# Package 'simcross'

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**Title** Simulate Experimental Crosses

**Description** Simulate and plot general experimental crosses. The focus is on simulating genotypes with an aim towards flexibility rather than speed. Meiosis is simulated following the Stahl model, in which chiasma locations are the superposition of two processes: a proportion p coming from a process exhibiting no interference, and the remainder coming from a process following the chi-square model.

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License GPL-3

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2 AILped

## **R** topics documented:

	Allped	2
	calc_Lstar	3
	CCcolors	4
	check_pedigree	4
	collapse_do_alleles	5
	convert2geno	6
	convert2geno_allchr	7
	create_parent	9
	cross	9
	get_geno	11
	mouseL_cox	11
	mouseL_mgi	12
	plot_crosslines	13
	plot_ind	14
	sim_4way_pedigree	15
	sim_ail_pedigree	16
	sim_ail_pedigree_fix_n	17
	sim_crossovers	19
	sim_dof1_pedigree	20
	sim_do_pedigree	21
	sim_do_pedigree_fix_n	23
	sim_from_pedigree	24
	sim_from_pedigree_allchr	25
	sim_meiosis	26
	sim_ril_pedigree	27
	where_het	28
Index		<b>3</b> 0

AILped

 ${\it Example AIL pedigree}$ 

## Description

Example matrix describing the pedigree for advanced intercross lines

## Usage

data(AILped)

## **Format**

A data frame with five columns: individual id, mom, dad, sex (0 for females and 1 for males) and generation.

calc\_Lstar 3

#### **Source**

Derived from the pedF8 dataset in the QTLRel package, https://cran.r-project.org/package=QTLRel

## **Examples**

```
data(AILped)
x <- sim_from_pedigree(AILped)</pre>
```

calc\_Lstar

Calculate adjusted chromosome length for obligate chiasma

## **Description**

Calculate the reduced chromosome length that will give the target expected number of chiasmata when conditioning on there being at least one chiasma on the four-strand bundle.

## Usage

```
calc_Lstar(L, m = 0, p = 0)
```

#### **Arguments**

L Length of chromosome (in cM); must be > 50

m Interference parameter for chi-square model

p Proportion of chiasmata coming from no-interference process

#### Value

Adjusted length of chromosome

#### See Also

```
cross(), sim_meiosis(), sim_crossovers()
```

```
calc_Lstar(100, 0, 0)
calc_Lstar(60, 10, 0.1)
```

check\_pedigree

**CCcolors** 

Collaborative Cross colors

## Description

Get the vector of colors for the Collaborative Cross

## Usage

```
CCcolors(palette = c("new", "original", "official"))
```

## **Arguments**

palette

Which version of the colors to use? (New or original)

#### Value

vector of eight colors

## **Examples**

CCcolors()

check\_pedigree

Check a pedigree for errors

## **Description**

Perform a series of checks on the tabular data for a pedigree, checking for problems

#### **Usage**

```
check_pedigree(pedigree, ignore_sex = FALSE)
```

#### **Arguments**

pedigree Numeric matrix or data frame with four columns: ID, mom ID, dad ID, sex. Sex

is coded as 0=female, 1=male. There can be additional columns, but they'll be

ignored.

ignore\_sex If TRUE, ignore the sex values completely (appropriate for hermaphroditic species.)

#### **Details**

The parents should be listed before any of their offspring. Founders should have 0's for mother and father; all others should have non-zero values for the parents, and the parents should appear in the pedigree. Father should be male and mothers should be female (unless ignore\_sex=TRUE). Individual identifiers should be unique and non-zero. There should be no missing values anywhere. (NAs are allowed in the sex column if ignore\_sex=TRUE.)

collapse\_do\_alleles 5

## Value

TRUE (invisibly) if everything is okay; otherwise gives an error.

#### See Also

```
sim_from_pedigree(), sim_ril_pedigree()
```

#### **Examples**

```
tab <- sim_ril_pedigree(7)
check_pedigree(tab)</pre>
```

collapse\_do\_alleles

Collapse alleles for simulated DO genotypes

## **Description**

When simulating Diversity Outbreds, we need to specify parents 1-16, with 9-16 being the males from strains 1-8. This function collapses replaces alleles 9-16 with 1-8, to make the result ordinary DO-type data.

## Usage

```
collapse_do_alleles(xodata)
```

#### **Arguments**

xodata

The sort of detailed genotype/crossover data generated by sim\_from\_pedigree().

#### Value

The input object, with alleles 9-16 replaced by 1-8.

