Package 'paramlink'

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Title Parametric Linkage and Other Pedigree Analysis in R

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Description A suite of tools for analysing pedigrees with marker data, including parametric linkage analysis, forensic computations, relatedness analysis and marker simulations. The core of the package is an implementation of the Elston-Stewart algorithm for pedigree likelihoods, extended to allow mutations as well as complex inbreeding. Features for linkage analysis include singlepoint LOD scores, power analysis, and multipoint analysis (the latter through a wrapper to the 'MERLIN' software). Forensic applications include exclusion probabilities, genotype distributions and conditional simulations. Data from the 'Familias' software can be imported and analysed in 'paramlink'. Finally, 'paramlink' offers many utility functions for creating, manipulating and plotting pedigrees with or without marker data (the actual plotting is done by the 'kinship2' package).

Depends R (>= 3.3)

Imports assertthat, graphics, kinship2, maxLik, stats, utils

Suggests identity, igraph

SystemRequirements For the MERLIN wrapper, MERLIN (http://csg.sph.umich.edu/abecasis/Merlin/) must be installed and pointed to in the PATH environment variable.

License GPL (>= 2)

LazyData Yes

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allGenotypes

Description

Auxiliary functions computing possible genotype combinations in a pedigree. These are not normally intended for end users.

Usage

```
allGenotypes(n)
fast.grid(argslist, as.list = FALSE)
```

geno.grid.subset(x, partialmarker, ids, chrom, make.grid = T)

Arguments

n	a positive integer.
argslist	a list of vectors.
as.list	if TRUE, the output is a list, otherwise a matrix.
x	a linkdat object.
partialmarker	a marker object compatible with x.
ids	a numeric with ID labels of one or more pedigree members.
chrom	a character, either 'X' or 'AUTOSOMAL'. If missing, the 'chrom' attribute of partialmarker is used. If this is also missing, then 'AUTOSOMAL' is taken as the default value.
make.grid	a logical. If FALSE, a list is returned, otherwise fast.grid is applied to the list before returning it.

Value

allGenotypes returns a matrix with 2 columns and n + n*n(n-1)/2 rows containing all possible (unordered) genotypes at a biallelic locus with alleles 1,2,...{},n. fast.grid is basically a stripped down version of expand.grid.

Author(s)

Magnus Dehli Vigeland

```
m = allGenotypes(2)
stopifnot(m == rbind(c(1,1), c(2,2), 1:2))
```

as.data.frame.linkdat linkdat to data.frame conversion

Description

Convert a linkdat object to data.frame for pretty printing.

Usage

```
## S3 method for class 'linkdat'
as.data.frame(x, ..., famid = F,
markers = seq_len(x$nMark), alleles = NULL, missing = NULL,
singleCol = FALSE, sep = "")
```

Arguments

х	a linkdat object.
	further arguments (not used).
famid	a logical indicating if the family identifier should be included as the first column.
markers	a numeric indicating which markers should be included/printed.
alleles	a character containing allele names, e.g. alleles=c('A', 'B').
missing	the character (of length 1) used for missing alleles. Defaults to '0'.
singleCol	a logical: Should the two alleles for each marker be pasted into one column or kept in separate columns?
sep	a single character to be used as allele separator if singleCol=TRUE.

Details

This function is mainly intended for pretty-printing linkdat objects (for instance it is called by print.linkdat). For direct manipulation of the pedigree and/or marker matrices, it is better to use as.matrix.linkdat.

Value

A data.frame.

Author(s)

Magnus Dehli Vigeland

See Also

as.matrix.linkdat

as.matrix.linkdat

Examples

```
x = linkdat(toyped)
x
# Printing x as above is equivalent to:
as.data.frame(x, sep = '/', missing = '-', singleCol = TRUE)
```

as.matrix.linkdat linkdat to matrix conversion

Description

Converts a linkdat object to a matrix (basically following a pre-makeped LINKAGE format), with marker annotations and other info attached as attributes.

Usage

```
## S3 method for class 'linkdat'
as.matrix(x, include.attrs = TRUE, ...)
```

restore_linkdat(x, attrs = NULL, checkped = TRUE)

Arguments

Х	a linkdat object. In restore_linkdat: A numerical matrix in LINKAGE format.
include.attrs	a logical indicating if marker annotations and other info should be attached as attributes. See value.
	not used.
attrs	a list containing marker annotations and other linkdat info compatible with x, in the format produced by as.matrix. If NULL, the attributes of x itself are used.
checkped	a logical, forwarded to linkdat. If FALSE, no checks for pedigree errors are performed.

Details

restore_linkdat is the reverse of as.matrix.

The way linkdat objects are created in paramlink, marker data are stored as a list of marker objects. Each of these is essentially a matrix with various attributes like allele frequencies, map info a.s.o.. This format works well for marker-by-marker operations (e.g. likelihoods and LOD scores), but makes it somewhat awkward to operate 'horizontally', i.e. individual-by-individual, for instance if one wants to delete all genotypes of a certain individual, or rearrange the pedigree in some way.

It is therefore recommended to convert the linkdat object to a matrix first, do the necessary manipulations on the matrix, and finally use restore_linkdat. Attributes are often deleted during matrix manipulation, so it may be necessary to store them in a variable and feed them manually to restore_linkdat using the attrs argument.

With default parameters, restore_linkdat(as.matrix(x)) should reproduce x exactly.

Value

For as.matrix: A matrix with xnInd rows and 6 + 2*xnMark columns. The 6 first columns describe the pedigree in LINKAGE format, and the remaining columns contain marker alleles, using the internal (numerical) allele coding and 0 for missing alleles. If include.attrs = TRUE the matrix has the following attributes:

- markerattr (a list of marker annotations)
- available (the availability vector)
- model (the disease model, if present)
- plot.labels (plot labels, if present)
- orig.ids (original individual IDs)

For restore_linkdat: A linkdat object.

Author(s)

Magnus Dehli Vigeland

See Also

linkdat, as.data.frame.linkdat

```
x = linkdat(toyped, model=1)
y = restore_linkdat(as.matrix(x))
stopifnot(all.equal(x,y))
# If attributes are lost during matrix manipulation: Use the 'attrs' argument.
xmatr = as.matrix(x)
newmatr = xmatr[-4, ] # NB: attributes are lost here
z = restore_linkdat(newmatr, attrs = attributes(xmatr))
# Should be the same as:
```

```
z2 = removeIndividuals(x, 4)
stopifnot(all.equal(z, z2))
```

dominant

Description

Medical pedigree with 23 individuals of which 15 are genotyped with 650 SNP markers. Eleven family members are affected by a disease, showing an autosomal dominant inheritance pattern.

Usage

dominant

Format

A data frame with 23 rows and 1306 columns, describing the pedigree and marker data in premakeped format. The first 6 columns contain the pedigree structure and affection status, while the final 1300 columns hold the marker alleles.

- FAMID. Family ID
- ID. Individual ID
- FID. Father ID
- MID. Mother ID
- SEX. Gender (male=1, female=2)
- AFF. Affection status (unaffected=1, affected=2, unkonwn=0)
- M1_1. First allele of marker 1
- M1_2. Second allele of marker 1
- ...
- M650_1. First allele of marker 650
- M650_2. Second allele of marker 650

All markers are SNPs, whose alleles are written as 1 and 2. Missing alleles are denoted by 0.

```
x = linkdat(dominant)
summary(x)
```

examineKinships

Description

This function provides a convenient way to check for pedigree errors in a linkage project or other situations where marker data is available for several members. The function calls IBDestimate to estimate IBD coefficients for all indicated pairs of pedigree members and produces a color-coded plot where wrong relationships are easy to spot.

Usage

```
examineKinships(x, who = "all", interfam = c("founders", "none", "all"),
makeplot = T, pch = 4, ...)
```

Arguments

х	A linkdat object, or a list of such.
who	A character vector of one or more of the words 'parents', 'siblings', 'hugs' (= halfsibs/uncles/grandparents), 'cousins' and 'unrelated'. Two additional single- word values are possible: 'all' (all of the above, plus 'other') and 'close' (= 'parents', 'siblings', 'hugs', 'cousins').
interfam	A character; either 'founders', 'none' or 'all', indicating which interfamiliar pairs of individuals should be included. Only relevant if x is a list of several linkdat objects.
makeplot	A logical.
pch	Plotting symbol (default: cross).
	Other plot arguments passed on to showInTriangle.

Value

A list of data.frames (one for each relation category) with IBD estimates.

Author(s)

Magnus Dehli Vigeland

See Also

IBDestimate, IBDtriangle, showInTriangle

exclusionPower

Examples

```
x = cousinsPed(1)
x = simpleSim(x, 500, alleles=1:2)
examineKinships(x)
# Pretend we didn't know the brothers (3 and 6) were related
x1 = branch(x, 3)
x2 = branch(x, 6)
x2$famid = 2
# Notice the error: An 'unrelated' dot close to the sibling point
examineKinships(list(x1, x2))
```

exclusionPower Power of exclusion

Description

Computes the power (of a single marker) of excluding a claimed relationship, given the true relationship.

Usage

```
exclusionPower(ped_claim, ped_true, ids, markerindex = NULL, alleles = NULL,
afreq = NULL, known_genotypes = list(), Xchrom = FALSE, plot = TRUE)
```

Arguments

ped_claim	a linkdat object, or a list of several linkdat and/or singleton objects, describing the claimed relationship. If a list, the sets of ID labels must be disjoint, that is, all ID labels must be unique.
ped_true	a linkdat object, or a list of several linkdat and/or singleton objects, describing the true relationship. ID labels must be consistent with ped_claim.
ids	individuals available for genotyping.
markerindex	NULL, or a single numeric indicating the index of a marker of ped_claim from which alleles, afreq and known_genotypes will be extracted.
alleles	a numeric or character vector containing marker alleles names. Ignored if marker index is non-NULL.
afreq	a numerical vector with allele frequencies. An error is given if they don't sum to 1 (rounded to 3 decimals). Ignored if markerindex is non-NULL.
known_genotypes	
	list of triplets (a, b, c), indicating that individual a has genotype b/c. Must be NULL if markerindex is non-NULL.

Details

This function computes the 'Power of exclusion', as defined and discussed in (Egeland et al., 2014).

Value

A single numeric value. If plot='plot_only', the function returns NULL after producing the plot.

Author(s)

Magnus Dehli Vigeland

References

T. Egeland, N. Pinto and M. D. Vigeland, *A general approach to power calculation for relationship testing*. Forensic Science International: Genetics 9 (2014): 186-190. DOI:10.1016/j.fsigen.2013.05.001

```
### A standard case paternity case:
### Compute the power of exclusion when the claimed father is in fact unrelated to the child.
claim = nuclearPed(noffs=1, sex=2)
                                 # Specifies individual 1 as the father of 3
true = list(singleton(id=1,sex=1), singleton(id=3, sex=2))
                                                     # Specifies 1 and 3 as unrelated
available = c(1, 3)  # Individuals 1 and 3 are available for genotyping
# Equifrequent autosomal SNP:
PE1 = exclusionPower(claim, true, available, alleles = 2, afreq=c(0.5,0.5))
# If the child is known to have genotype 1/1:
PE2 = exclusionPower(claim, true, available, alleles = 2, afreq=c(0.5,0.5),
                  known_genotypes=list(c(3,1,1)))
# Equifrequent SNP on the X chromosome:
PE3 = exclusionPower(claim, true, available, alleles = 2, afreq=c(0.5,0.5), Xchrom=TRUE)
stopifnot(PE1==0.125, PE2==0.25, PE3==0.25)
****
### Example from Egeland et al. (2012):
### Two females claim to be mother and daughter. Below we compute the power of various
### markers to reject this claim if they in reality are sisters.
*****
```

```
mother_daughter = nuclearPed(1, sex = 2)
sisters = relabel(nuclearPed(2, sex = c(2, 2)), c(101, 102, 2, 3))
# Equifrequent SNP:
PE1 = exclusionPower(ped_claim = mother_daughter, ped_true = sisters, ids = c(2, 3),
                     alleles = 2)
# SNP with MAF = 0.1:
PE2 = exclusionPower(ped_claim = mother_daughter, ped_true = sisters, ids = c(2, 3),
                     alleles = 2, afreq=c(0.9, 0.1))
# Equifrequent tetra-allelic marker:
PE3 = exclusionPower(ped_claim = mother_daughter, ped_true = sisters, ids = c(2, 3),
                     alleles = 4)
# Tetra-allelic marker with one major allele:
PE4 = exclusionPower(ped_claim = mother_daughter, ped_true = sisters, ids = c(2, 3),
                     alleles = 4, afreq=c(0.7, 0.1, 0.1, 0.1))
stopifnot(round(c(PE1,PE2,PE3,PE4), 5) == c(0.03125, 0.00405, 0.08203, 0.03090))
####### How does the power change if the true pedigree is inbred?
sisters_LOOP = addParents(sisters, 101, father = 201, mother = 202)
sisters_LOOP = addParents(sisters_LOOP, 102, father = 201, mother = 203)
# Equifrequent SNP:
PE5 = exclusionPower(ped_claim = mother_daughter, ped_true = sisters_LOOP,
                     ids = c(2, 3), alleles = 2)
# SNP with MAF = 0.1:
PE6 = exclusionPower(ped_claim = mother_daughter, ped_true = sisters_LOOP,
                     ids = c(2, 3), alleles = 2, afreq=c(0.9, 0.1))
stopifnot(round(c(PE5,PE6), 5) == c(0.03125, 0.00765))
## Not run:
# Equifrequent tetra-allelic marker:
PE7 = exclusionPower(ped_claim = mother_daughter, ped_true = sisters_LOOP,
                     ids = c(2, 3), alleles = 4)
# Tetra-allelic marker with one major allele:
PE8 = exclusionPower(ped_claim = mother_daughter, ped_true = sisters_LOOP,
                     ids = c(2, 3), alleles = 4, afreq=c(0.7, 0.1, 0.1, 0.1))
stopifnot(round(c(PE7,PE8), 5) == c(0.07617, 0.03457))
## End(Not run)
```

Familias2linkdat Convert 'Familias' output to linkdat objects

Description

Familias is a widely used program for computations in forensic genetics. The function documented here facilitates the use of paramlink for specialized computations which are not implemented in Familias, e.g. conditional simulations.

