# Package 'bqtl'

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<b>Description</b> QTL mapping toolkit for inbred crosses and recombinant inbred lines. Includes maximum likelihood and Bayesian tools.
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Some Introductory Comments

# Description

A Starting Point

Some pointers to a few key functions in BQTL

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#### New to R?

- Be sure to check out all of the free documentation that comes with R.
- The example function is very helpful in getting familiar with a new function. You type example(fun) and the examples in the documentation for fun are run, then you can read the documentation to get a bette sense of what is really going on. My personal favorite is to type par(ask=T), hit the 'enter' key, then example(image), and 'enter' again; after each display you hit the 'enter' key to get to the next one.
- library(bgtl) is needed to load the *BQTL* functions and data sets.

#### **Key Functions**

#### Data Input \

make.map.frame defines the map,

marker.levels The help page describes several functions that define the coding scheme for marker levels,

make.analysis.obj combines marker data, phenotype data, and the map.frame to create an object that can be used by data analysis functions.

#### Maximum Likelihood Methods \

bqtl does a host of things from marker regression and interval mapping to full maximum likelihood. The best way to get started is to run example(bqtl) and take a look at the resulting output.

locus is very helpful in specification of runs.

# **Approximate Bayesian Analysis** \

linear.bayes For a good starting point try example(linear.bayes)

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

# **Description**

The approximation provided by linear.bayes can be improved by performing Laplace approximations. This function is a development version of a wrapper to do that for all of the returned by linear.bayes.

# Usage

```
adjust.linear.bayes(lbo, ana.obj=lbo$call$ana.obj, ...)
```

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#### **Arguments**

1bo The object returned by linear.bayes

ana.obj The analysis.object used to create 1bo. This need not be given explicitly, iff

the original version is in the search path.

... currently unused

#### Value

A list of class "adjust.linear.bayes" containing:

odds A vector, typically of length k giving the odds for models of size 1, 2, ..., k under

a uniform posterior relative to a model with no genes.

loc.posterior The marginal posterior probabilities by locus coefficients The marginal posterior means of the coefficients

one.gene.adj Results of fits for one gene models

n.gene.adj Results of fits for modles with more than one gene

call the call to adjust.linear.bayes

#### Note

For large linear.bayes objects invloving many gene models, this can require a very long time to run.

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

# See Also

linear.bayes

bqtl

Bayesian QTL Model Fitting

# **Description**

Find maximum likelihood estimate(s) or posterior mode(s) for QTL model(s). Use Laplace approximation to determine the posterior mass associated with the model(s).

# Usage

```
bqtl(reg.formula, ana.obj, scope = ana.obj$reg.names, expand.specials = NULL, ...)
```

bqtl 5

# Arguments

reg.formula A formula.object like y ~ add.PVV4 \* add.H15C12. The names of the indepen-

dent variables on the right hand side of the formula are the names of loci or the names of additive and dominance terms associated with loci. In addition, one can use locus or configs terms to specify one or a collection of terms in a shorthand notation. See locus for more details. The left hand side is the name of a trait variable stored in the search path, as a column of the data frame data,

or y if the phenotype variable in ana. obj is used.

ana.obj The result of make.analysis.obj.

scope passed to lapadj

expand.specials

passed to lapadj

... Arguments to pass to lapadj, e.g. rparm and return.hess

#### **Details**

This function is a wrapper for lapadj. It does a lot of useful packaging through the configs terms. If there is no configs term, then the result is simply the output of lapadj with the call attribute replaced by the call to bqtl

#### Value

The result(s) of calling lapadj. If configs is used in the reg. formula, then the result is a list with one element for each formula. Each element is the value returned by lapadj

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

#### References

Tierney L. and Kadane J.B. (1986) Accurate Approximations for Posterior Moments and Marginal Densities. *JASA*, **81**,82–86.

#### See Also

```
locus, configs, lapadj
```

# **Examples**

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```
several.epi <-
                                             # 20 epistatic models
   bqtl( bc.phenotype ~ m.12 * locus(31:50), little.ana.bc)
                                             # main effects only
several.main <-
   bqtl( bc.phenotype ~ m.12 + locus(31:50), little.ana.bc)
max.loglik <- max( loglik(several.epi) - loglik(several.main) )</pre>
round(
     c( Chi.Square=2*max.loglik, df=1, p.value=1-pchisq(2*max.loglik,1))
      ,2)
five.gene <-
                                              ## a five gene model
 bqtl( bc.phenotype ~ locus( 12, 32, 44, 22, 76 ), little.ana.bc , return.hess=TRUE )
regr.coef.table <- summary(five.gene)$coefficients</pre>
round( regr.coef.table[,"Value"] + # coefs inside 95% CI
          qnorm(0.025) * regr.coef.table[,"Std.Err"] %0%
            c("Lower CI"=1, "Estimate"=0, "Upper CI"=-1),3)
```

bqtl-internal

Internal BQTL functions

#### **Description**

Internal bqtl functions and objects

# Usage

```
x %equiv% y
map.dx(lambda, theta, min.lambda)
rhs.bqtl(reg.terms, ana.obj, bqtl.specials, local.covar, scope,
        expand.specials = NULL, method, ...)
zero.dup(x,dig=6)
uniq.config(swap.obj)
```

# Arguments

lambda (2\*(recomb fraction-1/2) theta recomb fraction

min.lambda smallest map distance to use

reg.terms a formula

ana.obj an analysis.object

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bqtl.specials a vector of acceptable special names

local.covar a function

scope vector of strings

expand.specials

logical, whether to use expand.grid on the loci

method e'g' "F2", "BC!", etc

... not sure

swap.obj result of swap

x numeric vector or matrix
y numeric vector or matrix

dig how many significant digits to use

# **Details**

These are not to be called by the user.

bqtl.fitter

Get loglikelihoods for many models of a common form

# **Description**

For a single type of model, this function evaluates multiple models that differ only in terms of the loci involved. The looping is all done by internal C functions, so this is faster than simply using bqtl to do the same thing.

# Usage

```
bqtl.fitter(setup, loc.mat, ana.obj)
```

# **Arguments**

setup The object returned by bqtl(<...> , setup=TRUE )

loc.mat A matrix of locus numbers, s.t. nrow(loc.mat) equals the number of loci in

setup

ana.obj An analysis.object. Usually the one used in setup

# **Details**

In order to avoid the computational overhead of running large loops of very repetitive operations in R/S, bqtl.fitter used after the setup=TRUE option in bqtl will loop through the loci specified in loc.mat using internal C code. This is many times faster than running the same code via bqtl.

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#### Value

For now it only returns the loglikelihood. But it would be trivial to build an option that would allow other quantities computed to be returned, and this should probably be done. However, some care is needed to keep objects from becoming unmanageably large.

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

# See Also

bqtl

# **Examples**

```
data( little.ana.bc )
little.setup <-
bqtl( bc.phenotype~locus(1)*locus(2), little.ana.bc, setup=TRUE )
combos <- t( as.matrix( expand.grid( 1:21, 44:64 ) ) )
little.update <- bqtl.fitter(little.setup, combos, little.ana.bc)
little.res <- matrix( little.update, nr=21 )
image( 1:21, 44:64, little.res )
rm(little.ana.bc, little.update, little.res )</pre>
```

coef.bqtl

Extract Coefficients from fitted objects

#### **Description**

Return a vector or matrix of coefficients as appropriate

# Usage

```
## S3 method for class 'bqtl'
coef(object,...)
```

# **Arguments**

object The object returned by bqtl ... ignored

#### Value

A vector (if bqtl returned a single fit) or matrix (if bqtl returned a list with more than one fit)

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

configs 9

#### See Also

bqtl

configs	Lookup loci or effects for genetic model formulas	

# Description

Convert numeric indexes to names of regressors for a genetic model. One or many genetic models can be specified through the use of this function. It is used on the right hand side of a formula in the bqtl function.

#### Usage

