

Package ‘randomizeR’

March 19, 2023

Type Package

Title Randomization for Clinical Trials

Version 3.0.1

Maintainer Ralf-Dieter Hilgers <rhilgers@ukaachen.de>

Description This tool enables the user to choose a randomization procedure based on sound scientific criteria. It comprises the generation of randomization sequences as well the assessment of randomization procedures based on carefully selected criteria. Furthermore, ‘randomizeR’ provides a function for the comparison of randomization procedures.

License GPL (>= 3)

Encoding UTF-8

Depends R (>= 3.6.0), methods, ggplot2, plotrix, survival, mvtnorm

Imports pracma, cubature, mstate, purrr, rlang, magrittr, dplyr, coin, PwrGSD

Collate 'getDesign.R' 'randPar.R' 'pbrPar.R' 'rarPar.R' 'POC.R' 'OBF.R' 'GSD.R' 'abcdPar.R' 'randSeq.R' 'abcdSeq.R' 'getParameters.R' 'getExpectation.R' 'survEndp.R' 'expEndp.R' 'normEndp.R' 'endpoint.R' 'util.R' 'getStat.R' 'power.R' 'imbalance.R' 'corGuess.R' 'logRankDistribution.R' 'doublyF.R' 'testDec.R' 'doublyT.R' 'chronBias.R' 'selBias.R' 'bias.R' 'issue.R' 'assess.R' 'bbcdPar.R' 'bbcdSeq.R' 'ebcPar.R' 'bsdPar.R' 'bsdSeq.R' 'chenPar.R' 'chenSeq.R' 'chronBiasStepT.R' 'combinedBias.R' 'compare.R' 'crPar.R' 'crSeq.R' 'createParam.R' 'derFunc.R' 'desFunc.R' 'getDesFunc.R' 'derringerLs.R' 'derringerRs.R' 'derringerTs.R' 'desScores.R' 'desirability.R' 'ebcSeq.R' 'evaluate.R' 'gbcdPar.R' 'gbcdSeq.R' 'hadaPar.R' 'hadaSeq.R' 'maccombo.R' 'mpPar.R' 'mpSeq.R' 'pbrSeq.R' 'probUnDes.R' 'rtbdSeq.R' 'rpbrSeq.R' 'randomBlockSeq.R' 'randomizeROverview.R' 'randomizeRPackage.R' 'rarSeq.R' 'rpbrPar.R' 'tbdPar.R' 'rtbdPar.R' 'saveAssess.R' 'saveRand.R' 'stratifiedAnalysis.R' 'tbdSeq.R' 'udPar.R' 'udSeq.R'

Suggests testthat, knitr

RoxygenNote 7.2.1

NeedsCompilation no

Author David Schindler [aut],
 Diane Uschner [aut],
 Denis Razsolkov [ctb],
 Dimitar Mihaylov [ctb],
 Marcia Viviane Rueckbeil [ctb],
 Martin Manolov [ctb],
 Thi Mui Pham [ctb],
 Michael Martini [ctb],
 Ralf-Dieter Hilgers [aut, ths, cre],
 Nicole Heussen [aut, ths]

Repository CRAN

Date/Publication 2023-03-19 22:20:02 UTC

R topics documented:

randomizeR-package	4
a	5
abcdPar	5
analyse	6
assess	7
bbcdPar	8
blocks	9
bsdPar	10
chenPar	11
chronBias	12
coin	13
combineBias	14
compare	15
corGuess	16
createParam	17
createSeq	18
crPar	18
derFunc	19
desirability	20
ebcPar	21
evaluate	22
expEndp	24
gbcdPar	25
generateAllSequences	26
generateRandomSequences	28
genNcps_new	33
getCorGuesses	33
getDesFunc	34
getDesign	34

getDesScores	37
getDistributionPars	38
getExpectation	39
getProbabilities	41
getRandomizationList	42
get_p_values_new	43
GSD_OBF	44
GSD_POC	44
hadaPar	45
imbal	46
issue	47
K	48
lambda	48
method	49
mpPar	49
mti	50
mu	50
N	51
normEndp	51
overview	52
pbrPar	54
plot	55
plotDes	56
plotEv	57
plotSeq	58
probUnDes	58
randBlocks	60
randPar	60
randSeq-class	61
rarPar	61
ratio	62
rho	63
rpbrPar	63
rtbdPar	64
saveAssess	65
saveRand	66
scale	66
seed	67
selBias	67
setPower	69
shape	70
sigma	70
summary	70
survEndp	72
tbdPar	73
TV	73
type	74
udPar	74

randomizeR-package *Randomization for Clinical Trials*

Description

This tool enables the user to choose a randomization procedure based on sound scientific criteria. It comprises the generation of randomization sequences as well the assessment of randomization procedures based on carefully selected criteria. Furthermore, randomizeR provides a function for the comparison of randomization procedures.

Acknowledgement

This research is embedded in the [IDeAI project](#), which has received funding from the European Union's Seventh Framework Programme for research, technological development and demonstration under Grant Agreement no 602552.

Author(s)

David Schindler <dv.schindler@gmail.com>, Diane Uschner <Diane.Uschner@gmail.com>, Ralf-Dieter Hilgers, Nicole Heussen, Marcia Viviane Rueckbeil <marcia.rueckbeil@rwth-aachen.de>

References

- W. F. Rosenberger and J. M. Lachin (2002) *Randomization in Clinical Trials*. Wiley.
- #¹ D. Uschner, D. Schindler, R. D. Hilgers and N. Heussen (2018). "randomizeR: An R Package for the Assessment and Implementation of Randomization in Clinical Trials." *Journal of Statistical Software*, **85**(8), pp. 1-22. doi: 10.18637/jss.v085.i08 .
- D. Schindler (2016) *Assessment of Randomization Procedures in the Presence of Selection and Chronological Bias*. PhD Thesis.
- D. Uschner, R. D. Hilgers, N. Heussen (2018). "The impact of selection bias in randomized multi-arm parallel group clinical trials." *PLOS ONE*, 13(1): e0192065. doi: 10.1371/journal.pone.0192065.
- M. V. Rueckbeil, R. D. Hilgers, N. Heussen (2019). "Randomization in survival studies: An evaluation method that takes into account selection and chronological bias." *PLOS ONE*, 14(6): e0217946. doi: 10.1371/journal.pone.0217946.

See Also

For functionality for randomization procedures, see [randPar](#) and [genSeq](#). For the criteria for the assessment of randomization procedures, see [issues](#). For the assessment and comparison of randomization procedures, see [assess](#) and [compare](#).

a	<i>Function returning the adjusting parameter a slot of an S4 object</i>
---	--

Description

Function returning the adjusting parameter a slot of an S4 object

Usage

a(obj)

Arguments

obj object of class randPar

Value

the value of the adjusting parameter a of an S4 object

abcdPar	<i>Representing Accelerated Biased Coin Design</i>
---------	--

Description

Represents the randomization procedure Accelerated Biased Coin Design.

Usage

abcdPar(N, a, groups = LETTERS[1:2])

Arguments

N	integer for the total sample size of the trial.
a	nonnegative parameter which controls the degree of randomness: For decreasing a the allocations become deterministic, while for increasing a the randomization procedure tends to complete randomization.
groups	character vector of labels for the different treatments.

Details

This is a class of 'biased coins' where the probability of selecting the under-represented treatment is dependent from the absolute difference between the two treatment allocations up to the current step.

Value

S4 object of the class `abcdPar`.