Usage

Familias2linkdat(familiasped, datamatrix, loci)

```
readFamiliasLoci(loci)
```

connectedComponents(ID, FID, MID)

Arguments

familiasped	A FamiliasPedigree object or a list of such.
datamatrix	A data frame with two columns per marker (one for each allele) and one row per individual.
loci	A FamiliasLocus object or a list of such.
ID	An integer vector: Individual ID.
FID	An integer vector: ID of father.
MID	An integer vector: ID of mother.

Details

The Familias program represents pedigrees and marker data in a way that differs from paramlink in several ways, mostly because of paramlink's stricter definition of a 'pedigree'. In paramlink, a pedigree must be connected, have numerical IDs, and each member must have either 0 or 2 parents present in the pedigree. None of this is required by FamiliasPedigree objects. The conversion function Familias2linkdat takes care of all of these potential differences: It converts each FamiliasPedigree into a list of connected linkdat objects, additional parents are added where needed, and non-numerical ID labels are stored in the plot.labels slot of the linkdat object(s).

Value

A linkdat object, or a list of such.

Author(s)

Magnus Dehli Vigeland, Thore Egeland

References

Windows Familias is freely available from http://familias.name.

See Also

See Familias and linkdat.

IBDestimate

Examples

```
# Example
## Not run:
library(Familias)
data(NorwegianFrequencies)
locus1 = FamiliasLocus(NorwegianFrequencies$TH01)
persons = c('mother', 'daughter', 'AF')
ped1 = FamiliasPedigree(id=persons, dadid=c(NA, 'AF', NA), momid=c(NA, 'mother', NA),
                         sex=c('female', 'female', 'male'))
datamatrix = data.frame(TH01.1=c(NA, 8, NA), TH01.2=c(NA,9.3, NA))
rownames(datamatrix) = persons
x = Familias2linkdat(ped1, datamatrix, locus1)
plotPedList(list(x), new=TRUE, frametitles=c('H1'), available='shaded', marker = 1)
# Example
library(fam2r)
data(F21)
pedigrees = F21$pedigrees
datamatrix = F21$datamatrix
loci = F21$loci
x = Familias2linkdat(pedigrees, datamatrix, loci)
plotPedList(x, new=TRUE, frametitles=c('H1', 'H2'), available='shaded')
# Give dev.width explicitly to allow for long names
plotPedList(x, new=TRUE, frametitles=c('H1', 'H2'), available='shaded',
            dev.width=17)
# Numerical labels work better
plotPedList(x, new=TRUE, id.labels='num', frametitles=c('H1', 'H2'),
            available='shaded')
## End(Not run)
```

IBDestimate Relatedness estimation

Description

Estimate the pairwise IBD coefficients $(\kappa_0, \kappa_1, \kappa_2)$ for specified pairs of pedigree members, using maximum likelihood methods. The optimization machinery is imported from the maxLik package.

Usage

```
IBDestimate(x, ids, markers = NULL, start = c(0.99, 0.001), tol = 1e-07)
```

Arguments

х	A single linkdat object or a list of linkdat and/or singleton objects.
ids	Either a vector of length 2, or a matrix with two columns, indicating the the pair(s) of individuals for which IBD estimates should be computed. If a matrix, each row corresponds to a pair. The entries can be either characters (matching the plot.labels of the linkdat object(s)) or integers (matching the orig.ids identifiers of the linkdat object(s)).
markers	A numeric indicating which marker(s) to include. If NULL (default), all markers are used.
start	Numeric of length 2, indicating the initial value of (κ_0, κ_2) in the optimisation (passed on to maxLik).
tol	A single numeric: the optimising tolerance value; passed on to maxLik).

Details

This function optimises the log-likelihood function first described in (Thompson, 1975). Optimisation is done in the (κ_0, κ_2) -plane and restricted to the probability triangle defined by $\kappa_0 \ge 0, \kappa_2 \ge 0$ $0, \kappa_0 + \kappa_2 \le 1.$

Value

A data.frame with 8 columns: ID1, ID2 (numeric IDs), Name1, Name2 (plot labels, if present), N (#markers with no missing alleles), κ_0 , κ_1 , κ_2 .

Author(s)

Magnus Dehli Vigeland

References

E. A. Thompson (2000). Statistical Inferences from Genetic Data on Pedigrees. NSF-CBMS Regional Conference Series in Probability and Statistics. Volume 6.

See Also

examineKinships, IBDtriangle, maxLik

```
# Simulate marker data for two siblings
x = nuclearPed(2)
x = setPlotLabels(x, c("Sib1", "Sib2"), c(3,4))
x = simpleSim(x, 200, 1:2) # 200 equifrequent SNPs
# Estimate the IBD coefficients for the siblings
est1 = IBDestimate(x, ids=c(3,4))
```

IBDtriangle

```
xlist = list(branch(x, 3), branch(x, 4))
est2 = IBDestimate(xlist, ids=c(3,4))
stopifnot(identical(est1, est2))
# If the pedigree has plot.labels, they can be used as IDs
est3 = IBDestimate(x, ids=c("Sib1", "Sib2"))
stopifnot(identical(est1, est3))
```

IBDtriangle IBD triangle plot

Description

The IBD triangle is typically used to visualize the pairwise relatedness of non-inbred individuals. Various annotations are available, including points marking the most common relationships, contour lines for the kinship coefficients, and shading of the unattainable region.

Usage

```
IBDtriangle(relationships = c("UN", "PO", "MZ", "S", "H,U,G", "FC", "SC",
  "DFC", "Q"), kinship.lines = numeric(), shading = "lightgray", pch = 16,
  cex_points = 1.2, cex_text = 1, axes = FALSE)
```

Arguments

A character vector indicating relationships points to be included in the plot. By default all of the following are included: UN=unrelated; PO=parent/offspring;
MZ=monozygotic twins; S=full siblings; H=half siblings; U=uncle/niece and
similar; G=grandparent/grandchild; FC=first cousins; SC=second cousins; DFC=double first cousins; Q=quadruple first half cousins.
A numeric vector. (See Details.)
The shading color for the unattainable region.
Symbol used for the relationship points (see par).
A single numeric controlling the symbol size for the relationship points.
A single numeric controlling the font size for the relationship labels.
Draw surrounding axis box?

Details

For any pair of non-inbred individuals A and B, their genetic relationship can be summarized by the IBD coefficients ($\kappa_0, \kappa_1, \kappa_2$), where

 $\kappa_i = P(AandBshare ialleles IBD atrandom autosom allocus).$

Since $\kappa_0 + \kappa_1 + \kappa_2 = 1$, any relationship corresponds to a point in the triangle in the (κ_0, κ_2) -plane defined by $\kappa_0 \ge 0, \kappa_2 \ge 0, \kappa_0 + \kappa_2 \le 1$. The choice of κ_0 and κ_2 as the axis variables is done for reasons of symmetry and is not significant (other authors have used different views of the triangle).

As shown in (Thompson, 1976) points in the subset of the triangle defined by $4\kappa_0\kappa_2 > \kappa_1^2$ is unattainable for pairwise relationships. By default this region in shaded in a 'lightgray' color. The IBD coefficients are linearly related to the kinship coefficient ϕ by the formula

$$\phi = 0.25\kappa_1 + 0.5\kappa_2$$

By indicating values for ϕ in the kinship.lines argument, the corresponding contour lines are shown as dashed lines in the triangle plot.

Author(s)

Magnus Dehli Vigeland

References

E. A. Thompson (1975). The estimation of pairwise relationships. Annals of Human Genetics 39.

E. A. Thompson (1976). A restriction on the space of genetic relationships. Annals of Human Genetics 40.

See Also

examineKinships

Examples

IBDtriangle()
IBDtriangle(kinship=c(0.25, 0.125), shading=NULL, cex_text=0.8)

is.linkdat

Is an object a linkdat object?

Description

Functions for checking whether an object is a linkdat object, a singleton or a list of such.

Usage

is.linkdat(x)

is.singleton(x)

is.linkdat.list(x)

Arguments

x Any R object.

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likelihood

Details

Note that the singleton class inherits from linkdat, so if x is a singleton, is.linkdat(x) returns TRUE.

Value

For is.linkdat: TRUE if x is a linkdat (or singleton) object, and FALSE otherwise. For is.singleton: TRUE if x is a singleton object, and FALSE otherwise. For is.linkdat.list: TRUE if x is a list of linkdat/singleton objects.

Author(s)

Magnus Dehli Vigeland

See Also

linkdat

Examples

likelihood

likelihood(x, ...)

Pedigree likelihood

Description

Calculates various forms of pedigree likelihoods.

Usage

```
## S3 method for class 'linkdat'
likelihood(x, locus1, locus2 = NULL, theta = NULL,
  startdata = NULL, eliminate = 0, logbase = NULL, loop_breakers = NULL,
  ...)
## S3 method for class 'singleton'
likelihood(x, locus1, logbase = NULL, ...)
## S3 method for class 'list'
```

```
likelihood(x, locus1, locus2 = NULL, ..., returnprod = TRUE)
```

```
likelihood_LINKAGE(x, marker, theta = NULL, afreq = NULL, logbase = NULL,
TR.MATR = NULL, initialCalc = NULL, singleNum.geno = NULL,
loop_breakers = NULL)
```

Arguments

x	a linkdat object, a singleton object, or a list of such objects. In likelihood_LINKAGE, x must be a linkdat object, with x\$model different from NULL.
	further arguments.
locus1	a marker object compatible with x. If x is a list, then locus1 must be a list of corresponding marker objects.
locus2	either NULL, the character 'disease', or a marker object compatible with x. See Details.
theta	the recombination rate between locus1 and locus2 (in likelihood_LINKAGE: between the marker and the disease locus). To make biological sense theta should be between 0 and 0.5.
startdata	for internal use.
eliminate	mostly for internal use: a non-negative integer indicating the number of itera- tions in the internal genotype-compatibility algorithm. Positive values can save time if partialmarker is non-empty and the number of alleles is large.
logbase	a numeric, or NULL. If numeric the log-likelihood is returned, with logbase as basis for the logarithm.
loop_breakers	a numeric containing IDs of individuals to be used as loop breakers. If NULL, automatic selection of loop breakers will be performed. See breakLoops.
returnprod	a logical; if TRUE, the product of the likelihoods is returned, otherwise a vector with the likelihoods for each pedigree in the list.
marker	an integer between 0 and x\$nMark, indicating which marker to use in the calculation.
afreq	a numeric containing the marker allele frequencies.
TR.MATR, initia	lCalc, singleNum.geno
	for internal use, speeding up linkage computations with few-allelic markers.

Details

All likelihoods are calculated using the Elston-Stewart algorithm.

If locus2 = NULL, the result is simply the likelihood of the genotypes observed at the marker in locus1.

If locus2 = 'disease', the result is the likelihood of the marker genotypes in locus1, given the affection statuses of the pedigree members, the disease model and the recombination rate theta between the marker and disease loci. The main use of this is for computation of LOD scores in parametric linkage analysis.

If locus2 is a marker object, the result is the likelihood of the genotypes at the two markers, given the recombination rate theta between them.

linkage.power

Value

The likelihood of the data. If the parameter logbase is a positive number, the output is log(likelihood, logbase).

