

# Package ‘vhica’

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**Type** Package

**Title** Vertical and Horizontal Inheritance Consistence Analysis

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**Suggests** ape, plotrix, parallel, seqinr, gtools

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**Description** The “Vertical and Horizontal Inheritance Consistence Analysis” method is described in the following publication: “VHICA: a new method to discriminate between vertical and horizontal transposon transfer: application to the mariner family within *Drosophila*” by G. Wallau. et al. (2016) <[DOI:10.1093/molbev/msv341](https://doi.org/10.1093/molbev/msv341)>. The purpose of the method is to detect horizontal transfers of transposable elements, by contrasting the divergence of transposable element sequences with that of regular genes.

**License** GPL-2

**NeedsCompilation** no

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## Description

The package implements the VHICA method described in Wallau et al. (in prep). The purpose of the method is to detect horizontal transfers of transposable elements, by contrasting the divergence of transposable element sequences with that of regular genes. Two files should be provided, for both a set of reference genes and transposable element sequences: (i) pairwise divergence across species (e.g., dS), (ii) codon usage bias for all genes and elements in all species.

## Details

Package: vhica  
Type: Package  
License: GPL-v2

This package contains three main functions.

- `read.vhica`: reads sequence files and generates an object of class `vhica` that will be used for further analysis.
- `plot.vhica`: plots the VHICA regression between two species, and displays how far transposable elements (or any kind of other sequences) are from the reference genes.
- `image.vhica`: plots the consistency of a specific element across all species, which makes it possible to build evolutionary scenarios.

In addition, it provides tools to calculate divergence (`div`) and codon usage bias (`CUB`), which are necessary to apply the VHICA method.

## Author(s)

Implementation: Arnaud Le Rouzic <arnaud.le-rouzic@universite-paris-saclay.fr>

Scientists who designed the method: Gabriel Wallau, Aurélie Hua-Van, Arnaud Le-Rouzic.

Maintainer: Arnaud Le Rouzic <arnaud.le-rouzic@universite-paris-saclay.fr>

Repository: <https://github.com/lerouzic/vhica>

## References

Gabriel Luz Wallau, Arnaud Le Rouzic, Pierre Capy, Elgion Loreto, Aurélie Hua-Van. VHICA: A new method to discriminate between vertical and horizontal transposon transfer: application to the mariner family within *Drosophila*. *Molecular biology and evolution* 33 (4), 1094-1109.

## Examples

```
file.cb <- system.file("extdata", "mini-cbias.txt", package="vhica")
file.div <- system.file("extdata", "mini-div.txt", package="vhica")
file.tree <- if(require("ape")) system.file("extdata", "phylo.nwk", package="vhica") else NULL
vc <- read.vhica(cb.filename=file.cb, div.filename=file.div)
plot(vc, "dere", "dana")
im <- image(vc, "mellifera:6", treefile=file.tree, skip.void=TRUE)
summary(im)
```

---

CUB

*Computes the Codon Usage Bias of DNA sequences*

---

## Description

The function reads aligned sequences in a fasta file and estimates the codon usage bias for each sequence. Several methods exist to estimate CUB; so far, only the "Effective Number of Codons" (ENC) calculation is available.

## Usage

```
CUB(file = NULL, sequence = NULL, method = "ENC")
```

## Arguments

file	FASTA file in which aligned sequences are stored.
sequence	Alternatively, the result of <code>seqinr::read.fasta</code> .
method	The method used to compute CUB. "ENC": Effective Number of Codons, as described in Wright (1990).

## Value

A named vector of CUB scores. Names correspond to sequence names in the dataset.

## Author(s)

Aurelie Hua-Van and Arnaud Le Rouzic.

## References

Wright, F. (1990). The 'effective number of codons' used in a gene. *Gene*, 87(1), 23-29.

## See Also

[div](#)

## Examples

```
seq.file <- system.file("extdata/Genes", "Amd.fas", package="vhica")
CUB(seq.file)
```

---

`div`*Computation of the synonymous divergence between sequences*

---

**Description**

The divergence between DNA sequences can be synonymous (neutral) or non-synonymous. Synonymous differences are generally considered as a better proxy for evolutionary divergence, as it is not affected by selection. This function computes the synonymous divergence between sequences.

