Package 'DOQTL'

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Description DOQTL is a quantitative trait locus (QTL) mapping pipeline designed for Diversity Outbred mice and other multi-parent outbred populations. The package reads in data from genotyping arrays and perform haplotype reconstruction using a hidden Markov model (HMM). The haplotype probabilities from the HMM are then used to perform linkage mapping. When founder sequences are available, DOQTL can use the haplotype reconstructions to impute the founder sequences onto DO genomes and perform association mapping.
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R topics documented:
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 ${\sf add.missing.F1s}$

Add Missing F1 Samples

Description

Given a set of CC or DO founders, impute the genotypes or intensities of missing F1s.

Usage

```
add.missing.F1s(founders, snps)
```

Arguments

founders List, required: Contains either an element called 'geno' or two elements called

'x' and 'y'.

snps Data.frame, required: Data.frame containing the SNPs. SNP ID, chr, Mb and

cM locations in columna 1 - 4, respectively.

Value

List with the founders data structure updated to include missing FALSE1 samples.

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Author(s)

Daniel Gatti

Examples

```
## Not run:
    load(url("ftp://ftp.jax.org/MUGA/muga_snps.Rdata"))
    founders = add.missing.FALSE1s(founders, snps = muga_snps)
## End(Not run)
```

add.slash

Add a forward slash to a character string.

Description

If the argument does not end with a forward slash, add one.

Usage

```
add.slash(path)
```

Arguments

path

Character string containing a file path.

Value

Returns a character string with a forward slash added to the end, if the argument did not end with a forward slash already.

Author(s)

Daniel Gatti

```
add.slash("/dir")
```

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addLog

Add two log values.

Description

When two numbers that are on the log scale must be summed, transforming them back to the non-log scale and taking the sum is slow. This function takes the exp() of the values only when necessary.

Usage

```
addLog(x, y)
```

Arguments

x Numeric containing a value on the log scale.

y Numeric containing a value on the log scale.

Details

This function checks to see if the difference between the maximum value and the other value is less than the machine precision. If it is, then the exp() is taken for those values that differ by less than the machine precision, they are summed and returned to a log scale. If the maximum value is differs from the other values by greater than the machine precision, then return the maximum value.

Value

Numeric value containing the sum of the arguments on a log scale.

Author(s)

Daniel Gatti

See Also

```
addLogVector
```

```
addLog(log(10), log(1))
```

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 ${\it add} Log Vector \\$

Add a vector of log values.

Description

When a summation must be taken on a vector of numbers that are on the log scale, transforming them back to the non-log scale and taking the sum is slow. This function takes the exp() of the values only when neccessary.

Usage

```
addLogVector(x)
```

Arguments

Χ

Numeric vector containing values on a log scale to be summed on the untransformed scale.

Details

This function checks to see if the difference between the maximum values and the remaining values is less than the machine precision. If it is, then the exp() is taken for those values that differ by less than the machine precision, they are summed and returned to a log scale. If the maximum value is differs from the other values by greater than the machine precision, then return the maximum value.

Value

Numeric value containing the sum of the values on a log scale.

Author(s)

Daniel Gatti

See Also

 addLog

```
addLogVector(log(1:10))
```

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assoc.map	Perform association mapping on DO mice.	

Description

Given the phenotypes and genotype probabilities, impute the Sanger SNPs onto DO genomes and perform association mapping.

Usage

Arguments

pheno	Data.frame containing the phenotype data. Sample IDs must be in rownames. One of the columns should be called sex and contain M or FALSE to indicate the sex of each sample.
pheno.col	Either a numeric vector containing column IDs to map in pheno or a vector of column names in pheno.
probs	A 3 dimensional array of genotype probabilities as provided from condense.model.probs Samples, founder and markers in dims 1:3. All dimensions must have dimnames.
K	Numeric matrix containing kinship values for the samples in pheno and probs. Sample IDs must be in rownames and colnames.
addcovar	Numeric matrix of additive covariates to use in mapping. Sample IDs must be in rownames.
snps	Data.frame containing marker IDs, chromosomes, Mb and cM locations in columns 1:4.
chr	Character containing the chromosome on which to map.
start	Numeric value containing the proximal position for mapping. May be in Mb or base pairs (see Details).
end	Numeric value containing the distal position for mapping. May be in Mb or base pairs (see Details).
model	Character string that is one of "additive", "dominance" or "full". Indicates the type of model to fit. Note that the probs must match the type of model being fit. See condense.model.probs to output different probs.

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Character string that is either "one", or "two" indicating whether a single scan or a pairwise scan should be performed across the interval.

Output

Character string that is either "lod", "p-value" or "bic" indicating the mapping statistic to return.

snp.file

Character string containing the full path to the SNP file to use. Currently points to a location on the Jackson Laboratory FALSETP server

Integer indicating the number of permutations to perform. Default = 1000.

Details

FALSEor each interval between two markers, we take the average founder haplotype contribution for each sample. Then, using the proportion of each founder (8 in the case of DO mice), we impute the Sanger SNPs in this interval onto each DO sample.

The start and ending locations are assumed to be in Mb if they are below 200 and in bp if over 200.

Value

Data.frame containing the locations, SNPs and mapping statistic for the requested samples in the requested interval.

Author(s)

Daniel Gatti

References

Combined sequence-based and genetic mapping analysis of complex traits in outbred rats. Rat Genome Sequencing and Mapping Consortium, Baud A, Hermsen R, Guryev V, Stridh P, Graham D, McBride MW, FALSEoroud T, Calderari S, Diez M, Ockinger J, Beyeen AD, Gillett A, Abdelmagid N, Guerreiro-Cacais AO, Jagodic M, Tuncel J, Norin U, Beattie E, Huynh N, Miller WH, Koller DL, Alam I, FALSEalak S, Osborne-Pellegrin M, Martinez-Membrives E, Canete T, Blazquez G, Vicens-Costa E, Mont-Cardona C, Diaz-Moran S, Tobena A, Hummel O, Zelenika D, Saar K, Patone G, Bauerfeind A, Bihoreau MT, Heinig M, Lee YA, Rintisch C, Schulz H, Wheeler DA, Worley KC, Muzny DM, Gibbs RA, Lathrop M, Lansu N, Toonen P, Ruzius FALSEP, de Bruijn E, Hauser H, Adams DJ, Keane T, Atanur SS, Aitman TJ, FALSElicek P, Malinauskas T, Jones EY, Ekman D, Lopez-Aumatell R, Dominiczak AFALSE, Johannesson M, Holmdahl R, Olsson T, Gauguier D, Hubner N, FALSEernandez-Teruel A, Cuppen E, Mott R, FALSElint J. Nat Genet. 2013 Jul;45(7):767-75. doi: 10.1038/ng.2644. Epub 2013 May 26. PMID: 23708188 Using progenitor strain information to identify quantitative trait nucleotides in outbred mice. Yalcin B, FALSElint J, Mott R. Genetics. 2005 Oct;171(2):673-81. Epub 2005 Aug 5. PMID: 16085706 Mouse genomic variation and its effect on phenotypes and gene regulation. Keane TM, Goodstadt L, Danecek P, White MA, Wong K, Yalcin B, Heger A, Agam A, Slater G, Goodson M, FALSEurlotte NA, Eskin E, Nellaker C, Whitley H, Cleak J, Janowitz D, Hernandez-Pliego P, Edwards A, Belgard TG, Oliver PL, McIntyre RE, Bhomra A, Nicod J, Gan X, Yuan W, van der Weyden L, Steward CA, Bala S, Stalker J, Mott R, Durbin R, Jackson IJ, Czechanski A, Guerra-Assuncao JA, Donahue LR, Reinholdt LG, Payseur BA, Ponting CP, Birney E, FALSElint J and Adams DJ Nature 2011;477;7364;289-94 PUBMED: 21921910 Sequence-based characterization of structural variation in the mouse genome. Yalcin B, Wong K, Agam A, Goodson M, Keane TM, Gan X, Nellaker assoc.plot 9

C, Goodstadt L, Nicod J, Bhomra A, Hernandez-Pliego P, Whitley H, Cleak J, Dutton R, Janowitz D, Mott R, Adams DJ and FALSElint J Nature 2011;477;7364;326-9 PUBMED: 21921916

See Also

```
assoc.plot
```

Examples

assoc.plot

Plot association mapping results.

Description

After performing association mapping using assoc.map, plot the mapping statistic along with genes in the QTL interval.

Usage

```
assoc.plot(results,
mgi.file = "ftp://ftp.jax.org/SNPtools/genes/MGI.20130703.sorted.txt.gz",
highlight, highlight.col = "red", thr, show.sdps = FALSE, ...)
```

Arguments

results	Data.frame containing output from assoc.map.
mgi.file	Character string containing the full path to a Tabix indexed file of gene locations. Default points to a version of the MGI genome feature file.
highlight	Character vector containing gene symbols to highlight in the plot.
highlight.col	Vector of colors to use when highlighting genes.
thr	Numeric value above which data points should be colored red and SNPs with these points returned.
show.sdps	Logical value (default = FALSE) that is TRUE if the strain distribution pattern (SDP) for the SNPs should be shown. When used with thr, only plots the founder SDPs for SNPs above thr.
	Additional arguments passed to plot.

Details

Given the output from assoc.map, plot the LOD or difference in BIC values across the QTL interval in the top panel. Plot the genes in the interval in the lower panel. Make sure to use Sanger SNP and MGI feature files that are on the same genome build.

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Value

A plot with the mapping statistic in the top panel and genes in the lower panel. If thr is not missing, then filter the SNPs in the results argument and return only those with a mapping statistic greater than thr.

Author(s)

Daniel Gatti

See Also

```
assoc.map
```

Examples

```
## Not run:
    results = assoc.map(pheno = pheno, pheno.col = 1, probs = probs, K = K, addcovar = addcovar,
snps = snps, chr = 1, start = 40, end = 45)
    assoc.plot(results, thr = 3, show.sdps = TRUE)
## End(Not run)
```

batch.normalize

Batch normalize the X & Y intensity data.

Description

This function batch normalizes the X & Y intensity data by subtracting batch medians from the X & Y intensities.

Usage

```
batch.normalize(path = ".", snps)
quantilenorm(x1, y1, x2, y2)
```

Arguments

path	Character, the full path to the input files, which must be either "x.txt" and "y.txt" or "x.filt.txt" and "y.filt.txt".
snps	Data.frame, with three columns containing SNP ID, chromosome and Mb location in that order. May be obtained from ftp://ftp.jax.org/MUGA .
x1	Numeric matrix containing X intensities for batch 1 containing samples in rows and markers in columns. Number of samples should be larger than x2.
y1	Numeric matrix containing Y intensities for batch 1 containing samples in rows and markers in columns. Number of samples should be larger than y2.
x2	Numeric matrix containing X intensities for batch 2 containing samples in rows and markers in columns.

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y2

Numeric matrix containing Y intensities for batch 2 containing samples in rows and markers in columns.

Details

quantile.norm adjusts the intensities of samples in batch 2 to those of batch 1. The number of samples in batch 1 should be greater than the number of samples in batch 2. At each SNP, we form quantiles of the X1 (or Y1) intensity distribution, discarding the upper and lower 0.01

Value

FALSEor batch.normalize: returns value is returned. The batch normalized intensities are written to "x.filt.batch.norm.txt" and "y.filt.batch.norm.txt".

FALSEor quantilenorm: returns normalized X and Y values for batch 2.

Note

FALSEuture releases may include more sophisticated normalization algorithms.

Author(s)

Daniel Gatti

See Also

```
extract.raw.data, filter.samples
```

Examples

```
## Not run:
    load(url("ftp://ftp.jax.org.MUGA/muga_snps.Rdata"))
    batch.normalize(path = "/demo/MUGA/", snps = muga_snps)
## End(Not run)
```

bayesint

FALSEind a Bayesian Credible Interval around a QTL.

Description

This function normalizes the area under the QTL curve on the given chromosome and finds a region that is 95

Usage

```
bayesint(qtl, chr, prob = 0.95, expandtomarkers = FALSE)
```

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Arguments

qt1 data.frame: four columns with SNP ID, Chr, position, and LOD in each column.

chr character: the chromosome on which the QTL lies.

prob numeric: must be between 0 and 1

expandtomarkers

boolean: if TRUE, expand the QTL interval to the nearest flanking markers.

