_tree)
msaPath <- system.file('extdata', 'ZIKV.fasta', package = 'sitePath')
addMSA(zikv_tree, msaPath = msaPath, msaFormat = 'fasta')</pre>
```

plot.phyMSAmatched

Visualize the results

#### **Description**

The plot function to visualize the return of functions in the package. The underlying function applies ggplot2. The function name plot is used to keep the compatibility with previous versions, but they do not behave like the generic plot function since 1.5.4.

A phyMSAmatched object will be plotted as a tree diagram.

A lineagePath object will be plotted as a tree diagram and paths are black solid line while the trimmed nodes and tips will use gray dashed line.

A parallelSites object will be plotted as original phylogenetic tree marked with parallel mutations attached as dot plot.

A fixationSites object will be plotted as original phylogenetic tree marked with fixation substitutions

A sitePath object can be extracted by using extractSite on the return of fixationSites.

A fixationIndels object will be plotted as original phylogenetic tree marked with indel fixation.

A fixationPath object will be plotted as a phylo object. The tips are clustered according to the fixation sites. The transition of fixation sites will be plotted as a phylogenetic tree. The length of each branch represents the number of fixation mutation between two clusters.

plot.phyMSAmatched

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#### Usage

```
## S3 method for class 'phyMSAmatched'
plot(x, y = TRUE, ...)

## S3 method for class 'lineagePath'
plot(x, y = TRUE, showTips = FALSE, ...)

## S3 method for class 'parallelSites'
plot(x, y = TRUE, ...)

## S3 method for class 'fixationSites'
plot(x, y = TRUE, tipsGrouping = NULL, ...)

## S3 method for class 'sitePath'
plot(x, y = NULL, select = NULL, showTips = FALSE, ...)

## S3 method for class 'fixationIndels'
plot(x, y = TRUE, ...)

## S3 method for class 'fixationPath'
plot(x, y = TRUE, ...)
```

#### **Arguments**

X	The object to plot.
У	Whether to show the fixation mutation between clusters. For lineagePath object and sitePath object, it is deprecated and no longer have effect since 1.5.4.
• • •	Other arguments. Since 1.5.4, the function uses ggtree as the base function to make plots so the arguments in plot.phylo will no longer work.
showTips	Whether to plot the tip labels. The default is FALSE.
tipsGrouping	A list to hold the grouping of tips for how the tree will be colored.
select	For a sitePath object, it can have result on more than one evolution pathway. This is to select which path to plot. The default is NULL which will plot all the paths. It is the same as select in plotSingleSite.

# Value

A ggplot object to make the plot.

```
data(zikv_tree)
data(zikv_align)
tree <- addMSA(zikv_tree, alignment = zikv_align)
plot(tree)
paths <- lineagePath(tree)
plot(paths)
parallel <- parallelSites(paths)</pre>
```

20 plotFixationSites

```
plot(parallel)
fixations <- fixationSites(paths)
plot(fixations)
sp <- extractSite(fixations, 139)
plot(sp)
x <- fixationPath(fixations)
plot(x)</pre>
```

plotFixationSites

Plot the result of fixation sites

#### Description

Visualize the results of paraFixSites

# Usage

```
plotFixationSites(x, ...)
## S3 method for class 'fixationSites'
plotFixationSites(x, site = NULL, ...)
## S3 method for class 'paraFixSites'
plotFixationSites(x, site = NULL, ...)
```

#### **Arguments**

return from paraFixSites
 further arguments passed to or from other methods.
 the number of the site according to setSiteNumbering. If not provided, all sites will be plotted as labels on the tree

#### Value

A ggplot object.

```
data(zikv_tree_reduced)
data(zikv_align_reduced)
paraFix <- paraFixSites(zikv_tree_reduced, alignment = zikv_align_reduced)
plotFixationSites(paraFix)</pre>
```

plotMutSites 21

plotMutSites

Plot tree and mutation sites

# **Description**

The mutated sites for each tip in a phylogenetic tree will be represented as colored dots positioned by their site number.

# Usage

```
plotMutSites(x, ...)
## S3 method for class 'SNPsites'
plotMutSites(x, showTips = FALSE, ...)
## S3 method for class 'lineagePath'
plotMutSites(x, ...)
## S3 method for class 'parallelSites'
plotMutSites(x, ...)
## S3 method for class 'fixationSites'
plotMutSites(x, ...)
## S3 method for class 'paraFixSites'
plotMutSites(
  widthRatio = 0.75,
  fontSize = 3.88,
  dotSize = 1,
  lineSize = 0.5,
)
```

#### **Arguments**

X	An SNPsites object.
	Other arguments
showTips	Whether to plot the tip labels. The default is FALSE
widthRatio	The width ratio between tree plot and SNP plot
fontSize	The font size of the mutation label in tree plot
dotSize	The dot size of SNP in SNP plot
lineSize	The background line size in SNP plot

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#### Value

A tree plot with SNP as dots for each tip.

# **Examples**