#### See Also

```
sim_do_pedigree(), sim_do_pedigree_fix_n(), sim_from_pedigree()
```

```
# simulate DO pedigree
tab <- sim_do_pedigree(8)

# simulate genotypes for that pedigree
dat <- sim_from_pedigree(tab)
# collapse to alleles 1-8
dat <- collapse_do_alleles(dat)

# also works with data on multiple chromosomes</pre>
```

6 convert2geno

```
dat <- sim_from_pedigree(tab, c("1"=100, "2"=75, "X"=100), xchr="X") dat <- collapse_do_alleles(dat)
```

convert2geno

Convert continuous allele information into marker genotypes

## Description

Convert the continuous crossover location information produced by sim\_from\_pedigree to marker genotypes

#### Usage

```
convert2geno(xodat, map, founder_geno = NULL, shift_map = FALSE)
```

#### **Arguments**

xodat The sort of detailed genotype/crossover data generated by sim\_from\_pedigree()

map vector of marker locations; can also be a list of such vectors (one per chro-

mosome), in which case xodat and founder\_geno must be lists with the same

length.

founder\_geno Optional matrix (size n\_founders x length(map)) of founder genotypes. If

coded as 1/2 (or 1/3), results are 1/2/3 genotypes. If coded as A/T/G/C/N, results are A/T/G/C/N/H genotypes. If coded as letters A-H for the 8 founders, results

are two-letter genotypes AA-HH with 36 possible values.

shift\_map If TRUE, shift genetic map to start at 0

#### Value

If founder\_geno is provided or there are just two founders, the result is a numeric matrix of genotypes, individuals x markers, with genotypes 1/2/3 codes for 11/12/22 genotypes.

If founder\_geno is not provided and there are more than two founders, the result is a 3-dimensional array, individuals x markers x alleles, with the third dimensional corresponding to the maternal and paternal allele.

If the input map is a list (the components being chromosomes), then xodat and founder\_geno must be lists of the same length, and the result will be a list of matrices.

#### See Also

```
get_geno(), sim_from_pedigree()
```

convert2geno\_allchr 7

#### **Examples**

```
# simulate AIL pedigree
tab <- sim_ail_pedigree(12, 30)</pre>
# simulate data from that pedigree
dat <- sim_from_pedigree(tab)</pre>
# marker map (could also use sim.map in R/qtl)
map <- seq(0, 100, by=5)
names(map) <- paste0("marker", seq(along=map))</pre>
# convert data to marker genotypes
geno <- convert2geno(dat, map)</pre>
# AIL with multiple chromosomes
dat <- sim_from_pedigree(tab, c("1"=100, "2"=75, "X"=100), xchr="X")</pre>
# marker map
multmap <- list("1"=seq(0, 100, by=5),
                 "2"=seq(0, 75, by=5),
                 "X"=seq(0, 100, by=5))
for(i in 1:3)
  names(multmap[[i]]) <- paste0("marker", i, "_", 1:length(map[[i]]))</pre>
geno <- convert2geno(dat, multmap)</pre>
# simulate DO pedigree
tab <- sim_do_pedigree(8)</pre>
# simulate data from that pedigree
dat <- sim_from_pedigree(tab)</pre>
# simulate founder snp alleles
fg <- matrix(sample(1:2, 8*length(map), repl=TRUE), nrow=8)</pre>
# for DO, need female & male founders (to deal with X chr)
fg <- rbind(fg, fg)</pre>
# convert dat to SNP genotypes
geno <- convert2geno(dat, map, fg)</pre>
# if fg not provided, result is a 3d array
genoarray <- convert2geno(dat, map)</pre>
```

convert2geno\_allchr

Convert continuous allele information into marker genotypes for multiple chromosomes

## **Description**

Wrap up of convert2geno to adequate multiple chromosomes.

#### Usage

```
convert2geno_allchr(
  xodat,
  map,
  id = NULL,
```

convert2geno\_allchr

```
founder_geno = NULL,
return.matrix = TRUE,
shift_map = FALSE
)
```

#### **Arguments**

8

xodat The sort of detailed genotype/crossover data generated by sim\_from\_pedigree\_allchr()

map marker locations, a list with elements for each chromosome

id ids for which individuals genotypes is desired

founder\_geno Optional list of matrices (one per chromosome) of size n\_founders x n\_markers,

with the founder genotypes. If coded as 1/2 (or 1/3), results are 1/2/3 genotypes. If coded as A/T/G/C/N, results are A/T/G/C/N/H genotypes. If coded as letters A-H (in the case of 8 founders), results are two-letter genotypes AA-HH with

36 possible values.

return.matrix If FALSE, the result is a list of length n\_chrs, otherwise it is converted into a

matrix if size length(id) x n\_markers.

shift\_map If TRUE, shift genetic map to start at 0

#### Value

If founder\_geno is provided or there are just two founders, the result is a numeric matrix of genotypes, individuals x markers, with genotypes 1/2/3 codes for 11/12/22 genotypes. If there are more than two founders and founder\_geno are letters, the result is a character matrix, too.

If founder\_geno is not provided and there are more than two founders, the result is a 3-dimensional array, individuals x markers x alleles, with the third dimensional corresponding to the maternal and paternal allele.