```
configs(x,...,scope, method = NULL)
```

#### **Arguments**

Χ	Typically an integer, an integer vector, an array, or a list with a configs com-
	ponent such as returned by swapbc1. However, it can also be a character string,
	vector, et cetera, in which case the elements must belong to names(scope)

... Optional arguments to be used when is.atomic(x) is TRUE.

scope (Optional and)

Usually not supplied by the user. Rather bqtl fills this in automatically. A vector of regressor names, like the reg. names component returned by make.analysis.obj.

When mode(x) is "character", then names(scope) **must** be non-NULL

method (Optional and) Usually not supplied by the user. A method like "F2". Typically,

this is determined by internal code.

#### **Details**

configs is used in the model formula notation of bqtl, possibly more than once, and possibly with regressors named in the usual manner. configs is intended to speed up the specification and examination of genetic models by allowing many models to be specified in a shorthand notation in a single model formula. The names of genetic loci can consist of marker names, names that encode chromosome number and location, or other shorthand notations. The names of terms in genetic models will typically include the names of the locus and may prepend "add." or "dom." or similar abbreviations for the 'additive' and 'dominance' terms associated with the locus.

When used as in  $bqtl(y \sim configs(34), my.analysis.obj)$ , it will look up the term my.analysis.obj\$reg.names[34]. When this is passed back to bqtl, it get pasted into the formula and is subsequently processed to yield the fit for a one gene model.

When used as in bqtl(y ~ configs(34,75,172), my.analysis.obj) it looks up each term and returns a result to bqtl that results in fitting a 3 gene model (without interaction terms).

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When x is a vector, array, or list, the processing typically returns pieces of many model formulas. bqtl(y ~ configs(26:75),...) results in a list of 50 different one gene model fits from bqtl for the terms corresponding to the 26th through the 75th variables. bqtl(y ~ configs(cbind(c(15,45,192),c(16,46,193))) returns two four gene models. And more generally, whenever is.array(x) is TRUE, the columns (or slices) specify dim(x)[1]/length(x) different models. When x\$configs is an array, this also happens. This turns out to be useful when the result of running swapbc1 or swapf2 is treated as an importance sample. In such a case, bqtl(y ~ configs(my.swap),my.analysis.obj) will return a list in which element i is the ith sample drawn when my.swap <-swapbc1(...) was run.

#### Value

A character vector whose element(s) can be parsed as the right hand side of a model formula.

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

#### See Also

bqtl and the examples there for a sense of how to use configs, make.analysis.obj for the setup that encodes the marker map and the marker information, swapbc1 and swapf2 for generating samples to be screened by bqtl.

covar

Treat locus as covariate

# **Description**

Sometimes it is helps speed computations to linearize the likelihood or at least a part of it w.r.t. the locus allele values. Both 'Haley-Knott regression' and 'composite interval mapping' use this approach. covar provides a mechanism for creating formula objects that specify such linearizations.

# Usage

covar(x)

#### **Arguments**

Х

The name of a locus (except for F2 designs, when it is the name of an effect like 'add.m.32') or any argument of the sort that locus allows. If x evaluates to a single value, then additional atomic elements may be included as with locus.

#### Details

The function covar actually only returns x. The real work is done by a covar function that is hidden inside of bqtl, where the arguments are parsed as for locus. Each of the return values from locus is prefixed by "covar(" and suffixed by ")". If x is a name of a locus or effect, then paste("covar(",deparse(x),")") is returned. Later, when bqtl calls lapadj, terms like covar(PVV4.1) are recognized as requiring a linearization w.r.t. effect 'PVV4.1'.

formula.bqtl

# Value

a character string or vector

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

# References

HALEY, C. S. and S. A. KNOTT, 1992 A simple regression method for mapping quantitative trait loci in line crosses using flanking markers. Heredity 69:315-324.

Knapp SJ, Bridges WC, and Birkes D. Mapping quantitative trait loci using molecular marker linkage maps. Theoretical and Applied Genetics 79: 583-592, 1990.

ZENG, Z.-B., 1994 Precision mapping of quantitative trait loci. Genetics 136:1457-1468

#### See Also

locus, add, dom, configs

formula.bqtl

Extract formula from bqtl object

# **Description**

formula method for class bqt1

# Usage

```
## S3 method for class 'bqtl'
formula(x, ...)
```

#### **Arguments**

x The object returned by bqt1 ....

#### Value

a formula object

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

# See Also

bqtl

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lapadj

Approximate marginal posterior for chosen model

# **Description**

lapadj provides the Laplace approximation to the marginal posterior (over coefficients and dispersion parameter) for a given genetical model for a quantitative trait. A by-product is the parameter value corresponding to the maximum posterior or likelihood.

# Usage

# **Arguments**

reg.formula A formula, like  $y\sim add.X.3+dom.X.3+add.x.45*add.x.72$  ana.obj Seemake.analysis.obj, which returns objects like this

rparm One of the following:

A scalar that will be used as the ridge parameter for all design terms except for

the intercept ridge parameter which is set to zero

A vector who named elements can be matched by the design term names returned in \$reg.vec. If no term named "intercept" is provided,

rparm["intercept"] will be set to zero.

A vector with (q-1)\*k elements (this works when there are no interactions spec-

ified). If names are provided, these will be used for matching.

Positive entries are 'ridge' parameters or variance ratios in a Bayesian prior for the regression coefficients. Larger values imply more shrinkage or a more con-

centrated prior for the regresion coefficients.

tol Iteration control parameter

return.hess Logical, include the Hessian in the output?
mode.names names to use as dimnames(mode.mat)[[2]]

mode.mat Not usually set by the user. A matrix which indicates the values of regressor

variables corresponding to the allele states. If mode. mat is not given by the user,

ana.obj\$mode.mat is used.

maxit Maximum Number of iterations to perform

nem Number of EM iterations to use in reinitializing the pseudo-Hessian

setup.only If TRUE, do not run. Return an object that can be use for a direct call to .C

subset expression to evaluate using ana.obj\$data as the environment

casewt a vector of non-negative weights

start.parm Vector of starting values for the maximization

. . . other objects needed in fitting

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#### **Details**

The core of this function is a quasi-Newton optimizer due to Minami (1993) that has a computational burden that is only a bit more than the EM algorithm, but features fast convergence. This is used to find the mode of the posterior. Once this is in hand, one can find the Laplace approximation to the marginal likelihood. In addition, some useful quantities are provided that help in estimating the marginal posterior over groups of models.

#### Value

A list with components to be used in constructing approximations to the marginal posterior or a list that can be used to call the underlying C code directly. In the former case, these are:

adj The ratio of the laplace approximation to the posterior for the correct likelihood

to the laplace approximation to the posterior for the linearized likelihood

logpost The logarithm of the posterior or likelihood at the mode

parm the location of the mode

posterior The laplace approximation of the marginal posterior for the exact likelihood

hk.approx Laplace approximation to the linearized likelihood hk.exact Exact marginal posterior for the linearized likelihood

reg.vec A vector of the variables used

rparm Values of ridge parameters used in this problem.

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

#### References

Berry C.C.(1998) Computationally Efficient Bayesian QTL Mapping in Experimental Crosses. ASA Proceedings of the Biometrics Section. 164–169.

Minami M. (1993) Variance estimation for simultaneous response growth curve models. Thesis (Ph. D.)–University of California, San Diego, Department of Mathematics.

linear.bayes Bayesian QTL mapping via Linearized Likelihood

# **Description**

The Bayesian QTL models via a likelihood that is linearized w.r.t. a fixed genetic model. By default, all one and two gene models (without epistasis) are fitted and a MCMC sampler is used to fit 3,4, and 5 gene and (optionally) larger models.

#### Usage

```
linear.bayes(x, ana.obj, partial=NULL, rparm, specs,
scope, subset, casewt, ...)
```

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#### **Arguments**

x a formula giving the QTL and the candidate loci or a varcov object

ana.obj An analysis.object, see make.analysis.obj
partial a formula giving covariates to be controlled

rparm A ridge parameter. A value of 1 is suggested, but the default is 0.

specs An optional list with components gene.number (to indicate the model sizes),

burn.in (to indicate the number of initial MCMC cycles to discard), and n. cycles (to indicate how many MCMC cycles to perform for each model size). If no val-

ues are supplied, specs defaults to

list(gene.number=c(1,2,3,4,5),burn.in=1,n.cycles=c(0,0,200,100,100))

scope Not generally used. If supplied this will be passed to varcov.

Subset Not generally used. If supplied this will be passed to varcov.

Not generally used. If supplied this will be passed to varcov.

... optional arguments to pass to twohk and swap

#### **Details**

This function is a wrapper for varcov, twohk, swap, and summary. swap, and a better understanding of optional arguments and the object generated is gained from their documentation.

#### Value

hk The object returned by twohk

swaps A list of objects returned by calls to swap. Element i in swaps is for i gene

models.

smry A list of objects returned by calls to summary.swap. Some elements may be

NULL if no samples were requested or if the sampling process yielded degenerate results. Usually, this happens if no posterior is specified for the regression

coefficients, i.e. if rparm=0 was used or implied

odds A Vector of odds (relative to a no gene setup) for each model size evaluated.

The odds are computed under a prior that places equal weights on models of each size considered (and are, therefore, Bayes Factors). If models of size 1 and 2 are not evaluated or if some degenerate results were encountered, this will be

NULL

coefs A vector of posterior means of the regression coefficients. If models of size 1

and 2 are not evaluated or if some degenerate results were encountered, this will

be NULL

loc.posterior A vector of locus-wise posterior probabilities that the interval covered by this

locus contains a gene. If models of size 1 and 2 are not evaluated or if some

degenerate results were encountered, this will be NULL

call The call that generated this object

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

little.ana.bc

#### References

Berry C.C.(1998) Computationally Efficient Bayesian QTL Mapping in Experimental Crosses. *ASA Proceedings of the Biometrics Section*. 164–169.

#### **Examples**

```
data( little.ana.bc )
little.lin <- linear.bayes( bc.phenotype~locus(all), little.ana.bc, rparm=1 )
par(mfrow=c(2,3))
plot( little.ana.bc, little.lin$loc.posterior, type="h" )
little.lin$odds
par(mfrow=c(1,1))
plot(fitted(little.lin), residuals(little.lin))</pre>
```

little.ana.bc

A simulated dataset

# **Description**

A simulation of a BC1 cross of 150 organisms with a genome of around 500 cM consisting of 5 chromosomes. The format is that created by make.analysis.obj

#### **Details**

This dataset is built up from several others. The basic data are:

little.bc.pheno A vector of phenotype data

little.bc.markers A map.frame of marker data and

**little.dx** A data frame with 50 rows and 2 columns that specify the map locations of a simulated set of markers

These are used to construct