Author(s)

Magnus Dehli Vigeland

See Also

lod

Examples

linkage.power Power of a linkage study

Description

Power analysis of parametric linkage studies

Usage

```
linkage.power(x, N = 100, available = x$available, afreq = c(0.5, 0.5),
loop_breakers = NULL, threshold = NULL, seed = NULL, verbose = FALSE)
## S3 method for class 'powres'
summary(object, threshold = NULL, ...)
```

Arguments

х	a linkdat object with a valid model. (See setModel.)
Ν	an integer; the number of markers to simulate.
available	a vector containing IDs of the available individuals, i.e. those whose genotypes should be simulated.
afreq	a numerical vector with sum 1; the population frequencies for the marker alleles.
loop_breakers	a numeric containing IDs of individuals to be used as loop breakers. Relevant only if the pedigree has loops. See breakLoops.
threshold	NULL, or a single numeric. If numeric, the output includes the percentage of simulated markers having LOD larger than threshold.
seed	NULL, or a numeric seed for the random number generator.
verbose	a logical passed on to linkageSim. If TRUE, some details are shown during the marker simulation.
object	a powres object, normally produced by linkage.power.
	not used.

Value

The function prints a summary and returns invisibly a powres object, which is a list with the following entries:

sim	A linkdat object with the simulated markers
lod	The LOD scores (computed with recombination fraction theta=0) of the simulated markers
maxlod	The highest LOD score of the simulated markers
elod	The average LOD score for the simulated markers

the maximum LOD score for each element of values.

Author(s)

Magnus Dehli Vigeland

References

The marker simulation takes place in linkageSim, which implements the algorithm used by SLINK: http://linkage.rockefeller.edu/ott/SLINK.htm

See Also

linkdat,linkageSim

linkageSim

Examples

```
# Note: In the examples below N is set very low in order to reduce time consumption.
# Increase N to get more interesting results.
x = nuclearPed(3)
x = swapAff(x, c(1,3,4))
x = setModel(x, 1) # Autosomal dominant
linkage.power(x, N=1)
# X-linked recessive example:
y = linkdat(Xped, model=4)
linkage.power(y, N=1)
# Power of homozygosity mapping:
z = addOffspring(cousinPed(1), father=7, mother=8, noffs=1, aff=2)
z = setModel(z, 2) # Autosomal recessive model
pow = linkage.power(z, N=1, loop_breaker=7, seed=123)
stopifnot(round(pow$maxlod, 1) == 1.2)
```

linkageSim

Simulate markers linked to a disease locus.

Description

Simulates markers (with up to 4 alleles) conditional on the pedigree structure, affection statuses and disease model.

Usage

```
linkageSim(x, N = 1, available = x$available, afreq = NULL,
partialmarker = NULL, loop_breakers = NULL, unique = FALSE,
seed = NULL, verbose = TRUE)
```

Arguments

х	a linkdat object
Ν	a positive integer: the number of markers to be simulated
available	a vector containing IDs of the available individuals, i.e. those whose genotypes should be simulated.
afreq	a vector of length < 5 containing the population frequencies for the marker alleles.
partialmarker	Either NULL (indicating no given marker data), or a marker object.
loop_breakers	a numeric containing IDs of individuals to be used as loop breakers. Relevant only if the pedigree has loops. See breakLoops.

linkdat

unique	a logical indicating if duplicates among the simulated markers should be re- moved.
seed	NULL, or a numeric seed for the random number generator.
verbose	a logical.

Details

All markers are simulated under the condition that the recombination fraction between the marker and the disease locus is 0. This is an implementation of the algorithm used in SLINK of the LINK-AGE/FASTLINK suite.

Value

a linkdat object equal to x except its markerdata entry, which consists of the N simulated markers.

Author(s)

Magnus Dehli Vigeland

References

G. M. Lathrop, J.-M. Lalouel, C. Julier, and J. Ott (1984). *Strategies for Multilocus Analysis in Humans*, PNAS 81, pp. 3443-3446.

See Also

linkage.power

Examples

```
x = linkdat(toyped, model=1)
y = linkageSim(x, N=10, afreq=c(0.5, 0.5))
stopifnot(length(mendelianCheck(y))==0)
z = addOffspring(cousinPed(1), father=7, mother=8, noffs=1, aff=2)
z = setModel(z, 2)
linkageSim(z, N=1, afreq = c(0.1, 0.2, 0.7))
```

linkdat

Linkdat objects

Description

Functions to create and display 'linkdat' objects.

linkdat

Usage

```
linkdat(ped, model = NULL, map = NULL, dat = NULL, freq = NULL,
annotations = NULL, missing = 0, header = FALSE, checkped = TRUE,
verbose = TRUE, ...)
singleton(id, sex = 1, famid = 1, verbose = FALSE, ...)
## S3 method for class 'linkdat'
print(x, ..., markers)
## S3 method for class 'linkdat'
summary(object, ...)
write.linkdat(x, prefix = "", what = c("ped", "map", "dat", "freq",
    "model"), merlin = FALSE)
## S3 method for class 'linkdat'
subset(x, subset = x$orig.ids, ...,
    markers = seq_len(x$nMark))
```

Arguments

a matrix, data.frame or a character with the path to a pedigree file in standard LINKAGE format. (See details)
either a linkdat.model object (typically y\$model for some linkdat object y), or a single integer with the following meaning: $1 =$ autosomal dominant; $2 =$ auto- somal recessive; $3 =$ X-linked dominant; $4 =$ X-linked recessive. In each of these cases, the disease is assumed fully penetrant and the disease allele frequency is set to 0.00001. If model=NULL, no model is set.
a character with the path to a map file in MERLIN format, or NULL. If non-NULL, a dat file must also be given (next item).
a character with the path to a dat file in MERLIN format, or NULL. (Only needed if map is non-NULL.) $% \left({\left({{{\rm{NULL}}} \right)_{\rm{star}}} \right)$
a character with the path to a allele frequency file in MERLIN (short) format, or NULL. If NULL, all markers are interpreted as equifrequent.
a list (of the same length and in the same order as the marker columns in x) of marker annotations. If this is non-NULL, then all of map, dat, freq should be NULL.
the character (of length 1) used for missing alleles. Defaults to '0'.
a logical, relevant only if ped points to a ped file: If TRUE, the first line of the ped file is skipped.
a logical. If FALSE, no checks for pedigree errors are performed.
a logical: verbose output or not.
further arguments.
single numerics describing the individual ID and gender of the singleton.

famid	a numeric: the family ID of the singleton.
x, object	a linkdat object.
markers	a numeric indicating which markers should be included/printed.
prefix	a character string giving the prefix of the files. For instance, if $prefix='fam1'$ and what=c('ped', 'map'), the files 'fam1.ped' and 'fam1.map' will be created.
what	a character vector forming a subset of c('ped', 'map', 'dat', 'freq', 'model'), indicating which files should be created. All files are written in MERLIN style (but see the next item!)
merlin	a logical. If TRUE, the marker alleles are relabeled to 1,2,, making sure that the generated files are readable by MERLIN (which does not accept non- numerical allele labels in the frequency file.) If FALSE (the default) the allele labels are unchanged. In this case, x should be exactly reproducible from the files. (See examples.)
subset	a numeric containing the individuals in the sub-pedigree to be extracted. NB: No pedigree checking is done here, so make sure the subset form a meaningful, closed pedigree.

Details

The file (or matrix or data.frame) ped must describe one or several pedigrees in standard LINKAGE format, i.e. with the following columns in correct order:

1 Family id (optional) (FAMID)

2 Individual id (ID),

3 Father id (FID),

4 Mother id (MID),

5 Gender (SEX): 1 = male, 2 = female,

6 Affection status (AFF): 1 = unaffected, 2 = affected, 0 = unknown,

7 First allele of first marker,

8 Second allele of first marker,

9 First allele of second marker,

a.s.o.

Only columns 2-6 are mandatory. The first column is automatically interpreted as family id if it has repeated elements.

Internally the individuals are relabeled as 1,2,..., but this should rarely be of concern to the end user. Some pedigree checking is done, but it is recommended to plot the pedigree before doing any analysis.

Details on the formats of map, dat and frequency files can be found in the online MERLIN tutorial: http://csg.sph.umich.edu/abecasis/Merlin/

A singleton is a special linkdat object whose pedigree contains 1 individual. The class attribute of a singleton is c('singleton', 'linkdat')

linkdat

Value

A linkdat object, or a list of linkdat objects. A linkdat object is essentially a list with the following entries, some of which can be NULL.

pedigree	data.frame with 5 columns (ID, FID, MID, SEX, AFF) describing the pedigree in linkage format. (NB: Internal labeling used.)
orig.ids	the original individual id labels.
nInd	the number of individuals in the pedigree.
founders	vector of the founder individuals. (NB: Internal labeling used.)
nonfounders	vector of the nonfounder individuals (NB: Internal labeling used.)
hasLoops	a logical: TRUE if the pedigree is inbred.
subnucs	list containing all (maximal) nuclear families in the pedigree. Each nuclear fam- ily is given as a vector of the form c(pivot, father, mother, child1,), where the pivot is either the id of the individual linking the nuclear family to the rest of the pedigree, or 0 if there are none. (NB: Internal labeling used.)
markerdata	a list of marker objects.
nMark	the number of markers.
available	a numeric vector containing IDs of available individuals. Used for simulations and plots.
model	a linkdat.model object, essentially a list containing the model parameters. See <pre>setModel</pre> for details.
loop_breakers	a matrix with original loop breaker ID's in the first column and their duplicates in the second column. This is set by breakLoops.

Author(s)

Magnus Dehli Vigeland

See Also

pedCreate, pedModify, pedParts, setModel

```
x = linkdat(toyped, model=1)
x
summary(x)
##### test read/write:
x = modifyMarker(x, 1, alleles=c('B','C'), afreq=c(.9, .1), chrom=2, name='SNP1', pos=123)
write.linkdat(x, prefix='toy')
y = linkdat('toy.ped', map='toy.map', dat='toy.dat', freq='toy.freq', model=1)
unlink(c('toy.ped', 'toy.map', 'toy.dat', 'toy.freq', 'toy.model'))
stopifnot(isTRUE(all.equal(x,y)))
##### test singletons:
```

```
w = singleton(id=3, sex=2)
T1 = all.equal(w, linkdat(ped=rbind(c(3,0,0,2,1))))
w = markerSim(w, N=5, alleles=2, afreq=c(0.1,.9))
T2 = all.equal(w, relabel(relabel(w, 10), 3))
T3 = all.equal(w, swapSex(swapSex(w, 3), 3))
T4 = all.equal(w, swapAff(swapAff(w, 3), 3))
stopifnot(T1, T2, T3, T4)
#### several ways of creating the same linkdat object:
alleles = c(157, 160, 163)
afreq = c(0.3, 0.3, 0.4)
gt10 = c(160, 160)
gt14 = c(160, 163)
z1 = relabel(addOffspring(nuclearPed(1), father=3, noffs=1, aff=2), 10:14)
z1 = addMarker(z1, marker(z1, 10, gt10, 14, gt14, alleles=alleles, afreq=afreq))
z1 = setModel(z1, 2)
z2 = addParents(relabel(nuclearPed(1), 12:14), 12, father=10, mother=11)
z2 = addMarker(z2, rbind(gt10, 0, 0, 0, gt14), alleles=alleles, afreq=afreq)
z2 = setModel(swapAff(z2, 14), 2)
z3 = linkdat(data.frame(ID=10:14, FID=c(0,0,10,0,12), MID=c(0,0,11,0,13),
             SEX=c(1,2,1,2,1), AFF=c(1,1,1,1,2),
             M=c('160/160', '0/0', '0/0', '0/0', '160/163')), model=2)
z3 = modifyMarker(z3, 1, alleles=alleles, afreq=afreq)
write.linkdat(z1, prefix='test')
z4 = linkdat('test.ped', map='test.map', dat='test.dat', freq='test.freq',
             model=2)
z4 = modifyMarker(z4, 1, alleles=alleles, chrom=NA, pos=NA, name=NA)
write.linkdat(z1, prefix='test', merlin=TRUE)
z5 = linkdat('test.ped', map='test.map', dat='test.dat', freq='test.freq',
             model=2)
z5 = modifyMarker(z5, 1, alleles=alleles, chrom=NA, pos=NA, name=NA)
stopifnot(isTRUE(all.equal(z1,z2)), isTRUE(all.equal(z1,z3)),
          isTRUE(all.equal(z1,z4)), isTRUE(all.equal(z1,z5)))
unlink(c('test.ped', 'test.map', 'test.dat', 'test.freq', 'test.model'))
```

linkres

S3 methods for class 'linkres'.

Description

Functions for printing, summarizing and plotting the results of a linkage analysis.