#### See Also

```
convert2geno()
```

```
library(qtl)
# marker map
map <- sim.map(len=rep(100, 19), n.mar=10, include.x=FALSE)
# simulate AIL pedigree
tab <- sim_ail_pedigree(12, 30)
# simulate data from that pedigree
dat <- sim_from_pedigree_allchr(tab, map)
names(map) <- paste0("marker", seq(along=map))
# convert data to marker genotypes
id <- which(tab[, "gen"]==12)
geno <- convert2geno_allchr(dat, map, id)</pre>
```

create\_parent 9

create\_parent

Create a parent object

## Description

Create a parent object

## Usage

```
create_parent(L, allele = 1)
```

## Arguments

```
L chromosome length in cM allele vector of integers for alleles, of length 1 or 2
```

## Value

A list with two components, for the individual's two chromosomes. Each is a list with alleles in chromosome intervals (as integers) and locations of the right endpoints of those intervals.

#### See Also

```
cross(), sim_meiosis()
```

## **Examples**

```
create_parent(100, 1)
create_parent(100, 1:2)
```

cross

Cross two individuals

## **Description**

Simulate the cross of two individuals to create a single progeny

## Usage

```
cross(
   mom,
   dad,
   m = 10,
   p = 0,
   xchr = FALSE,
   male = FALSE,
```

10 cross

```
obligate_chiasma = FALSE,
  Lstar = NULL
)
```

## **Arguments**

mom	An individual object, as produced by create_parent() or this function.			
dad	An individual object, as produced by create_parent() or this function.			
m	interference parameter for chi-square model			
p	proportion of crossovers coming from no-interference process			
xchr	If TRUE, simulate X chromosome			
male	If TRUE, simulate a male (matters only if xchr=TRUE)			
obligate_chiasma				
	If TRUE, require an obligate chiasma on the 4-strand bundle at meiosis.			
Lstar	Adjusted chromosome length, if obligate_chiasma=TRUE. Calculated if not provided.			

#### **Details**

Simulations are under the Stahl model with the interference parameter being an integer. This is an extension of the chi-square model, but with chiasmata being the superposition of two processes, one following the chi-square model and the other exhibiting no interference.

## Value

A list with two components, for the individual's two chromosomes. Each is a list with alleles in chromosome intervals (as integers) and locations of the right endpoints of those intervals.

#### See Also

```
create_parent(), sim_meiosis(), sim_crossovers(), calc_Lstar()
```

```
mom <- create_parent(100, 1:2)
dad <- create_parent(100, 1:2)
child <- cross(mom, dad)</pre>
```

get\_geno 11

get\_geno

Get genotype at a single position

## Description

With data on the continuous crossover location information produced by sim\_from\_pedigree, grab the genotype at a given position.

#### Usage

```
get_geno(xodat, position)
```

## **Arguments**

xodat The sort of detailed genotype/XO data generated by sim\_from\_pedigree()

position Position (in cM) for which to obtain genotypes

#### Value

A numeric matrix with two columns: the maternal and paternal allele for each individual.

#### See Also

```
sim_from_pedigree(), convert2geno()
```

## **Examples**

```
# simulate AIL pedigree
tab <- sim_ail_pedigree(12, 30)
# simulate data from that pedigree
dat <- sim_from_pedigree(tab)
# get genotype at position 30 cM
geno <- get_geno(dat, 30)</pre>
```

mouseL\_cox

Mouse chromosome lengths

## Description

Mouse chromosome lengths in cM from the Cox et al. map

## Usage

```
data(mouseL_cox)
```

mouseL\_mgi

## **Format**

A numeric vector with lengths in cM for the 20 mouse chromosomes.

#### **Source**

Taken from Table 1 of Cox et al. (2009) A new standard genetic map for the laboratory mouse. Genetics 182:1335-1344. https://doi.org/10.1534/genetics.109.105486doi:10.1534/genetics.109.105486

#### See Also

```
mouseL_mgi
```

## **Examples**

```
data(mouseL_cox)
```

mouseL\_mgi

Mouse chromosome lengths

## Description

Mouse chromosome lengths in cM from the Mouse Genome Informatics (MGI) standard map.

## Usage

```
data(mouseL_mgi)
```

## **Format**

A numeric vector with lengths in cM for the 20 mouse chromosomes.

## **Source**

Taken from Table 1 of Cox et al. (2009) A new standard genetic map for the laboratory mouse. Genetics 182:1335-1344. https://doi.org/10.1534/genetics.109.105486doi:10.1534/genetics.109.105486

## See Also

```
mouseL_cox
```

```
data(mouseL_mgi)
```

plot\_crosslines 13

plot\_crosslines

Plot cross lines

## Description

Add lines for a cross

## Usage

```
plot_crosslines(
  momloc,
  dadloc,
  kidsloc,
  gap = 3,
  chrlength = 30,
  cex = 1.5,
  lwd = 2,
  arrow_length = 0.1,
  col = "white",
  ...
)
```

## **Arguments**

momloc An (x,y) vector with center location for mother dadloc An (x,y) vector with center location for mother Either an (x,y) vector with center location for a kid, or a list of such for multiple kidsloc kids gap Gap arrows and points/rectangles chrlength Length of chromosomes cex Character expansion for x point lwd Line width for points, segments, and arrows arrow\_length The length parameter in the call to graphics::arrows() col Color of lines and points

Additional arguments passed to arrows() and segments()

#### Value

None.