```
little.mf.5 A map.frame with 'pseudo-markers' at least every 5 cM made from
    little.mf.5 <- make.map.frame(little.map.frame, nint=marker.fill(little.map.frame,
    reso=5, TRUE))</pre>
```

Then phenotype, covariate, and marker data are combined

```
with little.mf.5
```

little.bc.pheno A data.frame with the variable bc.phenotype

little.bc.markers A data.frame with marker state information

#### See Also

The examples in make.analysis.obj

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little.ana.f2

A simulated dataset

# Description

A simulation of an F2 cross of 150 organisms with a genome of around 500 cM consisting of 5 chromosomes. The format is that created by make.analysis.obj

# Usage

```
data(little.ana.f2)
```

little.bc.markers

Simulated Marker Data

# **Description**

The little.bc.markers data frame has 150 rows and 50 columns with the simulated marker data from a BC1 cross of 150 organisms with a genome of around 500 cM consisting of 5 chromosomes. Some NA's have been intentionally introduced.

#### Usage

```
data(little.bc.markers)
```

#### **Format**

This data frame contains the following columns:

- m.1 a factor with levels AA Aa
- m.2 a factor with levels AA Aa
- m.3 ditto
- m.4 ditto
- m.5 ditto
- m.6 ditto
- m.7 ditto
- m.8 ditto
- m.9 ditto
- m.10 ditto
- m.11 ditto
- m.12 ditto
- m.13 ditto

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- m.14 ditto
- m.15 ditto
- m.16 ditto
- m.17 ditto
- m.18 ditto
- **m.19** ditto
- m.20 ditto
- m.21 ditto
- m.22 ditto
- m.23 ditto
- m.24 ditto
- m.25 ditto
- **m.26** ditto
- m.27 ditto
- m.28 ditto
- m.29 ditto
- \_\_\_\_\_
- m.30 ditto
- m.31 ditto
- m.32 ditto
- m.33 ditto
- m.34 ditto
- m.35 ditto
- m.36 ditto
- m.37 ditto
- m.38 ditto
- m.39 ditto
- m.40 ditto
- m.41 ditto
- m.42 ditto
- m.43 ditto
- m.44 dittom.45 ditto
- **m.46** ditto
- **m.47** ditto
- m.48 ditto
- m.49 ditto
- m.50 ditto

row.names row.names

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little.bc.pheno

Simulated Phenotype Data

# Description

The little.bc.pheno data frame has 150 rows and 1 columns.

# Usage

```
data(little.bc.pheno)
```

#### **Format**

This data frame contains the following columns:

bc.phenotype a numeric vector of simulated phenotype data

little.f2.markers

Simulated Marker Data

# Description

The little.f2.markers data frame has 150 rows and 50 columns with the simulated marker data from an F2 cross of 150 organisms with a genome of around 500 cM consisting of 5 chromosomes.

# Usage

```
data(little.f2.markers)
```

# **Format**

This data frame contains the following columns:

- m.1 a factor with levels AA Aa aa
- m.2 a factor with levels AA Aa aa
- m.3 ditto
- m.4 ditto
- m.5 ditto
- m.6 ditto
- m.7 ditto
- m.8 ditto
- m.9 ditto
- **m.10** ditto

little.f2.markers

- m.11 ditto
- m.12 ditto
- m.13 ditto
- m.14 ditto
- m.15 ditto
- m.16 ditto
- m.17 ditto
- m.18 ditto
- **m.19** ditto
- m.20 ditto
- m.21 ditto
- m.22 ditto
- m.23 ditto
- **....**
- m.24 ditto
- m.25 a factor with levels A- aa
- m.26 ditto
- m.27 ditto
- m.28 ditto
- m.29 ditto
- m.30 ditto
- m.31 ditto
- m.32 ditto
- m.33 ditto
- m.34 ditto
- m.35 ditto
- m.36 ditto
- m.37 ditto
- m.38 ditto
- m.39 ditto
- m.40 ditto
- m.41 ditto
- m.42 ditto
- m.43 ditto
- m.44 ditto
- m.45 a factor with levels a-
- m.46 ditto
- m.47 ditto
- m.48 ditto
- m.49 a factor with levels AA Aa aa
- m.50 a factor with levels AA Aa aa

row.names row names

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little.f2.pheno

Simulated Phenotype Data

# Description

The little.f2.pheno data frame has 150 rows and 1 columns.

# Usage

```
data(little.f2.pheno)
```

#### **Format**

This data frame contains the following columns:

**f2.phenotype** a numeric vector of simulated phenotype data

little.map.dx

Marker Map Description for Simulated Data

# Description

The little.map.dx data frame has 50 rows and 2 columns that specify the map locations of a simulated set of markers

# Usage

```
data(little.map.dx)
```

# **Format**

This data frame contains the following columns:

```
marker.names a factor with levels m.1...m.50
```

cM a numeric vector of map locations in centimorgans

little.map.frame 21

little.map.frame

Package of Simulated Marker Map Information

# **Description**

The little.map.frame data frame has 50 rows and 9 columns that describe the marker map of little.map.dx in the format produced by make.map.frame. little.map.dx has the minimal data needed to construct this.

# Usage

```
data(little.map.frame)
```

#### **Format**

This data frame contains the following columns:

marker.name a factor with levels m. 1 m. 2 ... m. 50

cM a vector of locations

**prior** weights to be used in sampling and Bayesian computations

pos.type a factor with levels left right center

is.marker always TRUE for these data

pos.plot a vector of plotting positions

lambda transformed recombination fractions

locus an abbreviated locus name

**chr.num** the chromosome number 1, 2, 3, 4, or 5.

little.mf.5

Package of Simulated Marker Map Information

# **Description**

The little.mf.5 data frame has 114 rows and 9 columns consisting of little.map.frame plus 64 'virtual' marker loci

#### Usage

```
data(little.mf.5)
```

22 locus

#### **Format**

This data frame contains the following columns:

marker.name The marker names taken from little.map.frame and those created to fill virtual markers in between actual markers.

cM a vector of locations

**prior** weights to be used in sampling and Bayesian computations

pos.type a factor with levels left right center

is.marker TRUE for the 50 markers, FALSE for the 'virtual' markers

pos.plot a vector of plotting positions

lambda transformed recombination fractions

locus an abbreviated locus name

**chr.num** the chromosome number 1, 2, 3, 4, or 5.

locus

Lookup loci or effects for genetic model formulas

#### Description

Convert numeric indexes to names of regressors for a genetic model. One or many genetic models can be specified through the use of this function. It is used on the right hand side of a formula in the bqtl function.

#### Usage

```
locus(x, ..., scope, method, chromo, cM, ana.obj)
add(x, ..., scope, method)
dom(x, ..., scope, method)
```

# **Arguments**

X	Typically an integer, an integer vector, or an array whose elements are integers.
	These index loci described in a map.frame object.

However, x can also be a character string, vector, et cetera, in which case the elements must belong to names(scope).

Optional arguments (usually integers) to be used when is.atomic(x) is TRUE.

chromo A chromosome number or 2 ordered numbers. The loci on the chromosome or in the range of chromosome numbers are used. If chromo is used, x must not be

used.

cM (Optional) map distance or two giving a location near a locus or range of lo-

cations from which loci will be included. If the one chromosome number is specified in chromo, cM must be ordered. If cM is omitted, all loci on the chro-

mosome(s) will be included.

locus 23

scope	(Optional and) Usually not supplied by the user. Rather bqtl fills this in automatically. A vector of regressor names, like the reg.names component returned by make.analysis.obj.
method	(Optional and) Usually not supplied by the user. Like scope, bqtl takes care of filling this in with "BC1", "F2", et cetera as appropriate.
ana.obj	Usually not specified by the user. This is the analysis.object to be used to lookup loci if a chromo argument is used.

#### **Details**

locus is used in the model formula notation of bqtl, possibly more than once, and possibly with regressors named in the usual manner. locus is intended to speed up the specification and examination of genetic models by allowing many models to be specified in a shorthand notation in a single model formula. The names of genetic loci can consist of marker names, names that encode chromosome number and location, or other shorthand notations. The names of terms in genetic models will typically include the names of the locus and may prepend "add." or "dom." or similar abbreviations for the 'additive' and 'dominance' terms associated with the locus.

When used as in  $bqtl(y \sim locus(34), my.analysis.obj)$ , it will look up the term or terms corresponding to the 34th locus. When this is passed back to bqtl, it is pasted into a text string that will become a formula and is subsequently processed to yield the fit for a one gene model.

When used as in bqtl( $y \sim locus(34,75,172)$ , my.analysis.obj) it looks up each term and returns a result to bqtl that results in fitting a 3 gene model (without interaction terms).

When x is a vector or array, the processing typically returns pieces character strings for many model formulas. bqtl(y ~ locus(26:75), ...) results in a list of 50 different one gene model fits from bqtl for the terms corresponding to the 26th through the 75th variables. bqtl(y ~ locus(cbind(c(15,45,192),c(16,46,1)) returns two three gene models. And more generally, whenever is.array(x) is TRUE, the columns (or slices) specify dim(x)[1]/length(x) different models.

The chromo argument performs a lookup of loci on the chromosome via the function map.index. If cM is also given, the locus nearest that location is used. If two values are given for cM all loci in the range are used.

add(x) and dom(x) are alternatives that specify that only the *additive* or *dominance* terms in an F2 intercross.

#### Value

A character vector whose element(s) can be parsed as the right hand side of a model formula(s).

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

#### See Also

configs, bqtl, and the examples there for a sense of how to use locus, make.analysis.obj for the setup that encodes the marker map and the marker information.

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loglik

Extract loglikelihood, log posterior, or posterior from fitted models

# Description

A fitted model or a list of such generated by bqtl has a maximum log likelihood or log posterior and a posterior. These functions simply extract them.

# Usage