Default = FALSEALSE.

Value

Data frame with the SNP ID, Chr, position and LOD for the left and right side of the interval and the maximum QTL.

Author(s)

Daniel Gatti

References

Saunak, S. (2001) A Statistical FALSEramework for Quantitative Trait Mapping. *Genetics*, **159** (1), 371–387.

See Also

```
scanone, scanone.perm
```

Examples

```
## Not run: bayesint(qtl, 1)
```

calc.genoprob

Calculate the founder genotype probabilities at each SNP.

Description

This function performs genome reconstruction using either allele calls or allele intensities. We recommend using allele intensities where available because they often produce better genotype reconstructions.

Usage

```
calc.genoprob(data, chr = "all", output.dir = ".", plot = TRUE,
array = c("megamuga", "muga", "other"),
sampletype = c("DO", "CC", "DOF1", "other"), method = c("intensity", "allele"),
founders, transprobs, snps)
```

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Arguments

data

A list with named elements containing the information needed to reconstruct genomes.

When method = intensity: x: Numeric matrix, num.samples x num.snps, with X intensities for all samples. Sample IDs and SNP IDs must be in rownames and colnames. y: Numeric matrix, num.samples x num.snps, with Y intensities for all samples. Sample IDs and SNP IDs must be in rownames and colnames. sex: Character vector, containing "M" or F indicating sex. Sample IDs must be in names. gen: Character matrix containing the generation of DO outbreeding for each sample. For the DO, this should be "DO" followed by a number with no space between them. For CC mice, this should be CC. Sample IDs must be in names.

When method = allele: geno: Character matrix, num.samples x num.snps, with allele calls (A,C,G,T,H or N) for all samples. Sample IDs and SNP IDs must be in rownames and colnames. sex: Character vector, containing "M" or F indicating sex. Sample IDs must be in names. gen: Character matrix containing the generation of DO outbreeding for each sample. For the DO, this should be "DO" followed by a number with no space between them. For CC mice, this should be CC. Sample IDs must be in names.

chr

Character vector containing chromosomes to run. Must match the chromosome IDs in the snps table. "all" (default) will run all chromosomes.

output.dir

Character string containing the full path where output should be written. The directory must exist already.

plot

Boolean that is true if the user would like to plot a sample chromosome as the model progresses. Default = TRUE.

array

Character string indicating the array type. Must be one of "megamuga", "muga" or "other". Default equals "megamuga".

sampletype

Character string containing the type of samples being run. Must be one of "DO", "CC", "DOF1", or "other". Default equals "DO".

method

Character string containing method of genome reconstruction. Must be one of "intensity" or "allele". Default equals "intensity".

founders

List containing founder information for non-DO or CC crosses. *Not required for DO*.

When method = intensity: x: Numeric matrix, num.samples x num.snps, with X intensities for all founders and F1s (if available). Sample IDs and SNP IDs must be in rownames and colnames. y: Numeric matrix, num.samples x num.snps, with Y intensities for all founders and F1s (if available). Sample IDs and SNP IDs must be in rownames and colnames. sex: Character vector, containing "M" or "F" indicating sex. Sample IDs must be in names. code: Character vector containing two letter genotype codes for each founder sample. Sample IDs must be in names.

When method = allele: geno: Character matrix, num.samples x num.snps, with allele calls for all founders and F1s (if available). Sample IDs and SNP IDs must be in rownames and colnames. sex: Character vector, containing "M" or "F" indicating sex. Sample IDs must be in names. code: Character vector

calc.genoprob.alleles

containing two letter genotype codes for each founder sample. Sample IDs must be in names.

When sampletype = DOF1 x: Numeric matrix, num.samples x num.snps, with X intensities for all founders and F1s (if available). Sample IDs and SNP IDs must be in rownames and colnames. y: Numeric matrix, num.samples x num.snps, with Y intensities for all founders and F1s (if available). Sample IDs and SNP IDs must be in rownames and colnames. sex: Character vector, containing "M" or "F" indicating sex. Sample IDs must be in names. code: Character vector containing two letter genotype codes for each founder sample. This should be "II" for an inbred mutant strain. Sample IDs must be in names. direction: Character string that is either "DOxMUT" if a female DO was crossed with a male mutant mouse or "MUTxDO" if a female mutant mouse was crossed with a male DO. This affects the genotyping of teh X chromosome.

transprobs

Function to call to estimate the transition probabilities between markers for non-DO samples. *Not required for DO*.

snps

Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM locations in columns 1 through 4, respectively. *Not required for DO*.

Value

No value is returned. The output files are written to output.dir.

Author(s)

Daniel Gatti

Examples

```
## Not run:
    calc.genoprob(cross, chr.to.run = 1:19, output.dir = "do.data", plot = FALSE,
init.means = NULL, init.covars = NULL)
## End(Not run)
```

calc.genoprob.alleles Calculate the founder genotype probabilities at each SNP using allele calls.

Description

This function performs genome reconstruction using allele calls. We recommend using allele intensities where available because they often produce better genotype reconstructions.

Usage

```
calc.genoprob.alleles(data, chr, founders, snps, output.dir = ".",
trans.prob.fxn = do.trans.probs, plot = FALSE)
```

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Arguments

data A list with named elements containing the information needed to reconstruct

genomes.

geno: Character matrix, num.samples x num.snps, with allele calls (A,C,G,T,H or N) for all samples. Sample IDs and SNP IDs must be in rownames and colnames. sex: Character vector, containing "M" or F indicating sex. Sample IDs must be in names. gen: Character matrix containing the generation of DO outbreeding for each sample. For the DO, this should be "DO" followed by a number with no space between them. For CC mice, this should be CC. Sample

IDs must be in names.

chr Character vector containing chromosomes to run. Must match the chromosome

IDs in the snps table. "all" (default) will run all chromosomes.

founders List containing founder information for non-DO or CC crosses. When method

= allele: geno: Character matrix, num.samples x num.snps, with allele calls (A,C,G,T,H or N) for all founders and FALSE1s (if available). Sample IDs and SNP IDs must be in rownames and colnames. sex: Character vector, containing "M" or F indicating sex. Sample IDs must be in names. code: Character vector containing two letter genotype codes for each founder sample. Sample IDs must

be in names.

snps Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM

locations in columns 1 through 4, respectively. Not required for DO.

output.dir Character string containing the full path where output should be written. The

directory must exist already.

plot Boolean that is true if the user would like to plot a sample chromosome as the

model progresses. Default = TRUE.

trans.prob.fxn Function to use when computing the transtion probabilities between markers.

Value

No value is returned. The output files are written to output.dir.

Author(s)

Daniel Gatti

```
## Not run:
    calc.genoprob.alleles(data, chr = 1:19, founders = founders,
snps = snps, output.dir = "do.data")
## End(Not run)
```

calc.genoprob.intensity

Calculate the founder genotype probabilities at each SNP.

Description

This function performs genome reconstruction using allele intensities. We recommend using allele intensities where available because they often produce better genotype reconstructions.

Usage

```
calc.genoprob.intensity(data, chr, founders, snps, output.dir = ".", trans.prob.fxn,
plot = FALSE)
```

Arguments

data

A list with named elements containing the information needed to reconstruct

When method = intensity: x: Numeric matrix, num.samples x num.snps, with X intensities for all samples. Sample IDs and SNP IDs must be in rownames and colnames. y: Numeric matrix, num.samples x num.snps, with Y intensities for all samples. Sample IDs and SNP IDs must be in rownames and colnames. sex: Character vector, containing "M" or "F" indicating sex. Sample IDs must be in names. gen: Character matrix containing the generation of DO outbreeding for each sample. For the DO, this should be "DO" followed by a number with no space between them. For CC mice, this should be CC. Sample IDs must be in names.

chr

Character vector containing chromosomes to run. Must match the chromosome IDs in the snps table. "all" (default) will run all chromosomes.

founders

List containing founder information for non-DO or CC crosses. Not required for

When method = intensity: x: Numeric matrix, num.samples x num.snps, with X intensities for all founders and F1s (if available). Sample IDs and SNP IDs must be in rownames and colnames. y: Numeric matrix, num.samples x num.snps, with Y intensities for all founders and F1s (if available). Sample IDs and SNP IDs must be in rownames and colnames. sex: Character vector, containing "M" or "F" indicating sex. Sample IDs must be in names. code: Character vector containing two letter genotype codes for each founder sample. Sample IDs must be in names.

Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM locations in columns 1 through 4, respectively. Not required for DO.

output.dir

snps

Character string containing the full path where output should be written. The directory must exist already.

trans.prob.fxn FALSEunction to call to estimate the transition probabilities between markers for non-DO samples. Not required for DO.

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plot

Boolean that is true if the user would like to plot a sample chromosome as the model progresses. Default = TRUE.

Value

No value is returned. The output files are written to output.dir.

Author(s)

Daniel Gatti

Examples

Description

This function intersects the given variants with the genes in that region and classifies them according to "intergenic", "3UTR", "exon", "intron" or "5UTR".

Usage

```
categorize.variants(variants,
    mgi.file = "http://cgd.jax.org/tools/SNPtools/MGI/MGI.20130305.sorted.txt.gz")
```

Arguments

```
variants data.frame, Variants as returned by get.variants{get.variants}.

mgi.file Character, full path to the MGI feature file. On the JAX campus, this defaults to "http://cgd.jax.org/tools/SNPtools/MGI/MGI.20130305.sorted.txt.gz".
```

Value

FALSEor SNPs and Indels: data.frame: with eight columns: ID, CHR, POS, REFALSE, ALT, symbol, id, type. The first four columns are simply copied over from the SNP file. The symbol column contains the Gene Symbol. The id column contains a gene ID (MGI, Ensembl, NCBI or VEGA). The type column contains "intergenic", "3UTR", "exon", "intron" or "5UTR", depending on the location of the variant in a gene. FALSEor SVs: data.frame: with eight columns: ID, CHR, POS, REFALSE, ALT, symbol, id, type. The first four columns are simply copied over from the SNP file. The symbol column contains the Gene Symbol. The id column contains a gene ID (MGI, Ensembl, NCBI or VEGA). The type column contains "intergenic", "3UTR", "exon", "intron" or "5UTR", depending on the location of the variant in a gene.

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Author(s)

Daniel Gatti

See Also

```
get.variants{get.variants}
```

Examples

```
data(example.snps)
snp.type = categorize.variants(variants = example.snps[1:50,])
```

cc.trans.probs

Transition probabilities for CC mice.

Description

Thie function returns the transition probabilities for fully inbred Collaborative Cross mice.

Usage

```
cc.trans.probs(states, snps, chr = c(1:19, "X"), sex = c("M", "F"))
```

Arguments

states	Character vector containing the two letter codes (i.e. AA, BB, CC, etc.) for the homozygous states.
snps	Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM locations in columns 1 through 4, respectively. <i>Not required for DO</i> .
chr	Character containing the chromosome.
sex	Character that is either M or FALSE, indicating the sex to use. Only relevant on X chromosome.

Details

This function calculates the transition probabilities for fully inbred CC mice between two markers. It uses the equations for eight way RILs by sib mating.

Value

A matrix of transition probabilities between genotype states.

Note

This function has not been fully tested.

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Author(s)

Daniel Gatti

References

The genomes of recombinant inbred lines. Broman KW. Genetics. 2005 FALSEeb;169(2):1133-46. Epub 2004 Nov 15. Erratum in: Genetics. 2006 Aug;173(4):2419. PMID: 15545647

Examples

```
## Not run: cc.trans.probs(states = states, snps = snps, chr = 1)
```

cluster.strains

cluster.strains

Description

Given a set of numeric SNPs, cluster the strains based on allele sharing. This function hierarchically clusters the strains based on the proportion of SNPs that share alleles between each strain. Note: numeric snps, not character snps, are the input to this function.

Usage

```
cluster.strains(variants)
```

Arguments

variants

data.frame, numeric variants as returned by convert.variants.to.numeric.