**Usage**

```
div(file = NULL, sequence = NULL, sqs = NULL, method = "LWL85",  
pairwise = TRUE, max.lim = 3)
```

**Arguments**

<code>file</code>	FASTA file in which aligned sequences are stored.
<code>sequence</code>	Alternatively, the result of <code>seqinr::read.fasta</code> .
<code>sqs</code>	Vector of sequence names to be compared. If not provided, all pairwise comparisons will be performed.
<code>method</code>	Method used to compute the divergence. So far, only the LWL85 method (from Li et al. 1985).
<code>pairwise</code>	Boolean: should the divergence be calculated for each pair of sequences or on the whole dataset? This is of particular importance when indels (gaps) are present in sequences, as codons with gaps are generally discarded by most methods. Setting this option to TRUE is thus more likely to give accurate results with multiple-gap sequences, but the calculation will also be slower.
<code>max.lim</code>	Maximum value for divergence. Depending on the algorithm, various corrections can bring the divergence value above 100%. Values larger than <code>max.lim</code> will be replaced by NAs, as they can be problematic for further statistical tests.

**Details**

The LWL85 method is a wrapper around the `kaks` function from the `seqinr` package.

**Value**

A 3-column data frame with the following fields:

- `div`: The divergence score
- `sq1`: The first sequence in the comparison
- `sq2`: The second sequence in the comparison

**Author(s)**

Aurelie Hua-Van and Arnaud Le Rouzic

## References

Li, W. H., Wu, C. I., & Luo, C. C. (1985). A new method for estimating synonymous and non-synonymous rates of nucleotide substitution considering the relative likelihood of nucleotide and codon changes. *Molecular biology and evolution*, 2(2), 150-174.

## See Also

[CUB](#)

## Examples

```
seq.file <- system.file("extdata/Genes", "Amd.fas", package="vhica")
div(seq.file)
```

---

image.vhica	<i>Consistency matrix for a transposable element in the VHICA analysis.</i>
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---

## Description

This function plots a composite figure summarizing the evolutionary properties of a transposable element in a group of related species. Discrepancies may indicate horizontal transfers.

## Usage

```
## S3 method for class 'vhica'
image(x, element = "", H1.test = "bilat", treefile = NULL,
      skip.void = FALSE, species = NULL, p.threshold = 0.05,
      p.adjust.method = "bonferroni", ncolors = 1024,
      main = element, threshcol=0.1, colsqueeze=1,
      species.font.family="mono", species.font.cex=1,
      max.spname.length=10, ...)
```

## Arguments

x	An object of class <code>vhica</code> , created by the function <a href="#">read.vhica</a> .
element	The name of the transposable element, as specified in the data files. If the element is not present in the data, the program halts.
H1.test	A value among "bilat", "lower", or "greater".
treefile	A Newick file containing a phylogenetic tree. Species names in the tree need to match the data. If absent, the figure will not display the phylogenetic relationship (which makes the interpretation impossible).
skip.void	Whether or not the figure should show species that do not contain the transposable element.
species	A named character vector to display pretty species names. The names of the vector are the real species names (as they will appear in the figure), the content of the vector are species codes as in the data files.

p.threshold	Threshold for the p-value (above which the color gradient increases).
p.adjust.method	As documented in <a href="#">p.adjust</a> .
ncolors	Number of colors in the gradient.
main	Main title of the figure (default: the name of the transposable element).
threshcol	Part of the color spectrum devoted to non-significant values.
colsqueeze	Values larger than 1 shrink the color gradient around the threshold.
species.font.family	Font family for the species names.
species.font.cex	Font size of the species names.
max.spname.length	Maximum length of species names. Longer labels are truncated.
...	Further arguments to the generic function <a href="#">image</a> .

### Details

The figure displays in blue TE copies that are more divergent than expected between species, and in red copies that are less divergent than expected. If several lineages of copies are present in a species, the table will be split to display both lineages. Keys for the interpretation of the pattern and the reconstruction of an evolutionary scenario are provided in the original publication.

### Value

The function returns (invisibly) a list of elements (object of class `vhicaimage`) which can be used for further analysis: `tree` contains the phylogenetic tree (object of class "phylo"), `species` is the vector of species, `stats` is a matrix of  $\log_{10}(P\text{-values})$  (positive elements are minus  $\log_{10}(P)$  corresponding to positive residuals), `codedS` is a matrix reminding the divergence rates from the data. Calling the method [summary.vhicaimage](#) on this object returns a nicely formatted data frame.

### Author(s)

Implementation: Arnaud Le Rouzic <lerouzic@legs.cnrs-gif.fr>  
 Scientists who designed the method: Gabriel Wallau, Aurélie Hua-Van, Arnaud Le-Rouzic.

### References

Gabriel Luz Wallau, Arnaud Le Rouzic, Pierre Capy, Elgion Loreto, Aurélie Hua-Van. VHICA: A new method to discriminate between vertical and horizontal transposon transfer: application to the mariner family within *Drosophila*. *Molecular biology and evolution* 33 (4), 1094-1109.

### See Also

[read.vhica](#), [plot.vhica](#), [summary.vhicaimage](#).