```
data(zikv_tree_reduced)
data(zikv_align_reduced)
tree <- addMSA(zikv_tree_reduced, alignment = zikv_align_reduced)
plotMutSites(SNPsites(tree))</pre>
```

plotParallelSites

Plot the result of fixation sites

#### **Description**

Visualize the results of paraFixSites

#### Usage

```
plotParallelSites(x, ...)
## S3 method for class 'parallelSites'
plotParallelSites(x, site = NULL, ...)
## S3 method for class 'paraFixSites'
plotParallelSites(x, site = NULL, ...)
```

# Arguments

```
    return from paraFixSites
    further arguments passed to or from other methods.
    the number of the site according to setSiteNumbering
```

#### Value

A ggplot object.

```
data(zikv_tree)
data(zikv_align)
paraFix <- paraFixSites(zikv_tree, alignment = zikv_align)
plotParallelSites(paraFix)</pre>
```

plotSingleSite 23

plotSingleSite	Color the tree by a single site

#### **Description**

Plot and color the tree according to amino acid/nucleotide of the selected site. The color scheme depends on the seqType set in addMSA function.

For lineagePath, the tree will be colored according to the amino acid of the site. The color scheme tries to assign distinguishable color for each amino acid.

For parallelSites, the tree will be colored according to the amino acid of the site if the mutation is not fixed.

For fixationSites, it will color the ancestral tips in red, descendant tips in blue and excluded tips in grey.

#### Usage

```
plotSingleSite(x, site, ...)
## S3 method for class 'lineagePath'
plotSingleSite(x, site, showPath = TRUE, showTips = FALSE, ...)
## S3 method for class 'sitesMinEntropy'
plotSingleSite(x, site, ...)
## S3 method for class 'parallelSites'
plotSingleSite(x, site, showPath = TRUE, ...)
## S3 method for class 'fixationSites'
plotSingleSite(x, site, select = NULL, ...)
```

#### Arguments

x	The object to plot.
site	For lineagePath, it can be any site within sequence length. For fixationSites and parallelSites, it is restrained to a predicted fixation site. The numbering is consistent with the reference defined by setSiteNumbering.
•••	Other arguments. Since 1.5.4, the function uses ggtree as the base function to make plots so the arguments in plot.phylo will no longer work.
showPath	If plot the lineage result from lineagePath. The default is TRUE.
showTips	Whether to plot the tip labels. The default is FALSE.
select	Select which fixation path in to plot. The default is NULL which will plot all the fixations.

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#### Value

Since 1.5.4, the function returns a ggplot object so on longer behaviors like the generic plot function

#### See Also

```
plot.sitePath
```

#### **Examples**

```
data(zikv_tree)
data(zikv_align)
tree <- addMSA(zikv_tree, alignment = zikv_align)
paths <- lineagePath(tree)
plotSingleSite(paths, 139)
fixations <- fixationSites(paths)
plotSingleSite(fixations, 139)</pre>
```

reexports

Objects exported from other packages

# Description

These objects are imported from other packages. Follow the links below to see their documentation.

```
ape as.phylo, read.tree
seqinr read.alignment
tidytree as.treedata
```

sars2\_align

Multiple sequence alignment of SARS-CoV-2 genome CDS

# Description

The raw sequences were downloaded from GISAID database (https://www.gisaid.org/) and aligned using MAFFT (https://mafft.cbrc.jp/alignment/software/) with default settings.

#### Usage

```
data(sars2_align)
```

#### **Format**

an alignment object

sars2\_tree 25

sars2_tree Phylogenetic tree of SARS-CoV-2 genome CDS	sars2_tree	Phylogenetic tree of SARS-CoV-2 genome CDS
---	------------	--

#### **Description**

Tree was built from sars2\_align using RAxML (http://www.exelixis-lab.org/) with default settings. The tip EPI\_ISL\_402125 was used as the outgroup to root the tree.

# Usage

```
data(sars2_tree)
```

#### **Format**

a phylo object

setSiteNumbering

Set site numbering to the reference sequence

# Description

A reference sequence can be used to define a global site numbering scheme for multiple sequence alignment. The gap in the reference sequence will be skipped for the numbering. Also, the site that is gap or amino acid/nucleotide for too many tips will be ignored but won't affect numbering.

#### Usage