Value

data.frame: of numeric variants with the strains clustered.

Author(s)

Daniel Gatti

See Also

```
convert.variants.to.numeric, variant.plot
```

```
data(example.snps)
variants = convert.variants.to.numeric(variants = example.snps[1:100,])
variants = cluster.strains(variants)
```

20 coefplot

coef.doqtl

Return the coefficients of a DOQTL object.

Description

Return the coefficients of a DOQTL object.

Usage

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

Arguments

object A DOQTL object as returned by scanone.
... Additional arguments.

Value

List containing matrices with QTL mapping model coefficients.

Author(s)

Daniel Gatti

See Also

scanone

Examples

```
head(coef(example.qtl[[1]]))
```

coefplot

Plot the QTL model coefficients

Description

Given a DOQTL object, plot the founder allele coefficients on one chromosome. The coefficients are centered around zero before plotting.

Usage

```
coefplot(doqtl, chr = 1, stat.name = "LOD", conf.int = TRUE, legend = TRUE,
colors = "DO", sex, ...)
```

colSumsLog 21

Arguments

doqtl	A DOQTL object as produced by scanone. A list containing two elements: lod and coef.
chr	Character containing the chromosome to plot.
stat.name	Character string containing the name of the mapping statistic.
conf.int	Boolean that is TRUE if the QTL support interval should be shaded in the plot. Default = TRUE.
legend	Boolean that is TRUE if the color legend for the DO founders should be drawn. Default = TRUE.
colors	Either "DO", in which case DO colors are supplied or a data.frame with three columns containing the founder letter code, founder strain name and founder color in columns 1:3.
sex	Character that is either FALSE or M, indicating the sex to use. Only used on X chromosome.
	Additional arguments to be passed to plot.

Value

No value is returned. A plot with the founder coefficients in the top panel and the LOD score in the bottom panel is drawn.

Author(s)

Daniel Gatti

See Also

```
scanone, plot.doqtl
```

Examples

```
## Not run:
    coefplot(qtl, chr = 1)
## End(Not run)
```

 ${\tt colSumsLog}$

Sum columns of log transformed data.

Description

Given a matrix of log transformed values, sum the rows or columns on the untransformed scale.

Usage

```
colSumsLog(logmat)
```

22 condense.model.probs

Arguments

logmat Numeric matrix of natural log transformed values.

Details

See addLog.

Value

Numeric vector with values summed on an untransformed scale.

Author(s)

Daniel Gatti

See Also

addLog

Examples

```
colSumsLog(matrix(log(runif(100)), nrow = 10, ncol = 10))
```

condense.model.probs Condense 36 state genotypes down to founder genotypes.

Description

Additive condenses the heterozygous genotype calls down to the founder allele contributions. Dominance will eventually provide additive and dominance values (method currently uncertain). FALSEull simply gathers all of the genotype probabilities together.

Usage

```
condense.model.probs(path = ".", write, model = c("additive", "dominance", "full"))
get.additive(files, samples)
get.dominance(files, samples)
get.full(files, samples)
```

Arguments

path	Character containing the path to the *.Rdata files that contain the genotype probabilities.
write	Character that is the filename to write the results to.
model	Character string that is one of "additive", "dominance" or "full". See details.
files	Vector of files to read.
samples	Character vector of sample IDs that match the files argument.

convert.allele.calls 23

Details

get.additive, get.dominance and get.full are helper functions.

Value

Three dimensional array of haplotype or genotype probabilities. Num. samples by num. founders (or genotypes) by num. SNPs. Writes out to a *.Rdata file.

Author(s)

Daniel Gatti

See Also

```
calc.genoprob
```

Examples

```
## Not run: condense.model.probs(write = "model.probs.Rdata")
```

convert.allele.calls Convert allele calls to numeric values.

Description

Converts allele calls in A,C,G,T,H,N format into numbers with 0: homozygous A, 1: heterozygous, 2: homozygous B, 3: no call.

Usage

```
convert.allele.calls(geno)
```

Arguments

geno

Character matrix containing A, C, G, T, H or N.

Value

Numeric matrix containing 0, 1, 2 or 3.

Author(s)

Daniel Gatti

```
## Not run: convert.allele.calls(geno)
```

24 convert.genotypes

```
convert.genes.to.GRanges
```

Convert MGI genes to GRanges.

Description

Given the output of get.mgi.features, convert the results to a GRanges object.

Usage

```
convert.genes.to.GRanges(mgi)
```

Arguments

mgi

Date.frame as returned by get.mgi.features.

Value

GRanges object containing the genes in the MGI argument.

Author(s)

Daniel Gatti

Examples

```
## Not run: convert.genes.to.GRanges(mgi)
```

convert.genotypes

Convert the genotype data from A,C,G,T format to A, H, B, N.

Description

Convert the genotype data from A,C,G,T format to A, H, B, N.

Usage

```
convert.genotypes(geno)
```

Arguments

geno

Character matrix containing A, C, G, T allele calls.

Value

Character matrix containing A, H, B, N.

Author(s)

Daniel Gatti

Examples

```
## Not run: convert.genotypes(geno)
```

```
convert.variants.to.GRanges
```

convert.variants.to.GRanges

Description

Given a data.frame of SNPs, convert the SNP locations to a GRanges object.

Usage

```
convert.variants.to.GRanges(variants)
```

Arguments

variants

Data frame with four header columns and SNPs in the remaining columns.

Details

This function creates a GRanges object from the CHR and POS columns of the SNP data.frame.

Value

GRanges object containing the SNP locations.

Author(s)

Daniel Gatti

```
data(example.snps)
gr = convert.variants.to.GRanges(example.snps[1:100,])
```

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```
convert.variants.to.numeric
```

convert.variants.to.numeric

Description

Given a matrix or data frame with character ACGT SNP values, convert them to numeric values with the major allele coded as 0 and the minor allele as 1.

Usage

```
convert.variants.to.numeric(variants)
```

Arguments

variants

Data frame with four header columns. The SNPs must be in ACGT format in columns 5 through ncol(snps).

Details

This function is used before calling plot.alleles() to convert the SNPs into a numeric form suitable for plotting.

Value

Data.frame of the same dimentions as the snps argument, but with the alleles converted to 0 or 1.

Author(s)

Daniel Gatti

Examples

```
data(example.snps)
numeric.snps = convert.variants.to.numeric(example.snps[1:100,])
```

```
create.genotype.states
```

Create genotype states.

Description

Given a set of founders, create all of the possible unphased genotype states between them.

Usage

```
create.genotype.states(founders)
```

create.html.page 27

Arguments

founders Character vector of letter codes indicating the founders.

Details

Given a set of founder IDs, create all possible unphased genotypes and sort them.

Value

Character vector of unphased genotypes that can be created from the given founders.

Author(s)

Daniel Gatti

Examples

```
create.genotype.states(founders = LETTERS[1:8])
```

create.html.page

Create an HTML QTL report

Description

Given a DOQTL object, create an HTML page that reports the QTL and creates QTL plots. Permutations for assessing significance thresholds can be supplied.

Usage

```
create.html.page(path, qtl, pheno.name, perms, assoc)
```

Arguments

path Character string containing the path to which to write the report.

qtl DOQTL object containing a list with two elemends: lod and coef.

pheno.name Character string containing the phenotype name.

perms Numeric vector containing the permutation LOD scores for this phenotype.

assoc Boolean, if TRUE, look for corresponding *.Rdata files containing the names of

the qtl in the current working directory and plot the association plots. If FALSE

(default), do not plot association analysis.

Details

The function creates an HTML page with a QTL plot, a table of significant QTL and coefficient plots for the significant loci.

28 create.Rdata.files

Value

Data.frame with the significant QTL from this phenotype.

Author(s)

Daniel Gatti

See Also

```
html.report
```

Examples

```
## Not run: create.html.page(path, qtl, pheno.name, perms)
```

create.Rdata.files

Convert *.txt files to *.Rdata files.

Description

This is used to convert the *.txt genotype probability files to *.Rdata files.

Usage

```
create.Rdata.files(prob.files)
```

Arguments

prob.files Character vector containing genotype probability file names.

Value

No value returned. Write out a *.Rdata file for each *.txt file.

Author(s)

Daniel Gatti

```
## Not run: create.Rdata.files(prob.files)
```

do.colors 29

do.colors

do.colors

Description

The letter codes, strain names and official colors for the DO founders.

Usage

do.colors

Format

A data frame with 8 rows and 3 columns.

CC_Designation a factor with levels A B C D E FALSE G H

Strain a factor with levels 129S1/SvImJ A/J C57BL/6J CAST/EiJ NOD/ShiLtJ NZO/H1LtJ PWK/PhJ WSB/EiJ

R_Color a factor with levels #00A000 #00A0FALSE0 #1010FALSE0 #808080 #9000FALSE4 #FALSE00000 #FALSE08080 #FALSE0FALSE000

Details

This contains the official colors that should be used when plotting data involving the DO founders.

Source

Copied from UNC Systems Genetics Website

Examples

do.colors

do.states

do.states

Description

The 36 unphased genotype states for the DO on the autosomes and X chromosome. Also the 8 DO founder letter codes.

Usage

do.states

do.trans.probs

Format

A list frame with 3 elements.

auto Character vector with two letter codes for each of the possible DO genotype states.

X List with two elements, FALSE and M, containing the genotype codes for the X chromosome.

founders Character vector containing the founder letter codes.

Details

This contains the letter codes for each of the 36 unphased genotype states in the DO. It also contains the founder letters.

Examples

do.states

do.trans.probs	Determine DO transition probabilities

Description

Determine the genotype state transition probability for DO mice of a specific generation between all of the markers on a given chromosome.

Usage

```
do.trans.probs(states, snps, chr = c(1:19, "X"), sex = c("M", "F"), do.gen)
```

Arguments

states	Character vector of possible genotype states.
snps	Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM locations in columns 1 through 4, respectively.
chr	Character containing the chromosome for which transition probabilities should be calculated.
sex	Character that is one of FALSE or M, indicating the sex to use on the X chromosome.
do.gen	Number vector indicating the DO outbreeding generations to calculate.

Details

This function is used to calculate the transition probabilities between markers for different DO outbreeding generations.

do2sanger 31

Value

List containing one element per unique DO generation supplied in the do.gen argument. Each list element contains a 3 dimensional array of transition probabilities between each pair of markers (num.states by num.states by num.markers - 1).

Author(s)

Daniel Gatti and Karl Broman

References

Haplotype probabilities in advanced intercross populations. Broman KW. G3 (Bethesda). 2012 FALSEeb;2(2):199-202. doi: 10.1534/g3.111.001818. Epub 2012 FALSEeb 1. PMID: 22384398 Genotype probabilities at intermediate generations in the construction of recombinant inbred lines. Broman KW. Genetics. 2012 FALSEeb;190(2):403-12. doi: 10.1534/genetics.111.132647. PMID: 22345609

Examples

```
## Not run: do.trans.probs(states, snps, chr = c(1:19, "X"), sex = c("M", F), do.gen)
```

do2sanger	Impute the Sanger SNPs onto DO genomes	

Description

Given a set of DO genotype probability files and the location of the Tabix indexed Sanger file, impute the Sanger SNPs on to DO genomes.

Usage