References

A. B. Antognini and A. Giovagnoli (2004) A new 'biased coin design' for the sequential allocation of two treatments. *Journal of the Royal Statistical Society. Series C (Applied Statistics)* **53**, No. 4, 651-664

See Also

Other randomization procedures: `bbcdPar`, `bsdPar`, `chenPar`, `crPar`, `createParam()`, `ebcPar`, `gbcdPar`, `hadaPar`, `mpPar`, `pbrPar`, `rarPar`, `rpbrPar`, `rtbdPar`, `tbdPar`, `udPar`

analyse

Creates stratified sequences Compares stratified sequences to their respective non-stratified version under the influence of bias.

Description

Creates stratified sequences Compares stratified sequences to their respective non-stratified version under the influence of bias.

Arguments

`endp` object of class `endpoint`.
`pr` at least one object of class `randPar` or just a list of objects of class `randPar`

Details

Stratified and Non-stratified versions of a randomization sequence behave differently with respect to issues like selection bias, chronological bias or combined bias. The `analyse` function creates both versions of a sequence for each of the specified randomization procedures and analyses them in relation to the bias created according to the `theta` and `eta` values. The first argument should specify the total sample size of patients. The second argument should be one of class `normEndp` describing a normally distributed endpoint. The third argument should be the allocation ratio for the different strata. The fourth argument should be the number of strata in the clinical trial. The fifth and sixth arguments should be the selection bias effect `eta` and the time trend `theta`. The seventh argument should be a vector of strings representing different randomization procedures. The strings should be given as described by the `getDesign` function. Any additional parameters should be given after the design name of the procedure encapsulated in parenthesis.

Value

The function returns a matrix that summarizes the performance of the randomization procedures. The values for each randomization procedure represent the percentage of sequences that kept the 5

assess

Assessing randomization sequences

Description

Assesses randomization sequences based on specified issues in clinical trials.

Usage

```
assess(randSeq, ..., endp)

## S4 method for signature 'randSeq,missing'
assess(randSeq, ..., endp)

## S4 method for signature 'randSeq,endpoint'
assess(randSeq, ..., endp)
```

Arguments

randSeq	object of class randSeq.
...	at least one object of class issue or just a list of objects of the class issue.
endp	object of class endpoint, or missing.

Details

Randomization sequences behave differently with respect to issues like selection bias, chronological bias, or loss in power estimation. The `assess` function evaluates the behavior of randomization sequences with respect to these issues. The first argument should be a result of one of the functions [genSeq](#) or [getAllSeq](#). The second argument should be any number of [issues](#) arising in a clinical trial. The last argument `endp` may be provided if the assessment should take the distribution of the treatment groups into account, e.g. for power evaluation.

Value

S4 object of class `assessment` summarizing the assessment of the randomization procedure.

See Also

Representation of randomization procedures: [randPar](#)
Generation of randomization sequences: [genSeq](#)
[issues](#) for the assessment of randomization sequences

Examples

```

# assess the full set of Random Allocation Rule for N=4 patients
sequences <- getAllSeq(rarPar(4))
issue1 <- corGuess("CS")
issue2 <- corGuess("DS")
issue3 <- imbal("imb")
issue4 <- imbal("maxImb")
assess(sequences, issue1, issue2, issue3, issue4)

# assess one sequence of the Big Stick Design with respect to correct guesses
sequence <- genSeq(bsdPar(10, 2), seed = 1909)
assess(sequence, issue1)

# assess the same sequence with respect to selection bias and power for a normal endpoint
endp <- normEndp(c(2, 2), c(1, 1))
issue5 <- selBias("CS", 4, "exact")
issue6 <- setPower(2, "exact")
assess(sequence, issue1, issue5, issue6, endp = endp)

# assess the same sequence with respect to selection bias for an exponential endpoint
endp <- expEndp(lambda = c(0.5, 0.5), cenRate=0.1, accrualTime=1, cenTime=5)
issue7 <- selBias("CS", 0.1, "exact")
assess(sequence, issue1, issue7, endp = endp)

# recommended plot for the assessment of rejection probabilities
RP <- getAllSeq(crPar(6))
cB <- chronBias(type = "linT", theta = 1/6, method = "exact")
sB <- selBias(type = "CS", eta = 1/4, method = "exact")
normEndp <- normEndp(c(0, 0), c(1, 1))
A <- assess(RP, cB, sB, endp = normEndp)
D <- A$D
desiredSeq <- round(sum(D[,2][D[,3] <= 0.05 & D[,4] <= 0.05]), digits = 4)
colnames(D) <- c("Seq", "Prob", "SB", "linT")
g <- ggplot(D, aes(x = SB, y = linT))
g <- g + annotate("rect", xmin = 0, xmax = 0.05, ymin = 0, ymax = 0.05,
alpha=0.2, fill="green")
g <- g + geom_point(alpha = 1/10, size = 3, col = "orange")
g <- g + geom_vline(xintercept = 0.05, col = "red")
g <- g + geom_hline(yintercept = 0.05, col = "red")
g <- g + geom_text(data = NULL, x = 0, y = 0,
label = paste("Proportion:", desiredSeq), hjust=0, vjust=0, size = 7)
g

```

Description

Represents the randomization procedure Bayesian Biased Coin Design.

Usage

```
bbcdPar(N, a, groups = LETTERS[1:2])
```

Arguments

N integer for the total sample size of the trial.

a nonnegative parameter which controls the degree of randomness: For decreasing a the allocations become deterministic, while for increasing a the randomization procedure tends to complete randomization.

groups character vector of labels for the different treatments.

Details

Extension of Efron's biased coin design.

Value

S4 object of the class `bbcdPar`.

References

A. B. Antognini and Maroussa Zagoraiou (2014) Balance and randomness in sequential clinical trials: the dominant biased coin design. *Pharmaceutical Statistics* **13(2)**, 119-127

See Also

Other randomization procedures: [abcdPar](#), [bsdPar](#), [chenPar](#), [crPar](#), [createParam\(\)](#), [ebcPar](#), [gbcdPar](#), [hadaPar](#), [mpPar](#), [pbrPar](#), [rarPar](#), [rpbrPar](#), [rtbdPar](#), [tbdPar](#), [udPar](#)

blocks *Function returning the block slot of an S4 object*

Description

Function returning the block slot of an S4 object

Usage

```
blocks(obj)
```

Arguments

obj object of class `pbrPar`

Value

a vector with the lengths of each block of a `pbrPar` object

bsdPar	<i>Representing Big Stick Design</i>
--------	--------------------------------------

Description

Represents the randomization procedure Big Stick Design.

Usage

```
bsdPar(N, mti, groups = LETTERS[1:2])
```

Arguments

N	integer for the total sample size of the trial.
mti	maximum tolerated imbalance in patient numbers during the trial.
groups	character vector of labels for the different treatments.

Details

Tossing a fair coin as long as the difference in group sizes does not exceed the mti. If the mti is reached a deterministic allocation is done, so that the difference in group sizes is reduced.

Value

S4 object of the class bsdPar.

References

J. F. Soares and C. F. Jeff Wu (1983) Some Restricted Randomization Rules in Sequential Designs. *Comm. in Stat.*, **12**, 2017-34.

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [chenPar](#), [crPar](#), [createParam\(\)](#), [ebcPar](#), [gbcdPar](#), [hadaPar](#), [mpPar](#), [pbrPar](#), [rarPar](#), [rpbrPar](#), [rtbdPar](#), [tbdPar](#), [udPar](#)

chenPar *Representing Chen's Design*

Description

Represents the randomization procedure Chen's Design.