**Examples**

```

file.cb <- system.file("extdata", "mini-cbias.txt", package="vhica")
file.div <- system.file("extdata", "mini-div.txt", package="vhica")
file.tree <- if(require("ape")) system.file("extdata", "phylo.nwk", package="vhica") else NULL
vc <- read.vhica(cb.filename=file.cb, div.filename=file.div)
plot(vc, "dere", "dana")
im <- image(vc, "mellifera:6", treefile=file.tree, skip.void=TRUE)
summary(im)

```

---

plot.vhica

*Plots a VHICA regression between two species.*


---

**Description**

The VHICA method is based on a contrast between gene divergence and codon usage bias. A regression between divergence and codon usage provides a reference, and sequences of interest (typically, transposable elements) will be compared to the reference genes.

**Usage**

```

## S3 method for class 'vhica'
plot(x, sp1 = NULL, sp2 = NULL, ...)

```

**Arguments**

x	An object of class <code>vhica</code> , created by <a href="#">read.vhica</a> .
sp1	Name of the first species, as in the data files.
sp2	Name of the second species, as in the data files.
...	Additional options for <a href="#">plot</a> .

**Details**

The resulting figure displays genes as circles, and transposable elements as symbols.

**Author(s)**

Implementation: Arnaud Le Rouzic <lerouzic@legs.cnrs-gif.fr>  
 Scientists who designed the method: Gabriel Wallau, Aurélie Hua-Van, Arnaud Le-Rouzic.

**References**

Gabriel Luz Wallau, Arnaud Le Rouzic, Pierre Capy, Elgion Loreto, Aurélie Hua-Van. VHICA: A new method to discriminate between vertical and horizontal transposon transfer: application to the mariner family within *Drosophila*. *Molecular biology and evolution* 33 (4), 1094-1109.

**See Also**

[read.vhica](#), [image.vhica](#)

## Examples

```
file.cb <- system.file("extdata", "mini-cbias.txt", package="vhica")
file.div <- system.file("extdata", "mini-div.txt", package="vhica")
file.tree <- if(require("ape")) system.file("extdata", "phylo.nwk", package="vhica") else NULL
vc <- read.vhica(cb.filename=file.cb, div.filename=file.div)
plot(vc, "dere", "dana")
image(vc, "mellifera:6", treefile=file.tree, skip.void=TRUE)
```

---

read.vhica

*Reads divergence and codon usage data files for the VHICA method.*

---

## Description

The VHICA method relies on two sources of information: (i) the divergence between sequences, and (ii) the codon usage bias. This function reads two data files and creates an object of class `vhica` that can be further explored by `plot.vhica` and `image.vhica`. Input can be either (1) two vectors of fasta file names (one for the genes, one for the putatively transferred genes), or (2) already processed files containing codon usage bias and divergence data (see Details).

## Usage

```
read.vhica(gene.fasta=NULL, target.fasta=NULL,
cb.filename=NULL, div.filename=NULL,
reference = "Gene", divergence = "dS",
CUB.method="ENC", div.method="LWL85", div.pairwise=TRUE,
div.max.lim=3, species.sep="_", gene.sep=".", family.sep=".", ...)
```

## Arguments

<code>gene.fasta</code>	Sequence files (FASTA format) containing the aligned sequences (respecting the translation phase) for all species of the reference genes.
<code>target.fasta</code>	Sequence files (FASTA format) containing the aligned sequence of the putatively transferred genes.
<code>cb.filename</code>	File name for the codon usage bias data. If FASTA files are provided, this file will be created.
<code>div.filename</code>	File name for the divergence data. If FASTA files are provided, this file will be created.
<code>reference</code>	Name of the reference type in the codon usage file. Default is "Gene".
<code>divergence</code>	Name of the divergence column in the divergence file. Default is "dS".
<code>CUB.method</code>	Method to be used for Codon Usage Bias calculation (see <code>CUB</code> ).
<code>div.method</code>	Method to be used for divergence calculation (see <code>div</code> ).
<code>div.pairwise</code>	Whether divergence should be calculated from the whole alignment of between pairs of sequences (see <code>div</code> ).
<code>div.max.lim</code>	Maximum divergence score. Estimated divergence much larger than 100% are likely to be problematic and should not be considered.



species.sep	Separator for species (or equivalent) labels in sequence names. Any character string following this separator will be disregarded – be careful about potential duplicates.
gene.sep	Separator for gene names from gene sequence files.
family.sep	Separator for target sequence sub-families.
...	Further parameters for the internal function .reference.regression.