```
do2sanger(do.files, snps, output.file = "do2sanger.txt", snp.file =
"ftp://ftp.jax.org/SNPtools/variants/cc.snps.NCBI38.txt.gz")
```

Arguments

do.files	Character vector of *.genotype.probs.Rdata files that contain the posterior probabilities.	
snps	Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM locations in columns 1 through 4, respectively.	
output.file	Character string to write the results to.	
snp.file	Character string with path to a Tabix indexed SNP file. Default is from JAX FALSETP site	

32 do2sanger

Details

We read in a single genotype probability file, which must have been saved as a *.Rdata file. The format is a matrix with markers in rows and states in columns and dimnames for both. FALSEor each pair of markers, we take the average genotype probability. Then we take the DO genotype with the highest probability and split it into the two founder haplotypes. We get the Sanger SNPs for each of the two founders, convert them to 1, 1 or 2 and insert them into the DO sample.

Value

No value is returned. The Sanger SNPs mapped onto the DO genomes are written out to the output.file. The file will contain 0, 1 or 2 as the allele calls.

Author(s)

Daniel Gatti

References

Combined sequence-based and genetic mapping analysis of complex traits in outbred rats. Rat Genome Sequencing and Mapping Consortium, Baud A, Hermsen R, Guryev V, Stridh P, Graham D, McBride MW, FALSEoroud T, Calderari S, Diez M, Ockinger J, Beyeen AD, Gillett A, Abdelmagid N, Guerreiro-Cacais AO, Jagodic M, Tuncel J, Norin U, Beattie E, Huynh N, Miller WH, Koller DL, Alam I, FALSEalak S, Osborne-Pellegrin M, Martinez-Membrives E, Canete T, Blazquez G, Vicens-Costa E, Mont-Cardona C, Diaz-Moran S, Tobena A, Hummel O, Zelenika D, Saar K, Patone G, Bauerfeind A, Bihoreau MT, Heinig M, Lee YA, Rintisch C, Schulz H, Wheeler DA, Worley KC, Muzny DM, Gibbs RA, Lathrop M, Lansu N, Toonen P, Ruzius FALSEP, de Bruijn E, Hauser H, Adams DJ, Keane T, Atanur SS, Aitman TJ, FALSElicek P, Malinauskas T, Jones EY, Ekman D, Lopez-Aumatell R, Dominiczak AFALSE, Johannesson M, Holmdahl R, Olsson T, Gauguier D, Hubner N, FALSEernandez-Teruel A, Cuppen E, Mott R, FALSElint J. Nat Genet. 2013 Jul;45(7):767-75. doi: 10.1038/ng.2644. Epub 2013 May 26. PMID: 23708188 Using progenitor strain information to identify quantitative trait nucleotides in outbred mice. Yalcin B, FALSElint J, Mott R. Genetics. 2005 Oct;171(2):673-81. Epub 2005 Aug 5. PMID: 16085706 Mouse genomic variation and its effect on phenotypes and gene regulation. Keane TM, Goodstadt L, Danecek P, White MA, Wong K, Yalcin B, Heger A, Agam A, Slater G, Goodson M, FALSEurlotte NA, Eskin E, Nellaker C, Whitley H, Cleak J, Janowitz D, Hernandez-Pliego P, Edwards A, Belgard TG, Oliver PL, McIntyre RE, Bhomra A, Nicod J, Gan X, Yuan W, van der Weyden L, Steward CA, Bala S, Stalker J, Mott R, Durbin R, Jackson IJ, Czechanski A, Guerra-Assuncao JA, Donahue LR, Reinholdt LG, Payseur BA, Ponting CP, Birney E, FALSElint J and Adams DJ Nature 2011;477;7364;289-94 PUBMED: 21921910 Sequence-based characterization of structural variation in the mouse genome. Yalcin B, Wong K, Agam A, Goodson M, Keane TM, Gan X, Nellaker C, Goodstadt L, Nicod J, Bhomra A, Hernandez-Pliego P, Whitley H, Cleak J, Dutton R, Janowitz D, Mott R, Adams DJ and FALSElint J Nature 2011;477;7364;326-9 PUBMED: 21921916

See Also

assoc.map

estimate.cluster.params 33

Examples

```
## Not run: do2sanger(do.files, snps, output.file = "do2sanger.txt",
    snp.file = "ftp://ftp.jax.org/SNPtools/variants/cc.snps.NCBI38.txt.gz")
## End(Not run)
```

estimate.cluster.params

Estimate genotype cluster means and variances

Description

Given the X and Y intensity data, perform model based clustering and estimate the genotype state cluster means and variances.

Usage

```
estimate.cluster.params(founders, data, chr)
keep.homozygotes(founders)
```

Arguments

founders

List containing founder information for non-DO or CC crosses. *Not required for DO*.

When method = intensity: x: Numeric matrix, num.samples x num.snps, with X intensities for all founders and FALSE1s (if available). Sample IDs and SNP IDs must be in rownames and colnames. y: Numeric matrix, num.samples x num.snps, with Y intensities for all founders and FALSE1s (if available). Sample IDs and SNP IDs must be in rownames and colnames. sex: Character vector, containing "M" or F indicating sex. Sample IDs must be in names. code: Character vector containing two letter genotype codes for each founder sample. Sample IDs must be in names.

data

A list with named elements containing the information needed to reconstruct genomes.

When method = intensity: x: Numeric matrix, num.samples x num.snps, with X intensities for all samples. Sample IDs and SNP IDs must be in rownames and colnames. y: Numeric matrix, num.samples x num.snps, with Y intensities for all samples. Sample IDs and SNP IDs must be in rownames and colnames. sex: Character vector, containing "M" or F indicating sex. Sample IDs must be in names. gen: Character matrix containing the generation of DO outbreeding for each sample. FALSEor the DO, this should be "DO" followed by a number with no space between them. FALSEor CC mice, this should be CC. Sample IDs must be in names.

chr

Character containing the current chromosome.

34 example.genes

Details

At each marker, use mclust to perform model based clustering on all of the data and get estimates of the means and variances for each cluster. Then assign each of the 36 genotype states to the nearest founder cluster.

keep.homozygotes is an internal helper function.

Value

List containing two elements:

r.t.means Three dimensional array containing rho and theta genotype cluster means.

r.t.covars Three dimensional array containing rho and theta genotype cluster variances.

Author(s)

Daniel Gatti

See Also

```
hmm.intensity
```

Examples

```
## Not run: estimate.cluster.params(founders, data, chr)
```

example.genes

example.genes

Description

A set of genes from the Mouse Genome Informatics that are used in the documentation examples. (http://www.sanger.ac.uk/resources/mouse/genomes/) FALSErom Chr 7: 103 - 105 Mb on NCBI Build 38.

Source

http://informatics.jax.org/

```
data(example.genes)
```

example.pheno 35

example.pheno

Example phenotypes.

Description

Example phenotypes.

Usage

example.pheno

Format

A data frame with 149 observations on the following 8 variables. Sample IDs in rownames and phenotype names in colnames.

Sample Character vector containing sample IDs.

Sex Character vector containing the sex of each animal.

Gen Character vector containing DO outbreeding generation and litter.

Diet Character vector containing diet, either chow or hf for high fat.

Coat.Color Character vector containing text description of mouse colors.

albino Numeric vector contining 1 if the mouse's coat color was white.

black Numeric vector contining 1 if the mouse's coat color was black.

HDLD2 Numeric vector of high density lipoprotein values.

Details

Data from Svenson et.al. paper below.

References

High-resolution genetic mapping using the Mouse Diversity outbred population. Svenson KL, Gatti DM, Valdar W, Welsh CE, Cheng R, Chesler EJ, Palmer AA, McMillan L, Churchill GA. Genetics. 2012 FALSEeb;190(2):437-47 PMID: 223445611

Examples

head(example.pheno)

36 example.qtl

example.qtl

Example QTL.

Description

Example QTL for the albino and HDLD2 traits in example.pheno. Albino is a binary, Mendelian trait that maps to the Tyrosinase locus on Chr 7. HDLD2 (high density lipoprotein) is a complex trait with many loci.

Usage

```
example.qtl
```

Format

A data frame with 149 observations on the following 8 variables. Sample IDs in rownames and phenotype names in colnames.

Sample Character vector containing sample IDs.

Sex Character vector containing the sex of each animal.

Gen Character vector containing DO outbreeding generation and litter.

Diet Character vector containing diet, either chow or hf for high fat.

Coat.Color Character vector containing text description of mouse colors.

albino Numeric vector contining 1 if the mouse's coat color was white.

black Numeric vector contining 1 if the mouse's coat color was black.

HDLD2 Numeric vector of high density lipoprotein values.

Details

Data from Svenson et.al. paper below.

References

High-resolution genetic mapping using the Mouse Diversity outbred population. Svenson KL, Gatti DM, Valdar W, Welsh CE, Cheng R, Chesler EJ, Palmer AA, McMillan L, Churchill GA. Genetics. 2012 FALSEeb;190(2):437-47 PMID: 223445611

```
names(example.qtl)
names(example.qtl[[1]])
```

example.snps 37

|--|--|

Description

A set of SNPs from the Sanger Mouse Genome project that are used in the documentation examples. (http://www.sanger.ac.uk/resources/mouse/genomes/) FALSErom Chr 7: 103 - 105 Mb on NCBI Build 38.

Source

http://www.sanger.ac.uk/resources/mouse/genomes/

Examples

```
data(example.snps)
```

extract.raw.data	Extract intensities, genotypes and call rates from from raw MUGA or MegaMUGA data files

Description

This function accepts a vector of input directories containing the raw MUGA or MegaMUGA raw data files. FALSEor each directory, the function reads the X and Y intensities, call rates and allele calls for all samples. It then combines all samples and writes the data to "x.txt", "y.txt", "geno.txt" and "call.rate.batch.txt" in the user specified output directory.

Usage

```
extract.raw.data(in.path = ".", prefix, out.path = ".", array = c("megamuga", "muga"))
```

Arguments

in.path	character vector, the full path to all MUGA directories from which data should be extracted.
prefix	character vector of same length as in.path containing a prefix to add to each sample ID in data sets being processed.
out.path	character, the full path to the directory where the output files should be written.
array	character, default = "megamuga", the type of array, either "muga" or "megamuga".

38 fast.qtlrel

Details

This function searches each directory for files with names containing "Sample_Map.txt" and "*_FAL-SEinalReport.txt". This has been the format that GeneSeek has consistently produced. The call rates are extracted and are written, along with a batch ID, to "call.rate.batch.txt". The X and Y intensities are extracted from the "FALSEinalReport" file and written to "x.txt" and "y.txt" respectively. The allele calls are extracted from the "FALSEinalReport" files and are written to "geno.txt". The prefix argument may be used to add a prefix to the sample IDs in order to distinguish different data sets.

Value

No return value. The files are written to the out.path directory.

Note

Do not change the names of the output files. They are required for downstream processing.

Author(s)

Daniel M. Gatti

See Also

```
filter.samples, batch.normalize
```

Examples

fast.qtlrel

QTL mapping using QTLRel

Description

This extracts teh core of the QTLRel algorithm for additive covariates.

Usage

```
fast.qtlrel(pheno, probs, K, addcovar, snps)
```

fast.qtlrel 39

Arguments

pheno Data.frame containing the phenotype data. Sample IDs in rownames.

probs Three dimensional numeric array containing the founder halplotype contribu-

tions. Num.samples by num.founder by num.markers. Sample IDs, founder

letters and SNP IDs must be in dimnames.

K Numeric matrix containing the kinship between samlpes. Sample IDs must be

in rownames and colnames.

addcovar Numeric matrix containing additive covariates to run in the mapping model.

Sample IDs must be in rownames.

snps Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM

locations in columns 1 through 4, respectively.

Details

We extracted code from QTLRel, but removed several options to speed up the pipeline for QTL mapping with additive covariates and a kinship matrix.

Value

List containing two elements:

lod List containing two elements: A: Numeric matrix containing the SNP ID, chro-

mosome, Mb, cM, percent variance explained, likelihood ratio statistic, log of the odd ratio, p-value and -log10(p-value) for autosomes. X: Numeric matrix containing the SNP ID, chromosome, Mb, cM, percent variance explained, likelihood ratio statistic, log of the odd ratio, p-value and -log10(p-value) for X

chromosome.

coef List containing two elements: A: Numeric matrix containing QTL model co-

efficients on autosomes. Markers in rows. X: List containing two elements for the X chromosome: FALSE: Numeric matrix containing QTL model coefficients for females. Markers in rows. M: Numeric matrix containing QTL model

coefficients for males. Markers in rows.

Author(s)

Daniel Gatti

References

Cheng R, Abney M, Palmer AA, Skol AD. QTLRel: an R package for genome-wide association studies in which relatedness is a concern. BMC Genet. 2011 Jul 27;12:66.

See Also