Usage

```
chenPar(N, mti = N, p = 0.5, groups = LETTERS[1:2])
```

Arguments

N	integer for the total sample size of the trial.
mti	maximum tolerated imbalance in patient numbers during the trial.
p	success probability of the biased coin (e.g. in Efron's Biased Coin Design).
groups	character vector of labels for the different treatments.

Details

Flip a biased coin with probability p in favor of the treatment which is allocated less frequently as long as the difference in group sizes does not exceed the `mti`. If the `mti` is reached a deterministic allocation is done, so that the difference in group sizes is reduced. If both treatments have been assigned equally often a fair coin is tossed.

Value

S4 object of the class `chenPar`.

References

Chen Yung-Pin (1999) Biased coin design with imbalance tolerance. *Comm. in Stat.*, **15**, 953-975.

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [bsdPar](#), [crPar](#), [createParam\(\)](#), [ebcPar](#), [gbcdPar](#), [hadaPar](#), [mpPar](#), [pbrPar](#), [rarPar](#), [rpbrPar](#), [rtbdPar](#), [tbdPar](#), [udPar](#)

chronBias *Representing chronological bias*

Description

Represents the issue of chronological bias in a clinical trial.

Usage

```
chronBias(type, theta, method, saltus, alpha = 0.05)
```

Arguments

type	character string, should be one of "linT", "logT", or "stepT", see Details.
theta	factor of the time trend for further details see type.
method	character string, should be one of "sim" or "exact", see Description.
saltus	integer or missing specifying the patient index (i.e. position) of the step in case of step time trend.
alpha	significance level

Details

Chronological bias can be an issue in the design of a clinical trial. The `chronBias` function is a constructor function for an S4 object of the class `chronBias` representing the issue of chronological bias, s.a. time trends, in a clinical trial. It supports two possible modes, `method="sim"` and `method="exact"`, and three different types of trend.

If `method="sim"`, the object represents the simulated type-I-error rate given the level `alpha`, the selection effect `eta` and the biasing strategy `type`. When calling `assess` for a `chronBias` object with `method="sim"`, one test decision is computed for each sequence of `randSeq`. The type-I-error rate (power) is the proportion of falsely (correctly) rejected null hypotheses.

If `method="exact"`, the object represents the exact type-I-error probability given the level `alpha`, the selection effect `eta` and the biasing strategy `type`. When calling `assess` for a `chronBias` object with `method="exact"`, the p -value of each randomization sequence is computed. For normal endpoints and two treatment groups these p -values are exact values which can be calculated from the sum of the corresponding quantiles of the doubly noncentral t-distribution. For more than two treatment groups, exact p -values are computed using a doubly noncentral F distribution. For exponential endpoints the p -values are obtained using an approximation formula.

Types of chronological bias:

`type = "linT"` Represents linear time trend. Linear time trend means that the time trend function of the patients, i.e. expected response for normal endpoints, increases evenly by $\theta/(N-1)$ with every patient included in the study, until reaching `theta` after `N` patients. Linear time trend may occur as a result of gradually relaxing in- or exclusion criteria throughout the trial. It can be represented by the formula:

$$f(i) = (i - 1)/(N - 1)\theta$$

type = "logT" Represents logarithmic time trend. Logarithmic time trend means that the time trend function of the patients, i.e. expected response for normal endpoints, increases logarithmically in the patient index by $\theta/\log(N)$ with every patient included in the study, until reaching θ after N patients. Logarithmic time trend may occur as a result of a learning curve, i.e. in a surgical trial. It can be represented by the formula:

$$\log(i)/\log(N)\theta$$

type = "stepT" Represents step trend. Step trend means that the expected response of the patients increases by θ after a given point ("saltus") in the allocation process. Step trend may occur if a new device is used after the point $c = \text{"saltus"}$, or if the medical personal changes after this point. Step time trend can be represented by the formula:

$$f(i) = 1_{c < i \leq N} \theta$$

Value

S4 object of class `chronBias`, a formal representation of the issue of chronological bias in a clinical trial.

References

G. K. Rosenkranz (2011) The impact of randomization on the analysis of clinical trials. *Statistics in Medicine*, **30**, 3475-87.

M. Tamm and R.-D. Hilgers (2014) Chronological bias in randomized clinical trials under different types of unobserved time trends. *Methods of Information in Medicine*, **53**, 501-10.

See Also

Other issues: `combineBias()`, `corGuess`, `imbal`, `issue`, `selBias`, `setPower()`

Examples

```
# create a linear time trend with theta = 0.5 for which the exact rejection probabilities
# are calculated
cbias <- chronBias("linT", 0.5, "exact")

# create a stepwise time trend with theta = 1 after 10 allocations for which the test
# decision is simulated
cbias <- chronBias("stepT", 1, "sim", 10)
```

coin

Function returning the coin slot of an S4 object

Description

Function returning the coin slot of an S4 object

Usage

```
coin(obj)
```

Arguments

obj object extending class randPar or randSeq

Value

The success probability of the biased coin

combineBias	<i>Combined bias criterion</i>
-------------	--------------------------------

Description

This class combines a selBias object and a chronBias object to a new object. In the analysis within the new object the two types of bias are treated as additive effect for normal endpoints and as multiplicative effect for exponential endpoints.

Usage

```
combineBias(selBias, chronBias)
```

Arguments

selBias object of class selBias
chronBias object of class chronBias

Value

A combined bias object that combines a selBias and a chronBias object

See Also

Other issues: [chronBias](#), [corGuess](#), [imbal](#), [issue](#), [selBias](#), [setPower\(\)](#)

Examples

```
chronBias <- chronBias(type="linT", theta=1, method="sim")  
selBias <- selBias(type="CS", eta=1, method="sim")  
combineBias(selBias, chronBias)
```

`compare`*Comparison of randomization procedures*

Description

Compares randomization procedures based on a specified issue in clinical trials.

Usage

```
compare(issue, ..., endp)

## S4 method for signature 'issue,missing'
compare(issue, ..., endp)

## S4 method for signature 'issue,endpoint'
compare(issue, ..., endp)
```

Arguments

<code>issue</code>	object of class <code>issue</code> .
<code>...</code>	at least one object of class <code>randSeq</code> or a list of objects of class <code>randSeq</code> .
<code>endp</code>	object of class <code>endpoint</code> , or <code>missing</code> .

Details

Randomization procedures behave differently with respect to issues like selection bias, chronological bias, or loss in power estimation. The `compare` function evaluates the behavior of randomization procedures with respect to one issue. Its first argument should represent one of the implemented [issues](#). The second argument should be any number of objects of the class `randSeq`. These objects represent the randomization procedures for the planned comparison. The last argument `endp` may be provided if the assessment should take the distribution of the treatment groups into account, e.g. for power evaluation.

Value

S4 object of class `comparison` summarizing the comparison of the randomization procedures.

See Also

Representation of randomization procedures: [randPar](#)
Generation of randomization sequences: [genSeq](#)
[issues](#) for the assessment of randomization sequences

Examples

```

# compare Random Allocation Rule and Big Stick for N = 4 with respect to
# correct guesses
RAR <- getAllSeq(rarPar(4))
BSD <- getAllSeq(bsdPar(4, mti = 2))
corGuess <- corGuess("CS")
(comp <- compare(corGuess, RAR, BSD))
plot(comp)

# compare the same procedures with respect to selection bias for a normal endpoint
endp <- normEndp(c(2, 2), c(1, 1))
selBias <- selBias("CS", 4, "exact")
(comp <- compare(selBias, RAR, BSD, endp = endp))
plot(comp)

# compare the same procedures with respect to selection bias for an exponential endpoint
endp <- expEndp(lambda = c(0.5, 0.5), cenRate=0.1, accrualTime=1, cenTime=5)
selBias <- selBias("CS", 0.1, "exact")
(comp <- compare(selBias, RAR, BSD, endp = endp))
plot(comp)

```

corGuess

Representing the expected number of correct guesses

Description

Represents the expected number of correct guesses of randomization sequences.