## Details

Details about CUB and divergence calculations can be found in [CUB](#) and [div](#). If CUB and/or divergence need to be calculated by an external program, it is possible to provide them in the following format:

- Codon usage bias Example of data file:

	Type	sp1	sp2	sp3
CG4231	Gene	42.3	51.1	47.2
CG2214	Gene	47.2	44.9	53.2
Pelem1	TE	36.2	47.0	44.4
...				

- Row names (or first column)sequence index
- Type whether the sequence is a reference (default: Gene) or a focal sequence (transposable element, ...)
- Following columns a measurement of codon bias (ENC, CBI...) for every species

- Divergence Example of data file:

seq	dS	sp1	sp2
CG4231	0.84	Dmel	Dsim
CG4231	0.46	Dmel	Dana
CG4231	0.58	Dsim	Dana
CG2214	0.10	Dmel	Dsim
...			

- First column (or row names): sequence index
- Second column: divergence measurement
- Columns 3 and 4: the pair of species on which the divergence is calculated
- Row names and Col names are allowed but disregarded

## Value

The function returns an object of class `vhica`, a list containing:

- `cbias`: A codon bias array
- `div`: The divergence matrix
- `reg`: The result of all pairwise regressions
- `reference`: The reference option
- `target`: The sequence type that is not the reference
- `divergence`: The divergence option
- `family.sep`: The character used to indicate TE sub-families

**Author(s)**

Implementation: Arnaud Le Rouzic  
 Scientists who designed the method: Gabriel Wallau, Aurelie Hua-Van, Arnaud Le Rouzic.

**References**

Gabriel Luz Wallau, Arnaud Le Rouzic, Pierre Cappy, Elgion Loreto, Aurelie Hua-Van. VHICA: A new method to discriminate between vertical and horizontal transposon transfer: application to the mariner family within Drosophila. *Molecular biology and evolution* 33 (4), 1094-1109.

**See Also**

[plot.vhica](#), [image.vhica](#), [CUB](#), [div](#)

**Examples**

```
file.cb <- system.file("extdata", "mini-cbias.txt", package="vhica")
file.div <- system.file("extdata", "mini-div.txt", package="vhica")
file.tree <- if(require("ape")) system.file("extdata", "phylo.nwk", package="vhica") else NULL
vc <- read.vhica(cb.filename=file.cb, div.filename=file.div)
plot(vc, "dere", "dana")
image(vc, "mellifera:6", treefile=file.tree, skip.void=TRUE)
```

---

summary.vhicaimage	<i>Provides a data.frame that nicely displays the information returned by <a href="#">image.vhica</a>.</i>
--------------------	--

---

**Description**

The [image.vhica](#) routine displays visually the statistical support for horizontal transfers, and can return an object of class `vhicaimage`. The current summary method reorganizes this object into a dataframe that can be displayed or reused in further analysis.

**Usage**

```
## S3 method for class 'vhicaimage'
summary(object, divrate=NA, p.thresh=1, ...)
```

**Arguments**

object	An object of class <code>vhicaimage</code> , created by <a href="#">image.vhica</a> .
divrate	Optional divergence rate (in neutral substitutions per Myr).
p.thresh	Optional p-value threshold. By default, all data is returned.
...	Additional options for <a href="#">summary</a> (unused).

**Value**

The resulting data.frame has 4 or 5 columns. The two first columns are sp1 and sp2, the two species between which the horizontal transfer is tested, in an arbitrary order. The column p.value contains the p-value calculated as in [image.vhica](#) (including the possible correction for multiple testing). The dS column is a copy of the corresponding divergence from the original data. The last, optional column Time(MyA) is a molecular clock estimate of the time of divergence between the two sequences, based on the divergence rate (when provided).

**Author(s)**

Implementation: Arnaud Le Rouzic <lerouzic@legs.cnrs-gif.fr> and Gabriel Wallau  
Scientists who designed the method: Gabriel Wallau, Aurélie Hua-Van, Arnaud Le-Rouzic.

**References**

Gabriel Luz Wallau, Arnaud Le Rouzic, Pierre Capy, Elgion Loreto, Aurélie Hua-Van. VHICA: A new method to discriminate between vertical and horizontal transposon transfer: application to the mariner family within Drosophila. *Molecular biology and evolution* 33 (4), 1094-1109.

**See Also**

[read.vhica](#), [image.vhica](#)

**Examples**

```
file.cb <- system.file("extdata", "mini-cbias.txt", package="vhica")
file.div <- system.file("extdata", "mini-div.txt", package="vhica")
file.tree <- if(require("ape")) system.file("extdata", "phylo.nwk", package="vhica") else NULL
vc <- read.vhica(cb.filename=file.cb, div.filename=file.div)
plot(vc, "dere", "dana")
im <- image(vc, "mellifera:6", treefile=file.tree, skip.void=TRUE)
summary(im)
```

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