```
loglik(x, ...)
logpost(x, ...)
posterior(x, ...)
```

# **Arguments**

x The object produced by bqt1... Currently unused

#### Value

A vector of numbers whose length equals the number of fitted models in x

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

#### See Also

bqtl

make.analysis.obj

Set up data for QTL mapping

# **Description**

Create commonly used objects for the analysis of a backcross or intercross experiment or of recombinant inbred lines.

# Usage

```
make.analysis.obj(data, map.frame, marker.frame, marker.levels=NULL,
method="F2", casewt=NULL,varcov=FALSE,mode.mat=NULL)
```

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# **Arguments**

data A data. frame (or vector) of phenotype and (optionally) covariate information

A map.frame.object (see make.map.frame) encoding the map information map.frame

and other details of the study

marker.frame A marker.frame.object. A matrix or data.frame of marker state information.

marker.levels A vector of length six or NULL. If NULL then the defaults for the elements are:

Element	F2.default	BC.default	RI.default
1	"AA"	"AA"	"AA"
2	"Aa"	"Aa"	"aa"
3	"aa"	"nil"	"nil"
4	"A-"	"nil"	"nil"
5	"a-"	"nil"	"nil"
6	"-"	"_"	"_"

NA's are allowed in marker. frame as well as the sixth element("--" by default) to denote missing data. To use other coding schemes replace "AA" and "aa" by codes for homozygous states, "Aa" by the code for heterozygotes, "A-" by the code for 'not aa', "a-" by the code for 'not AA', and "--" by the missing code. Positions 3:5 are just placeholders if method!="F2", but must be present.

One of "F2", "BC1", "RI.self", or "RI.sib" method

casewt If there are multiple observations on one genotype (such as in recombinant in-

breds) this can be used to assign a weight to each observation. The wisdom of

doing this is debatable.

If FALSE, don't create a varcov.object. Otherwise give an index into data to varcov

select a dependent variable. See varcov

mode.mat If NULL use the default. For method=="F2" ( and the default marker.levels of

AA, Aa, and aa ), this is a 3 by 2 matrix:

Genotype	add	dom
AA	1	-1
Aa	0	1
aa	-1	-1

For method=="BC1" (and the default marker.levels of AA and Aa),it is

Genotype AA 1 Aa -1

and for RIL methods (and the default marker.levels of AA and aa),it is

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Genotype	
AA	1
aa	-1

Other choices of marker.levels will relabel the corresponding rows.

#### **Details**

A lot of stuff is bundled together in one object. The function is really just a wrapper calling other make.\* functions.

#### Value

#### A list with components

data data.frame of phenotype, covariate information, and regressors created by

make.regressor.matrix

varcov A varcov.object. See make.varcov

reg.names The names of the regressors from make.regressor.matrix

method The method argument in the call.

state.matrix See make.state.matrix
loc.right See make.loc.right
map.frame See make.map.frame
casewt The casewt argument
mode.mat The mode.mat used

version A string giving the version of BQTL from qhich the objects was created

call The function call

#### Note

This can be quite a LARGE object.It might be better in crosses with lots (say, thousands) of markers, or in which many 'virtual' markers are used, or on computers with limited RAM to store each component separately. Not all components are used in every type of analysis.

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

# See Also

make.map.frame for definition of the marker map, The internally used functions are: make.loc.right, make.state.matrix, make.regressor.matrix, make.varcov, and make.marker.numeric

make.loc.right 27

# **Examples**

make.loc.right

Keep track of fully informative markers or states

# **Description**

Helps speed computations in multigene models by allowing a quick assessment of whether two loci are independent given the marker information for the individual.

#### Usage

```
make.loc.right(marker.frame, marker.distances)
```

# **Arguments**

```
marker.frame A marker.frame.object marker.distances
```

Actually a misnomer, this is a vector with a zero in the last position of each chromosome.

#### Value

A matrix of the same dimension as marker. frame whose elements index the column on the next (right) fully informative marker.

# Author(s)

```
Charles C. Berry <cberry@ucsd.edu>
```

28 make.map.frame

make.location.prior

Provide a default prior

# **Description**

Uses the map distances as a means of assigning a prior for chromosomal location. Basically, this function attempts to assign equal weight according to the spacing or markers and 'virtual' markers.

# Usage

```
make.location.prior( x, add.2.end=0, normalize=TRUE )
```

# **Arguments**

 $x = e^{-mgd}$  ,where mgd is the map distance in Morgans

add. 2. end How many Morgans to extend the first and last interval on each chromosome

normalize If TRUE, let the result sum to 1.0

# Value

A vector of length(x) whose sum is one, if normalize==TRUE

#### Author(s)

Charles C. Berry, <cberry@ucsd.edu>

make.map.frame

Create marker map specifications

# **Description**

A map.frame.object describes a marker map and additional loci that may be used in a QTL study. Each row pertains to one locus. Names of markers, abbreviated names, distances, and other necessary and useful information are bundled.

# Usage

```
make.map.frame(dx,chr.num = NULL, prior = make.location.prior(lambda),
    morgan = 100, nint = NULL, reso = NULL)
```

make.map.frame 29

# Arguments

dx An object of class "map.frame" or class "data.frame" or a vector or a data.frame

with a column named cM, M, or dx or whose first column gives location on each chromosome in centiMorgans (from start of chromosome or Morgans if M was the column name). It is best if names(dx) (for vector arguments) or row.names(dx) (for data.frame arguments) give names of markers for later ref-

erence, but this isn't really necessary.

chr. num (Optional) Vector of chromosome numbers

prior (Optional) Vector of Prior probabilities for the loci

morgan 100 if centiMorgans, 1 if Morgans

nint (Optional) Vector of one plus number of 'virtual' markers to be inserted after

each locus

reso Maximum distance between loci. If necessary fill in with 'pseudo-markers'

#### **Details**

The QTL analysis depends on information about the marker map and on specifications of the loci to be studied. The 'map.frame' contains this information.

#### Value

A data frame with components:

marker.name The full text identifier for this marker, e.g. "HH.360L.Col" is a marker on chro-

mosome 1 of arabidopsis thaliana, and names like this can be used for reference purposes. 'Virtual' markers have a suffix appended to the name of the previous

marker.

cM Location on the chromosome. If this is a marker of a locus that was input via

dx, then it is just the value of dx.

pos.type "left" if it is the first locus on this chromosome, "right" if it is last, or "center"

otherwise.

is.marker TRUE if this was actually a marker, FALSE if it is a 'virtual' marker

pos.plot Plotting position for this locus. Typically the same as dx.

lambda Twice the recombination fraction minus one.

locus An abbreviation for the locus of the form "C.<chr.num>.<cM>"

chr.num The chromosome number.

#### Note

The idea in having all of this bundled together is to make it easier for plot and summary methods to be implemented and to allow convenient references in formula based methods.

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

30 make.marker.numeric

#### **Examples**

```
data( little.map.dx )
little.map.frame <- make.map.frame( little.map.dx )
plot( little.map.frame ) # there is a plot method
# add 'virtual' markers to map
little.mf.5 <- make.map.frame(little.map.frame,reso=5)
print(little.mf.5[1:10,],digits=1) # show a few rows
plot( little.mf.5 ) # notice the 'virtual' markers added</pre>
```

make.marker.numeric

Translate a marker.frame.object to numeric matrix

# **Description**

Not to be called directly by users. This utility function simply returns the coded numeric values corresponding to the allele states.

#### Usage

```
make.marker.numeric(marker.frame, level.names=NULL)
```

# **Arguments**

marker.frame A data.frame.object consisting of factors or character vectors that encode the

allele states.

level.names A vector of length 6 to translate the levels attribute or character codes into allele

states that make.state.matrix understands. If necessary, dummy codes are

used to fill the vector.

#### Value

A matrix, for which column i is match(as.character(marker.frame[,i]),level.names)

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

make.regressor.matrix 31

make.regressor.matrix Create regressors using expected marker values

#### **Description**

Create regression variables for markers and loci between or near markers by imputation conditional on known marker states.

# Usage

```
make.regressor.matrix(state.matrix, mode.mat=NULL)
```

#### **Arguments**

state.matrix A state.matrix.object - see make.state.matrix for more details

mode.mat A matrix which indicates the values of regressor variables corresponding to the

allele states. If mode.mat=NULL (the default) a mode.mat is inferred from the dimensions of state.matrix. For the F2 intercross these are typically additive and dominance codes like (-1,0,1) and (1,-1,1). For BC1 backcross and RI lines,

typically the values are (-1,1).

#### Value

A matrix with variables suitable for use as regressors.

#### References

Haley C.S. and Knott S.A. (1992) A simple regression method for mapping quantitative trait loci in line crosses using flanking markers. *Heredity* **69**,315-324.

#### See Also