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linkres

Usage

```
## S3 method for class 'linkres'
print(x, ...)
## S3 method for class 'linkres'
summary(object, ...)
## S3 method for class 'linkres'
as.data.frame(x, ..., sort = TRUE)
peakSummary(x, threshold, width = 1, physmap = NULL)
## S3 method for class 'linkres'
plot(x, chrom = NULL, ylim = NULL, ...)
```

Arguments

x, object	a linkres object (normally produced by lod or merlin).
	further arguments.
sort	a logical, indicating if the data frame should be sorted according to map position.
threshold	a single numeric. A peak is defined as a regions of at least width consecutive markers LOD score above threshold.
width	a single numeric.
physmap	a matrix or data frame with three columns: Marker name, chromosome and physical position. This argument is optional.
chrom	NULL, or a numeric containing chromosome numbers. In the latter case only results for the markers on the indicated chromosomes will be plotted.
ylim	NULL, or a numeric of length 2: to be passed on to plot.default.

Author(s)

Magnus Dehli Vigeland

See Also

lod, merlin

```
x = linkdat(toyped, model=1)
lods = lod(x, theta='max')
summary(lods)
as.data.frame(lods)
```

Description

Calculates the two-point LOD scores of a pedigree for the specified markers. The recombination ratio between the disease and marker loci can be either fixed at specific values, or optimized.

Usage

```
lod(x, markers = seq_len(x$nMark), theta = 0, loop_breakers = NULL,
max.only = FALSE, verbose = FALSE, tol = 0.01)
```

Arguments

Х	a linkdat object.
markers	an integer vector denoting which markers to use.
theta	either a numeric containing specific recombination ratio(s), or the word 'max', indicating that the recombination ratio should be optimized by the program.
loop_breakers	a numeric containing IDs of individuals to be used as loop breakers. Relevant only if the pedigree has loops. See breakLoops.
max.only	a logical indicating whether only the maximum LOD score should be returned.
verbose	a logical: verbose output or not.
tol	a numeric passed on to optimize as its tolerance parameter.

Details

The LOD score of a marker is defined as

$$LOD(\theta) = \log[10] \frac{L(\theta)}{L(0.5)}$$

where $L(\theta)$ denotes the likelihood of the observed marker genotypes given a recombination ratio θ between the marker and the disease locus.

Value

If max.only=TRUE, the highest computed LOD score is returned, as a single number.

Otherwise a linkres object, which is essentially a matrix containing the LOD scores. The details depend on the other parameters:

If theta is numeric, the matrix has dimensions length(theta) * length(markers), and the entry in row t, column m is the lod score of the pedigree for marker m assuming a recombination rate of t.

If theta='max', the linkres matrix has one column per marker and two rows: The first containing the LOD score and the second the optimal recombination ratio for each marker.

If a marker has incompatible values (i.e. if a child of homozygous 1/1 parents has a 2 allele), the corresponding output entries are NaN.

lod

lod.peaks

Author(s)

Magnus Dehli Vigeland

See Also

likelihood, optimize, breakLoops

Examples

```
x = linkdat(toyped, model=1)
res = lod(x)
res_theta = lod(x, theta=c(0, 0.1, 0.2, 0.5))
res_max = lod(x, theta='max')
stopifnot(all(0.3 == round(c(res, res_theta['0',], res_max['LOD',]), 1)))
# bigger pedigree with several markers
y = linkdat(dominant)
y = setModel(y, model=1, penetrances=c(.001, .9, .99))
lod(y, markers=305:310)
lod(y, markers=305:310, theta='max')
# Example with pedigree with loops:
z = linkdat(twoloops, model=2) # fully penetrant autosomal recessive model.
# add SNP for which individuals 15 and 16 are homozygous for the rare allele.
m = marker(z, 15:16, c(1,1), alleles=1:2, afreq=c(0.001, 0.999))
z = addMarker(z, m)
res1 = lod(z)
# manual specification of loop breakers gives same result
res2 = lod(z, loop_breakers=c(8,12))
# making the marker triallelic and adding some genotypes.
z = modifyMarker(z, marker=1, ids=c(7,9,11,13), genotype=3, alleles=1:3, afreq=c(0.001, 0.499, 0.5))
plot(z, 1)
res3 = lod(z)
z = modifyMarker(z, marker=1, alleles=1:4, afreq=c(0.001, 0.499, 0.25, 0.25))
res4 = lod(z)
stopifnot(all(3 == round(c(res1, res2, res3, res4), 1)))
```

lod.peaks

LOD score peaks

Description

Identify LOD score peaks

Usage

lod.peaks(x, threshold, width = 1)

Arguments

х	a linkres object
threshold	a single numeric
width	a positive integer

Details

The function first transforms x to a data frame (using as.data.frame.linkres with sort=T. A peak is defined a run of at least width consecutive markers with LOD score above or equal to threshold. If possible, one flanking marker is included on each side of the peak.

Value

A list of data frames.

Author(s)

Magnus Dehli Vigeland

See Also

linkres, lod, merlin,

Examples

```
## minimal example
x = linkdat(toyped, model=1)
res = lod(x)
peak1 = lod.peaks(res, threshold=0)
peak2 = lod.peaks(res, threshold=0, width=2)
peak3 = lod.peaks(res, threshold=1)
stopifnot(length(peak1)==1, nrow(peak1[[1]])==1, length(peak2)==0, length(peak3)==0)
```

LR

Likelihood ratios of pedigree hypotheses

Description

This function computes likelihood ratios for a given a list of pedigrees (linkdat/singletons objects), one of which is the 'reference', with genotype data from the same set of markers. Data exported from the 'Familias' software can be analysed by using Familias2linkdat prior to calling this function.

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Usage

LR(x, ref, markers)

Arguments

x	A list of pedigrees. Each pedigree is either a single linkdat/singleton object, or a list of such objects (the latter is necessary if the pedigree is disconnected).
ref	A single integer, indicating the index of the reference pedigree. This is used in the denominator of each LR.
markers	A vector of integers, indexing which markers should be included. If NULL (the default) all markers are used.

Value

A list with entries

LR	Likelihood ratios
LRperMarker	Likelihood ratios for each marker
likelihoodsPerS	System
	Likelihoods for each marker
time	user, system and elapsed time

Author(s)

Magnus Dehli Vigeland and Thore Egeland

See Also

IBDtriangle, examineKinships

```
# Simulate genotypes for 5 tetraallelic markers for a pair of full sibs
set.seed(123)
sibs = simpleSim(nuclearPed(2), N=5, alleles=1:4, available=3:4)
# Create two alternative hypotheses and transfer the simulated genotypes to them
halfsibs = addOffspring(nuclearPed(1),father=1,noffs=1,id=4)
halfsibs = transferMarkerdata(sibs, halfsibs)
unrel = list(singleton(3), singleton(4))
unrel = transferMarkerdata(sibs, unrel)
# Compute LR with 'unrelated' as reference
LR(list(sibs, halfsibs, unrel), ref=3)
## Not run:
data(adoption)
```

markers

```
x = Familias2linkdat(adoption$pedigrees, adoption$datamatrix, adoption$loci)
result = LRparamlink(x, ref=2)
result33 = LRparamlink(x, ref=2, marker=c(11,33))
## End(Not run)
```

markers

Marker functions

Description

Functions for setting and manipulating marker genotypes for 'linkdat' objects.

Usage

```
marker(x, ..., allelematrix, alleles = NULL, afreq = NULL, missing = 0,
  chrom = NA, pos = NA, name = NA, mutmat = NULL)
addMarker(x, m, ...)
setMarkers(x, m, annotations = NULL, missing = 0)
modifyMarker(x, marker, ids, genotype, alleles, afreq, chrom, name, pos)
getMarkers(x, markernames = NULL, chroms = NULL, fromPos = NULL,
  toPos = NULL)
removeMarkers(x, markers = NULL, markernames = NULL, chroms = NULL,
  fromPos = NULL, toPos = NULL)
swapGenotypes(x, ids)
modifyMarkerMatrix(x, ids, new.alleles)
```

Arguments

х	a linkdat object
	an even number of vectors, indicating individuals and their genotypes. See examples.
allelematrix	a matrix with one row per pedigree member and two columns per marker, con- taining the alleles for a single marker.
alleles	a numeric or character vector containing allele names.
afreq	a numerical vector with allele frequencies. An error is given if they don't sum to 1 (rounded to 3 decimals).
missing	a numeric - or character - of length 1, indicating the code for missing alleles.

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markers

chrom	NA or an integer (the chromosome number of the marker).
pos	NA or a non-negative real number indicating the genetic position (in cM) of the marker.
name	NA or a character (the name of the marker).
mutmat	a mutation matrix, or a list of two such matrices named 'female' and 'male'. The matrix/matrices must be square, with the allele labels as dimnames, and each row must sum to 1 (after rounding to 3 decimals).
m	a marker object or a matrix with alleles. (In setMarkers this matrix can contain data of several markers.)
annotations	a list of marker annotations.
marker, markers	
	a numeric indicating which marker(s) to use/modify.
ids	a numeric indicating individual(s) to be modified. In swapGenotypes this must have length 2.
genotype	a vector of length 1 or 2, containing the genotype to be given the ids individuals. See examples.
markernames	NULL or a character vector.
chroms	NULL or a numeric vector of chromosome numbers.
fromPos, toPos	NULL or a single numeric.
new.alleles	a numerical matrix of dimensions length(ids),2*x\$nMark. Entries refer to the internal allele numbering.

Value

The marker function returns an object of class marker: This is a numerical 2-column matrix with one row per individual, and attributes 'alleles' (a character vector with allele names), 'nalleles' (the number of alleles) and 'missing' (the input symbol for missing marker alleles), 'chrom' (chromosome number), 'name' (marker identifier), 'pos' (chromosome position in cM).

For addMarker, setMarkers, removeMarkers, modifyMarker, modifyMarkerMatrix and swapGenotypes, a linkdat object is returned, whose markerdata element has been set/modified.

For getMarkers a numeric vector containing marker numbers (i.e. their indices in x\$markerdata) for the markers whose 'name' attribute is contained in markernames, 'chrom' attribute is contained in chroms, and 'pos' attribute is between from and to. NULL arguments are skipped, so getMarkers(x) will return seq_len(x\$nMark) (i.e. all markers).

Author(s)

Magnus Dehli Vigeland

See Also

linkdat

Examples

```
x = linkdat(toyped)
x = removeMarkers(x, 1) # removing the only marker.
х
# Creating and adding a SNP marker with alleles 'a' and 'b', for which
# individual 1 is heterozygous, individuals 2 and 4 are homozygous for the
# 'b' allele, and individual 3 has a missing genotype.
m1 = marker(x, 1, c('a', 'b'), c(2,4), c('b', 'b'))
x = addMarker(x, m1)
# A rare SNP for which both children are heterozygous.
# The 'alleles' argument can be skipped, but is recommended to ensure
# correct order of the frequencies.
m2 = marker(x, 3:4, 1:2, alleles=1:2, afreq=c(0.99, 0.01))
x = addMarker(x, m2)
# Modifying the second marker:
# Making it triallelic, and adding a genotype to the father.
x = modifyMarker(x, marker=2, alleles=1:3, ids=1, genotype=2:3)
# Adding an empty SNP (all genotypes are missing):
x = addMarker(x, 0, alleles=c('A', 'B'))
# Similar shortcut for creating a marker for which all
# pedigree members are homozygous for an allele (say 'b'):
x = addMarker(x, 'b')
# Alternative: m = marker(x, 'b'); addMarker(x, m)
```

markerSim

Marker simulation

Description

Simulates marker genotypes conditional on the pedigree structure, affection statuses and disease model.

Usage

```
markerSim(x, N = 1, available = NULL, alleles = NULL, afreq = NULL,
partialmarker = NULL, loop_breakers = NULL, eliminate = 0,
seed = NULL, verbose = TRUE)
```

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markerSim

Arguments

x	a linkdat object
Ν	a positive integer: the number of markers to be simulated
available	a vector containing IDs of the available individuals, i.e. those whose genotypes should be simulated. By default, all individuals are included.
alleles	a vector containing the alleles for the marker to be simulation. If a single integer is given, this is interpreted as the number of alleles, and the actual alleles as 1:alleles. Must be NULL if partialmarker is non-NULL.
afreq	a vector of length 2 containing the population frequencies for the marker alleles. Must be NULL if partialmarker is non-NULL.
partialmarker	Either NULL (resulting in unconditional simulation), a marker object (on which the simulation should be conditioned) or the index of an existing marker of x.
loop_breakers	a numeric containing IDs of individuals to be used as loop breakers. Relevant only if the pedigree has loops, and only if partialmarker is non-NULL. See breakLoops.
eliminate	A non-negative integer, indicating the number of iterations in the internal genotype- compatibility algorithm. Positive values can save time if partialmarker is non-NULL and the number of alleles is large.
seed	NULL, or a numeric seed for the random number generator.
verbose	a logical.