. . .

## See Also

```
plot_ind()
```

14 plot\_ind

#### **Examples**

plot\_ind

Plot an individual

## Description

Add an individual, as a pair of chromosomes, to a plot

## Usage

```
plot_ind(
   ind,
   center,
   chrlength = 30,
   chrwidth = 3,
   gap = 3,
   col = CCcolors(),
   border = "black",
   lend = 1,
   ljoin = 1,
   allborders = FALSE,
   ...
)
```

## **Arguments**

ind An individual object, as output by create\_parent() or cross()

center (x,y) vector for the center of the individual

chrlength Length of chromosomes (Can be a vector of length 2, in which case the two

chromosomes will be different lengths, aligned at the top. This is for the X

chromosome.)

chrwidth Width of chromosomes
gap Gap between chromosomes

col Vector of colors

sim\_4way\_pedigree 15

border Color for border

lend Passed to graphics::rect()
ljoin Passed to graphics::rect()

allborders If TRUE, put borders around all segments
... Additional arguments passed to rect()

#### Value

None.

#### See Also

```
plot_crosslines()
```

## **Examples**

sim\_4way\_pedigree

Simulate pedigree for 4-way intercross

## Description

Simulate a 4-way cross, among four inbred lines (a table of individual, mom, dad, sex)

## Usage

```
sim_4way_pedigree(ngen = 1, nsibs = 100)
```

## Arguments

ngen Number of intercross generations (1 or 2)

nsibs Vector with number of siblings in the sibships in the last generation.

sim\_ail\_pedigree

#### **Details**

We start with a set of 4 individuals (representing four inbred lines), and make a pair of crosses to generate a pair of heterozygous individuals. These are then crosses to generate a set of F1 individuals. If ngen==1, we stop there, with sum(nsibs) individuals in this last generation. If gen==2, we generate length(nsibs) male/female pairs of F1 offspring; these are intercrossed to generate a set of sibships, with lengths defined by the values in nsibs. Individuals in the last generation are alternating female/male.

#### Value

A data frame with five columns: individual ID, mother ID, father ID, sex, and generation. Founders have 0 for mother and father ID. Sex is coded 0 for female and 1 for male.

#### See Also

```
sim_from_pedigree(), sim_ril_pedigree(), sim_do_pedigree(), sim_ail_pedigree()
```

## **Examples**

```
# 100 F1s between heterozygous parents
tab <- sim_4way_pedigree(1, 100)
# could also do this
tab2 <- sim_4way_pedigree(1, rep(10, 10))
# 120 F2s in 10 sibships each of size 12
tab3 <- sim_4way_pedigree(ngen=2, rep(12, 10))</pre>
```

sim\_ail\_pedigree

Simulate AIL pedigree

#### **Description**

Simulate a pedigree for advanced intercross lines (a table of individual, mom, dad, sex)

## Usage

```
sim_ail_pedigree(
  ngen = 12,
  npairs = 30,
  nkids_per = 5,
  design = c("nosib", "random")
)
```

sim\_ail\_pedigree\_fix\_n

#### **Arguments**

ngen Number of generations of outbreeding

npairs Number of breeding pairs at each generation

nkids\_per Number of offspring per pair for the last generation

design How to choose crosses: either random but avoiding siblings, or completely at

17

random

#### **Details**

Advanced intercross lines (AIL) are generated from a pair of inbred lines. We cross them and then cross the F1 to generate npair breeding pairs. The subsequent ngen outbreeding generations then proceed by crossing a male and female from the preceding generation (mated completely at random, with design="random", or avoiding siblings, with design="nosib"). Each breeding pair gives a single female and a single male to the next generation, except at the last generation nkids\_per offspring are mated, in equal numbers male and female. (If nkids\_per is an odd number, the number of males and females in each sibship will differ by one, alternating between sibships, with one additional female and then one additional male.

#### Value

A data frame with five columns: individual ID, mother ID, father ID, sex, and generation. Founders have 0 for mother and father ID. Sex is coded 0 for female and 1 for male.

#### See Also

```
sim_from_pedigree(), sim_ril_pedigree(), sim_do_pedigree(), sim_4way_pedigree()
```

#### **Examples**

```
tab <- sim_ail_pedigree(12, 30)</pre>
```

```
sim\_ail\_pedigree\_fix\_n
```

Simulate AIL pedigree with fixed n

#### **Description**

Simulate a pedigree for advanced intercross lines (a table of individual, mom, dad, sex) so that the last generation reaches a desired sample size n

#### Usage

```
sim_ail_pedigree_fix_n(
  ngen = 12,
  nkids_per = 5,
  nsample_ngen = 150,
  npairs = NULL,
  method = c("last2", "sub2"),
  design = c("nosib", "random")
)
```

## Arguments

ngen	Number of generations of outbreeding
nkids_per	Number of offspring per pair for the last generation
nsample_ngen	Number of individuals desired at the last generation
npairs	Number of breeding pairs at each generation. If missing, we use 30 when method="last2" and 300 when method="sub2".
method	Method used to generate pedigree: either expand at the last two generations or generate a pedigree with a large number of pairs and select a subset to have the desired sample size.
design	How to choose crosses: either random but avoiding siblings, or completely at random

#### **Details**

The default value for npairs depends on the choice of method. For method="last2", we use a default of npairs=30; for method="sub2", we use a default of npairs=300.