Usage

```
corGuess(type)
```

Arguments

type character string, should be one of "CS" or "DS", see Details.

Details

Selection bias can be an issue in the design of a clinical trial. The expected number of correct guesses is one measure for selection bias. The corGuess function is a constructor function for an S4 object of the class corGuess representing the issue of correct guesses in a clinical trial. The parameter type takes the following values:

"CS" refers to "convergence strategy", i.e. the investigator predicts the treatment which has hitherto occurred less often.

"DS" refers to "divergence strategy", i.e. the investigator predicts the treatment which has hitherto occurred more often.

Value

S4 object of class corGuess, a formal representation of the issue of correct guesses in a clinical trial.

References

D. Blackwell and J.L. Hodges Jr. (1957) Design for the control of selection bias. *Annals of Mathematical Statistics*, **25**, 449-60.

See Also

Other issues: [chronBias](#), [combineBias\(\)](#), [imbal](#), [issue](#), [selBias](#), [setPower\(\)](#)

 createParam

Representing any randomization procedure

Description

Represents any randomization procedure for a two-armed clinical trial.

Usage

```
createParam(method, N, mti, bc, rb, p, ini, add, filledBlock)
```

Arguments

method	method that is used to generate the (random) allocation sequence. It can take values PBR, RAR, HAD, PWR, EBC, BSD, CR, TBD, UD, and MP.
N	integer for the total sample size of the trial.
mti	maximum tolerated imbalance in patient numbers during the trial.
bc	vector which contains the lengths k_1, \dots, k_l of each block. This means that the vector bc will have one entry for each block.
rb	block lengths of the blocks that can be selected equiprobable at random.
p	success probability of the biased coin (e.g. in Efron's Biased Coin Design).
ini	integer representing the initial urn composition.
add	integer representing the number of balls that are added to the urn in each step.
filledBlock	logical whether the last block should be filled or not.

Details

Depending on the input of the user, createParam creates an object representing a randomization procedures for a two-armed clinical trial (see also [randPar](#)).

Value

S4object of the corresponding randomization procedure class.

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [bsdPar](#), [chenPar](#), [crPar](#), [ebcPar](#), [gbcdPar](#), [hadaPar](#), [mpPar](#), [pbrPar](#), [rarPar](#), [rpbrPar](#), [rtbdPar](#), [tbdPar](#), [udPar](#)

createSeq	<i>Query to create a randomization sequence of a particular randomization procedure</i>
-----------	---

Description

This function is a query to create an corresponding randomization sequence for a two-armed clinical trial. If `file` is defined, the generated sequence is automatically saved to the corresponding path.

Usage

```
createSeq(file)
```

Arguments

`file` A connection, or a character string naming the file to write to.

Value

an object `Param`, which is available

crPar	<i>Representing Complete Randomization</i>
-------	--

Description

Represents the randomization procedure Complete Randomization.

Usage

```
crPar(N, K = 2, ratio = rep(1, K), groups = LETTERS[1:K])
```

Arguments

`N` integer for the total sample size of the trial.

`K` number of treatment groups (e.g. $K=2$ if we compare one experimental against one control treatment).

`ratio` vector of length K . The total sample number N and all used block lengths (`bc`) have to be divisible by `sum(ratio)`.

`groups` character vector of labels for the different treatments.

Details

Toss a fair coin N times in case K=2 and assign the treatments according to the result of the coin. In case of K>2, replace the coin by a die with K sides.

Value

S4 object of the class crPar.

References

W. F. Rosenberger and J. M. Lachin (2002) *Randomization in Clinical Trials*. Wiley.

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [bsdPar](#), [chenPar](#), [createParam\(\)](#), [ebcPar](#), [gbcdPar](#), [hadaPar](#), [mpPar](#), [pbrPar](#), [rarPar](#), [rpbrPar](#), [rtbdPar](#), [tbdPar](#), [udPar](#)

 derFunc

Representing Derringer-Suich desirability functions

Description

Represents the Derringer-Suich desirability approach.

Usage

```
derFunc(TV, SLs, b)
```

Arguments

TV	numeric specifying the optimal desired value called the target value.
SLs	numeric vector of length at most 2 specifying the lower and/or upper specified border.
b	numeric vector of length at most 2 specifying the weight(s) for the punishment of deviations from the target value.

Details

derFunc represents the framework for left, right and two-sided desirability functions introduced by Derringer and Suich (1980). For all three different kinds of desirability functions the parameter TV must be specified. If the parameter SLs has length 1, either the left- or right-sided desirability function is created depending from whether the value is smaller (left-sided) or greater (right-sided) than the target value. By specifying SLs as a vector of length 2 a two-sided desirability function is created where the lower specified border is determined as the smaller value of SLs and thus the upper specified border is determined as the greater value. If there are no values specified for the weights, then they are automatically set to 1 (linear loss).

Value

S4 object of class `derFunc`, a formal representation of desirability functions introduced by Derringer and Suich.

References

Derringer, G., and Suich, R., (1980) Simultaneous Optimization of Several Response Variables. *Journal of Quality Technology*, **12**, 214-219.

See Also

Other desirability topics: [evaluate\(\)](#), [getDesScores\(\)](#), [plotDes\(\)](#), [plotEv\(\)](#), [probUnDes\(\)](#)

Examples

```
# create an object of a left-sided desirability function
dLeft <- derFunc(0.5, 0.3, 2)

# create an object of a right-sided desirability function
dRight <- derFunc(0.5, 0.8, 1)

# create an object of a two-sided desirability function
dLR <- derFunc(0.5, c(0.3, 0.9), c(3, 1))
```

desirability

Desirability functions within the scope of clinical trials

Description

Illustrates the interplay between functions related to desirability indices.

Details

Currently, `randomizeR` encompasses the class of desirability functions introduced by Derringer and Suich (1980) and corresponding functions to evaluate and compare randomization sequences which have been assessed on the basis of desirability indices of specific issues:

- **derFunc** represents the class of desirability functions according to Derringer-Suich (1980).
- **getDesScores** can be applied to an object of class `assessment` together with prespecified desirability functions to compare the behavior of randomization sequences (on a common scale $\in [0, 1]$).
- **plotDes** plots a `desScores` object on a radar chart.
- **evaluate** performs a comparison of sequences from different randomization sequences on the basis of object of the class `desScores`.
- **plotEv** plots an `evaluation` object on a radar chart.
- **probUnDes** computes the probability of undesired randomization sequences with respect to certain issues and desirability functions.