```
make.state.matrix
```

make.state.matrix

Create state.matrix.object

# **Description**

Create a state.matrix.object to be used encode marker information in a form in which it can be used in subsequent calculations.

# Usage

```
make.state.matrix(marker.frame, marker.distances, method="F2")
```

32 make.varcov

#### **Arguments**

marker.frame

Actually, this is a misnomer. This is NOT amarker.frame.object. Rather it is obtained by by call like make.marker.numeric(marker.frame.object) (see make.marker.numeric) and it is coerced to a matrix. It encodes marker allele states. One column is used for each marker or pseudo-marker (basically a placeholder with all missing values). The entries are in 1:6, if NA's are present, they are recoded to 6. The columns are arranged in linkage groups with presumed order reflected in the actual order of the columns.

marker.distances

Distances between the markers in the 'lambda' metric. -log(lambda)/2 is the Haldance map distance. Linkage groups are separated by values of 0.0.

method

method = "F2" is the default, and "BC1", "RI.self", and "RI.sib" are other options. The assumed setup is as follows (strains are A and a):

marker state	F2.code	BC.code	RI.code
"AA"	1	1	1
"Aa"	2	2	
"aa"	3		2
"A-" (not aa)	4		
"a-" (not AA)	5		
"-" (unknown)	6	6	6

#### Value

n by k by q array. q is 3 for method="F2" and 2 for others methods. Each element encodes the probability of the allele state conditional on the marker states.

#### Note

It might have been better to design this array so that the third subscript moves fastest. In large problems, the current structure may involve excessive memory access.

# References

Lander E.S. and Green P. (1987) Construction of multilocus genetic linkage maps in humans. *Proceedings of the National Academy of Sciences of the United States of America*, **84**(8), 2363–7.

Jiang C. and Zeng Z-B. (1997) Mapping quantitative trait loci with dominant and missing markers in various crosses from tow inbred lines. *Genetica* **101**, 47-58.

make.varcov	Create moment matrices

#### **Description**

Create a moment matrix of the marker variables and of the regressors by the phenotype variable. For use in regression modelling on the markers.

map.index 33

#### Usage

```
make.varcov(regressor.matrix, y, subset=is.finite(y), casewt=NULL)
```

#### **Arguments**

regressor.matrix

The object produced by make.regressor.matrix

y A vector of phenotype information with the same number of elements as there

are rows in regressor.matrix

subset Logical vector with the same number of elements as there are rows in regressor.matrix

to indicate which rows to keep.

casewt Optional vector of case weights.

#### Value

#### A list with components

var.x	Moment matrix of the marker regressor variables
cov.xy	Moment matrix of the marker regressor variables versus the phenotype variable
var.y	The Second central moment of the phenotype variable
df	sum(subset==TRUE) - 1

#### Note

It is generally NOT a good idea to do regressions on ill-conditioned designs using the moment matrices like this. The excuse for doing so here is twofold. First, calculations using this method are used to perform importance sampling, so minor numerical inaccuracies in computing the probabilites used in sampling get straightened out by the importance weights. Second, it will typically be the case that a prior is set on the regression coefficients and this results in a positive constant (aka a 'ridge' parameter) being added to diagonal of varcov\$var.x and this reduces the ill-conditioning. Of course the rational for using the method is to speed the sampling, and it is very effective at doing so.

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

map.index Look up numerical index(es) of map locations	
--	--

# **Description**

One way to index a locus (loci) in a genetic map is by the numerical index of its row (their rows). map.index performs a lookup in a specific map.frame given one (or two) chromosome number(s) and one (or two) map distance(s).

map.index

# Usage

```
map.index(x, ...)
```

#### **Arguments**

x A map.frame or analysis.object

For methods that look up a location in a map.frame the following named arguments may be used: chromo A chromosome number or 2 ordered numbers cM (Optional) map distance or two. If the same chromosome number is used twice in chromo, cM must be ordered. If cM is omitted, all loci on the chromosome will be included.

#### **Details**

It is often convenient to refer to genetic loci or regions by the numerical index(es) in a map.frame. map.index allows lookups according to the approximate map location.

#### Value

A numerical vector of one or more row numbers. If only chromo is specified, all row numbers on the specified chromosome are returned. If chromo has two elements, then all row numbers on those chromosomes with numbers in range(chromo) will be returned. If one of each of chromo and cM are specified, then the row number of the closest locus will be returned. For two of each, row numbers in the range of the closest matches will be returned.

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

#### See Also

make.map.frame for a description of how map information is organized.

# **Examples**

```
data(little.ana.bc)
map.index(little.ana.bc,chromo=1,cM=25) # locus nearest 1,25
index.chr.1 <- map.index(little.ana.bc,chromo=1)
fit.on.1 <- bqtl(bc.phenotype~locus(index.chr.1),little.ana.bc)
summary( loglik( fit.on.1 ) )</pre>
```

map.location 35

ap.location Report map location
---------------------------------

# **Description**

Report the chromosome number and location of loci in a genetic map.

# Usage

```
map.location(x,...)
map.loc(x, ...)
```

# **Arguments**

x A object of class map.frame,analysis.object, bqtl, or bqtl.list

Other arguments usage depend on the class of x: y A vector of row numbers or map.names specifying which subset of the map.frame of x is to be returned chromo: A vector of chromosome numbers cM (Optional) map distance vector. If the same chromosome number is used twice in chromo, cM must be ordered. If cM is omitted, all loci on each chromosome listed in chromo will be included.

map.namesA vector of map.names

# **Details**

It is often helpful to refer to genentic loci by their locations. The methods of map.location (alias map.loc) will extract the row index, chromosome number and location, and the name for specified loci. For direct lookups of the loci in a map.frame or analysis.object, one must specify y or chromo or map.names. When class(x)=="bqtl" map.locations of terms used in a call to bqtl are returned. When cM is used, an attempt will be made to match the location; if the match fails, the nearest locus will be used. When there are two elements in chromo and two in cM, all the map locations in between the matching loci will be returned.

#### Value

An object of class map.location which inherits from map. frame. It has columns:

```
chr.num The chromosome number

cM The location in centiMorgans on that chromosome.

marker.name The name by which that marker is known

attr(, "row.names")

An index of the locations
```

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

36 map.names

#### See Also

```
make.map.frame
```

# **Examples**

```
data(little.ana.bc)
map.loc(little.ana.bc, c(1,15,45))
map.loc(little.ana.bc,chromo=3,cM=22)
map.loc(little.ana.bc,"m.12")
rm(little.ana.bc)
```

map.names

Look up names of markers or loci

# Description

This is a generic helper function with methods that will return the names of markers or loci.

# Usage

```
map.names(x,...)
```

#### **Arguments**

x An object that has marker names in it. Methods for objects of the map.frame, analysis.object,bqtl, and bqtl.list class.

For class(x)=="analysis.object" or class(x)=="map.frame", arguments chromo and cM can be used as in map.index

#### **Details**

When applied to an object of class bqtl

```
map.names(x, \dots, ana.obj )
```

can be used to specify where to find the data.

# Value

A character vector

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

marker.fill 37

## See Also

```
map.index, map.location
```

## **Examples**

```
data(little.ana.bc)
map.names(little.ana.bc,chromo=1,cM=24)
map.names(little.ana.bc,chromo=c(1,1),cM=c(40,55))
fit <- bqtl( bc.phenotype ~ locus(23,42) , little.ana.bc )
map.names( fit )</pre>
```

marker.fill

Map Positions Between Markers

## **Description**

Given a set of markers, one wants to create a finer map at a given resolution. marker fill takes a a collection of marker distances and a desired resolution and finds positions that are intermediate and at that resolution.

## Usage

```
marker.fill(map.frame, reso, return.nint = FALSE)
```

## **Arguments**

map.frame A map.frame.object.

reso The desired interval between loci in the same metric as map.frame\$cM

return.nint Whether to output a vector of number of intervals to produce in each existing

interlocus interval

## Value

If return.nint is TRUE, a vector of integers is returned. It indicates how many intervals to place between this marker and the next to achive the desired minimum distance.

If return.nint is FALSE, a vector of distances is returned. The names attribute has suffixes added to indicate positions filled to the 'right' of existing markers. Thus if markers 'mark.01' and 'mark.02' are in succession at a distance of 3 and reso==1, then the value associated with 'mark.01' (which was 3) becomes 1, a value of 1 is associated with new loci called 'mark.01.1' and 'mark.01.2' in created with values of 1 each. The returned vector is ordered by chromosome, then marker or filled locus.

38 marker.levels

## See Also

```
make.map.frame
```

## **Examples**

```
data( little.map.frame )
little.nint <- marker.fill( little.map.frame, reso=5, TRUE )
cbind(nint=little.nint,cM=little.map.frame$cM)[1:10,]
rm( little.map.frame, little.nint )</pre>
```

marker.levels

Define marker level codes

## **Description**

The coding scheme used to define marker.levels is set up by these functions. *BQTL* has defaults that these functions can help the user to redefine.

# Usage