## Value

A data frame with five columns: individual ID, mother ID, father ID, sex, and generation. Founders have 0 for mother and father ID. Sex is coded 0 for female and 1 for male.

#### See Also

```
sim\_from\_pedigree(), sim\_ril\_pedigree(), sim\_ail\_pedigree(), sim\_do\_pedigree(), sim\_do\_pedigree(), sim\_do\_pedigree(), sim\_do\_pedigree_fix\_n()
```

```
tab <- sim_ail_pedigree_fix_n(12)</pre>
```

sim\_crossovers 19

sim_crossovers Simulate crossover locations using the Stahl model	sim_crossovers	Simulate crossover locations using the Stahl model	
---	----------------	--	--

## **Description**

Simulate crossover locations on a single meiotic product using the Stahl model.

## Usage

```
sim_crossovers(L, m = 10, p = 0, obligate_chiasma = FALSE, Lstar = NULL)
```

## **Arguments**

1	1 41-	_ C		:	-1/
L	length	OI	cnr	ın	CIVI

m Interference parameter (m=0 is no interference)

p Proportion of chiasmata from no-interference mechanism (p=0 gives pure chi-

square model)

obligate\_chiasma

If TRUE, require an obligate chiasma on the 4-strand bundle at meiosis.

Lstar Adjusted chromosome length, if obligate\_chiasma=TRUE. Calculated if not

provided.

#### Details

Chiasma locations are a superposition of two processes: a proportion p exhibiting no interference, and a proportion (1-p) following the chi-square model with interference parameter m. Crossover locations are derived by thinning the chiasma locations with probability 1/2.

Simulations are under the Stahl model with the interference parameter being an integer. This is an extension of the chi-square model, but with chiasmata being the superposition of two processes, one following the chi-square model and the other exhibiting no interference.

#### Value

Numeric vector of crossover locations, in cM

## References

Copenhaver, G. P., Housworth, E. A. and Stahl, F. W. (2002) Crossover interference in arabidopsis. *Genetics* **160**, 1631–1639.

Foss, E., Lande, R., Stahl, F. W. and Steinberg, C. M. (1993) Chiasma interference as a function of genetic distance. *Genetics* **133**, 681–691.

Zhao, H., Speed, T. P. and McPeek, M. S. (1995) Statistical analysis of crossover interference using the chi-square model. *Genetics* **139**, 1045–1056.

20 sim\_dof1\_pedigree

#### **Examples**

```
x <- sim_crossovers(200, 10, 0)
x <- sim_crossovers(200, 10, 0.04)
x <- sim_crossovers(100, 0, 0, obligate_chiasma=TRUE)</pre>
```

sim\_dof1\_pedigree

Simulate pedigree for F1 between diversity outbreds and another inbred line

## **Description**

Simulate a pedigree for a set of DOF1 individuals: the F1 offspring of a set of diversity outbred mice and another inbred strain (such as a mutant line).

#### Usage

```
sim_dof1_pedigree(
  ngen = 12,
  npairs = 144,
  ccgen = rep(4:12, c(21, 64, 24, 10, 5, 9, 5, 3, 3)),
  nkids_per = 5,
  design = c("nosib", "random")
)
```

#### **Arguments**

ngen	Number of generations of outbreeding
npairs	Number of breeding pairs at each generation
ccgen	Vector of length npairs, with the number of generations for each CC line. If length 1, it is repeated to a vector of length npairs.
nkids_per	Number of offspring per pair for the last DO generation (each will be crossed to produce one F1)
design	How to choose crosses: either random but avoiding siblings, or completely at random

#### **Details**

Diversity outbred (DO) mice are generated from a set of 8 inbred lines. We need two individuals from each line (one female and one male) as the order of the initial crosses will be randomized; for example, sometimes the individual from line 1 will be a mother and sometimes a father. The founders are numbered 1-8 for the females from the 8 lines, and 9-16 for the corresponding males.

Diversity Outbred mice are generated by first creating a panel of partially-inbred 8-way RIL (the so-called pre-CC, for pre-Collaborative Cross). The ccgen argument specifies the number of inbreeding generations for each of the CC lines. We generate a pre-CC line for each of the npairs breeding pairs, and generate a sibling pair from each as the starting material.

sim\_do\_pedigree 21

The subsequent ngen outbreeding generations then proceed by crossing a male and female from the preceding generation (mated completely at random, with design="random", or avoiding siblings, with design="nosib"). Each breeding pair gives a single female and a single male to the next generation, except at the last generation nkids\_per offspring are mated, in equal numbers male and female. (If nkids\_per is an odd number, the number of males and females in each sibship will differ by one, alternating between sibships, with one additional female and then one additional male.

The default for ccgen is taken from Figure 1 of Svenson et al. (2012).