Examples

```

# perform a comparison of randomization sequences from different randomization procedures
# with the help of desirability functions

issue1 <- corGuess("CS")
issue2 <- chronBias(type = "linT", theta = 1/4, method = "exact")
RAR <- getAllSeq(rarPar(4))
BSD <- getAllSeq(bsdPar(4, mti = 2))
A1 <- assess(RAR, issue1, issue2, endp = normEndp(c(0,0), c(1,1)))
A2 <- assess(BSD, issue1, issue2, endp = normEndp(c(0,0), c(1,1)))

d1 <- derFunc(TV = 0.5, 0.75, 2)
d2 <- derFunc(0.05, c(0, 0.1), c(1, 1))

# apply the getDesScores function to the assessment output with the specified desirability
# functions to evaluate the behaviour of randomization sequences on a [0,1] scale

DesScore <- getDesScores(A1, d1, d2, weights = c(5/6, 1/6))
DesScore2 <- getDesScores(A2, d1, d2, weights = c(5/6, 1/6))

# plotting the desScores objects
plotDes(DesScore, quantiles = TRUE)
plotDes(DesScore2, quantiles = TRUE)

# summarize the results of getDesScore with respect to the statistic "mean"
evaluate(DesScore, DesScore2)

# plot the evaluation objects for a visualized comparison
plotEv(evaluate(DesScore, DesScore2))

# display which randomzation procedure produces more undesired randomization sequences
# with respect to certain issues and desirability functions
probUnDes(DesScore)
probUnDes(DesScore2)

```

 ebcPar

Representing Efron's Biased Coin Design

Description

Represents the randomization procedure Efron's Biased Coin Design.

Usage

```
ebcPar(N, p, groups = LETTERS[1:2])
```

Arguments

N	integer for the total sample size of the trial.
p	success probability of the biased coin (e.g. in Efron's Biased Coin Design).
groups	character vector of labels for the different treatments.

Details

Flip a biased coin with probability p in favor of the treatment which is allocated less frequently. If both treatments have been assigned equally often a fair coin is tossed.

Value

S4 object of the class `ebcPar`.

References

B. Efron (1971) Forcing a sequential experiment to be balanced. *Biometrika*, **58**, 403-17.

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [bsdPar](#), [chenPar](#), [crPar](#), [createParam\(\)](#), [gbcdPar](#), [hadaPar](#), [mpPar](#), [pbrPar](#), [rarPar](#), [rpbrPar](#), [rtbdPar](#), [tbdPar](#), [udPar](#)

evaluate

Evaluation of several randomization procedures with respect to certain desirability functions applied to specified issues.

Description

Evaluation of several randomization procedures with respect to certain desirability functions applied to specified issues.

Usage

```
evaluate(..., statistic)
```

```
## S4 method for signature 'missing'
evaluate(..., statistic)
```

```
## S4 method for signature 'character'
evaluate(..., statistic)
```

Arguments

...	at least one object of the class <code>desScores</code> or a list of objects of the class <code>desScores</code> .
statistic	character string that specifies on the basis of which statistic the evaluate function should be applied. The statistic can be chosen from "mean", "median", "min" or "max".

Details

The `evaluate` function allows the user to compare and evaluate different randomization procedures. It expects a number of objects that result when applying the `getDesScores` function to an `assess` object and specified desirability functions. The `evaluate` function summarizes the desirability scores of each randomization procedure on the basis of a prespecified statistic and incorporates them into a data frame. If no statistic is specified then it is automatically set to mean. If the function is applied to only one object it corresponds simply to `summary(getDesScores(...))`.

Value

S4 object of class `evaluation` Comparison of randomization procedures with respect to desirability functions applied to specified issues, summarized by a prespecified statistic.

References

D. Schindler *Assessment of Randomization Procedures in the Presence of Selection and Chronological Bias*. PhD Thesis.

See Also

Representation of randomization procedures: [randPar](#)

Generation of randomization sequences: [genSeq](#)

[issues](#) for the desirability of randomization sequences

Other desirability topics: [derFunc](#), [getDesScores\(\)](#), [plotDes\(\)](#), [plotEv\(\)](#), [probUnDes\(\)](#)

Examples

```
# compare Random Allocation Rule to Big Stick Design with respect to different issues
# and their corresponding desirability functions
issue1 <- corGuess("CS")
issue2 <- corGuess("DS")
RAR <- getAllSeq(rarPar(4))
BSD <- getAllSeq(bsdPar(4, mti = 2))
A1 <- assess(RAR, issue1, issue2)
A2 <- assess(BSD, issue1, issue2)

d1 <- derFunc(TV = 0.1, 0.7, 2)
d2 <- derFunc(0.5, c(0.3, 0.8), c(1, 1))
DesScore <- getDesScores(A1, d1, d2, weights = c(5/6, 1/6))
DesScore2 <- getDesScores(A2, d1, d2, weights = c(5/6, 1/6))

evaluate(DesScore, DesScore2)
evaluate(DesScore, DesScore2, statistic = "max")
```

`expEndp`*Representation of exponentially distributed endpoints*

Description

Represents exponentially distributed endpoints in clinical trials.

Usage

```
expEndp(lambda, cenRate, accrualTime = 0, cenTime)
```

Arguments

<code>lambda</code>	vector of the exponential rate parameters in each treatment group.
<code>cenRate</code>	exponential censoring rate in a survival study.
<code>accrualTime</code>	duration of the accrual period in a survival study.
<code>cenTime</code>	total duration of a survival study (maximum length of followup).

Details

The `expEnd` function is a constructor function for an S4 object of the class `expEnd` representing an exponentially distributed endpoint in a clinical trial. In conjunction with the `assess` function, exponential endpoints admit the calculation of the 'exact' type-I-error probability and power using an approximation formula.

Value

A S4 object representing an exponentially distributed endpoint in a clinical trial.

See Also

Compute exact or simulated type-I-error: [assess](#).

Other endpoint types: [normEndp](#), [survEndp](#)

Examples

```
# set the parameters of two exponentially distributed endpoints  
endp <- expEndp(lambda = c(1, 2), cenTime = 10, cenRate = 0.01)
```

`gbcdPar`*Representing Generalized Biased Coin Design*

Description

Represents the randomization procedure Generalized Biased Coin Design.

Usage

```
gbcdPar(N, rho, groups = LETTERS[1:2])
```

Arguments

<code>N</code>	integer for the total sample size of the trial.
<code>rho</code>	nonnegative parameter which may be adjusted according to how strongly it is desired to balance the experiment. If $\rho = 1$, we have Wei's urn design with $\alpha = 0$. If $\rho = 0$, we have complete randomization.
<code>groups</code>	character vector of labels for the different treatments.

Details

Generalization of Wei's urn and Efron's biased coin design.

Value

S4 object of the class `gbcdPar`.

References

R. L. Smith (1984) Sequential treatment allocation using biased coin designs. *Journal of the Royal Statistical Society B*, **46**, 519-543.
W. F. Rosenberger and J. M. Lachin (2002) Randomization in Clinical Trials. *Wiley*, 64-65

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [bsdPar](#), [chenPar](#), [crPar](#), [createParam\(\)](#), [ebcPar](#), [hadaPar](#), [mpPar](#), [pbrPar](#), [rarPar](#), [rpbrPar](#), [rtbdPar](#), [tbdPar](#), [udPar](#)

generateAllSequences *Complete set of randomization sequences*

Description

Computes all randomization sequences for the given randomization procedure, and stores them in an object along with the parameters belonging to the randomization procedure.