```
setSiteNumbering(x, reference, gapChar, ...)
## S3 method for class 'phyMSAmatched'
setSiteNumbering(x, reference = NULL, gapChar = "-", minSkipSize = NULL, ...)
```

# **Arguments**

X	The object to set site numbering. It could be a phyMSAmatched or a lineagePath object.
reference	Name of reference for site numbering. The name has to be one of the sequences' name. The default uses the intrinsic alignment numbering
gapChar	The character to indicate gap. The numbering will skip the gapChar for the reference sequence.
	Further arguments passed to or from other methods.
minSkipSize	The minimum number of tips to have gap or ambiguous amino acid/nucleotide for a site to be ignored in other analysis. This will not affect the numbering. The default is 0.8.

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#### Value

The input x with numbering mapped to reference.

# Examples

```
data(zikv_tree)
msaPath <- system.file('extdata', 'ZIKV.fasta', package = 'sitePath')
tree <- addMSA(zikv_tree, msaPath = msaPath, msaFormat = 'fasta')
setSiteNumbering(tree)</pre>
```

similarityMatrix

Similarity between sequences

#### **Description**

Get similarity between aligned sequences with gap ignored.

#### Usage

```
similarityMatrix(tree)
```

#### **Arguments**

tree

The return from addMSA function.

#### Value

A diagonal matrix of similarity between sequences.

```
data(zikv_tree)
data(zikv_align)
tree <- addMSA(zikv_tree, alignment = zikv_align)
simMatrix <- similarityMatrix(tree)</pre>
```

sitePath-deprecated 27

sitePath-deprecated

Deprecated functions in package 'sitePath'

#### **Description**

These functions are provided for compatibility with older versions of 'sitePath' only, and will be defunct at the next release.

#### **Details**

The following functions are deprecated and will be made defunct; use the replacement indicated below:

• multiFixationSites: fixationSites

sitesMinEntropy

Fixation sites prediction

#### **Description**

After finding the lineagePath of a phylogenetic tree, sitesMinEntropy perform entropy minimization on every site of the sequence to group the tips according to amino acid/nucleotide.

#### Usage

```
sitesMinEntropy(x, ...)
## S3 method for class 'lineagePath'
sitesMinEntropy(
    x,
    minEffectiveSize = NULL,
    searchDepth = 1,
    method = c("compare", "insert", "delete"),
    ...
)
```

#### **Arguments**

x A lineagePath object returned from lineagePath function.

... further arguments passed to or from other methods.

minEffectiveSize

The minimum number of tips in a group.

searchDepth The function uses heuristic search but t

The function uses heuristic search but the termination of the search cannot be intrinsically decided. searchDepth is needed to tell the search when to stop.

method The strategy for predicting the fixation. The basic approach is entropy minimiza-

tion and can be achieved by adding or removing fixation point, or by comparing

the two.

28 SNPsites

#### Value

A sitesMinEntropy object.

# **Examples**

```
data(zikv_tree_reduced)
data(zikv_align_reduced)
tree <- addMSA(zikv_tree_reduced, alignment = zikv_align_reduced)
sitesMinEntropy(lineagePath(tree))</pre>
```

**SNPsites** 

Finding sites with variation

#### **Description**

Single nucleotide polymorphism (SNP) in the whole package refers to variation of amino acid. SNPsite will try to find SNP in the multiple sequence alignment. A reference sequence and gap character may be specified to number the site.

#### Usage

```
SNPsites(tree, ...)
## S3 method for class 'phyMSAmatched'
SNPsites(tree, minSNP = NULL, ...)
```

#### **Arguments**

tree A phyMSAmatched object.

... Other arguments

minSNP Minimum number of a mutation to be a SNP. The default is 10th of the total tree

tips.

#### Value

A SNPsites object.

```
data(zikv_tree_reduced)
data(zikv_align_reduced)
tree <- addMSA(zikv_tree_reduced, alignment = zikv_align_reduced)
SNPsites(tree)</pre>
```

zikv\_align 29

zikv\_align

Multiple sequence alignment of Zika virus polyprotein

## **Description**

The raw protein sequences were downloaded from ViPR database (https://www.viprbrc.org/) and aligned using MAFFT (https://mafft.cbrc.jp/alignment/software/). with default settings.

zikv\_align\_reduced is a truncated version of zikv\_align

#### Usage

```
data(zikv_align)
data(zikv_align_reduced)
```

#### **Format**

an alignment object an alignment object

zikv\_tree

Phylogenetic tree of Zika virus polyprotein

#### **Description**

Tree was built from zikv\_align using RAxML (http://www.exelixis-lab.org/) with default settings. The tip ANK57896 was used as outgroup to root the tree.

zikv\_tree\_reduced is a truncated version of zikv\_tree

#### Usage

```
data(zikv_tree)
data(zikv_tree_reduced)
```

#### **Format**

```
a phylo object
a phylo object
```

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