We assume that the F1 offspring are all from a cross DO female x line 17 male, and so the last generation of the DO is taken to be all females.

#### Value

A data frame with seven columns: individual ID, mother ID, father ID, sex, generation, a TRUE/FALSE indicator for whether DO or pre-DO, and a TRUE/FALSE indicator for whether DOF1. Founders have 0 for mother and father ID. Sex is coded 0 for female and 1 for male.

#### References

Svenson KL, Gatti DM, Valdar W, Welsh CE, Cheng R, Chesler EJ, Palmer AA, McMillan L, Churchill GA (2012) High-resolution genetic mapping using the mouse Diversity Outbred population. Genetics 190:437-447

#### See Also

```
sim_from_pedigree(), sim_ril_pedigree(), sim_ail_pedigree(), sim_4way_pedigree()
```

## **Examples**

```
tab <- sim_dof1_pedigree(8)
```

sim\_do\_pedigree

Simulate a pedigree for Diversity Outbred mice

#### **Description**

Simulate a pedigree for generating Diversity Outbred (DO) mice (a table of individual, mom, dad, sex).

## Usage

```
sim_do_pedigree(
  ngen = 12,
  npairs = 144,
  ccgen = rep(4:12, c(21, 64, 24, 10, 5, 9, 5, 3, 3)),
  nkids_per = 5,
  design = c("nosib", "random")
)
```

22 sim\_do\_pedigree

#### **Arguments**

ngen Number of generations of outbreeding
npairs Number of breeding pairs at each generation

ccgen Vector of length npairs, with the number of generations for each CC line. If

length 1, it is repeated to a vector of length npairs.

nkids\_per Number of offspring per pair for the last generation

design How to choose crosses: either random but avoiding siblings, or completely at

random

#### **Details**

Diversity outbred (DO) mice are generated from a set of 8 inbred lines. We need two individuals from each line (one female and one male) as the order of the initial crosses will be randomized; for example, sometimes the individual from line 1 will be a mother and sometimes a father. The founders are numbered 1-8 for the females from the 8 lines, and 9-16 for the corresponding males.

Diversity Outbred mice are generated by first creating a panel of partially-inbred 8-way RIL (the so-called pre-CC, for pre-Collaborative Cross). The ccgen argument specifies the number of inbreeding generations for each of the CC lines. We generate a pre-CC line for each of the npairs breeding pairs, and generate a sibling pair from each as the starting material.

The subsequent ngen outbreeding generations then proceed by crossing a male and female from the preceding generation (mated completely at random, with design="random", or avoiding siblings, with design="nosib"). Each breeding pair gives a single female and a single male to the next generation, except at the last generation nkids\_per offspring are mated, in equal numbers male and female. (If nkids\_per is an odd number, the number of males and females in each sibship will differ by one, alternating between sibships, with one additional female and then one additional male.

The default for ccgen is taken from Figure 1 of Svenson et al. (2012).

#### Value

A data frame with six columns: individual ID, mother ID, father ID, sex, generation, and TRUE/FALSE indicator for whether DO or pre-DO. Founders have 0 for mother and father ID. Sex is coded 0 for female and 1 for male.

#### References

Svenson KL, Gatti DM, Valdar W, Welsh CE, Cheng R, Chesler EJ, Palmer AA, McMillan L, Churchill GA (2012) High-resolution genetic mapping using the mouse Diversity Outbred population. Genetics 190:437-447

#### See Also

```
sim_from_pedigree(), sim_ril_pedigree(), sim_ail_pedigree(), sim_4way_pedigree()
```

```
tab <- sim_do_pedigree(8)</pre>
```

sim\_do\_pedigree\_fix\_n Simulate a pedigree for Diversity Outbreds for a target sample size

## **Description**

Simulate a pedigree for Diversity Outbred (DO) mice (a table of individual, mom, dad, sex) so that the last generation reaches a desired sample size.

## Usage

```
sim_do_pedigree_fix_n(
   ngen = 12,
   nkids_per = 5,
   nccgen = 15,
   nsample_ngen = 150,
   npairs = NULL,
   method = c("last2", "sub2", "fixcc"),
   design = c("nosib", "random"),
   selc.method = c("byfamily", "byindiv")
)
```

## Arguments

ngen	Number of generations of outbreeding
nkids_per	Number of offspring per pair for the last generation
nccgen	The number of generations for each CC line, only used when method is not "fixcc".
nsample_ngen	Number of individuals desired at the last generation
npairs	Number of breeding pairs at each generation. If missing, we use 30 when method="last2" and 300 when method="sub2".
method	Method used to generate the pedigree: either expand at the last two generations or generate a pedigree with a large number of pairs and then select a subset to have the desired sample size. With method="fixce", we use the pre-CC generations as performed at the Jackson Lab.
design	How to choose crosses: either random but avoiding siblings, or completely at random
selc.method	Method used to select the individuals from last generation.

## **Details**

The default number of breeding pairs depends on the chosen method. With method="last2", the default is npairs=30; with method="sub2", the default is npairs=300; with method="fixcc", npairs is ignored and is fixed at 144.