```
scanone, scanone
```

Examples

```
## Not run: fast.qtlrel(pheno, probs, K, addcovar, snps)
```

40 filter.geno.probs

fill.in.snps

Interpolate between SNPs at the same cM value.

Description

Go through the SNPs and look for stretches where the differences in cM values from one SNP to the next equals 0. Interpolate from the first SNP to the last SNP in such a stretch.

Usage

```
fill.in.snps(snps)
```

Arguments

snps

Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM locations in columns 1 through 4, respectively.

Details

On each chromosome, look for sets of contiguous markers that have the same cM location. Interpolate evenly spaced markers spanning the markers proximal and distal to this set.

Value

Data.frame containing the marker locations. SNP ID, chromosome, Mb and cM locations in columns 1 through 4, respectively.

Author(s)

Daniel Gatti

Examples

```
## Not run: fill.in.snps(snps)
```

filter.geno.probs

Remove SNPs where the genotype probabilities are too low for one founder state

Description

This function accetps the 8 state founder probabilities and searches for SNPs where the founder probabilities are too low for one founder. It removes these SNPs. The QTL mapping model becomes numerically unstable if these SNPs are not removed.

filter.samples 41

Usage

```
filter.geno.probs(geno)
```

Arguments

geno

Numeric 3D array, with samples in dim[1], states = dim[2] and SNPs in dim[3].

Value

Numeric 3D array, with samples in dim[1], states = dim[2] and SNPs in dim[3]. The SNPs with low probabilities for a single founder have been removed.

Author(s)

Daniel Gatti

See Also

```
scanone, scanone.perm
```

Examples

```
## Not run:
    filter.geno.probs(geno)
## End(Not run)
```

filter.samples

FALSEilter X, Y and genotype data by call rate

Description

This function reads in "x.txt", "y.txt", "geno.txt" and "call.rate.batch.txt" from the user specified input directory and removes samples with a call rate less than the supplied threshold (default = 0.9). It then writes the files out to "x.filt.txt", "y.filt.txt", "geno.filt.txt" and "call.rate.batch.filt.txt".

Usage

```
filter.samples(path = ".", thr = 0.9)
```

Arguments

path Character, the full path to the directory where the files reside.

thr Numeric, call rate threshold below which samples will be removed. Default =

0.9.

Value

data.frame with the sample IDs and call rates of the removed samples. The intensity and genotype files are written to the input directory.

Author(s)

Daniel Gatti

See Also

```
extract.raw.data, batch.normalize
```

Examples

```
## Not run:
    filter.samples(path = "/tmpdir/output")
## End(Not run)
```

find.overlapping.genes

find.overlapping.genes

Description

FALSEind genes that intersect with a given set of variants.

Usage

```
find.overlapping.genes(variants,
mgi.file = "http://cgd.jax.org/tools/SNPtools/MGI/MGI.20130305.sorted.txt.gz",
type = c("gene", "exon"))
```

Arguments

variants Data.frame with variants. The type attribute must be set to one of "snp", "indel",

or "sv".

mgi.file Character. FALSEull path to the MGI gene file. Defaults to "http://cgd.jax.org/tools/SNPtools/MGI/MGI.

type Character. One of "gene" or "exon". Indicates whether to intersect SNPs with

genes (from beginning to end, including introns) or exons only.

Details

Gets the gene locations from MGI and variants.

Value

Data.frame with gene locations and symbols.

gene.plot 43

Author(s)

Daniel Gatti

See Also

```
get.mgi.features{get.mgi.features}
```

Examples

```
## Not run:
    data(example.snps)
    x = find.overlapping.genes(variants = example.snps[19478:19506,])
## End(Not run)
```

gene.plot

gene.plot

Description

Given a genomic region, plot the genes in the interval. line.up.genes and resolve.collisions are internal functions.

Usage

```
gene.plot(mgi, rect.col = "grey30", text.col = "black", ...)
```

Arguments

mg1	Data.frame with MGI gene locations as returned by get.mgi.features{get.mgi.features}.
rect.col	Color vector that is the color to use to plot gene rectangles. May be a single color for all genes or a vector with colors for each gene. Default = "grey30".
text.col	Color vector that is the color to use to plot gene names. May be a single color for all genes or a vector with colors for each gene. Default = "black".
	Other arguments to be passed into plot.

Details

The spacing algorithm attempts to organize the genes in such a way that they do not collide. The rect.col and text.col arguments are recycled if they are shorter than the number of genes. They can be used to highlight specific genes.

Value

Data.frame with gene locations and symbols.

44 generic.trans.probs

Author(s)

Daniel Gatti

See Also

```
get.mgi.features{get.mgi.features}
```

Examples

```
## Not run:
  data(example.genes)
  g = gene.plot(mgi = example.genes)
## End(Not run)
```

generic.trans.probs

Generic transition probabilities

Description

Generic function to provide starting transition probabilities. Not ready for use in this version.

Usage

```
generic.trans.probs(states, snps, chr = c(1:19, "X"), sex = c("M", "F"))
```

Arguments

states	Character vector containing the genotype states coded as letters.
snps	Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM locations in columns 1 through 4, respectively.
chr	Character containing the chromosome to use.
sex	Character that is either FALSE or M indicating the sex. Only used on x chromo-
	some.

Details

This function is a placeholder for a more sophisticated algorithm to be developed later. It is an effort to provide a set of starting transition probabilities for mapping populations other than the DO.

Value

A matrix of transition probabilities between genotype states.

Note

This function is still in development.

genome.summary.plots 45

Author(s)

Daniel Gatti

See Also

```
do.trans.probs, cc.trans.probs
```

Examples

```
## Not run: generic.trans.probs(states, snps, chr = 1, sex = "M")
```

```
genome.summary.plots Genome summary plots
```

Description

These plots show summaries of the founder haplotype proportions across SNPs or samples.

Usage

```
genotype.by.sample.barplot(results)
genotype.by.snp.barplot(results, snps)
```

Arguments

results Data.frame as produced by summarize.by.snps or summarize.by.samples.

snps Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM

locations in columns 1 through 4, respectively.

Details

Barplots across samples or SNPs are produced from the relevant summary files. This is intended to be run after DO genomes have been reconstructed for a large set of samples to verify that founder allele frequencies are consistent across SNPs and samples.

Author(s)

Daniel Gatti

See Also

```
summarize.by.snps, summarize.by.samples
```

46 get.chr.lengths

Examples

```
## Not run:
    load(url("ftp://ftp.jax.org/MUGA/muga_snps.Rdata"))
    results = summarize.by.samples(path = ".", snps = muga_snps)
    genotype.by.sample.barplot(results)
## End(Not run)
```

get.chr.lengths

Get chromosome lengths for the mouse

Description

Using the org.Mm.eg.db package, get the chromosome lengths.

Usage

```
get.chr.lengths()
```

Details

Keeps only the lengths of the chromosomes, not random or unmapped.

Value

Named vector of chromosome lengths.

Author(s)

Daniel Gatti

Examples

```
get.chr.lengths()
```

get.do.states 47

get.do.states

Get the 36 genotype states for the DO

Description

Get the 36 genotype states for the DO

Usage

```
get.do.states()
```

Value

Character vector containing the two letter genotype codes for the 36 genotype states.

Author(s)

Daniel Gatti

Examples

```
get.do.states()
```

get.gene.name

Get the gene symbol

Description

Given an MGI ID, get the gene symbol from an MGI data.frame.

Usage

```
get.gene.name(value, mgi)
```

Arguments

value Character string containing an MGI ID.

mgi Data.frame containing MGI data from get.mgi.features.

Value

Character string containing the gene symbol.

Author(s)

Daniel Gatti

48 get.max.geno

See Also

```
get.mgi.features
```

Examples

```
## Not run: get.gene.name(value, mgi)
```

get.machine.precision Get the machine precsion

Description

Get the machine precision from .Machine\$double.eps on a log10 scale.

Usage

```
get.machine.precision()
```

Value

Numeric value containing the machine precision.

Author(s)

Daniel Gatti

Examples

```
get.machine.precision()
```

get.max.geno

Get the genotype with the highest probability

Description

FALSEor each sample at each marker, return the genotype state with the highest probability.

Usage

```
get.max.geno(probs)
```

Arguments

probs

Three dimensional numeric matrix containing the founder haplotype contributions.

get.mgi.features 49

Details

If the maximum probability is greater than 0.75, it is called homozygous. Otherwise, we take the two highest probabilities and call heterozygous.

Value

Character matrix containing the genotype with the highest probability at each locus.

Author(s)

Daniel Gatti

Examples

```
## Not run: get.max.geno(probs)
```

get.mgi.features

get.mgi.features

Description

Retrieve the MGI features within a genomic region. FALSEeatures include genes, gene models, non-coding RNA, etc., but not SNPs. This allows the user to filter by source and type of feature.

Usage

```
get.mgi.features(file = "ftp://ftp.jax.org/SNPtools/genes/MGI.20130703.sorted.txt.gz",
chr = NULL, start = NULL, end = NULL, source = c("all", "MGI", "VEGA", "ENSEMBL",
   "Blat", "NCBI_Gene"), type = c("all", "gene", "pseudogenic_transcript",
   "pseudogenic_exon", "pseudogene", "match", "match-part", "transcript", "exon",
   "mRNA", "five_prime_UTR", "start_codon", "CDS", "stop_codon", "three_prime_UTR",
   "pseudogenic_mRNA", "pseudogenic_start_codon", "pseudogenic_CDS",
   "pseudogenic_stop_codon", "pseudogenic_five_prime_UTR", "pseudogenic_three_prime_UTR",
   "sequence_feature"))
```

Arguments

file	character, the full path to the Tabix indexed and zipped MGI feature file. Default = "http://cgd.jax.org/tools/SNPtools/MGI/MGI.20130305.sorted.txt.gz" for internal JAX use.
chr	Numeric vector, chr for each start and end position. Chr, start and end must all have the same length.
start	Numeric vector, start position in Mb or bp for each chr. Chr, start and end must all have the same length.
end	Numeric vector, end position in Mb or bp for each chr. Must be greater than or equal to the corresponding start value. Chr, start and end must all have the same length.

50 get.num.auto

source Character vector, the source of the annotation. Options are ("all", "MGI", "VEGA",

"ENSEMBL", "Blat", "NCBI_Gene"). "all" returns all features.

type Character vector, the type of feature. Options are ("all", "gene", "pseudogenic_transcript",

"pseudogenic_exon", "pseudogene", "match", "match-part", "transcript", "exon",

"mRNA", "five_prime_UTR", "start_codon", "CDS", "stop_codon", "three_prime_UTR",

"pseudogenic_mRNA", "pseudogenic_start_codon", "pseudogenic_CDS", "pseu-

dogenic_stop_codon", "pseudogenic_five_prime_UTR", "pseudogenic_three_prime_UTR",

"sequence_feature"). "all" returns all features.

Details

This function is designed to return features from the MGI gene feature file (GFALSEFALSE). You can select multiple regions on different chromosomes.

Value

A list of data.frames for each region requested. Each data.frame will contain 14 columns; seqid, source, type, start, stop, score, strand, phase, ID, Name, Parent, Dbxref, mgiName, bioType. If there is only one requested region, a single data frame is returned.

Author(s)

Daniel Gatti

References

The MGI GFALSEFALSE file is at ftp://ftp.informatics.jax.org/pub/mgigff/.

Examples

```
## Not run:
genes = get.mgi.features(chr = 7, start = 103 end = 105 source = "MGI", type = "gene")
## End(Not run)
```

get.num.auto

Get the number of autosomes

Description

Given a data.frame of SNPs, return the number of autosomes by counting the number of numeric chromosomes.

Usage

```
get.num.auto(snps)
```

get.pattern.variants 51

Arguments

snps

Data.frame containing the marker locations. SNP ID, chromosome, Mb and cM locations in columns 1 through 4, respectively.

Value

Numeric value containing the number of autosomes.

Author(s)

Daniel Gatti

Examples

```
## Not run: get.num.auto(snps)
```

```
get.pattern.variants get.pattern.variants
```

Description

Given a vector of strains names and a set of varaints, return the variants for which the given strains have one allele and the remaining strains have the other allele.

Usage