Usage

```
getAllSeq(obj)

## S4 method for signature 'pbrPar'
getAllSeq(obj)

## S4 method for signature 'rarPar'
getAllSeq(obj)

## S4 method for signature 'abcdPar'
getAllSeq(obj)

## S4 method for signature 'bbcdPar'
getAllSeq(obj)

## S4 method for signature 'ebcPar'
getAllSeq(obj)

## S4 method for signature 'bsdPar'
getAllSeq(obj)

## S4 method for signature 'chenPar'
getAllSeq(obj)

## S4 method for signature 'crPar'
getAllSeq(obj)

## S4 method for signature 'gbcdPar'
getAllSeq(obj)

## S4 method for signature 'hadaPar'
getAllSeq(obj)

## S4 method for signature 'mpPar'
getAllSeq(obj)

## S4 method for signature 'tbdPar'
getAllSeq(obj)
```

```
## S4 method for signature 'udPar'  
getAllSeq(obj)
```

Arguments

obj object specifying the randomization procedure, see [randPar](#) or [createParam](#).

Details

getAllSeq is a generic function which dispatches different methods depending on the type of input. The set of sequences of a procedure is computed by enumerating all possible sequences and eliminating those that are not possible in the randomization procedure specified by obj. The parameters of the randomization procedure are saved along with the sequences to ensure reproducibility of the results.

Value

An object inheriting from [randSeq](#), representing the set of randomization sequences for the given parameters. The output consists of the parameters used for the generation of the randomization sequences (see [createParam](#)) and the matrix M that stores the randomization sequences in its rows.

See Also

[createParam](#)

Examples

```
# all randomization sequences of Efron's Biased Coin Design with p = 0.667 for N = 6  
myPar <- ebcPar(6, 0.667)  
getAllSeq(myPar)  
  
# all randomization sequences of Big Stick Design with mti = 2 for N = 6  
myPar <- bsdPar(6, 2)  
getAllSeq(myPar)  
  
# all randomization sequences of Permuted Block Randomization with block sizes 4 and 2  
myPar <- pbrPar(c(4, 2))  
getAllSeq(myPar)
```

generateRandomSequences

Generate random sequences

Description

Generates randomization sequences from a given randomization procedure.

Usage

```
genSeq(obj, r, seed)
```

```
## S4 method for signature 'pbrPar,missing,numeric'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'pbrPar,numeric,numeric'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'pbrPar,missing,missing'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'pbrPar,numeric,missing'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'rarPar,numeric,numeric'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'rarPar,missing,numeric'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'rarPar,numeric,missing'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'rarPar,missing,missing'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'abcdPar,numeric,numeric'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'abcdPar,numeric,missing'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'abcdPar,missing,numeric'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'abcdPar,missing,missing'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'bbcdPar,numeric,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'bbcdPar,numeric,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'bbcdPar,missing,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'bbcdPar,missing,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'ebcPar,numeric,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'ebcPar,missing,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'ebcPar,numeric,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'ebcPar,missing,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'bsdPar,numeric,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'bsdPar,numeric,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'bsdPar,missing,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'bsdPar,missing,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'chenPar,numeric,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'chenPar,numeric,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'chenPar,missing,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'chenPar,missing,missing'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'crPar,numeric,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'crPar,missing,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'crPar,numeric,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'crPar,missing,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'gbcdPar,numeric,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'gbcdPar,numeric,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'gbcdPar,missing,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'gbcdPar,missing,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'hadaPar,numeric,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'hadaPar,missing,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'hadaPar,numeric,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'hadaPar,missing,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'mpPar,numeric,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'mpPar,missing,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'mpPar,numeric,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'mpPar,missing,missing'  
genSeq(obj, r, seed)
```

```
## S4 method for signature 'rpbrPar,missing,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'rpbrPar,numeric,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'rpbrPar,missing,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'rpbrPar,numeric,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'tbdPar,numeric,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'tbdPar,missing,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'tbdPar,numeric,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'tbdPar,missing,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'rtbdPar,numeric,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'rtbdPar,missing,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'rtbdPar,numeric,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'rtbdPar,missing,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'udPar,numeric,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'udPar,missing,numeric'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'udPar,numeric,missing'  
genSeq(obj, r, seed)  
  
## S4 method for signature 'udPar,missing,missing'  
genSeq(obj, r, seed)
```

Arguments

obj	object specifying the randomization procedure, see randPar or createParam .
r	numeric indicating the number of random sequences to be generated at random, or missing.
seed	a single value, interpreted as an integer, that specifies the seed for the random number generation.

Details

genSeq generates randomization sequences for a randomization procedure as defined by the input parameters. genSeq has two modes, according to the input.

1. genSeq(obj, r): gives r random sequences from the design specified by obj, along with the parameters stored in obj.
2. genSeq(obj): gives one random sequences from the design specified by obj, along with the parameters stored in obj.

The sequences are generated by using the Monte-Carlo sampling technique to sample from the true distribution of the sequences according to the randomization procedure specified by obj. The parameters of the randomization procedure are saved along with the sequences to ensure reproducibility of the results.

Value

An object inheriting from [randSeq](#), representing the r randomization sequences generated at random for the specified randomization procedure. The output consists of the parameters used for the generation of the randomization sequences (see [createParam](#)) and the matrix M that stores the randomization sequences in its r rows. If r is missing, one sequence is generated by default.

Examples

```
# generate randomization sequences using Complete Randomization for N = 10
myPar <- crPar(10)
genSeq(myPar, 4)
genSeq(myPar)

# generate randomization sequences using the Random Allocation Rule for N = 10
myPar <- rarPar(10)
genSeq(myPar, 4)
genSeq(myPar)

# generate randomization sequences using the Maximal Procedure with mti = 2 and N = 10
myPar <- mpPar(10, 2)
genSeq(myPar, 4)
genSeq(myPar)
```

genNcps_new	<i>Calculation of the NCPs of each randomization sequence for the doubly noncentral t-distribution</i>
-------------	--

Description

Computes the noncentrality parameters delta and lambda for the doubly noncentral t-distribution of each randomization sequence.

Usage

```
genNcps_new(randSeq, bias, endp, weight = FALSE)
```

Arguments

randSeq	a list of randSeq(rCrSeq or others) with possible varying N's
bias	a list of biases - corresponding to the different randSeq's
endp	object of the class endpoint.
weight	if set to TRUE the weight will be set to 1, according to the paper

Value

a list containing the noncentrality parameters delta and lambda of all randomization sequences.

getCorGuesses	<i>Matrix of the guesses of the investigator</i>
---------------	--

Description

Calculates the guesses of the investigator of a randomization list following the specified guessing strategy.

Usage

```
getCorGuesses(randSeq, guessing)
```

Arguments

randSeq	object of the class randSeq.
guessing	object of the class corGuess.

Value

Matrix of the guesses of the investigator following the specified guessing strategy. No guess is abbreviated with "nG".

Examples

```
myPar <- bsdPar(10, 2)
M <- genSeq(myPar, 2)
type <- corGuess("CS")
getCorGuesses(M, type)
```

getDesFunc

Type of Desirability function

Description

Generates a character vector which specifies the used desirability function and its parameters

Usage

```
getDesFunc(obj)

## S4 method for signature 'derringerLs'
getDesFunc(obj)

## S4 method for signature 'derringerRs'
getDesFunc(obj)

## S4 method for signature 'derringerTs'
getDesFunc(obj)
```

Arguments

obj object of the class desFunc.