24 sim\_from\_pedigree

## Value

A data frame with six columns: individual ID, mother ID, father ID, sex, generation, and TRUE/FALSE indicator for whether DO or pre-DO. Founders have 0 for mother and father ID. Sex is coded 0 for female and 1 for male.

#### See Also

```
sim_from_pedigree(), sim_ril_pedigree(), sim_ail_pedigree(), sim_do_pedigree(), sim_4way_pedigree(),
sim_ail_pedigree_fix_n()
```

## **Examples**

```
tab <- sim_do_pedigree_fix_n(8)</pre>
```

sim\_from\_pedigree

Simulate genotypes for pedigree

## Description

Simulate genotypes along one chromosome for a pedigree

## Usage

```
sim_from_pedigree(
  pedigree,
  L = 100,
  xchr = FALSE,
  m = 10,
  p = 0,
  obligate_chiasma = FALSE
)
```

## Arguments

pedigree	Matrix or data frame describing a pedigree, with first four columns being individual ID, mom ID, dad ID, and sex (female as 0, male as 1).
L	Length of chromosome in cM (or a vector of chromosome lengths)
xchr	If TRUE, simulate X chromosome. (If L is a vector, this should be a vector of TRUE/FALSE values, of the same length as L, or a character string with the name of the X chromosome, in L.)
m	Crossover interference parameter, for chi-square model (m=0 corresponds to no interference).
p obligate_chiasm	proportion of crossovers coming from no-interference process

If TRUE, require an obligate chiasma on the 4-strand bundle at meiosis.

#### Value

A list with each component being the data for one individual, as produced by the cross() function. Those results are a list with two components, corresponding to the maternal and paternal chromosomes. The chromosomes are represented as lists with two components: an integer vector of alleles in chromosome intervals, and a numeric vector of locations of the right-endpoints of those intervals; these two vectors should have the same length.

If the input L is a vector, in order to simulate multiple chromosomes at once, then the output will be a list with length length(L), each component being a chromosome and having the form described above.

#### See Also

```
check_pedigree(), sim_ril_pedigree(), sim_ail_pedigree(), sim_ril_pedigree()
```

#### **Examples**

```
# simulate AIL pedigree
tab <- sim_ail_pedigree(12, 30)
# simulate data from that pedigree
dat <- sim_from_pedigree(tab)
# simulate multiple chromosomes
dat <- sim_from_pedigree(tab, c("1"=100, "2"=75, "X"=100), xchr="X")</pre>
```

```
sim_from_pedigree_allchr
```

Simulate genotypes for pedigree for multiple chromosomes

#### **Description**

Simulate genotypes along all chromosomes for a pedigree. This is a wrap up of sim\_from\_pedigree.

#### Usage

```
sim_from_pedigree_allchr(
  pedigree,
  map,
  m = 10,
  p = 0,
  obligate_chiasma = FALSE
)
```

#### Arguments

pedigree Matrix or data frame describing a pedigree, with first four columns being individual ID, mom ID, dad ID, and sex (female as 0, male as 1).

map marker locations, a list with elements for each chromosome

26 sim\_meiosis

m Crossover interference parameter, for chi-square model (m=0 corresponds to no interference).

p proportion of crossovers coming from no-interference process obligate\_chiasma

If TRUE, require an obligate chiasma on the 4-strand bundle at meiosis.

#### Value

A list with each component being the result from sim\_from\_pedigree, of length same as map.

## See Also

```
check_pedigree(), sim_ril_pedigree(), sim_ail_pedigree() sim_from_pedigree()
```

## **Examples**

```
library(qtl)
# marker map
map <- sim.map(len=rep(100, 19), n.mar=10, include.x=FALSE)
# simulate AIL pedigree
tab <- sim_ail_pedigree(12, 30)
# simulate data from that pedigree
dat <- sim_from_pedigree_allchr(tab, map)</pre>
```

sim\_meiosis

Simulate meiosis

#### **Description**

Output a random meiotic product from an input individual.

## Usage

```
sim_meiosis(parent, m = 10, p = 0, obligate_chiasma = FALSE, Lstar = NULL)
```

#### **Arguments**

parent An individual object, as output by create\_parent() or cross()

m interference parameter for chi-square model

p Proportion of chiasmata coming from no-interference process.

obligate\_chiasma

If TRUE, require an obligate chiasma on the 4-strand bundle at meiosis.

Lstar Adjusted chromosome length, if obligate\_chiasma=TRUE. Calculated if not provided.

sim\_ril\_pedigree 27

#### **Details**

Simulations are under the Stahl model with the interference parameter being an integer. This is an extension of the chi-square model, but with chiasmata being the superposition of two processes, one following the chi-square model and the other exhibiting no interference.

#### Value

A list with alleles in chromosome intervals (as integers) and locations of the right endpoints of those intervals.

#### References

Copenhaver, G. P., Housworth, E. A. and Stahl, F. W. (2002) Crossover interference in arabidopsis. *Genetics* **160**, 1631–1639.