```
get.pattern.variants(variants, strain.subset = NULL)
```

Arguments

variants

Matrix of data.frame of variants with four header columns containing annotation

and snps in the remaining columns.

strain.subset

character vector of strain names that comprise one subset.

Details

Use this function to obtain a subset of SNPs for which one set of strains contain one allele and the rest contain the other allele. This might come up in the course of QTL mapping where several strains have a high allele and others have a low allele.

Value

Data.frame of variants that match the requested pattern.

Author(s)

Daniel Gatti

52 get.strains

See Also

```
get.variants
```

Examples

```
data(example.snps)
snp.subset = get.pattern.variants(example.snps[1:1000,],
strain.subset = c("A/J", "NOD/ShiLtJ", "NZO/HlLtJ", "AKR/J"))
```

get.strains

get.strains

Description

Get available strain names for a set of strains from a variant file.

Usage

```
get.strains(
file = "http://cgd.jax.org/tools/SNPtools/Build38/sanger.snps.NCBI38.txt.gz")
```

Arguments

file

Character, full path to the variants file to use. Default is the file at the Center for Genome Dynamics at The Jackson Laboratory.

Value

Character vector of strain names in the variant file.

Author(s)

Daniel Gatti

Examples

```
## Not run:
  available.strains = get.strains()
  strains = available.strains[c(2, 5, 8, 9, 13:15, 17)]
  snps = get.variants(chr = 7, start = 103, end = 105, strains = strains)
## End(Not run)
```

get.trans.probs 53

get.trans.probs	Get the transition probabilities between markers.	

Description

Based on the DO founders and the recombination fraction, calculate the transition probability between markers on one chromosome.

Usage

```
get.trans.probs(r, do.gen, alpha, chr = c(1:19, "X"), sex = c("M", "F"))
```

Arguments

r	Double vector of recombination fractions between SNPs.
do.gen	Integer containing the outbreeding generation of DO.
alpha	Double vector, proportion of preCC progenitors at generation k. Generation numbers in the names.
chr	Character, one of 1:19, X.
sex	Character, either M or FALSE. Only used on X chromosome.

Value

Numeric three dimensional array of transition probabilities between each pair of markers (num.states by num.states by num.markers - 1).

Author(s)

Daniel Gatti

References

Haplotype probabilities in advanced intercross populations. Broman KW. G3 (Bethesda). 2012 FALSEeb;2(2):199-202. doi: 10.1534/g3.111.001818. Epub 2012 FALSEeb 1. PMID: 22384398 Genotype probabilities at intermediate generations in the construction of recombinant inbred lines. Broman KW. Genetics. 2012 FALSEeb;190(2):403-12. doi: 10.1534/genetics.111.132647. PMID: 22345609

Examples

```
## Not run: get.trans.probs(r, do.gen, alpha, chr = c(1:19, "X"), sex = c("M", "F"))
```

54 get.variants

Description

Get SNPs for a set of strains for a specific regions of chromosomes.

Usage

```
get.variants(file = "ftp://ftp.jax.org/SNPtools/variants/cc.snps.NCBI38.txt.gz",
chr, start, end, type = c("snp", "indel", "sv"), strains, polymorphic = TRUE,
quality)
```

Arguments

Character, full path to the SNP file to use. Default is the file at the Center for Genome Dynamics at The Jackson Laboratory.
Numeric vector, chr for each start and end position. Chr, start and end must all have the same length.
Numeric vector, start position in Mb or bp for each chr. Chr, start and end must all have the same length.
Numeric vector, end position in Mb or bp for each chr. Must be greater than or equal to the corresponding start value. Chr, start and end must all have the same length.
Character, with one of "snp", "indel", "sv". Indicates the type of variantes being queried.
Character vector, listing the strains to retrieve. The names can be obtained from get.strains(). Default returns all strains.
Boolean. Default = TRUE. If TRUE, retrieve only SNPs that are polymorphic among the requested strains. If FALSEALSE, return all SNPs in the requested interval(s).
Integer, denoting the confidence levels to return. This looks at the quality column of all requested strains and takes values greater than or equal to the given number. Look at the documentation for the SNP file you are using for details on the quality score. Default returns all SNPs.

Value

A list of data frames for each region requested. Each data frame will contain five header columns; ID, CHROM, POS, REFALSE & ALT corresponding to the SNP ID, chromosome, bp position, reference and alternate alleles. There follows two columns for each requested strain containing the allele calls and quality scores for each strain. If there is only one requested region, a single data frame is returned.

html.report 55

Author(s)

Daniel Gatti

See Also

```
get.strains
```

Examples

```
## Not run:
  available.strains = get.strains()
  strains = available.strains[c(2, 5, 8, 9, 13:15, 17)]
  snps = get.variants(chr = 7, start = 103, end = 105, strains = strains)
## End(Not run)
```

html.report

Create an HTML report for a set of QTL

Description

Given a list of QTL objects, create a set of HTML pages summarizing the QTL.

Usage

```
html.report(path, qtl, perms, assoc = FALSE)
```

Arguments

path Character string with path to which to write.

qt1 List containing DOQTL objects as produced by scanone.

perms Numeric vector containing permutation results for the phenotypes.

assoc Boolean, if TRUE, look for corresponding *.Rdata files containing the names of

the qtl in the current working directory and plot the association plots. If FALSE

(default), do not plot association mapping analysis.

Details

This function summarizes a set of QTL scans made using scanone. FALSEor each QTL in the qtl list, it creates an HTML page with a QTL plot and coefficient plots for the significant QTL. The permutations are used to assess significance. It then summarizes all of these in a table and creates a summary HTML page and a *.csv file with the QTL results.

Value

No value is returned. HTML report of QTL is written out to the specified directory.

56 intensity.plots

Author(s)

Daniel Gatti

Examples

```
## Not run: html.report(path, qtl, perms, merge = FALSE)
```

intensity.plots Plot founders and F1 hybrids or genotype state means and variances on an intensity plot.

Description

Given the X and Y (or rho and theta) array intensities, plot the values for one SNP and color and mark the founders and FALSE1s.

Usage

```
founder.F1.intensity.plot(theta, rho, s = 1, is.founder.F1, plotNew = TRUE, ...) intensity.mean.covar.plot(s, states, theta, rho, r.t.means, r.t.covars, sample, ...)
```

Arguments

theta	Numeric matrix containing theta (or X) intensity values.
rho	Numeric matrix containing rho (or Y) intensity values.
S	Numeric integer containing the SNP index in rho and theta to plot.
is.founder.F1	Boolean vector containing TRUE for samples that are DO founders or FALSE1s and FALSEALSE for those that are not.
plotNew	Boolean that is TRUE if a new plot should be made and FALSEALSE if the points should be added to the existing plot.
states	Character vector containing genotypes state letter codes.
r.t.means	Numeric three dimensional array containing genotype state means in rho and theta coordinates.
r.t.covars	Numeric three dimensional array containing genotype state variances in rho and theta coordinates.
sample	Numeric value containing the sample index into the rho and theta matrices to plot in orange.
	Additional arguments to be passed to plot.

Value

No return value. A scatter plot is made of the rho and theta (or X and Y) intensities at the requested SNP. Founders and F1 hybrid samples are colored.

kinship.probs 57

Author(s)

Daniel Gatti

Examples

```
## Not run: founder.F1.intensity.plot(theta, rho, s = 1, is.founder.F1, plotNew = TRUE, ...)
```

kinship.probs Create a kinship matrix.

Description

Read in the genotypes and produce a kinship matrix based on either allele sharing or haplotype contributions.

Usage

```
kinship.probs(probs, snps, bychr = FALSE)
kinship.alleles(geno)
```

Arguments

probs	Three dimensional numeric array containing the founder haplotype contributions. Num.samples by num.founder by num.markers.
snps	Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM locations in columns 1 through 4, respectively.
bychr	Boolean that is true if the current chromosome should be excluded from the calculation of kinship for the current chromosome. See Details.
geno	Character matrix of allele calls.

Details

Allele based method: This simply calculates the mean allele sharing between two individuals. Intensity based method: FALSEor each pair of samples, at each marker, calculate the cosine of the angle between the two founder contribution vectors. Take the average across the genome.

When bychr = FALSE, we calculate one kinship matrix for all chromosomes. When bychr TRUE, we calculate a different kinship matrix for each chromosome. FALSEor each chromosome, we remove use only the SNPs on the remaining chromosomes to calculate the kinship matrix for that chromosome. This is motivated by the Cheng et.al. reference below.

Value

Numeric matrix, with rows and columns equal to the number of samples in the genotype file.

Author(s)

Daniel Gatti

58 muga.snps.to.keep

References

Practical Considerations Regarding the Use of Genotype and Pedigree Data to Model Relatedness in the Context of Genome-Wide Association Studies. Cheng R, Parker CC, Abney M, Palmer AA. G3 (Bethesda). 2013 Oct 3;3(10):1861-1867. PMID: 23979941

See Also

```
extract.raw.data, filter.samples, batch.normalize, calc.genoprob
```

Examples

```
## Not run:
   K = kinship.probs(probs)
## End(Not run)
```

muga.snps.to.keep

SNPs to use for genotyping and mapping on the MUGA

Description

SNPs to use for genotyping and mapping on the MUGA. These are a combination of one culled by UNC due to poor performance or uncertain location and ones culled by JAX based on low intensity values.

Usage

```
muga.snps.to.keep
```

Format

A character vector with MUGA SNP IDs to use when genotyping or mapping.

Examples

```
muga.snps.to.keep
```

plot.doqtl 59

plot.dogtl	Plot a QTL
	- · · · · · · · · · · · · · · · · · · ·

Description

Given a genome scan produced by scanone, create a plot of the LOD score across all chromosomes.

Usage

```
## S3 method for class doqtl
plot(x, stat.name = c("lod", "neg.log10.p"), sig.thr = NULL, sig.col = "red", ...)
```

Arguments

X	DOQTL object containing the LOD score and model coefficients.
stat.name	Character string containing the name of the mapping statistic.
sig.thr	Numeric vector containing significance thresholds.
sig.col	Colors to use when plotting the significance thresholds above.
	Additional arguments to pass to plot.

Value

Creates a QTL plot.

Author(s)

Daniel Gatti

Examples

```
## Not run:
    qtl = scanone(pheno = pheno, pheno.col = 1, probs = probs, snps = snps)
plot(qtl)
## End(Not run)
```

plot.genoprobs

plot.genoprobs	Plot the genome of a DO sample.

Description

This function plots the genotype of a DO sample by taking the genotype with the maximum probability at each marker.

Usage

```
## S3 method for class genoprobs
plot(x, snps, colors = "D0", chrlen = "D0", ...)
  chr.skeletons(chr, chrlen = "D0", ...)
  genomic.points(chr = NULL, loc = NULL)
  write.genoprob.plots(path = ".", snps)
```

Arguments

X	Numeric matrix containing the 36 genotype state probabilities generated by calc.genoprob.
snps	Data.frame containing the SNP locations in prsmth. SNP ID, chromosome, Mb and cM locations in columns 1 through 4, respectively.
colors	Character string containing "DO" for the DO. FALSEor all other crosses, a data frame containing single letter codes, strain names and colors for the founders in columns 1 through 3, respectively.
chrlen	Character string containing "DO" for mice. FALSEor all other organisms, the lengths of each chromosome in Mb.
chr	Vector of chromosome IDs.
path	Character string containing the path to the posterior genotype probability files in R binary format (i.e. *.Rdata).
loc	Numeric vector of genome locations in Mb of same length as chr.
	Additional arguments to be passed to plot.

Value

plot.genoprobs plots the reconstructed DO genome in terms of founder haplotypes. chr.skeletons plots the chromosome skeletons. This is useful for further plotting. genomic.points plots locations on a genome such as might be drawn by chr.skeletons. write.genoprob.plots loops through all of the files supplied in the argument and writes out genotype plots.

Author(s)

Daniel Gatti

pxg.plot 61

See Also

```
calc.genoprob
```

Examples

```
## Not run: plot.genoprobs(prsmth, snps, main = "plot title")
```

pxg.plot	Phenotype by genotype plot at a single marker.	

Description

This function plots the phenotype versus the most probable genotype at a single marker.

Usage