```
bc1.levels( AA="AA", Aa="Aa", miss.val="--")
ri.levels( AA="AA", aa="aa", miss.val="--")
f2.levels( AA="AA", Aa="Aa", aa="aa", not.aa="A-", not.AA="a-",
miss.val="--")
```

# Arguments

AA	Always used: the code for the homozygous state from one parent line
Aa	F2 and BC1 setups: the code for the heterozygous state
aa	F2 and RI setups: the code for the homozygous state for the other parent line
not.aa	F2 only: the code for a dominant marker that rules out aa
not.AA	F2 only: the code for a dominant marker that rules out AA
miss.val	The character string for a missing (unknown) allele state. NAs are automatically detected, so this is only needed if string values are used to denote missing values.

## **Details**

It is essential that the codes intended by the user be clearly understood by *BQTL*. It is hoped that thees functions provide a bridge between the internals of *BQTL* and the user's view of the marker codes. Numeric values can be used, but they will be coerced to character values.

## Value

A vector with 6 elements corresponding to the values of AA, Aa, aa, not.aa, not.AA, and miss.val. For RI and BC1 setups, those that do not apply will be unnamed and set to "nil"

plot.map.frame 39

## Author(s)

Charles C. Berry <cberry@ucsd.edu>

#### See Also

```
make.analysis.obj
```

# **Examples**

```
### show the defaults:
f2.levels()
bc1.levels()
ri.levels()

### suppose that 1,2,3 are codes used in F2:
f2.levels(1,2,3)

### show what would happen changing "Aa" to "H"
f2.levels(Aa="H")
bc1.levels(Aa="H")
```

plot.map.frame

plots by chromosome location

## Description

Multiple x-y plots are formed using chromosome numbers (chr.num) and positions (pos.plot) specified in a object of the sort created by make.map.frame

## Usage

#### **Arguments**

x A map.frame.object or an analysis.object

y (optional) A vector with as many elements or a matrix with as many rows as nrow(x). If omitted, a plot will be drawn in a single frame representing the chromosomes as horizontal bars and giving tic marks to show the locations markers and virtual markers (if any).

40 plot.map.frame

... more args

fun A plotting function to be used *after* the plot axes and labels have been drawn.

The current default if (y.type == "matrix") matlines else lines usually is

good enough. But a fancier function could be used for a fancier plot.

type "1" for lines, "p" for points, et cetera. see par

include.rug if TRUE place a tick on the x-axis at each marker location

rug.lwd size of ticks

title.string (optional) label to prepend to each title

y.range range for y limits

ylab plot label for y-axis, see par xlab plot label for x-axis, see par

## **Details**

This function enables drawing graphs that depend on chromosome and chromosome location. Typically, one will use a command like par(mfrow=c(nrows,ncols)) first to set up a page on which multiple plots will be drawn. However, one can draw one plot per page on postscript devices by leaving par(mfrow=c(1,1))

#### Value

NULL - this function is called only for its side effects

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

## See Also

plot, lines, and matlines for general information on plotting functions; par for optional arguments to add as arguments; and make.map.frame for the details on the object the drives this function.

## **Examples**

```
data( little.ana.bc )
null.llk <- loglik(bqtl(bc.phenotype~1,little.ana.bc))
llk <- loglik( bqtl( bc.phenotype~locus(all), little.ana.bc) ) - null.llk
.old.par <- par(mfrow=c(2,3))
plot.map.frame(little.ana.bc$map.frame,llk)
par(.old.par)</pre>
```

predict.bqtl 41

## **Description**

The estimated coefficients and expected locus values are used to find fitted values for the QTL model

# Usage

```
## S3 method for class 'bqtl'
predict(object, newdata, ...)
## S3 method for class 'bqtl'
fitted(object, newdata, ...)
```

## **Arguments**

object An object of class bqt1

newdata An optional data.frame for which fitted values are to be found. If not specified,

the a search for the original data frame for the fit will be made.

... unused

## Details

The estimated coefficients for a specific QTL model fit are used along with the expected locus values (conditionally on the marker values) are used to find fitted values for the QTL model. This is *not* the only way in which such fits could be obtained; one could condition the expect marker values on *both* the trait value and the marker values. One could also define fitted values for specific genotype combinations, e.g. for a backcross with k animals and a two gene model 4 fitted values could be determined for each animal leading to 2\*2\*k values. In fact, using newdata one can do this.

## Value

A vector with as many elements as rows in newdata (after removing missing data) or in the original model.frame.

# Author(s)

```
Charles C. Berry <cberry@ucsd.edu>
```

#### See Also

bqtl

42 predict.linear.bayes

## **Examples**

```
data(little.ana.bc)
fit.pheno <- bqtl(bc.phenotype~locus(15)+locus(42),little.ana.bc)
summary(predict(fit.pheno))
genotype.grid <- expand.grid( c(-1,1), c(-1,1) )  # set up a grid
names(genotype.grid) <- map.names( fit.pheno )  # use matching names
fit.vals <- predict( fit.pheno, genotype.grid )  # make predictions
cbind( genotype.grid, fit.vals )  # print them!</pre>
```

predict.linear.bayes Residuals or Predicted Values for linear.bayes objects

## **Description**

The linear bayes object returns fitted coefficients. These are used to construct predicted values. Since the fitting process for linear bayes objects is based on moments of centered variables, the 'intercept' is lost; see 'Details' below.

## Usage

## **Arguments**

object An object returned by linear.bayes
... possibly the following
newdata Optional data.frame in which to do the calculations
return.resids Not usually set by the user.

#### Details

Since the linear bayes object is based on a moment matrix, some information is lost that must be reconstructed or assumed. The intercept and possibly the coefficients for control variates are aong these. Also, when the call to linear bayes supplied the moment matrix rather than formulae with which to create one, then it is unclear what variable was used as the regressand and hence which variable to use in ofrming residuals. So, in that case, residuals will report an error

residuals.bqtl 43

## Value

A vector of predicted values or residuals

## Author(s)

```
Charles C. Berry <cberry@ucsd.edu>
```

#### See Also

```
linear.bayes
```

residuals.bqtl

Residuals from QTL models

# **Description**

The phenotype data, estimated coefficients, and expected locus values are used to find fitted values for the QTL model

## Usage

```
## S3 method for class 'bqtl'
residuals(object,...)
```

## **Arguments**

object An object of class bqt1

... ignored

## **Details**

The estimated coefficients for a specific QTL model fit are used along with the expected locus values (conditionally on the marker values) are used to find fitted values for the QTL model; these are subtracted from the original trait values to get residuals. This is *not* the only way in which such fits could be obtained; one could condition the expected marker values on *both* the trait value and the marker values. One could also define fitted values for specific genotype combinations, e.g. for a backcross with k animals and a two gene model 4 fitted values could be determined for each animal leading to 2\*2\*k residuals.

# Value

A vector with as many elements trait values used in the original fitted model.

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

44 summary.adj

## See Also

bqtl

# **Examples**

```
data(little.ana.bc)
fit.pheno <- bqtl(bc.phenotype~locus(15)+locus(42),little.ana.bc)
summary(residuals(fit.pheno))
plot( fitted( fit.pheno ), residuals( fit.pheno) )</pre>
```

summary.adj

Summarize Laplace approximations

## **Description**

The linear approximations of swap are much improved by the use a Laplace approximations for loci that are not markers. This function combines the results of a call like bqtl(y~configs(swap.obj),...) with the data in swap.obj to provide improved posteriors, et cetera

# Usage

```
## S3 method for class 'adj'
summary(object, n.loc, coef.znames, mode.names=c("add",
"dom"), imp.denom=NULL, swap.obj=NULL,...)
```

## **Arguments**

object	Typically, this is the result of a call like bqtl(y~configs(swap.obj),)
n.loc	The number of genes in this model
coef.znames	map.names for the sample space
mode.names	NULL except for "F2", in which case it is uusally c("add", "dom")
imp.denom	Optional, and only used when some sampling scheme other than the default $\ensuremath{MCMC}$ generates object
swap.obj	The result of a call to swap
	unused

# **Details**

There are a lot of details. This sections nneds to be revised to reflect them.

summary.bqtl 45

## Value

A list with components

adj This multiplier adjusts the posterior odds for k vs k-1 gene models

var An estimate of the variance of adj
coef Posterior means of coefficients

loc Marginal Posterior for location for k gene model

hk.ratio.mean argh! I need to look this up

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

## References

Berry C.C. (1998) Computationally Efficient Bayesian QTL Mapping in Experimental Crosses. *ASA Proceedings of the Biometrics Section*, 164-169.

summary.bqtl

Summarize bqtl object

# **Description**

Extract coefficients (and related stats), loglikelihood, and residual standard error of the trait.

# Usage

```
## S3 method for class 'bqtl'
summary(object,...)
```

## **Arguments**

object The result of link{bqtl}
... Currently not used

# Value

A list containing

coefficients Either a vector of regression coefficents, or if object was created via bqtl(..., return.hess=TRUE),

then a matrix with coefficients, standard errors, t-statistics, and p-values

loglik the loglikelihood or log posterior

std.res The residual standard deviation of the trait

N The counts of all observations, the number omitted, and the number used in the

fit

46 summary.map.frame

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

# **Examples**

```
data(little.ana.bc)
fit <- bqtl( bc.phenotype~locus(4)*locus(45), little.ana.bc,
return.hess=TRUE )
summary(fit)</pre>
```

summary.map.frame

Summary methods for basic data objects

# Description

Provide a simple report on the data structure

# Usage