Value

A character vector which specifies the used desirability function and its parameters

getDesign

Design of a randomization procedure

Description

Generates a character vector which specifies the used randomization method

Usage

```
getDesign(obj)

## S4 method for signature 'pbrPar'
getDesign(obj)

## S4 method for signature 'rarPar'
getDesign(obj)

## S4 method for signature 'abcdPar'
getDesign(obj)

## S4 method for signature 'abcdSeq'
getDesign(obj)

## S4 method for signature 'bbcdPar'
getDesign(obj)

## S4 method for signature 'bbcdSeq'
getDesign(obj)

## S4 method for signature 'ebcPar'
getDesign(obj)

## S4 method for signature 'bsdPar'
getDesign(obj)

## S4 method for signature 'bsdSeq'
getDesign(obj)

## S4 method for signature 'chenPar'
getDesign(obj)

## S4 method for signature 'chenSeq'
getDesign(obj)

## S4 method for signature 'crPar'
getDesign(obj)

## S4 method for signature 'crSeq'
getDesign(obj)

## S4 method for signature 'ebcSeq'
getDesign(obj)

## S4 method for signature 'gbcdPar'
getDesign(obj)
```

```
## S4 method for signature 'gbcdSeq'  
getDesign(obj)  
  
## S4 method for signature 'hadaPar'  
getDesign(obj)  
  
## S4 method for signature 'hadaSeq'  
getDesign(obj)  
  
## S4 method for signature 'mpPar'  
getDesign(obj)  
  
## S4 method for signature 'mpSeq'  
getDesign(obj)  
  
## S4 method for signature 'pbrSeq'  
getDesign(obj)  
  
## S4 method for signature 'rRtbdSeq'  
getDesign(obj)  
  
## S4 method for signature 'rRpbrSeq'  
getDesign(obj)  
  
## S4 method for signature 'rarSeq'  
getDesign(obj)  
  
## S4 method for signature 'rpbrPar'  
getDesign(obj)  
  
## S4 method for signature 'tbdPar'  
getDesign(obj)  
  
## S4 method for signature 'rtbdPar'  
getDesign(obj)  
  
## S4 method for signature 'tbdSeq'  
getDesign(obj)  
  
## S4 method for signature 'udPar'  
getDesign(obj)  
  
## S4 method for signature 'udSeq'  
getDesign(obj)
```

Arguments

`obj` object of the class `randSeq` or `randPar`.

Value

the name of the randomization procedure used

getDesScores	<i>Applying desirability functions on issues of individual randomization sequences</i>
--------------	--

Description

Applying desirability function on issues of individual randomization sequences.

Usage

```
getDesScores(assess, ..., weights)

## S4 method for signature 'assessment,missing'
getDesScores(assess, ..., weights)

## S4 method for signature 'assessment,numeric'
getDesScores(assess, ..., weights)
```

Arguments

assess	object of class <code>assessment</code> .
...	at least one object of class <code>derFunc</code> or a list of objects of the class <code>derFunc</code> .
weights	weights for computing the geometric mean of several desirability scores. If missing, the issues are automatically equally weighted.

Details

Randomization sequences behave differently with respect to issues like selection bias, chronological bias, or loss in power estimation. The `getDesScores` function evaluates the behavior of randomization sequences with respect to these issues. The difference to the `assess` function is that it scales them to $[0,1]$ and makes them easier interpretable. The first argument should be a result of the `assess` function. The second argument should be any number of `derFunc` objects that represent the desirability functions. The last argument `weights` may be provided if the desirability functions should be weighted differently.

Value

S4 object of class `desirability` summarizing the desirability of the randomization procedure.

See Also

Representation of randomization procedures: [randPar](#)
 Generation of randomization sequences: [genSeq](#)
[issues](#) for the desirability of randomization sequences
 Other desirability topics: [derFunc](#), [evaluate\(\)](#), [plotDes\(\)](#), [plotEv\(\)](#), [probUnDes\(\)](#)

Examples

```
# compute the desire-function for the full set of Random Allocation Rule for N = 4
sequences <- getAllSeq(rarPar(4))
issue1 <- corGuess("CS")
issue2 <- chronBias("linT", 0.25, "exact")
endp <- normEndp(mu = c(0,0), sigma = c(1,1))
A <- assess(sequences, issue1, issue2, endp = endp)
d1 <- derFunc(0.5, 0.75, 1)
d2 <- derFunc(0.05, 0.1, 1)

D1 <- getDesScores(A, d1, d2)
summary(D1)

D2 <- getDesScores(A, d1, d2, weights = c(3/4, 1/4))
summary(D2)
```

getDistributionPars *Get distribution parameters of a randomization list*

Description

Generates a matrix of the distribution parameters of the included patients in the clinical trial.

Usage

```
getDistributionPars(randSeq, issue, endp)

## S4 method for signature 'randSeq,missing,survEndp'
getDistributionPars(randSeq, endp)

## S4 method for signature 'randSeq,chronBias,survEndp'
getDistributionPars(randSeq, issue, endp)

## S4 method for signature 'randSeq,selBias,survEndp'
getDistributionPars(randSeq, issue, endp)

## S4 method for signature 'randSeq,combinedBias,survEndp'
getDistributionPars(randSeq, issue, endp)

## S4 method for signature 'randSeq,combinedBiasStepTrend,survEndp'
getDistributionPars(randSeq, issue, endp)
```

Arguments

randSeq	object of the class randSeq.
issue	object of the class issue (optional).
endp	object of the class endpoint.

Value

a matrix of the distribution parameters of the included patients in the clinical trial.

Examples

```
# return the shape and scale parameters of a Weibull distribution
endp <- survEndp(shape = c(1,1), scale = c(0.5,1), cenTime = 10, cenRate = 0.01)
biasSB <- selBias("CS", log(2), "exact")
randSeq <- genSeq(rpbrPar(rb = 2, N = 12))
getDistributionPars(randSeq,biasSB,endp)
```

getExpectation	<i>Get expectations of a randomization list</i>
----------------	---

Description

Generates a matrix of the expectations of the included patients in the clinical trial.

Usage

```
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,missing,survEndp'
getExpectation(randSeq, endp)

## S4 method for signature 'randSeq,missing,expEndp'
getExpectation(randSeq, endp)

## S4 method for signature 'randSeq,missing,normEndp'
getExpectation(randSeq, endp)

## S4 method for signature 'randSeq,power,normEndp'
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,chronBias,normEndp'
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,chronBias,expEndp'
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,chronBias,survEndp'
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,chronBias,missing'
getExpectation(randSeq, issue)

## S4 method for signature 'randSeq,selBias,normEndp'
```

```
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,selBias,expEndp'
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,selBias,survEndp'
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,selBias,missing'
getExpectation(randSeq, issue)

## S4 method for signature 'randSeq,combinedBias,normEndp'
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,combinedBias,expEndp'
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,combinedBias,survEndp'
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,combinedBiasStepTrend,normEndp'
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,combinedBiasStepTrend,expEndp'
getExpectation(randSeq, issue, endp)

## S4 method for signature 'randSeq,combinedBiasStepTrend,survEndp'
getExpectation(randSeq, issue, endp)
```

Arguments

<code>randSeq</code>	object of the class <code>randSeq</code> .
<code>issue</code>	object of the class <code>issue</code> (optional).
<code>endp</code>	object of the class <code>endpoint</code> (optional).

Details

It is assumed that the expectations of the included patients in a clinical trial can be influenced in three different ways:

- The strength of selection bias and the guessing strategy of the investigator (see [selBias](#)).
- The strength of a linear time trend, which is described by an object of the class [chronBias](#).
- The expectations of the investigated treatment groups can be different (see e.g. [normEndp](#)).

Value

A matrix of the expectations of the included patients in the clinical trial.