Details

This implements (with various time savers) the algorithm used in SLINK of the LINKAGE/FASTLINK suite. If partialmarker is NULL, genotypes are simulated by simple gene dropping, using simpleSim.

Value

a linkdat object equal to x except its markerdata entry, which consists of the N simulated markers.

Author(s)

Magnus Dehli Vigeland

References

G. M. Lathrop, J.-M. Lalouel, C. Julier, and J. Ott, *Strategies for Multilocus Analysis in Humans*, PNAS 81(1984), pp. 3443-3446.

See Also

simpleSim,linkage.power

Examples

```
x = nuclearPed(2)
partial = marker(x, 3, 1, alleles=1:3)
markerSim(x, N=1, alleles=1:3)
markerSim(x, N=1, partialmarker=partial)
markerSim(x, N=1, partialmarker=partial)
markerSim(x, N=1, available=4, partialmarker=partial)
```

mendelianCheck Check for Mendelian errors

Description

Check marker data for Mendelian inconsistencies

Usage

```
mendelianCheck(x, remove = FALSE, verbose = !remove)
```

Arguments

remove a logical. If FALSE, the function returns the indices of markers found to rect. If TRUE, a new linkdat object is returned, where the incorrect ma have been deleted.	х	a linkdat object
	remove	a logical. If FALSE, the function returns the indices of markers found to incor- rect. If TRUE, a new linkdat object is returned, where the incorrect markers have been deleted.
verbose a logical. If TRUE, details of the markers failing the tests are shown.	verbose	a logical. If TRUE, details of the markers failing the tests are shown.

Value

A numeric containing the indices of the markers that did not pass the tests, or (if remove=TRUE) a new linkdat object where the failing markers are removed.

Author(s)

Magnus Dehli Vigeland

Examples

```
x = nuclearPed(3)
# Adding a SNP with a mendelian error:
# Individual 3 has an allele 'c' not carried by either parents
m1 = marker(x, 1, c('a','a'), 2, c('a','b'), 3, c('a','c'))
```

Another erroneous marker: The siblings carry more than 4 different alleles.

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mergePed

```
m2 = marker(x, 3, c(1,2), 4, c(3,4), 5, c(1,5))
# Another marker with incosistent genotypes among the siblings:
m3 = marker(x, 3, c(1,1), 4, c(2,2), 5, c(3,3))
# Another marker with incosistent genotypes among the siblings:
m4 = marker(x, 3, c(1,1), 4, c(2,3), 5, c(1,4))
# A correct marker (all homozygous for allele 'A')
m5 = marker(x, 1:5, 'A')
# An empty marker
m6 = marker(x)
x = setMarkers(x, list(m1,m2,m3,m4,m5,m6))
# Finding the errors
err_index = mendelianCheck(x, remove=FALSE)
stopifnot(all.equal(err_index, 1:4))
x_remove = mendelianCheck(x, remove=TRUE)
stopifnot(x_remove$nMark == 2)
```

mergePed

Merge two pedigrees

Description

This function merges two linkdat objects, joining them at the individuals with equal ID labels. This is especially useful for building 'top-heavy' pedigrees. Only linkdat objects without marker data are supported.

Usage

mergePed(x, y, quick = FALSE)

Arguments

х, у	linkdat objects
quick	a single logical. If TRUE, no pedigree checks are performed, and the individual
	ordering may be unfortunate.

Value

A linkdat object.

Author(s)

Magnus Dehli Vigeland

Examples

```
# Creating a trio where each parent have first cousin parents.
# (Alternatively, this could be built using many calls to addParents().)
x = cousinPed(1)
x = addOffspring(x, father=7, mother=8, noffs=1, id=9)
x = addOffspring(x, father=9, mother=10, noffs=1, id=11)
y = relabel(cousinPed(1), 101:108)
y = addOffspring(y, father=107, mother=108, noffs=1, sex=2, id=10)
y = addOffspring(y, father=9, mother=10, noffs=1, id=11)
# Joining x and y at the common individuals 9,10,11:
z = mergePed(x,y)
# plot all three pedigrees
par(mfrow=c(1,3))
plot(x); plot(y); plot(z)
```

merlin

MERLIN wrappers

Description

Wrappers for the MERLIN software, providing multipoint LOD scores and other computations on pedigrees with marker data. These functions require MERLIN to be installed and correctly pointed to in the PATH environment variable.

Usage

```
merlin(x, markers = seq_len(x$nMark), model = TRUE, theta = NULL,
    options = "", verbose = FALSE, generate.files = TRUE,
    cleanup = generate.files, logfile = "")
```

```
merlinUnlikely(x, remove = FALSE, verbose = !remove)
```

Arguments

Х	a linkdat object
markers	an integer vector indicating which markers to use (default: all).
model	a logical: If TRUE (and x\$model is not NULL), the file 'merlin.model' is created and '-model merlin.model' is included to the MERLIN command.
theta	a numeric with values between 0 and 0.5: The recombination value(s) for which the LOD score is computed. The values of theta are converted to centiMor- gan positions using the Haldane map function and included in the MERLIN command using theposition parameter. Works only for single markers (i.e. markers must consist of a single integer).

merlin

options	a character with additional options to the MERLIN command. See details.
verbose	a logical: Show MERLIN output and other information, or not.
generate.files	a logical. If TRUE, the files 'merlin.ped', 'merlin.dat', 'merlin.map', 'merlin.freq' and (if model=TRUE) 'merlin.model' are created in the working directory.
cleanup	a logical: Should the MERLIN files be deleted automatically?
logfile	a character. If this is given, the MERLIN screen output will be written to a file with this name.
remove	a logical. If FALSE, the function returns the indices of markers found to un- likely. If TRUE, a new linkdat object is returned, where the unlikely markers have been deleted.

Details

For these functions to work, MERLIN must be installed and the path to merlin.exe included in the PATH variable. The merlin function is first and foremost a wrapper to the parametric linkage functionality of MERLIN.

By default the following MERLIN command is run (via a call to system) after creating appropriate files in the current working directory:

The resulting multipoint LOD scores are extracted from the output and returned in R as a linkres object.

Additional command parameters can be passed on using the options argument (this is simply pasted onto the MERLIN command, so dashes must be included). For example, to obtain single-point LOD scores instead of multipoint, set options='-singlepoint'. (The singlepoint scores should agree with the results of lod(x), except in cases where some individuals have partial genotypes (see Examples).)

If model=FALSE the --model merlin.model part is removed from the MERLIN command above. This is necessary for some calculations, e.g. likelihoods (see Examples).

The merlinUnlikely function is a wrapper for MERLIN's '-error' command. The syntax is similar to that of mendelianCheck.

Value

If model=TRUE, a linkres object. Otherwise a character containing the complete MERLIN output.

For merlinUnlikely, a numeric containing the indices of the unlikely, or (if remove=TRUE) a new linkdat object where the unlikely markers are removed.

Author(s)

Magnus Dehli Vigeland

References

http://csg.sph.umich.edu/abecasis/Merlin/

Examples

```
## Not run:
x = linkdat(toyped, model=1)
x
# MERLIN treats partial genotypes (i.e. one known and one unknown allele) as missing:
lod_merlin = merlin(x)
lod_partial = lod(x)
x = modifyMarker(x, marker=1, ids=1, genotype=0)
lod_missing = lod(x)
stopifnot(lod_merlin == round(lod_missing, 4))
# Likelihood computation by MERLIN:
merlin(x, model=F, options='--lik')
## End(Not run)
```

oneMarkerDistribution Genotype probability distribution

Description

Computes the (joint) genotype probability distribution of one or several pedigree members, possibly conditional on partial marker data.

Usage

```
oneMarkerDistribution(x, ids, partialmarker, theta = NULL,
  grid.subset = NULL, loop_breakers = NULL, eliminate = 0,
  ignore.affection.status = FALSE, verbose = TRUE)
```

Arguments

х	A linkdat object.
ids	A numeric with ID labels of one or more pedigree members.
partialmarker	Either a marker object compatible with x, or the index (a single integer) of an existing marker of x.
theta	The recombination fraction between marker and disease locus. Only relevant if at least one individual is affected by disease. In that case an error is raised if theta is NULL, and if x does not have a disease model.

grid.subset	(Not relevant for most end users.) A numeric matrix describing a subset of all marker genotype combinations for the ids individuals. The matrix should have one column for each of the ids individuals, and one row for each combination: The genotypes are described in terms of the matrix $M = allGenotypes(n)$, where n is the number of alleles for the marker. If the entry in column j is the integer k, this means that the genotype of individual ids[j] is row k of M.	
loop_breakers	A numeric containing IDs of individuals to be used as loop breakers. Relevant only if the pedigree has loops. See breakLoops.	
eliminate	A non-negative integer, indicating the number of iterations in the internal genotype- compatibility algorithm. Positive values can save time if partialmarker is non-empty and the number of alleles is large.	
ignore.affection.status		
	A logical indicating if the 'AFF' column should be ignored (only relevant if some family members are marked as affected).	
verbose	A logical.	

Value

A named array (of dimension length(ids)) giving the joint marker genotype distribution for the ids individuals, conditional on 1) the marker allele frequencies given in partialmarker, 2) non-missing alleles in partialmarker, and 3) the disease model of x (if the pedigree is affected).

Author(s)

Magnus Dehli Vigeland

See Also

twoMarkerDistribution, allGenotypes

Examples

```
x = nuclearPed(2)
x_aff = swapAff(x, c(1,3))
x_aff = setModel(x_aff, model=1) # dominant model
snp = marker(x, 1, c(1,1), 2, c(1,0), alleles=1:2, afreq=c(0.1, 0.9))
res1 = oneMarkerDistribution(x, ids=3:4, partialmarker=snp)
res2 = oneMarkerDistribution(x_aff, ids=3:4, partialmarker=snp, theta=0.5)
# should give same result, since theta=0.5 implies that marker is independent of disease.
stopifnot(all.equal(res1, res2))
##### Different example for the same pedigree. A marker with 4 alleles:
m2 = marker(x, 3:4, c('C', 'D'), alleles=LETTERS[1:4])
oneMarkerDistribution(x, ids=1, partialmarker=m2)
```

```
# Same as above, but computing only the cases where individual 1 is heterozygous.
# (The numbers 5:10 refer to the 6 last rows of allGenotypes(4),
```

pedCreate

```
# which contain the heterozygous genotypes.)
oneMarkerDistribution(x, ids=1, partialmarker=m2, grid.subset=matrix(5:10, ncol=1))
#### Expanding on the previous example:
# Joint genotype probabilities of the parents, but including only the combinations
# where the father is heterozygous and the mother is homozygous:
grid = expand.grid(5:10, 1:4)
oneMarkerDistribution(x, ids=1:2, partialmarker=m2, grid.subset=grid)
#### Something else:
# The genotype distribution of an individual whose half cousin is homozygous
# for a rare allele.
y = halfCousinPed(degree=1)
snp = marker(y, 9, c('a', 'a'), alleles=c('a', 'b'), afreq=c(0.01, 0.99))
oneMarkerDistribution(y, ids=8, partialmarker=snp)
#### X-linked example:
z = linkdat(Xped, model=4) # X-linked recessive model
z2 = swapAff(z, 1:z$nInd, 1) # disease free version of the same pedigree
snpX = marker(z, c(5,15), c('A', 'A'), alleles=c('A', 'B'), chrom=23)
r1 = oneMarkerDistribution(z, ids=13, partialmarker=snpX, theta=0.5) # results: A - 0.8; B - 0.2
r2 = oneMarkerDistribution(z2, ids=13, partialmarker=snpX)
                                                                # should be same as above
r3 = oneMarkerDistribution(z, ids=13, partialmarker=snpX, theta=0) # results: A - 0.67; B - 0.33
stopifnot(all.equal(r1,r2), round(r1[1], 2)==0.8, round(r3[1], 2) == 0.67)
```

pedCreate

Create simple pedigrees

Description

These are utility functions for creating some common pedigree structures as linkdat objects.

Usage

```
nuclearPed(noffs, sex)
cousinsPed(degree, removal = 0, degree2 = NULL, child = FALSE)
halfCousinsPed(degree, removal = 0, degree2 = NULL, child = FALSE)
doubleCousins(degree1, degree2, removal1 = 0, removal2 = 0, child = FALSE)
doubleFirstCousins()
quadHalfFirstCousins()
```

fullSibMating(generations)

halfSibStack(generations)

cousinPed(degree)

halfCousinPed(degree)

Arguments

noffs	A positive integer, the number of offspring in the nuclear family.	
sex	A vector of length noffs; indicating the genders (1=male, 2=female) of the offspring. If missing, all offspring are taken to be males.	
degree, degree1	, degree2	
	Non-negative integers, indicating the degree of cousin-like relationships: 0=sib- lings, 1=first cousins; 2=second cousins, a.s.o. See Details and Examples.	
removal, removal1, removal2		
	Non-negative integers, indicating removals of cousin-like relationships. See De- tails and Examples.	
child	A logical: Should an inbred child be added to the two cousins?	
generations	A positive integer indicating the number of crossings.	