```
pxg.plot(pheno, pheno.col, probs, snp.id, snps, legend = TRUE, sex = NA, covar, ...)
```

Arguments

pheno	Data.frame containing the phenotype data with samples in rows and phenotypes in columns. Sample IDs in rownames and phenotype names in colnames.
pheno.col	Numeric or character vector: Either a vector of number that indicate columns to use or a set of column names in pheno.
probs	3D numeric array, containing the founder haplotype contributions or genotype probabilities. The sample IDs, founder letter codes and markers IDs must be in dimnames.
snp.id	Character string containing the marker to plot at. Must be in SNPs and dimnames(probs)[[3]].
snps	Data.frame containing 4 columns with marker location information. SNP ID, chr, Mb, cM in columns 1 through 4, respectively.
legend	Boolean that is true if a legend explaining the founder letter codes should be plotted.
sex	Character that is either FALSE or M, indicating sex. This is used only when the SNP ID is on the X chromosome and all of the samples are male. In this case, there are only 8 genotype states.
covar	Vector of categories for each sample (i.e. diet, treatement, etc.) which will be plotted as different symbols. Optional.
	Additional arguments passed along to plot.

Details

The most probable genotype is inferred from the eight founder allelic contributions at the marker. If a founder has a value > 0.75, it is assumed to be homozygous.

62 qtl.heatmap

Value

Creates a plot with the phenotype on the y-axis and the 36 DO genotypes on the x-axis.

Author(s)

Daniel Gatti

See Also

```
plot.dogtl, coefplot
```

Examples

```
## Not run: pxg.plot(pheno, pheno.col, founder.probs, snp.id, snps)
```

qtl.heatmap

Plot a Heatmap of all QTL

Description

This function accepts LOD scores and SNP locations and plots a heatmap of the QTL. Each QTL is scaled to between 0 and 1 and the QTL are hierarchically clustered. A tile plot is then created with QTL colored from white (strong) to black (weak). The motivation for this plot is to visualize QTL that may be coincident across different phenotypes.

Usage

```
qtl.heatmap(lod, ...)
```

Arguments lod

Data frame, containing the SNP names, chromosome an dbp locations in columns 1 through 3 and LOD scores for all phenotyped in the remaining columns.

Additional arguments to be passed to plot.

Value

No value is returned. A tile plot is plotted from the data.

Note

The QTL the appear in white are *not* neccessarily significant. They are simply the strongest QTL for each phenotype.

Author(s)

Daniel Gatti

qtl.LRS 63

See Also

```
plot.doqtl, scanone.perm
```

Examples

```
## Not run:
    qtl.heatmap(lod)
## End(Not run)
```

qtl.LRS

QTL mapping with no kinship.

Description

These are two functions that perform QTL mapping without a kinship matrix. They use the QR decomposition to speed up the computation. Other than for a quick screen or for assessing significance thresholds, we do not recommend mapping without a kinship matrix. They are included for historical reasons

Usage

```
qtl.LRS(pheno, probs, snps, addcovar = NULL)
permutations.qtl.LRS(pheno, probs, snps, addcovar, nperm = 1000,
return.val = c("lrs", "p"))
```

Arguments

pheno	Data.frame containing the phenotype data with samples in rows and phenotypes in columns. Sample IDs in rownames and phenotype names in colnames.
probs	Numeric three dimenaional array, containing the founder haplotype contributions or genotype probabilities. The sample IDs, founder letter codes and markers IDs must be in dimnames.
snps	data.frame containing 4 columns with marker location information. SNP ID, chr, Mb, cM in columns 1 through 4, respectively.
addcovar	data.frame or numeric matrix, containing any additive covariates. Sample IDs must be in rownames.
nperm	Numeric value containing the number of permutations to run.
return.val	Character string containing either "LRS" or "p", indicating the type of return statistic.

Details

The function performs Haley-Knott regression at the markers using the founder haplotype contributions in probs.

64 qtl.qtlrel

Value

List containing two elements. LRS: a data.frame containing the LOD scores. coef: numeric matrix containing model coefficients.

Author(s)

Daniel Gatti

See Also

scanone

Examples

```
## Not run: qtl.LRS(pheno, probs, snps, addcovar = NULL)
```

qtl.qtlrel

Use QTLRel to map a set of traits

Description

This function accepts phenotypes, genotype probabilities and a sample kinship matrix and maps the requested traits using an eight state linear model. FALSEixed covariates may be passed in as well. The output is written to two files: *.LOD.txt (containing the LOD scores for each SNP) and *.coef.txt. (containing the model coefficient at each SNP).

Usage

```
qtl.qtlrel(pheno, probs, K, addcovar, intcovar, snps)
```

Arguments

pheno	Data frame, containing the sample IDs, phenotype data and covariates.
probs	3D numeric array, containing the genotype probabilities for all samples at each SNP. Dimensions must be samples by states by SNPs and all dimensions must be named.
К	Numeric matrix, containing the kinship between individuals as computed by QTLRel.
addcovar	Numeric matrix containing additive covariates.
intcovar	Numeric matrix containing covariates that interact with the QTL.
snps	Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM locations in columns 1 through 4, respectively.

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Value

A list containing two elements:

lod Data.frame containing the SNP locations and LOD and p-values.

coef Data.frame containing the model coefficients.

Author(s)

Daniel Gatti

See Also

```
plot.doqtl, scanone, scanone.perm
```

Examples

```
## Not run:
   qtl.qtlrel(pheno, prob, K, covar = NULL, pheno.name = "")
## End(Not run)
```

qtl.simulate

Simulate a QTL in the DO

Description

Using a set of real DO genomes, simulate a QTL with a given minor allele frequency (MAFALSE), sample size and effect size.

Usage

```
qtl.simulate(probs, snps, K, sample.size = dim(probs)[[1]], effect.size = 1,
maf = 4, num.poly = 18, num.sims = 1000)
```

Arguments

probs	Numeric three dimensional array, containing the founder haplotype contribu- tions or genotype probabilities. The sample IDs, founder letter codes and mark- ers IDs must be in dimnames.
snps	Data.frame containing 4 columns with marker location information. SNP ID, chr, Mb, cM in columns 1 through 4, respectively.
K	Numeric matrix, containing the additive kinship matrix. The samples IDs must be in rownames and colnames.
sample.size	Numeric vector sample sizes. Must be less than or equal to the number of samples in probs.
effect.size	Numeric vector containing the effect sizes as the number of standard deviations from the phenotype mean.

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maf	Numeric value containing the minor allele frequecy as the number of founders. FALSEor DO, the value must be between 1 and 4.
num.poly	Numeric value containing the number of autosomes on which to simulate a polygenic background.
num.sims	Numeric value containing the number of simulations to run.

Details

This function will simulate a phenotype with a QTL. It will output a phenotype vector with sample IDs and a data.frame called qtl describing the simulated loci.

Value

Writes pheno and qtl lists to an R binary file.

Author(s)

Daniel Gatti

See Also

scanone

Examples

```
## Not run: qtl.simulate(probs, snps, K, sample.size = dim(probs)[[1]],
effect.size = 1, maf = 4, num.poly = 18, num.sims = 1000)
## End(Not run)
```

rankZ

Rank Z transformation

Description

This is also called the inverse normal transformation. We rank the data in x, divide by n-1, and take quantiles from the normal distribution using qnorm.

Usage

rankZ(x)

Arguments

Χ

Numeric vector of values to be transformed.

Details

We often use this when there are hundreds or thousands of phenotypes. This allows us to calculate permutation derived thresholds once and use them for all phenotypes.

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Value

Numeric vector with a normal distribution.

Author(s)

Daniel Gatti

Examples

```
rankZ(rlnorm(20))
```

read.vcf

Read and parse VCFALSE data

Description

*** CURRENTLY UNDER CONSTRUCTION ***. This function is designed to read in the Sanger VCFALSE files for SNPs, Indels and structural variants.

Usage

Arguments

vcf.file	Character string containing full path to a Sanger SNP, Indel or structural variant VCFALSE file.
chr	Character or numeric vector containing mouse chromosome IDs to query.
start	Numeric vector containing the start position on each chromosome to query. Must be the same length as chr. If the value is less than 200, it is assumed to be in Mb. Values greater than 200 are assumed to be in bp.
end	Numeric vector containing the end position on each chromosome to query. Must be the same length as chr. If the value is less than 200, it is assumed to be in Mb. Values greater than 200 are assumed to be in bp.
strains	Character vector containing strain names, as listed in the vcf.file, to retrieve. If missing, then all strains are returned. Use get.vcf.strains to get the strain names from the VCFALSE file.
return.val	Character string that is either "allele", which returns the nucleotide allele calls, or "number", which returns numeric values.
return.qual	Boolean that is TRUE if the quality columns should be returned.
csq	Boolean that is TRUE if the consequence column should be returned.

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Details

There is a very nice package called vcf2geno, but it is designed to work with the 1000 Genomes VC-FALSE format. The Sanger format is slightly different, hence the need for this function. FALSEuture plans include the addition of variant consequence selection. The Sanger variant files have 'snp', 'indel' or 'SV' in them and we use this to determine the type of file being processed.

Value

If one chromosome location is requested, a data.frame containing the requested variants and quality scores. The variant locations and reference alleles are in the first 5 columns and the variants and quality scores for each strain are in the remaining columns. If more than one location is requested, a list of data.frames containing the variants.

Note

!!! This function is still under development !!!

Author(s)

Daniel Gatti

References

The Sanger Mouse Genomes Project generated these variants. Raw data is at the Sanger FALSETP site.

Next-generation sequencing of experimental mouse strains. Yalcin B, Adams DJ, FALSElint J and Keane TM Mammalian genome: official journal of the International Mammalian Genome Society 2012;23;9-10;490-8 PUBMED: 22772437

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Examples

```
## Not run: read.vcf(vcf.file =
  "ftp://ftp-mouse.sanger.ac.uk/current_snps/mgp.v3.snps.rsIDdbSNPv137.vcf.gz",
  chr = 1, start = 5, end = 5.5)
## End(Not run)
```

scanone

Perform a genome scan.

Description

This function is the main QTL mapping function for point mapping at each marker. The user must supply phenotypes, genotype probabilities and marker locations. Optional kinship, additive and interactive covariates may be passed in. Scanone regresses the phenotype on the genotype probabilities and produces a LOD score and founder allele effects at each marker.

Usage

```
scanone(pheno, pheno.col = 1, probs, K, addcovar, intcovar, snps, model =
c("additive", "full"))
```

Arguments

pheno	data.frame containing phenotype data. Required. rownames must contain sample IDs and there must be a column labelled 'sex' to perform correct mapping on the X chromosome.
pheno.col	numeric or character vector: Either a vector of number that indicate columns to use or a set of column names in pheno.
probs	Numeric three dimensional array, containing the founder haplotype contributions or genotype probabilities. The sample IDs, founder letter codes and markers IDs must be in dimnames.
K	numeric matrix, containing the additive kinship matrix. The samples IDs must be in rownames and colnames.
addcovar	data.frame or numeric matrix, containing any additive covariates. Sample IDs must be in rownames.
intcovar	data.frame or numeric matrix, containing any covariates that interact with the QTL. Sample IDs must be in rownames.
snps	data.frame containing 4 columns with marker location information. SNP ID, chr, Mb, cM in columns 1 through 4, respectively.
model	character string, containing one of "additive" or "full", indicating the type of model to fit.

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Value

A list containing two elements:

lod Data.frame with nine columns containing the marker locations, LOD scores and

p-values for each phenotype. Column names are SNP_ID, Chr, Mb_NCBI38,

cM, perc.var, lrs, lod, p and neg.log10.p.

coef Numeric matrix containing the founder allele effects at each locus. Colnames

are the additive covariates plus the founder terms.

Author(s)

Daniel Gatti

See Also

scanone.perm

Examples

```
## Not run: scanone(pheno, pheno.col = 1, probs)
```

scanone.eqtl

Mapping using the Matrix EQTL algorithm.

Description

MatrixEQTL uses a series of matrix operations to greatly accelerate QTL mapping. It can accomodate additive covariates and a common kinship matrix for all phenotypes.

Usage