Foss, E., Lande, R., Stahl, F. W. and Steinberg, C. M. (1993) Chiasma interference as a function of genetic distance. *Genetics* **133**, 681–691.

Zhao, H., Speed, T. P. and McPeek, M. S. (1995) Statistical analysis of crossover interference using the chi-square model. *Genetics* **139**, 1045–1056.

#### See Also

```
create_parent(), cross(), sim_crossovers(), calc_Lstar()
```

## **Examples**

```
ind <- create_parent(100, 1:2)
prod <- sim_meiosis(ind)</pre>
```

sim\_ril\_pedigree

Generate a ril pedigree

#### **Description**

Generate a pedigree for multi-way recombinant inbred lines (a table of individual, mom, dad, sex)

#### Usage

```
sim_ril_pedigree(
  ngen = 20,
  selfing = FALSE,
  parents = 1:2,
  firstind = max(parents) + 1
)
```

28 where\_het

## **Arguments**

ngen Number of generations of inbreeding

selfing If TRUE, use selfing

parents Vector of the parents' IDs. Should be integers, and length must be a power of 2

(i.e., 2, 4, 8, ...)

firstind Positive integer to assign to the first child. Must be greater than max(parents).

#### Value

A data frame with five columns: individual ID, mother ID, father ID, sex, and generation. Founders have 0 for mother and father ID. Sex is coded 0 for female and 1 for male.

#### See Also

```
sim_from_pedigree(), sim_ail_pedigree(), sim_do_pedigree(), sim_4way_pedigree()
```

## **Examples**

```
tab <- sim_ril_pedigree(7)</pre>
```

where\_het

Find heterozygous regions

#### **Description**

Find regions of heterozygosity in an individual

#### Usage

```
where_het(ind)
```

## **Arguments**

ind

An individual object, as output be create\_parent() or cross()

## Value

A matrix with two columns; each row indicates the start and end of a region where the individual is heterozygous

#### See Also

```
sim_from_pedigree(), convert2geno()
```

where\_het 29

```
mom <- create_parent(100, 1:2)
dad <- create_parent(100, 1:2)
child <- cross(mom, dad)
where_het(child)</pre>
```

# **Index**

* color	convert2geno_allchr,7		
CCcolors, 4	create_parent, 9		
* datagen	create_parent(), 10, 14, 26–28		
create_parent, 9	cross, 9		
cross, 9	cross(), 3, 9, 14, 25–28		
sim_4way_pedigree, 15	0.000(), 0, 3, 1 1, 20 20		
sim_ail_pedigree, 16	get_geno, 11		
sim_ail_pedigree_fix_n, 17	get_geno(), 6		
sim_crossovers, 19	graphics::arrows(), 13		
sim_do_pedigree, 21	<pre>graphics::rect(), 15</pre>		
sim_dof1_pedigree, 20	<b>5</b> • <b>F</b> • • • • • • • • • • • • • • • • • • •		
sim_from_pedigree, 24	mouseL_cox, 11		
sim_from_pedigree_allchr, 25	mouseL_mgi, 12		
sim_meiosis, 26	- <b>G</b> ,		
sim_ril_pedigree, 27	plot_crosslines, 13		
* datasets	plot_crosslines(), 15		
AILped, 2	plot_ind, 14		
mouseL_cox, 11	plot_ind(), <i>13</i>		
mouseL_mgi, 12			
* hplot	sim_4way_pedigree, 15		
plot_crosslines, 13	sim_4way_pedigree(), 17, 18, 21, 22, 24, 28		
plot_ind, 14	<pre>sim_ail_pedigree, 16</pre>		
* utilities	sim_ail_pedigree(), 16, 18, 21, 22, 24-26,		
calc_Lstar, 3	28		
check_pedigree, 4	<pre>sim_ail_pedigree_fix_n, 17</pre>		
collapse_do_alleles, 5	<pre>sim_ail_pedigree_fix_n(), 24</pre>		
convert2geno, 6	sim_crossovers, 19		
convert2geno_allchr,7	$sim_crossovers(), 3, 10, 27$		
get_geno, 11	sim_do_pedigree, 21		
88	sim_do_pedigree(), 5, 16-18, 24, 28		
AILped, 2	<pre>sim_do_pedigree_fix_n, 23</pre>		
	<pre>sim_do_pedigree_fix_n(), 5, 18</pre>		
calc_Lstar, 3	sim_dof1_pedigree, 20		
calc_Lstar(), 10, 27	<pre>sim_from_pedigree, 24</pre>		
CCcolors, 4	sim_from_pedigree(), 5, 6, 11, 16–18, 21,		
check_pedigree, 4	22, 24, 26, 28		
check_pedigree(), 25, 26	<pre>sim_from_pedigree_allchr, 25</pre>		
collapse_do_alleles, 5	$sim_from_pedigree_allchr(), 8$		
convert2geno, 6	sim_meiosis, 26		
convert2geno(), 8, 11, 28	sim_meiosis(), <i>3</i> , <i>9</i> , <i>10</i>		

INDEX 31