Details

All individuals are created as unaffected. Use swapAff to edit this (see Examples). Use swapSex to change gender of pedigree members.

The call cousinsPed(degree=n, removal=k) creates a pedigree with two n'th cousins, k times removed. By default, removals are added on the right side. To override this, the parameter degree2 can be used to indicate explicitly the number of generations on the right side of the pedigree. When degree2 is given removal is ignored. (Similarly for halfCousinsPed.)

The function doubleCousins creates two individuals whose fathers are cousins (degree1, removal1) as well as their mothers (degree2, removal2). For simplicity, a wrapper doubleFirstCousins is provided for the most common case, double first cousins. Finally quadHalfFirstCousins produces a pedigree with quadruple half first cousins.

fullSibMating crosses full sibs continuously for the indicated number of generations.

halfSibStack produces a breeding scheme where the two individuals in the final generation are simultaneously half siblings and half n'th cousins, where n=1,...,generations.

cousinPed and halfCousinPed (written without the 's') are depreciated functions kept for backwards compatibility. They create cousin pedigrees, but without possibility for removals, and with a different ordering than their replacements cousinsPed and halfCousinsPed.

Value

A linkdat object.

Author(s)

Magnus Dehli Vigeland

See Also

swapAff, swapSex, removeIndividuals, addOffspring, relabel

Examples

```
# A nuclear family with 2 boys and 3 girls,
# where the father and the two boys are affected.
x = nuclearPed(noffs=5, sex=c(1,1,2,2,2))
x = swapAff(x, ids=c(1,3,4))
# Half sibs:
halfCousinsPed(degree=0)
# Grand aunt:
cousinsPed(degree=0, removal=2)
# Second cousins once removed.
cousinsPed(degree=2, removal=1)
# Again second cousins once removed,
# but with the 'removal' on the left side.
cousinsPed(degree=3, degree2=2)
# A child of first cousin parents.
cousinsPed(degree=1, child=TRUE)
# Consecutive brother-sister matings.
fullSibMating(3)
# Simultaneous half siblings and half first cousins
halfSibStack(2)
# Double first cousins
doubleFirstCousins()
# Quadruple half first cousins
# Weird plotting behaviour for this pedigree.
x = quadHalfFirstCousins()
#plot(x)
```

pedigreeLoops

Pedigree loops

pedigreeLoops

Description

Functions for identifying, breaking and restoring loops in pedigrees.

Usage

```
pedigreeLoops(x)
breakLoops(x, loop_breakers = NULL, verbose = TRUE)
tieLoops(x)
findLoopBreakers(x)
findLoopBreakers2(x)
```

Arguments

x	a linkdat object.
loop_breakers	either NULL (resulting in automatic selection of loop breakers) or a numeric containing IDs of individuals to be used as loop breakers.
verbose	a logical: Verbose output or not?

Details

Most of paramlink's handling of pedigree loops is done under the hood - using the functions described here - without need for explicit action from end users. When a linkdat object x is created, an internal routine detects if the pedigree contains loops, in which case x\$hasLoops is set to TRUE. In analyses of x where loops must be broken (e.g. lod score computation or marker simulation), this is done automatically by calling breakLoops.

In some cases with complex inbreeding, it can be instructive to plot the pedigree after breaking the loops. Duplicated individuals are plotted with appropriate labels (see examples).

The function findLoopBreakers identifies a set of individuals breaking all inbreeding loops, but not marriage loops. These require more machinery for efficient detection, and paramlink does this is a seperate function, findLoopBreakers2, utilizing methods from the igraph package. Since this is rarely needed for most users, igraph is not imported when loading paramlink, only when findLoopBreakers2 is called.

In practice, breakLoops first calls findLoopBreakers and breaks at the returned individuals. If the resulting linkdat object still has loops, findLoopBreakers2 is called to break any marriage loops.

Value

For breakLoops, a linkdat object in which the indicated loop breakers are duplicated. The returned object will also have a non-null loop_breakers entry, namely a matrix with the IDs of the original loop breakers in the first column and the duplicates in the second.

For tieLoops, a linkdat object in which any duplicated individuals (as given in the x\$loop_breakers entry) are merged. For any linkdat object x, the call tieLoops(breakLoops(x)) should return x.

For pedigreeLoops, a list containing all inbreeding loops (not marriage loops) found in the pedigree. Each loop is represented as a list with elements 'top', a 'bottom' individual, 'pathA' (individuals forming a path from top to bottom) and 'pathB' (creating a different path from top to bottom, with no individuals in common with pathA). Note that the number of loops reported here counts all closed paths in the pedigree and will in general be larger than the genus of the underlying graph.

For findLoopBreakers and findLoopBreakers2, a numeric vector of individual ID's.

Author(s)

Magnus Dehli Vigeland

Examples

```
x = cousinsPed(1, child=TRUE)
# Make the child affected, and homozygous for rare allele.
x = swapAff(x, 9)
x = setMarkers(x, marker(x, 9, c(2,2), alleles=1:2, afreq=c(0.99, 0.01)))
# Compute the LOD score under a recessive model. Loops are automatically broken in lod().
x = setModel(x, 2)
LOD1 = lod(x, theta=0.1)
stopifnot(round(LOD1, 2) == 0.88)
# Or we can break the loop manually before computing the LOD:
loopfree = breakLoops(x, loop_breaker=8)
plot(loopfree)
LOD2 = lod(loopfree, theta=0.1)
stopifnot(all.equal(x, tieLoops(loopfree)))
stopifnot(all.equal(LOD1, LOD2))
# Pedigree with marriage loop: Double first cousins
if(require('igraph')) {
    y = doubleCousins(1, 1, child=TRUE)
    findLoopBreakers(y) # --> 9
    findLoopBreakers2(y) \# --> 9 and 4
    breakLoops(y) # uses both 9 and 4
}
```

pedModify

Modify the pedigree of 'linkdat' objects

Description

Functions to modify the pedigree of a 'linkdat' object.

pedModify

Usage

```
swapSex(x, ids, verbose = TRUE)
swapAff(x, ids, newval = NULL)
addOffspring(x, father, mother, noffs, ids = NULL, sex = 1, aff = 1,
verbose = TRUE)
addSon(x, parent, id = NULL, aff = 1, verbose = TRUE)
addDaughter(x, parent, id = NULL, aff = 1, verbose = TRUE)
addParents(x, id, father, mother, verbose = TRUE)
removeIndividuals(x, ids, verbose = TRUE)
branch(x, id)
trim(x, keep = c("available", "affected"), return.ids = FALSE,
verbose = TRUE)
```

relabel(x, new, old)

Arguments

х	A linkdat object
verbose	A logical: Verbose output or not.
newval	A numeric, indicating affection status values for the ids individuals: 1=unaffected, 2=affected, 0=unknown. If NULL, the affection statuses are swapped 1 <-> 2, hence the main use of the newval argument is to assign 0's.
father, mother	Integers indicating the IDs of parents. If missing, a new founder individual is created (whose ID will be 1+the largest ID already in the pedigree).
noffs	A single integer indicating the number of offspring to be created.
sex, aff	Integer vectors indicating the gender and affection statuses of the offspring to be created (recycled if less than noffs elements).
parent	Integer ID of any pedigree member, which will be the father or mother (depending on its gender) of the new child.
id, ids	Individual ID label(s). In addOffspring the (optional) ids argument is used to specify ID labels for the offspring to be created.
keep	A character, either 'available' (trimming the pedigree for unavailable members) or 'affected' (trimming for unaffected members).
return.ids	A logical. If FALSE, the trimmed pedigree is returned as a new linkdat object. If TRUE, a vector containing the IDs of 'removable' individuals is returned
new	a numeric containing new labels to replace those in old.
old	a numeric containing ID labels to be replaced by those in new. If missing, old is set to x\$orig.ids, i.e. all members in their original order.

Details

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When removing an individual, all descendants are also removed as well as founders remaining without offspring.

The branch() function extracts the pedigree subset consisting of all descendants of id, including id itself and all relevant spouses.

Value

The modified linkdat object.

Author(s)

Magnus Dehli Vigeland

See Also

linkdat, nuclearPed

Examples

```
x = linkdat(toyped)
# To see the effect of each command below, use plot(x) in between.
x = addParents(x, id=2, father=5, mother=6)
x = swapSex(x, c(1,5))
x = swapSex(x, c(2,6))
x = addOffspring(x, mother=6, noffs=2, id=c(7,10))
x = removeIndividuals(x, 3)
x = swapAff(x, c(4,10))
stopifnot(setequal(x$orig.ids, c(1,2,4,5,6,7,10,11)))
# Trimming a pedigree
x = linkdat(dominant)
x_affectedOnly = trim(x, keep='affected')
unavail = trim(x, keep='available', return.ids=TRUE)
```

stopifnot(setequal(unavail, c(5, 19:23)), setequal(nonaff, c(6:7, 12:13, 19:23)))

nonaff = trim(x, keep='affected', return.ids=TRUE)

pedParts

Description

Utility functions for 'linkdat' objects, mainly for extracting various pedigree information.

Usage

```
offspring(x, id, original.id = TRUE)
spouses(x, id, original.id = TRUE)
related.pairs(x, relation = c("parents", "siblings", "grandparents",
    "nephews_nieces", "cousins", "spouses", "unrelated"), available = F,
    interfam = c("none", "founders", "all"), ...)
unrelated(x, id, original.id = TRUE)
leaves(x)
parents(x, id, original.id = TRUE)
grandparents(x, id, degree = 2, original.id = TRUE)
siblings(x, id, half = NA, original.id = TRUE)
cousins(x, id, degree = 1, removal = 0, half = NA, original.id = TRUE)
nephews_nieces(x, id, removal = 1, half = NA, original.id = TRUE)
ancestors(x, id)
descendants(x, id, original.id = TRUE)
```

Arguments

х	a linkdat object. In related.pairs possibly a list of linkdat objects.
id	a numerical ID label.
original.id	a logical indicating whether 'id' refers to the original ID label or the internal labeling.
relation	one of the words (possibly truncated) parents, siblings, grandparents, nephews_nieces, cousins, spouses, unrelated.
available	a logical, if TRUE only pairs of available individuals are returned.

interfam	one of the words (possibly truncated) none, founders or all, specifying which interfamiliar pairs should be included as unrelated in the case where x is a list of several pedigrees. If none, only intrafamiliar pairs are considered; if founders all interfamiliar pairs of (available) founders are included; if all, all interfamil- iar (available) pairs are included.
	further parameters
degree	a non-negative integer.
half	a logical or NA. If TRUE (resp FALSE), only half (resp. full) siblings/cousins/nephews/nieces are returned. If NA, both categories are incluced.
removal	a non-negative integer

Value

For ancestors(x,id), a vector containing the ID's of all ancestors of the individual id. For descendants(x,id), a vector containing the ID's of all descendants (i.e. children, grandchildren, a.s.o.) of individual id.

The functions cousins, grandparents, nephews_nieces, offspring, parents, siblings, spouses, unrelated, each returns an integer vector containing the ID's of all pedigree members having the specified relationship with id.

For related.pairs a matrix with two columns. Each row gives of a pair of pedigree members with the specified relation. If the input is a list of multiple pedigrees, the matrix entries are characters of the form 'X-Y' where X is the family ID and Y the individual ID of the person.

For leaves, a vector of IDs containing all pedigree members without children.

Author(s)

Magnus Dehli Vigeland

Examples

```
p = cbind(ID=2:9, FID=c(0,0,2,0,4,4,0,2), MID=c(0,0,3,0,5,5,0,8),
        SEX=c(1,2,1,2,1,2,2,2), AFF=c(2,1,2,1,2,1,1,2))
x = linkdat(p)
stopifnot(setequal(spouses(x, 2), c(3,8)),
        setequal(offspring(x, 2), c(4,9)),
        setequal(descendants(x, 2), c(4,6,7,9)),
        setequal(leaves(x), c(6,7,9)))
# Creating a loop and detecting it with 'pedigreeLoops'
# (note that we get two loops, one for each inbred child):
    loopx = addOffspring(x, father=4, mother=9, noffs=2)
    lps = pedigreeLoops(loopx)
    stopifnot(lps[[1]]$top == 2, setequal(sapply(lps, '[[', 'bottom'), 10:11))
# We add genotypes for a single SNP marker and compute a LOD score under a dominant model.
    loopx = setMarkers(loopx, cbind(1,c(2,1,2,1,2,1,1,2,1,1)))
    loopx = setModel(loopx, 1)
```

```
# Loops are automatically broken in lod():
LOD1 = lod(loopx, theta=0.1)
stopifnot(round(LOD1, 3) == 1.746)
# Or we can break the loop manually before computing the LOD:
loopfree = breakLoops(loopx, loop_breaker=4)
LOD2 = lod(loopfree, theta=0.1)
stopifnot(all.equal(loopx, tieLoops(loopfree)))
stopifnot(all.equal(LOD1, LOD2))
```

plot.linkdat

Plot pedigrees with genotypes

Description

This is the main function for pedigree plotting, with many options for controlling the appearance of pedigree symbols, labels and marker genotypes. Most of the work is done by the plotting functionality in the 'kinship2' package.