```
scanone.eqtl(expr, probs, K, addcovar, snps, sex)
```

Arguments

expr	Numeric matrix of phenotype values with samples in rows and phenotypes in columns. Rownames must contain sample IDs and colnames must contain phenotype names.
probs	Numeric three dimensional array containing the founder haplotype contributions. Num.samples by num.founders by num.markers.
K	Numeric matrix of kinship values for all samples. Num.samples by num.samples.
addcovar	Numeric matrix of additive covariates.
snps	Data.frame containing 4 columns with marker location information. SNP ID, chr, Mb, cM in columns 1 through 4, respectively.
sex	Character vector containing either "M" or F, indicating the sex of each sample. Used for mapping on the X chromosome.

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Details

Matrix EQTL centers and rotates the phenotype and genotype matrices using matrix operations. It only calculates the LOD score at each marker and does not provide coefficients.

Value

Numeric matrix of LOD scores for all phenotypes and markers.

Author(s)

Daniel Gatti

References

Matrix eQTL: ultra fast eQTL analysis via large matrix operations. Shabalin AA. Bioinformatics. 2012 May 15;28(10):1353-8.

See Also

scanone

Examples

```
## Not run: scanone.eqtl(expr, probs, K, addcovar, snps)
```

scanone.perm

Perform a genome scan.

Description

This function is the main QTL mapping function for point mapping at each marker. The user must supply phenotypes, genotype probabilities and marker locations. Optional kinship, additive and interactive covariates may be passed in. scanone.perm regresses the phenotype on the genotype probabilities and produces a LOD score and founder allele effects at each marker.

Usage

```
scanone.perm(pheno, pheno.col = 1, probs, addcovar, intcovar, snps, model =
c("additive", "full"), path = ".", nperm = 1000)
```

Arguments

pheno	data.frame containing phenotype data. Required. rownames must contain sam-
	1 10 1.1 .1 1 11 11 12 2. 6

ple IDs and there must be a column labelled 'sex' to perform correct mapping

on the X chromosome.

pheno.col numeric or character vector: Either a vector of number that indicate columns to

use or a set of column names in pheno.

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3D numeric array, containing the founder haplotype contributions or genotype probs probabilities. The sample IDs, founder letter codes and markers IDs must be in dimnames. addcovar data.frame or numeric matrix, containing any additive covariates. Sample IDs must be in rownames. intcovar data.frame or numeric matrix, containing any covariates that interact with the QTL. Sample IDs must be in rownames. snps data.frame containing 4 columns with marker location information. SNP ID, chr, Mb, cM in columns 1 through 4, respectively. Character string, containing one of "additive" or "full", indicating the type of mode1 model to fit. path Character string containing the location where permutation results should be

nperm Numeric, containing the number of permutations to run.

Value

A list containing two elements:

written.

lod Data.frame with seven columns containing the marker locations and LOD scores

for each phenotype.

coef Numeric matrix containing the founder allele effects at each locus.

Author(s)

Daniel Gatti

See Also

```
scanone.perm
```

Examples

```
## Not run: scanone.perm(pheno, pheno.col = 1, probs)
```

sdp.plot Plot association mapping results.

Description

Plot the founder strain distribution patterns (SDP) of the SNPs in results.

Usage

```
sdp.plot(results, ...)
```

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Arguments

results Data.frame containing output from assoc.map.

... Additional arguments passed to plot.

Details

Given the output from assoc.map, plot the SDPs for the SNPs in results. This will show which strains have the minor allele at each SNP in results. You may also add this plot to the top of assoc.plot.

Value

Produces a plot. There is no return value.

Author(s)

Daniel Gatti

See Also

```
assoc.map, assoc.plot
```

Examples

```
## Not run:
    results = assoc.map(pheno = pheno, pheno.col = 1, probs = probs, K = K, addcovar = addcovar,
snps = snps, chr = 1, start = 40, end = 45)
    sdp.plot(results[results[,12] > 3,])
## End(Not run)
```

sex.predict

Determine the sex of each sample

Description

This function uses the mean X and Y chromosome intensities to predict the sex of each sample. Linear discriminant analysis from the MASS package is used.

Usage

```
sex.predict(x, y, snps, plot = FALSE)
```

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Arguments

Х	Numeric matrix, num.samples x num.snps, containing X intensities for all samples.
У	Numeric matrix, num.samples x num.snps, containing Y intensities for all samples.
snps	Data.frame containing SNP IDs, chromosomes, Mb and cM locations in columns 1 through 4, respectively.
plot	Boolean that will create a plot of mean X and Y Chr intensities if TRUE. Default = FALSEALSE.

Value

Character vector with sex assignments.

Author(s)

Daniel Gatti

Examples

```
## Not run:
    data(snps)
    sex.predict(x, y, sex = rep(F, ncol(x)))
## End(Not run)
```

snp.plot snp.plot

Description

Lower level function that makes a tile plot the given SNPs with the major allele colored dark and the minor allele light. Optionally, add SNPs that match a certain pattern and genes and a QTL score.

Usage

```
snp.plot(variants, col = c("black", "grey50", "white"), cluster = TRUE, ref, highlight,
pattern.snps, mgi, qtl)
```

Arguments

variants	Data.frame with variants as returned by get.variants.	
col	Color vector with SNP colors for no call, alternate allele and reference allele.	
cluster	Boolean that indicates if the strains should be clustered.	
ref	Chracter that is the reference strain to use. Must be present in the strains in variants.	

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highlight	Character vector with strain names to highlight in the plot. Strain names must be present in the strains in variants.	
pattern.snps	Data.frame with SNPs that match some pattern. These will be plotted below the SNPs.	
mgi	Data.frame with genes in the interval to be plotted.	
qtl	data.frame with QTL values containing the chromosome, bp position and QTL score in column 1 to 3. Default = NULL.	

Details

Different strains can be used as the reference strain by using the ref argument. Otherwise, the major allele is plotted dark and the minor allele lighter. The QTL values will be scaled within the plotting interval and drawn with black (low) and red (high). If a strain pattern is provided, the SNPs matching the pattern are plotted in orange and any genes that they intersect with are also colored orange. Otherwise, genes are colored blue.

Value

Produces a tile plot with the locations along the horizontal axis and the strains in the SNP matrix along the vertical axis. Also, if a set of strains is given in the pattern argument, the SNPs that match that pattern are returned, categorized according to which gene they lie within.

Author(s)

Daniel Gatti

See Also

```
variant.plot, get.mgi.features, categorize.variants
```

Examples

```
## Not run:
data(qt1)
strains = get.strains()
variants = get.variants(chr = 7, start = 103, end = 105,
strains = strains[c(2,4,8,10,15:18)])
pattern.snps = get.pattern.variants(variants, strain.subset = c("C57BL/6J", "NOD/ShiLtJ",
    "NZO/HlLtJ"))
mgi = get.mgi.features(chr = 7, start = 103, end = 105, source = "MGI", type = "gene")
variants = convert.variants.to.numeric(variants)
snp.plot(variants = variants, pattern.snps = pattern.snps, mgi = mgi, qtl = qtl)
## End(Not run)
```

```
{\tt summarize.genotype.transitions}
```

Summarize the genotype data output by the genotyping HMM.

Description

These functions read in all of the individual genotype data files and summarizes the founder allele frequency by sample and SNP.

Usage

```
summarize.genotype.transitions(path = ".", snps)
summarize.by.snps(path = ".", snps)
summarize.by.samples(path = ".", snps)
num.recomb.plot(results, gen)
```

Arguments

path	Character, full path to the genotype directory where the *.genotype.probs.Rdata files are stored.
snps	Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM locations in columns 1 through 4, respectively.
results	Data.frame containing results as summarized by summarize.genotype.transitions.
gen	Numeric vector containing DO outbreeding generation for all samples in results. Must have sample IDs in names.

Value

Data.frame with six columns: sample, proximal genotype, distal genotype, chr, proximal Mb, distal Mb.

Author(s)

Daniel Gatti

Examples

```
## Not run: geno.sum = summarize.genotype.transitions()
```

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|--|

Description

Convenience function that makes a tile plot the given SNPs with the major allele colored dark and the minor allele light. Optionally, add SNPs that match a certain pattern and genes and a QTL score. This calls sdp.plot after gathering all of the data.

Usage

```
variant.plot(var.file = "ftp://ftp.jax.org/SNPtools/variants/cc.snps.NCBI38.txt.gz",
mgi.file = "ftp://ftp.jax.org/SNPtools/genes/MGI.20130703.sorted.txt.gz", chr,
start, end, type = c("snp", "indel", "sv"), strains = c("A/J", "C57BL/6J",
"129S1/SvImJ", "CAST/EiJ", "NOD/ShiLtJ", "NZO/HlLtJ", "PWK/PhJ", "WSB/EiJ"),
ref = "C57BL/6J", pattern, qtl)
```

Arguments

var.file	Character string with the full path to the tabix indexed Sanger/UNC SNP file. Default is a file at the Center for Genome Dynmaics at the Jackson Laboratory.
mgi.file	Character string with the full path to the tabix indexed MGI feature file. Default is a file at the Center for Genome Dynmaics at the Jackson Laboratory.
chr	Character with the chromosome ID to plot on. Required.
start	Numeric value with the start position to plot at. May be in bp or Mb. If the value is 200 or less, it is assumed to be in Mb. Otherwise it is assumed to be in bp. Required.
end	Numeric value with the end position to plot at. May be in bp or Mb. If the value is 200 or less, it is assumed to be in Mb. Otherwise it is assumed to be in bp. Required.
type	Character indicating the type of variant to retrieve. One of "snp", "indel", or "sv". Default = "snp".
strains	Character vector with the strains to use. Must be from the set of strains returned by get.strains{get.strains}. Default = Collaborative Cross/Diversity Outbred founders.
ref	Character string with the reference strain. Must be one of the names in the strains vector argument. Default = C57BL/6J.
pattern	Character vector with strain names in the strains argument. We search for SNPs for which these strains have one allele and all other strains have the opposite allele. A more sophisticated pattern matching method may be included in future releases. Default = NULL.
qtl	data.frame with QTL values containing the chromosome, bp position and QTL score in column 1 to 3. Default = NULL.

Details

Different strains can be used as the reference strain by using the ref argument. Otherwise, the major allele is plotted dark and the minor allele lighter. The QTL values will be scaled within the plotting interval and drawn with black (low) and red (high). If a strain pattern is provided, the SNPs matching the pattern are plotted in orange and any genes that they intersect with are also colored orange. Otherwise, genes are colored blue.

Value

Produces a tile plot with the locations along the horizontal axis and the strains in the SNP matrix along the vertical axis. Also, if a set of strains is given in the pattern argument, the SNPs that match that pattern are returned, categorized according to which gene they lie within.

Author(s)

Daniel Gatti

See Also

```
convert.variants.to.numeric, get.mgi.features, categorize.variants, snp.plot
```

Examples

```
## Not run:
data(qtl)
strains = get.strains()[c(2,4,8,10,15:18)]
variant.plot(chr = 7, start = 103, end = 105, strains = strains,
pattern = c("C57BL/6J", "NOD/ShiLtJ", "NZO/HlLtJ"), qtl = qtl)
## End(Not run)
```

write.founder.genomes Write out the genotypes of DO samples

Description

Given a directory containing files generated by the DOQTL HMM (i.e. that end with "genotype.probs.Rdata"), write out two files for each sample containing the founder haplotype blocks.

Usage

```
write.founder.genomes(filenames = dir(path = ".",
pattern = "genotype.probs.Rdata"), snps)
```

Arguments

snps

filenames Character vector of posterior genotype probability files in R binary format.

Data.frame containing the marker locations. SNP ID, chromosome, Mb anc cM

locations in columns 1 through 4, respectively.

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Details

FALSEor each sample, we take the genotype state with the highest probability and write it out.

Value

No value is returned. FALSEiles are written to the current working directory.

Author(s)

Daniel Gatti

Examples

```
## Not run: write.founder.genomes(filenames = dir(path = ".",
   pattern = "genotype.probs.Rdata"))
## End(Not run)
```

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