```
## S3 method for class 'map.frame'
summary(object,...)
```

# Arguments

```
object A map.frame or analysis.object ... ignored
```

# Value

a list

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

summary.swap 47

summary.swap	Summarize Gibbs samples for a k-gene model	

# Description

Calculate marginal posteriors for location, posterior means for coefficients, and the Bayes Factor for k vs k-1 genes

# Usage

```
## S3 method for class 'swap'
summary(object, method=NULL, ncoef=length(object$alt.coef), nloc=object$nloc,...)
```

# Arguments

object	The result of swap
method	Optional. One of the supported methods, see make.analysis.obj
ncoef	Optional. The number of coefficients in the class of models. Typically, 2*nloc for method=="F2" and nloc for all other methods
nloc	Optional. The number of loci in the sample space.
	ignored

# Value

# A list with components:

loc.posterior	A vector of (marginal) posterior odds for each locus compared to a no gene model
coefs	Posterior means of coefficients.
ratio	A list with components mean, an estimate of the Bayes Factor for $k$ versus $k$ -1 gene models, and $var$ , an estimate of its variance.

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

48 swap

swa	р		

MCMC sampling of multigene models

# **Description**

Given a k-gene model as a starting point, one gene is deleted and another is sampled in its place. This is done using an approximation to the posterior. Then another gene is deleted and another sampled,...

## Usage

```
swap(varcov, invars, rparm, nreps, ana.obj, ...)
```

# **Arguments**

varcov	The result of make.varcov
invars	Vector of numerical indexes of ana.obj\$reg.names telling which variables to start in the model. The first of these is immediately removed, so it is merely a placeholder. The number of genes in the model is therefore k <- length(invars) (except when ana.obj\$method=="F2" when it is k <- length(unique(col(ana.obj\$reg.names)[invars])
rparm	Scalar or vector with nrow(varcov\$var.x) elements; the 'ridge' parameters for the independent variables - larger values imply more shrinkage or a more concentrated prior for the regresion coefficients.
nreps	How many cycles (of k samples each) to perform.
ana.obj	An analysis.object — see make.analysis.obj
	Additional arguments override the default choices of candidate loci (locs), prior for locus (locs.prior), or method specified by ana.obj. Also, the default prior for model (combo.prior) when ana.obj\$method=="F2" can be overridden.  See swapbc1 and swapf2 for details.

## **Details**

An MCMC sampler for loci using the object of make.varcov is executed. This sampler uses the exact posterior probability under the assumed correctness of the regression model using expected genotypes given marker values. This amounts to linearizing the likelihood with respect to the (possibly unknown) locus states. For models where the loci are fully informative markers this is the true posterior.

The chain is implemented as follows: given a set of regressor variables to start, one variable is removed, all regressor variables not in the model are examined to determine the effect of each on the posterior. One variable is sampled. The process is repeated until each variable has been removed and a new one sampled in its place (possibly the same variable that was removed is sampled). And this whole cycle is repeated nreps times.

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#### Value

A list with components:

config A k by k by nreps array (or, for ana.obj\$method=="F2", a 2k by k by nreps

array) of the locations (variables) sampled in each iteration.

posteriors A vector of length k\*nreps with the posteriors of the models.

coefs A k by k matrix of the regression coefficients(or, for ana.obj\$method=="F2",

a 2k by nreps matrix).

call The call to swap

cond The k\*nreps posterior probabilities of the k-1 gene models.

marg The k\*nreps marginal posteriors for all k gene models that could be formed

using the current k-1 gene model

alt.marginal A vector with length(locs) (or 2\*length(locs)) elements. At each step, the

posterior associated with each candidate locus is added to an element of this vector. After all steps are finished, the result is normalized to sum to one. This

turns out to be a stable estimate of the marginal posterior.

alt.coef A vector with length(locs) (or 2\*length(locs)) elements. At each step, the

product of each posterior times the coefficient(s) associated with a candidate locus is added to an element of this vector. After all steps are finished, the result is normalized by the total marginal posterior. This turns out to be a stable estimate of the marginal (over all models) posterior mean of the regression coefficients.

## Author(s)

Charles C. Berry <cberry@ucsd.edu>

## References

Berry C.C. (1998) Computationally Efficient Bayesian QTL Mapping in Experimental Crosses. *ASA Proceedings of the Biometrics Section*, 164-169.

## **Examples**

```
data( little.ana.bc )
little.vc <- varcov( bc.phenotype~locus(all), little.ana.bc)
little.4 <- swap( little.vc, c(1,15,55,75), rparm=1, 50, little.ana.bc )
little.4.smry <- summary( little.4 )
print(c("Bayes Factor (3 vs 4)"=little.4.smry$ratio$mean))
par(mfrow=c(3,2))
plot( little.ana.bc, little.4.smry$loc.posterior, type="h",
    ylab="E(genes)" )
rm(little.4,little.vc,little.ana.bc)</pre>
```

50 swapbc1

swapbc1	Sample BC1 or Recombinant Inbred loci via approximate posterior.

## **Description**

An MCMC sampler for loci using precomputed dispersion matrices, various priors, and a preselected set of variables. For use with BC1 (backcross) designs and recombinant inbred lines.

# Usage

```
swapbc1(varcov, invars, rparm, nreps, ana.obj, locs=NULL,
locs.prior=NULL, tol=1e-10)
```

## **Arguments**

varcov	The result of make.varcov
rparm	Scalar or vector with nrow(varcov\$var.x) elements; the 'ridge' parameters for the independent variables - larger values imply more shrinkage or a more concentrated prior for the regresion coefficients.
nreps	How many cycles of MCMC to perform
ana.obj	A object produced by make.analysis.obj
invars	Which variables to start in the model. The first of these is immediately removed, so it is merely a placeholder. The number of genes in the model is therefore k <- length(invars)
locs	The columns of varcov\\$var.x to use. The default uses all of them.
locs.prior	The prior mass to associate with each variable. Typically, these sum to one, but sometimes they might each be set to one (as in computing lod scores).
tol	Used in forming QR decomposition. Let it be.

# **Details**

An MCMC sampler for loci using the object of make.varcov is executed. This sampler uses the exact posterior probability under the assumed correctness of the regression model using expected genotypes given marker values. This amounts to linearizing the likelihood with respect to the (possibly unknown) locus states. For models where the loci are fully informative markers this is the true posterior.

The chain is implemented as follows: given a set of regressor variables to start, one variable is removed, all regressor variables not in the model are examined to determine the effect of each on the posterior. One variable is sampled. The process is repeated until each variable has been removed and a new one sampled in its place (possibly the same variable that was removed is sampled). And this whole cycle is repeated nreps times.

swapf2 51

#### Value

A list with components:

config A k by k by nreps array of the locations sampled in each iteration.

posteriors A vector of length k\*nreps with the posteriors of the models.

coefs A k by k matrix of the regression coefficients.

call The call to swapbc1

cond The k\*nreps posterior probabilities of the k-1 gene models.

marg The k\*nreps marginal posteriors for all k gene models that could be formed

using the current k-1 gene model

alt.marginal A vector with length(locs) elements. At each step, the posterior associated

with each candidate locus is added to an element of this vector. After all steps are finished, the result is normalized to sum to one. This turns out to be an

exceedingly stable estimate of the marginal posterior.

alt.coef A vector with length(locs) elements. At each step, the product of each poste-

rior times the coefficient associated with a candidate locus is added to an element of this vector. After all steps are finished, the result is normalized by the total marginal posterior. This turns out to be an exceedingly stable estimate of the

marginal (over all models) posterior mean of the regression coefficients.

#### Author(s)

Charles C. Berry <cberry@ucsd.edu>

# References

Berry C.C. (1998) Computationally Efficient Bayesian QTL Mapping in Experimental Crosses. *ASA Proceedings of the Biometrics Section*, 164-169.

#### See Also

swapf2

swapf2 Sample F2 loci via approximate posterior

## **Description**

An MCMC sampler for loci using precomputed dispersion matrices, various priors, and a preselected set of variables. For use with F2 intercross design.

Using precomputed dispersion matrices, various priors, and a pre-selected set of variables, one locus is removed, all other loci are examined to determine the effect of each on the posterior. One locus is sampled. The process is repeated until each locus has been removed and a new one sampled in its place (possibly the same one that was removed is sampled).

52 swapf2

## Usage