Usage

```
## S3 method for class 'linkdat'
plot(x, marker = NULL, alleles = NULL, sep = "/",
   missing = "-", skip.empty.genotypes = FALSE, id.labels = NULL,
   title = NULL, available = FALSE, col = 1, deceased = numeric(0),
   starred = numeric(0), aff2 = NULL, margins = c(0.6, 1, 4.1, 1), ...)
## S3 method for class 'singleton'
plot(x, marker = NULL, alleles = NULL, sep = "/",
   missing = "-", skip.empty.genotypes = FALSE, id.labels = NULL,
   title = NULL, available = FALSE, col = 1, deceased = numeric(0),
   starred = numeric(0), aff2 = NULL, margins = c(8, 0, 0, 0), ...)
```

Arguments

Х	a linkdat object.
marker	either NULL, a vector of positive integers, a marker object, or a list of marker objects. If NULL, no genotypes are plotted. If a marker object (or a list of such), the genotypes are written below each individual in the pedigree, in the format determined by alleles, sep and missing. If a vector of integers is given, the corresponding marker objects are extracted from x\$markerdata.
alleles	a character vector with allele names.
sep	a character of length 1 separating alleles for diploid markers.
missing	the symbol (integer or character) for missing alleles.

skip.empty.genotypes		
	a logical. If TRUE, and marker is non-NULL, empty genotypes (which by default looks like '-/-') are not printed.	
id.labels	a vector with labels for each pedigree member. This defaults to x\$plot.labels is this is set (see setPlotLabels), otherwise to as.character(x\$orig.ids).	
title	the plot title. If NULL or ", no title is added to the plot.	
available	either a logical, a color name, or the word 'shaded'. If a color name is given, the available individuals (as defined by x\$available) are plotted in this color. If available=F no coloring is used, while (for backwards compatibility) available=T is equivalent to available='red'. The 'shaded' option results in diagonal shading.	
col	a vector with color indicators for the pedigree members. Recycled if necessary. By default everyone is drawn black.	
deceased	a numeric containing ID's of deceased pedigree members.	
starred	a numeric containing ID's of pedigree members that should be marked with a star in the pedigree plot.	
aff2	NULL, or a numeric with affection statuses (2=affected, 1=unaffected, 0=un-known) for a second trait.	
margins	a numeric of length 4 indicating the plot margins. For singletons only the first element (the 'bottom' margin) is used.	
	arguments passed on to plot.pedigree in the kinship2 package. In particular symbolsize and cex can be useful.	

Details

plot.linkdat is in essence a wrapper for plot.pedigree in the kinship2 package.

Author(s)

Magnus Dehli Vigeland, Guro Doerum

See Also

plot.pedigree, setPlotLabels

Examples

```
data(toyped)
x = linkdat(toyped)
plot(x, marker=1, alleles=c('a1','a2'), sep=' | ', deceased=2)
y = singleton(id=1)
m = marker(y, 1, c('A',0), alleles=c('A','B'))
plot(y, marker=m, id='indiv 1', title='Singleton', available=TRUE)
```

plotPedList

Description

This function creates a row of pedigree plots, each created by plot.linkdat. Each parameter accepted by plot.linkdat can be applied here. Some effort is made to guess a reasonable window size and margins, but in general the user must be prepared to do manual resizing of the plot window.

Usage

```
plotPedList(plot.arg.list, widths = NA, frames = T, frametitles = NULL,
fmar = NA, newdev = F, dev.height = NA, dev.width = NA, ...)
```

Arguments

plot.arg.list	A list of lists. Each element of plot.arg.list is a list, where the first element is the linkdat object to be plotted, and the remaining elements are passed on to plot.linkdat. These elements must be correctly named. See examples below.	
widths	A numeric vector of relative widths of the subplots. Recycled to length(plot.arg.list) if necessary, before passed on to layout. Note that the vector does not need to sum to 1.	
frames	Either a single logical (FALSE = no frames; TRUE = automatic framing) or a list of numeric vectors: Each vector must consist of consecutive integers, indicating subplots to be framed together. By default the framing follows the list structure of plot.arg.list.	
frametitles	A character vector of titles for each frame. If this is non-NULL, titles for indi- viduals subplots are ignored.	
fmar	A single number in the interval $[0,0.5)$ controlling the position of the frames.	
newdev	A logical, indicating if a new plot window should be opened.	
dev.height, dev.width		
	The dimensions of the new device (only relevant if newdev is TRUE). If these are NA suitable values are guessed from the pedigree sizes.	
	Further arguments passed on to each call to plot.linkdat.	

Details

See various examples in the Examples section below.

Note that for tweaking dev.height and dev.width the function dev.size is useful to determine the size of the active device.

Author(s)

Magnus Dehli Vigeland

See Also

plot.linkdat

Examples

```
# Simplest use: Just give a list of linkdat objects.
# To guess suitable plot window dimensions, use 'newdev=T'
peds = list(nuclearPed(3),cousinPed(2), singleton(12), halfCousinsPed(0))
plotPedList(peds, newdev=TRUE)
## Not run:
# Modify the relative widths (which are not guessed)
widths = c(2, 3, 1, 2)
plotPedList(peds, widths=widths)
# In most cases the guessed dimensions are not perfect.
# Resize plot window manually, and then plot again with newdev=F (default)
# plotPedList(peds, widths=widths)
## Remove frames
plotPedList(peds, widths=widths, frames=F)
# Non-default frames
frames = list(1, 2:3)
plotPedList(peds, widths=widths, frames=frames, frametitles=c('First', 'Second'))
# To give *the same* parameter to all plots, it can just be added at the end:
margins=c(2,4,2,4)
title='Same title'
id.labels=''
symbolsize=1.5 # note: doesn't work as expected for singletons
plotPedList(peds, widths=widths, frames=frames, margins=margins, title=title,
            id.labels=id.labels, symbolsize=symbolsize)
# For more control of individual plots, each plot and all its parameters
# can be specified in its own list:
x1 = nuclearPed(3)
x1$available = 3:5
m1 = marker(x1, 3, 1:2)
marg1 = c(5, 4, 5, 4)
plot1 = list(x1, marker=m1, margins=marg1, title='Plot 1', deceased=1:2)
x^2 = cousinsPed(2)
x2$available = leaves(x2)
m2 = marker(x2, leaves(x2), 'A')
marg2 = c(3, 4, 2, 4)
plot2 = list(x2, marker=m2, margins=marg2, title='Plot 2', symbolsize=1.2,
             skip.empty.genotypes=T)
x3 = singleton(12)
```

randomPed

randomPed

Random pedigree

Description

Creates a random medical pedigree with specified number of generations.

Usage

```
randomPed(gen, lambda = 2, penetrances = c(0, 1, 1), naff = "last.gen",
founder.mut = 1)
```

Arguments

gen	an integer in the interval [2,5] indicating the number of generations.
lambda	a positive numeric. For each descendant of the first generation, the number of offspring is sampled from a Poisson distribution with parameter lambda.
penetrances	a numeric of length 3, as in setModel.
naff	an integer specifying a lower bound on the number of affected individuals, or the character 'last.gen'. The latter produce a pedigree where at least one in the youngest generation is affected.
founder.mut	an integer, the number of disease alleles to be distributed among the founders.

Details

The function produces a random simple pedigree. Each founder is given at most one disease allele. At least one of the two top founders carries a disease allele.

Value

A linkdat object.

Author(s)

Magnus Dehli Vigeland

See Also

linkdat

Examples

```
plot(randomPed(3))
# gives error message: Not enough founder mutations
## Not run:
randomPed(gen=4, penetrances=c(0,0,1), naff=2, founder.mut=1)
## End(Not run)
```

readDatfile Read dat file in LINKAGE format

Description

Converts dat files in LINKAGE format to dat/map/freq files in MERLIN format

Usage

readDatfile(datfile, chrom, write_to = NULL)

Arguments

datfile	character. The path to the dat file.
chrom	integer chromosome number (needed to create the MERLIN map).
write_to	a character prefix used for naming the output files, or NULL if no files should be written.

Value

If write_to is NULL, a list of data.frames named dat, map and freq.

Author(s)

Magnus Dehli Vigeland

relatednessCoeff

Examples

No example given.

relatednessCoeff Relatedness coefficients

Description

Computes inbreeding coefficients for all pedigree members, and Jaquard's condensed identity coefficients for any pair of members. These are simple wrappers for functions in other packages or external programs.

Usage

```
inbreeding(x)
## S3 method for class 'coeff'
kinship(x, ids = NULL)
jacquard(x, ids)
jacquard2(x, ids, verbose = FALSE, cleanup = TRUE)
```

Arguments

х	a linkdat object.
ids	a integer vector of length 2.
verbose	a logical, indicating if messages from IdCoefs should be printed.
cleanup	a logical: If TRUE, the pedfile and sample file created for the IdCoefs run are deleted automatically.

Details

Both inbreeding and kinship.coeff are thin wrappers of kinship. Similarly, jacquard wraps identity::identity.coefs, which is an R implementation of the C program IdCoefs written by Mark Abney. The function identity::identity.coefs sometimes causes R to crash, hence as an alternative wrapper, jacquard2 executes an external call to the original C program IdCoefs (version 2.1.1). Of course, this requires IdCoefs to be installed on the computer (see link in the References section below) and the executable to be in a folder included in the PATH variable. The jacquard2 wrapper works by writing the necessary files to disk and calling IdCoefs via system.

For inbreeding, a numerical vector with the inbreeding coefficients, with names according to the ID labels x\$orig.ids.

For kinship.coeff, either a single numeric (if ids is a pair of pedigree members) or the whole kinship matrix, with x\$orig.ids as dimnames.

For jaquard and jaquard2, a numerical vector of length 9 (in the standard order of Jacquard's condensed identity coefficients).

Author(s)

Magnus Dehli Vigeland

References

The IdCoefs program: Abney, Mark (2009). A graphical algorithm for fast computation of identity coefficients and generalized kinship coefficients. Bioinformatics, 25, 1561-1563. http://home.uchicago.edu/~abney/abney_web/Software.html

See Also

kinship, identity.coefs

Examples

```
# Offspring of first cousins
x = cousinsPed(1, child=TRUE)
inb = inbreeding(x)
stopifnot(inb[9] == 1/16)
# if ID labels are not 1:9, care must be taken in extracting correct elements.
set.seed(1357)
y = relabel(x, sample(1:9))
child = leaves(y)
inbreeding(y)[child] #wrong
inb = inbreeding(y)[as.character(child)] #correct
inb
# the inbreeding coeff of the child equals the kinship coeff of parents
kin = kinship.coeff(y, parents(y, child))
stopifnot(inb==kin, inb==1/16)
```

relationLR

Relationship Likelihood Ratio

Description

Computes likelihood for two pedigrees and their ratio, the likelihood ratio (LR).

relationLR

Usage

```
relationLR(ped_numerator, ped_denominator, ids, alleles, afreq = NULL,
known_genotypes = list(), loop_breakers = NULL, Xchrom = FALSE,
plot = TRUE, title1 = "", title2 = "")
```

Arguments

ped_numerator	a linkdat object, or a list of several linkdat and/or singleton objects, describing the relationship corresponding to the hypothesis H1 (numerator). If a list, the sets of ID labels must be disjoint, that is, all ID labels must be unique.
ped_denominator	
	a linkdat object, or a list of several linkdat and/or singleton objects, describing the relationship corresponding to the hypothesis H2 (denominator). ID labels must be consistent with ped_claim.
ids	genotyped individuals.
alleles	a numeric or character vector containing marker alleles names
afreq	a numerical vector with allele frequencies. An error is given if they don't sum to 1 (rounded to 3 decimals).
known_genotypes	
	list of triplets (a, b, c), indicating that individual a has genotype b/c. Missing value is 0.
loop_breakers	Not yet implemented, only default value NULL currently handled
Xchrom	a logical: Is the marker on the X chromosome?
plot	either a logical or the character 'plot_only', controlling if a plot should be pro- duced. If 'plot_only', a plot is drawn, but no further computations are done.
title1	a character, title of leftmost plot.
title2	a character, title of rightmost plot.