```
swapf2(varcov, invars, rparm, nreps, ana.obj, locs,
      locs.prior, combo.prior, tol = 1e-10)
```

#### **Arguments**

varcov The result of make.varcov. The columns of varcov\$var.x must alternate 'ad-

ditive' and 'dominance' terms.

rparm The 'ridge' parameters for the independent variables - larger values imply more

shrinkage or a more concentrated prior for the regresion coefficients.

nreps How many cycles of MCMC to perform
ana.obj A object produced by make.analysis.obj

invars A vector of variable indexes. This determines which variables to start in the

model. If both additive and dominance terms are to be used, they should occupy adjacent locations in invars. The variable(s) associated with the first locus is (are) immediately removed, serving only as placeholder(s). If there are k loci associated with the variables, then all subsequent models have k loci, although the nuimber of variables may vary according to the selection of one or both of

the 'additive' or 'dominance' terms.

locs The pairs of columns of varcov\$var.x to use. The default uses all of them.

locs prior Vector whose elements are the prior masses to associate with each locus. Typ-

ically, these sum to one, but sometimes they might each be set to one (as in

computing lod scores). The default value sets them all to 1.0.

combo.prior The prior probability for each term or combination of terms for the phenotypic

effect at a locus. Typically, there will be three of these - one for the 'additive' term (linear in number of alleles from one parent strain), the 'dominance' term (quadratic in allele number), or both terms. The default sets them all to 1/3.

tol Used in forming QR decomposition. Let it be.

## **Details**

A call to swapf2 is used to obtain the results. This function is really just a wrapper.

## Value

A list with components:

configs A 2k by k by nreps array of indexes of variables sampled in each of the nreps

iterations. Models using less than 2k variables configs[,i,j] will contain one

or more zeroes in the last position(s)

posteriors A vector of length k\*nreps with the posteriors of the models sampled.

coefs A 2k by k by nreps matrix of the regression coefficients. Models using less

than 2k variables configs[,i,j] will contain one or more zeroes in the last

position(s)

call The call to swapf2

twohk 53

cond The k\*nreps posterior probabilities of the k-1 gene models.

marg The k\*nreps marginal posteriors for all k gene models that could be formed

using the current k-1 gene model)

alt.marginal A vector with length(locs) elements. At each step, the posterior associated

with each candidate locus is added to an element of this vector. After all steps are finished, the result is normalized to sum to one. This turns out to be an

exceedingly stable estimate of the relative marginal posterior.

alt.coef A vector with 2\*length(locs) elements. At each step, the product of each

posterior times the coefficient associated with a candidate variable is added to an element of this vector. After all steps are finished, the result is normalized by the total marginal posterior. This turns out to be an rather stable estimate of the

marginal (over all models) posterior mean of the regression coefficients.

## Author(s)

Charles C. Berry <cberry@ucsd.edu>

#### References

Berry C.C. (1998) Computationally Efficient Bayesian QTL Mapping in Experimental Crosses. *ASA Proceedings of the Biometrics Section*, 164-169.

## See Also

swapbc1

twohk

One and Two Gene Models Using Linearized Posterior

## **Description**

Fits all one and two gene models (without interactions aka 'epistasis') in an intercross, backcross, or recombinant inbred line. Uses a linear approximation to the likelihood, i.e. the expected allele states are used.

## Usage

```
twohk(varcov, ana.obj, ...)
```

## **Arguments**

varcov An object produced by make.varcov

ana.obj An analysis.object — see make.analysis.obj

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. . .

Additional arguments override the default choices of candidate loci (locs), prior for locus (locs.prior), or method specified by ana.obj: locs A vector indexing the loci to use. locs.prior The prior mass to associate with each locus. Typically, these sum to one, but sometimes they might each be set to one (as in computing lod scores). combo.prior Only valid for ana.obj\$method=="F2". The prior probability for each term or combination of terms for the phenotypic effect at a locus. Typically, there will be three of these - one for the 'additive' term (linear in number of alleles from one parent strain), the 'dominance' term (quadratic in allele number), or both terms. The default sets them all to 1/3.

## **Details**

The marginal posterior (integrating over regression parameters and dispersion) is calculated for each one and two gene model under the assumed correctness of the regression model using expected genotypes given marker values. This amounts to linearizing the likelihood with respect to the (possibly unknown) locus states. For models where the loci are fully informative markers this is the true posterior.

#### Value

A list with components:

loc.1	The marginal posterior for each one gene model relative to a no gene model. For
	twohkf2 this is a matrix of 3 columns; the first for models with additive terms,
	the second for dominance terms, and the third for both. The sum over all three
	columns yields the marginal posterior for the locus.

- The marginal posterior for each locus obtained by summing over all two gene models that include that locus—relative to a no gene model. For twohkf2 this is a matrix of 3 columns; the first for models with additive terms, the second for dominance terms, and the third for both.
- coefs.1 The regression coefficients for the genetic effect for each locus. For twohkf2, this is a matrix with two rows; the first is for the 'additive effect' and the second is for the 'dominance' effect.
- coefs.2 The marginal posterior mean of regression coefficients for the genetic effect for each locus obtained by averaging over all two gene models that include that locus according to the posterior masses. For twohkf2, this is a matrix with two rows; the first is for the 'additive effect' and the second is for the 'dominance' effect.

# Author(s)

Charles C. Berry <cberry@ucsd.edu>

## References

Haley C.S. and Knott S.A. (1992) A simple regression method for mapping quantitative trait loci in line crosses using flanking markers. *Heredity* **69**,315-324.

twohkbc1 55

## **Examples**

```
data(little.ana.bc)
little.vc<-make.varcov(little.ana.bc$data[,little.ana.bc$reg.names],
    little.ana.bc$data$bc.phenotype)
little.2<- twohk(little.vc,little.ana.bc,rparm=1)
print( c(odds.1=sum(little.2$loc.1),odds.2=sum(little.2$loc.2)) )
par(mfrow=c(3,2))
little.pe <- 2 * little.2$loc.2 / sum(little.2$loc.2)  #locus-wise posterior expectation
plot(little.ana.bc,little.pe,type="h",ylab="E(genes")
rm(little.2,little.vc,little.pe,little.ana.bc)</pre>
```

twohkbc1

One and Two Gene Models Using Linearized Posterior

## **Description**

Fits all one and two gene models (without interactions aka 'epistasis') in an intercross, backcross, or recombinant inbred line. Uses a linear approximation to the likelihood, i.e. the expected allele states are used.

## Usage

## **Arguments**

varcov	An object produced by make.varcov
ana.obj	An object produced by make.analysis.obj
rparm	The 'ridge' parameters for the independent variables - larger values imply more shrinkage or a more concentrated prior for the regresion coefficients.
locs	The columns (or pairs of columns for twohkf2) of $\cvent{var.x}$ to use. The default uses all of them.
locs.prior	The prior mass to associate with each locus. Typically, these sum to one, but sometimes they might each be set to one (as in computing lod scores).
combo.prior	Only valid for twohkf2. The prior probability for each term or combination of terms for the phenotypic effect at a locus. Typically, there will be three of these - one for the 'additive' term (linear in number of alleles from one parent strain), the 'dominance' term (quadratic in allele number), or both terms. The default sets them all to 1/3.

56 varcov

# **Details**

The marginal posterior (integrating over regression parameters and dispersion) is calculated for each one and two gene model under the assumed correctness of the regression model using expected genotypes given marker values. This amounts to linearizing the likelihood with respect to the (possibly unknown) locus states. For models where the loci are fully informative markers this is the true posterior.

#### Value

A list with components:

loc.1	The marginal posterior for each one gene model. For twohkf2 this is a matrix of 3 columns; the first for models with additive terms, the second for dominance terms, and the third for both. The sum over all three columns yields the marginal posterior for the locus.
loc.2	The marginal posterior for each locus - obtained by summing over all two gene models that include that locus. For twohkf2 this is a matrix of 3 columns; the first for models with additive terms, the second for dominance terms, and the third for both.
coefs.1	The regression coefficients for the genetic effect for each locus. For twohkf2, this is a matrix with two rows; the first is for the 'additive effect' and the second is for the 'dominance' effect.
coefs.2	The marginal posterior mean of regression coefficients for the genetic effect for each locus - obtained by averaging over all two gene models that include that

each locus - obtained by averaging over all two gene models that include that locus according to the posterior masses. For twohkf2, this is a matrix with two rows; the first is for the 'additive effect' and the second is for the 'dominance' effect.

## Author(s)

Charles C. Berry <cberry@ucsd.edu>

#### References

Haley C.S. and Knott S.A. (1992) A simple regression method for mapping quantitative trait loci in line crosses using flanking markers. *Heredity* **69**,315-324.

varcov	Create moment matrices

# **Description**

Create a moment matrix of the marker variables and of the regressors by the phenotype variable. For use in regression modelling on the markers.

varcov 57

## Usage

```
varcov(x, ana.obj, partial=NULL, scope,...)
```

## **Arguments**

X	A formula to specify the dependent and independent variables to be used in subsequent calculations e.g trait $\sim$ locus(.)
ana.obj	An analysis.object, seemake.analysis.obj
partial	A formula whose right hand side specifies variables to be treated as covariates.
scope	Usually not explicitly used. Optional vector of variable names.
	ignored

## **Details**

This is just a wrapper for make.varcov.

#### Value

A list with components

var.x	Moment matrix of the marker regressor variables
cov.xy	Moment matrix of the marker regressor variables versus the phenotype variable
var.y	The Second central moment of the phenotype variable
df	The degrees of freedom, when no variables are specified in partial it is sum(subset==TRUE)
	_ 1

## Note

It is generally NOT a good idea to do regressions on ill-conditioned designs using the moment matrices. The excuse for doing so here is twofold. First, calculations using this method are used to perform importance sampling, so minor numerical inaccuracies in computing the probabilites used in sampling get straightened out by the importance weights. Second, it will typically be the case that a prior is set on the regression coefficients and this results in a positive constant (aka a 'ridge' parameter) being added to diagonal of varcov()\$var.x and this reduces the ill-conditioning. Of course the rational for using the method is to speed the sampling, and it is very effective at doing so.

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## See Also

The examples in swap and twohk.

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