Details

This function computes the likelihood of two pedigrees (each corresponding to a hypothesis describing a family relationship). The likelihood ratio is also reported. Unlike other implementations we are aware of, partial DNA profiles are allowed here. For instance, if the genotype of a person is reported as 1/0 (0 is 'missing') for a triallelic marker with uniform allele frequencies, the possible ordered genotypes (1,1), (2,1), (1,2), (1,3) and (3,1) are treated as equally likely. (For general allele frequencies, genotype probabilities are obtained by assuming Hardy-Weinberg equilibrium.) A reasonable future extension would be to allow the user to weigh these genotypes; typically (1,1) may be more likely than the others. If plot='plot_only', the function returns NULL after producing the plot.

Value

lik.numerator	likelihood of data given ped_numerator
lik.denominator	
	likelihood of data given ped_denominator
LR	likelihood ratio lik.numerator/lik.denominator

Author(s)

Thore Egeland, Magnus Dehli Vigeland

See Also

exclusionPower

Examples

```
# A partial DNA profile is obtained from the person
# denoted 4 in the figure produced below
# There are two possibilities:
# H1: 4 is the missing relative of 3 and 6 (as shown to the left) or
# H2: 4 is unrelated to 3 and 6.
*****
p = c(0.2, 0.8)
alleles = 1:length(p)
g3 = c(1,1); g4 = c(1,0); g6 = c(2,2)
x1 = nuclearPed(2)
x1 = addOffspring(x1, father = 4, sex = 1, noff = 1)
m = marker(x1, 3, g3, 4, g4, 6, g6, alleles = alleles, afreq = p)
x1 = addMarker(x1, m)
x^2 = nuclearPed(2)
x^2 = addOffspring(x^2, father = 4, sex = 1, noff = 1)
m = marker(x2, 3, g3, 6, g6, alleles = alleles, afreq = p)
x^2 = addMarker(x^2, m)
missing = singleton(4, sex = 1)
m.miss = marker(missing, g4, alleles = alleles, afreq = p)
missing = addMarker(missing, m.miss)
x2 = relabel(x2, c(1:3, 99, 5:6), 1:6)
known = list(c(3, g3), c(4, g4), c(6, g6))
LR = relationLR(x1, list(x2, missing), ids = c(3,4,6),
               alleles = alleles, afreq = p, known = known,
               title1 = 'H1: Missing person 4 related',
               title2 = 'H2:Missing person 4 unrelated')$LR
# Formula:
p = p[1]
LR.a = (1+p)/(2*p*(2-p))
stopifnot(abs(LR - LR.a) < 1e-10)</pre>
```

setAvailable

Functions for modifying availability vectors

Description

Functions to set and modify the availability vector of a 'linkdat' object. This vector is used in 'linkage.power' and 'linkageSim', indicating for whom genotypes should be simulated.

setModel

Usage

setAvailable(x, available)

swapAvailable(x, ids)

Arguments

х	a linkdat object
available	a numeric containing the IDs of available individuals.
ids	the individual(s) whose availability status should be swapped.

Value

The modified linkdat object.

Author(s)

Magnus Dehli Vigeland

See Also

plot.linkdat,linkage.power,linkageSim

Examples

```
data(toyped)
x = linkdat(toyped)
x = setAvailable(x, 3:4)
x = swapAvailable(x, 2:3)
x$available
```

setModel

Set, change or display the model parameters for 'linkdat' objects

Description

Functions to set, change and display model parameters involved in parametric linkage analysis.

Usage

```
setModel(x, model = NULL, chrom = NULL, penetrances = NULL,
  dfreq = NULL)
## S3 method for class 'linkdat.model'
print(x, ...)
```

Arguments

х	<pre>in setModel: a linkdat object. In print.linkdat.model: a linkdat.model object.</pre>
model	NULL, or an object of class linkdat.model, namely a list with elements chrom, penetrances and dfreq. In the setModel function, the model argument can be one of the integers 1-4, with the following meanings:
	1 = autosomal dominant; fully penetrant, dfreq=1e-5
	2 = autosomal recessive; fully penetrant, dfreq=1e-5
	3 = X-linked dominant; fully penetrant, dfreq=1e-5
	4 = X-linked recessive; fully penetrant, dfreq=1e-5
chrom	a character, either 'AUTOSOMAL' or 'X'. Lower case versions are allowed and will be converted automatically.
penetrances	if chrom=='AUTOSOMAL': a numeric of length 3 - (f0, f1, f2) - where fi is the probability of being affected given i disease alleles.
	<pre>If chrom=='X': a list of two vectors, containing the penetrances for each sex: penetrances = list(male=c(f0, f1), female=c(f0, f1, f2)).</pre>
dfreq	the population frequency of the disease allele.
	further parameters

Value

setModel returns a new linkdat object, whose model entry is a linkdat.model object: A list containing the given chrom, penetrances and dfreq.

Author(s)

Magnus Dehli Vigeland

See Also

linkdat

Examples

```
data(toyped)
x = linkdat(toyped)
x1 = setModel(x, model=1)
summary(x1)
# The shortcut 'model=1' above is equivalent to
x2 = setModel(x, chrom='AUTOSOMAL', penetrances=c(0,1,1), dfreq=1e-5)
stopifnot(all.equal(x1, x2))
# X-linked recessive model:
y1 = setModel(x, model=4, dfreq=0.01)
summary(y1)
```

setPlotLabels

setPlotLabels Attach plot labels to a linkdat object

Description

This function attaches (or modifies) a character vector of plotting labels for the pedigree members of a linkdat object. This is useful since only numerical ID's are allowed in defining pedigrees in paramlink.

Usage

setPlotLabels(x, labels, ids = x\$orig.ids)

Arguments

x	A linkdat object.
labels	A character vector of the same length as ids.
ids	A numeric vector of numerical IDs. Must be a subset of x\$orig.ids.

Value

A new linkdat object, differing from x only in x\$plot.labels.

Author(s)

Magnus Dehli Vigeland

See Also

plot.linkdat

Examples

```
x = nuclearPed(1)
x = setPlotLabels(x, labels=c('Father', 'Mother', 'Son'))
plot(x)
```

showInTriangle

Description

Utility function for plotting points in the IBD triangle.

Usage

```
showInTriangle(k0, k2 = NULL, new = T, col = "blue", cex = 1, pch = 4,
lwd = 2, labels = NULL, col_labels = col, cex_labels = 0.8, pos = 1,
adj = NULL, ...)
```

Arguments

k0, k2	Numerical vectors giving coordinates for points to be plotted in the IBDtriangle.	
new	Logical indicating if a new IBDtriangle should be drawn.	
col, cex, pch,	lwd	
	Parameters passed onto points.	
labels	A character of same length as k0, or NULL.	
col_labels, cex_labels, pos, adj		
	Parameters passed onto text (if labels is non-NULL).	
	Plot arguments passed on to IBDtriangle.	

Author(s)

Magnus Dehli Vigeland

See Also

IBDtriangle, examineKinships

Examples

showInTriangle(k0=3/8, k2=1/8, label="3/4 siblings", pos=1)

simpleSim

Description

Unconditional simulation of unlinked markers

Usage

```
simpleSim(x, N, alleles, afreq, available, Xchrom = FALSE, mutmat = NULL,
seed = NULL, verbose = T)
```

Arguments

х	a linkdat object
Ν	a positive integer: the number of markers to be simulated
alleles	a vector containing the allele names. If missing, the alleles are taken to be $seq_along(afreq)$.
afreq	a vector of length 2 containing the population frequencies for the alleles. If missing, the alleles are assumed equifrequent.
available	a vector containing IDs of the available individuals, i.e. those whose genotypes should be simulated.
Xchrom	a logical: X linked markers or not?
mutmat	a mutation matrix, or a list of two such matrices named 'female' and 'male'. The matrix/matrices must be square, with the allele labels as dimnames, and each row must sum to 1 (after rounding to 3 decimals).
seed	NULL, or a numeric seed for the random number generator.
verbose	a logical.

Details

This simulation is done by distributing alleles randomly to all founders, followed by unconditional gene dropping down throughout the pedigree (i.e. for each non-founder a random allele is selected from each of the parents). Finally the genotypes of any individuals not included in available are removed.

Value

a linkdat object equal to x in all respects except its markerdata entry, which consists of the N simulated markers.

Author(s)

Magnus Dehli Vigeland

toyped

See Also

markerSim, linkageSim

Examples

```
x = nuclearPed(1)
simpleSim(x, N=3, afreq=c(0.5, 0.5))
```

```
y = addOffspring(cousinPed(1), father=7, mother=8, noffs=1)
simpleSim(y, N=3, alleles=LETTERS[1:10])
```

toyped

Toy pedigree for linkage analysis

Description

Toy pedigree with 4 individuals typed at a single SNP marker. Individual 1 is missing one allele.

Usage

toyped

Format

A data frame with 4 rows and 8 columns

Details

The format is standard LINKAGE (pre-makeped) format, with columns as follows:

- FAMID. Family ID
- ID. Individual ID
- FID. Father ID
- MID. Mother ID
- SEX. Gender (male=1, female=2)
- AFF. Affection status (unaffected=1, affected=2, unkonwn=0)
- M_A1. First allele of marker 1
- M_A2. Second allele of marker 1

Examples

toyped
linkdat(toyped)

Description

Transfer marker data between pedigrees (in the form of linkdat objects). Both the source and target can be lists of linkdat and/or singleton objects (these must have disjoint ID sets). Any previous marker data of the target is overwritten.

Usage

transferMarkerdata(from, to)

Arguments

from	a linkdat or singleton object, or a list of such objects.
to	a linkdat or singleton object, or a list of such objects.

Value

A linkdat object (or a list of such) similar to to, but where all individuals also present in from have marker genotypes copied over. Any previous marker data is erased.

Author(s)

Magnus Dehli Vigeland

See Also

linkdat

Examples

```
x = list(singleton(id=5), nuclearPed(noffs=2))
x = markerSim(x, N=5, alleles=1:5, verbose=FALSE, available=4:5)
y = nuclearPed(noffs=3)
y = transferMarkerdata(x, y)
stopifnot(all.equal(x[[1]], branch(y,5)))
stopifnot(all.equal(x[[2]], subset(y,1:4)))
```

twoloops

Description

A consanguineous pedigree with two inbreeding loops.

Usage

twoloops

Format

A data frame with 17 rows and 6 columns. See toyped for details about the format.

Examples

```
x = linkdat(twoloops)
plot(x)
```

twoMarkerDistribution Genotype probability distribution

Description

Computes the joint genotype distribution of two markers for a specified pedigree member, conditional on existing genotypes and pedigree information.

Usage

```
twoMarkerDistribution(x, id, partialmarker1, partialmarker2, theta,
loop_breakers = NULL, eliminate = 99, verbose = TRUE)
```

Arguments

Х	A linkdat object.
id	The individual in question.
partialmarker1,	partialmarker2
	Either a single integer indicating the number of one of x's existing markers, or a marker object.
theta	A single numeric in the interval $[0, 0.5]$ - the recombination fraction between the two markers.

Xped

loop_breakers	A numeric containing IDs of individuals to be used as loop breakers. Relevant only if the pedigree has loops. See breakLoops.
eliminate	A non-negative integer, indicating the number of iterations in the internal genotype compatibility algorithm. Positive values can save time if partialmarker is non-empty and the number of alleles is large.
verbose	A logical.

Value

A named matrix giving the joint genotype distribution.

Author(s)

Magnus Dehli Vigeland

See Also

oneMarkerDistribution

Examples

```
x = nuclearPed(2)
emptySNP = marker(x, alleles=c('a', 'b'))
SNP1 = marker(x, 1, c(1,1), 2, c(1,0), alleles=1:2, afreq=c(0.1, 0.9))
twoMarkerDistribution(x, id=2, emptySNP, SNP1, theta=0)
twoMarkerDistribution(x, id=2, emptySNP, SNP1, theta=0.5)
twoMarkerDistribution(x, id=3, emptySNP, SNP1, theta=0.5)
# X-linked example
SNPX_1 = marker(x, 2, c('a', 'b'), 3, 'b', alleles=c('a', 'b'), chrom=23)
SNPX_2 = marker(x, 2, c('a', 'b'), 3, 'b', alleles=c('a', 'b'), chrom=23)
r1 = twoMarkerDistribution(x, id=4, SNPX_1, SNPX_2, theta=0)
r2 = twoMarkerDistribution(x, id=4, SNPX_1, SNPX_2, theta=0.5)
stopifnot(all(r1==c(.5,0,0,.5)), all(r2==c(.25,.25,.25,.25)))
```

Xped

Example pedigree with X-linked disease pattern.

Description

A complex pedigree with an X-linked recessive disease pattern

Usage

Xped

A data frame with 15 rows and 6 columns. See toyped for details about the format.

Details

The format is standard LINKAGE (pre-makeped) format.

Examples

Xped

```
# Convert to a 'linkdat' object and set a recessive X-linked model:
x = linkdat(Xped, model=4)
summary(x)
```

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