Examples

```

# get Expectation for a normal endpoint
myPar <- bsdPar(10, 2)
M <- genSeq(myPar, 2)
cs <- selBias("CS", 2, "sim")
endp <- normEndp(mu = c(2, 2), sigma = c(1, 1))
getExpectation(M, cs, endp)

# get Expectation for an exponential endpoint
cs <- selBias("CS", 0.1, "sim")
endp <- expEndp(lambda = c(0.5, 1), cenTime = 10, cenRate = 0.01)
getExpectation(M, cs, endp)

```

getProbabilities	<i>Theoretical probability for randomization sequences</i>
------------------	--

Description

Calculate theoretical probability for observed randomization sequences

Usage

```

getProb(obj)

## S4 method for signature 'abcdSeq'
getProb(obj)

## S4 method for signature 'bbcdSeq'
getProb(obj)

## S4 method for signature 'bsdSeq'
getProb(obj)

## S4 method for signature 'chenSeq'
getProb(obj)

## S4 method for signature 'crSeq'
getProb(obj)

## S4 method for signature 'ebcSeq'
getProb(obj)

## S4 method for signature 'gbcdSeq'
getProb(obj)

## S4 method for signature 'hadaSeq'

```

```
getProb(obj)

## S4 method for signature 'mpSeq'
getProb(obj)

## S4 method for signature 'pbrSeq'
getProb(obj)

## S4 method for signature 'rarSeq'
getProb(obj)

## S4 method for signature 'tbdSeq'
getProb(obj)

## S4 method for signature 'udSeq'
getProb(obj)
```

Arguments

obj object of a class inheriting from `randSeq`. Formal representation of a randomization sequences together with the parameters that belong to the procedure that generated the sequences.

Value

a matrix with theoretical probabilities for observed randomization sequences

Examples

```
myPar <- bsdPar(10, 2)
M <- genSeq(myPar, 2)
getProb(M)

# all Sequences
par <- pbrPar(bc=c(2,2))
refSet <- getAllSeq(myPar)
probs <- getProb(refSet)

# sequences with probabilities
cbind(probs, refSet$M)
```

`getRandomizationList` *Accessor function for the randomization list*

Description

Get the randomization list coded in its groups.

Usage

```
getRandList(obj)
```

Arguments

obj object specifying the randomization procedure, see [randPar](#) or [createParam](#).

Value

A matrix with all randomization sequences of a S4 object

Examples

```
myPar <- bsdPar(10, 2)
M <- genSeq(myPar, 2)
getRandList(M)
```

get_p_values_new *Calculating p values*

Description

Computes the p values based on the noncentrality parameters delta and lambda for the doubly noncentral t-distribution

Usage

```
get_p_values_new(delta, lambda, N, alpha = 0.05, df = sum(N - 2))
```

Arguments

delta	The first noncentrality parameter
lambda	The second noncentrality parameter
N	the amount of patients in the trial
alpha	significance level
df	degrees of freedom

Value

a p value

GSD_OBF *Simulate Group Sequential Design according to O'Brien and Fleming*

Description

Calculates the mean type-I-error and go-no-go value for each stage of a group sequential design study according to O'Brien and Fleming's design.

Usage

```
GSD_OBF(n, norep, K, RP, approach)
```

Arguments

n	total sample size
norep	number of simulations to be conducted
K	number of stages. (Currently available: 2, 3 and 4)
RP	the randomization procedure used.(Currently available: 'RAR', 'PBD2', 'BSD' - BSD(3),BSD(5) and BSD(10), 'EBC' - EBC(0.5), EBC(2/3) and EBC(0.8), 'CHEN' - CHEN(2/3,3), CHEN(2/3,5) and CHEN(2/3,10))
approach	the approach used, where 1 corresponds to allocating all patients at the beginning of the trial and 2 corresponds to allocating patients for each stage separately (Only needed for 'RAR', 'BSD' and 'CHEN')

Value

A Dataframe with the mean type-I-error and go-no-go values for each stage of a group sequential design study according to O'Brien and Fleming's design

Examples

```
# Simulate a GSD according to O'Brien and Fleming's design with 30 patients, 10 simulation runs,
# 3 Stages using Random Allocation Rule as a randomization procedure and applying it to
# all patients at the beginning
GSD_OBF(30,10,2,'RAR',1)
```

GSD_POC *Simulate Group Sequential Design according to Pocock*

Description

Calculates the mean type-I-error and go-no-go value for each stage of a group sequential design study according to Pocock's design.

Usage

```
GSD_POC(n, norep, K, RP, approach)
```

Arguments

n	total sample size
norep	number of simulations to be conducted
K	number of stages. (Currently available: 2, 3 and 4)
RP	the randomization procedure used.(Currently available: 'RAR', 'PBD2', 'BSD' - BSD(3),BSD(5) and BSD(10), 'EBC' - EBC(0.5), EBC(2/3) and EBC(0.8), 'CHEN' - CHEN(2/3,3), CHEN(2/3,5) and CHEN(2/3,10))
approach	the approach used, where 1 corresponds to allocating all patients at the beginning of the trial and 2 corresponds to allocating patients for each stage separately (Only needed for 'RAR', 'BSD' and 'CHEN')

Value

A Dataframe with the mean type-I-error and go-no-go values for each stage of a group sequential design study according to Pocock's design/

Examples

```
# Simulate a GSD according to Pocock's design with 30 patients, 10 simulation runs,
# 2 Stages using Random Allocation Rule as a randomization procedure and applying it to
# all patients at the beginning
GSD_POC(30,10,2,'RAR',1)
```

hadaPar

Representing Hadamard Randomization

Description

Represents the randomization procedure Hadamard Randomization.

Usage

```
hadaPar(N, groups = LETTERS[1:2])
```

Arguments

N	integer for the total sample size of the trial.
groups	character vector of labels for the different treatments.

Details

Hadamard randomization has been proposed by R.A. Bailey. The key idea is to use the columns of a special Hadamard Matrix as a randomization scheme. The implemented algorithm uses the Hadamard Matrix with N=12 columns proposed in the paper, see references.

Value

S4 object of the class hadaPar.

Note

getProb and getAllSeq are currently only supported for hadaPar with total sample size N=12.

References

R.A. Bailey and P.R. Nelson (2003) Hadamard Randomization: A valid restriction of random permuted blocks. *Biometrical Journal*, **45**, 554-60.

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [bsdPar](#), [chenPar](#), [crPar](#), [createParam\(\)](#), [ebcPar](#), [gbcdPar](#), [mpPar](#), [pbrPar](#), [rarPar](#), [rpbrPar](#), [rtbdPar](#), [tbdPar](#), [udPar](#)

 imbal

Representing the allocation imbalance

Description

Represents the imbalance of the treatment assignments of patients in a clinical trial.

Usage

```
imbal(type)
```

Arguments

type character string, should be one of "imb", "absImb", "loss", or "maxImb", see Details.

Details

Balance of the treatment assignment of patients can be an issue in the design of a clinical trial. The imbal function is a constructor function for an S4 object of class imbal representing the issue of imbalance of a clinical trial. The parameter type can take the following values: The type

"imb" the final imbalance, i.e. difference in group sizes at the end of a trial

"absImb" the absolute value of the final imbalance

"loss" the loss in power estimation, i.e. imb^2/N

"maxImb" the maximal attained imbalance during the trial

Value

S4 object of class `imbal`, a formal representation of the issue of imbalance in a clinical trial.

References

A.C. Atkinson (2014) Selecting a biased coin design. *Statistical Science*, **29**, Vol. 1, 144-163.

See Also

Other issues: [chronBias](#), [combineBias\(\)](#), [corGuess](#), [issue](#), [selBias](#), [setPower\(\)](#)

issue

Assessment criteria for clinical trials

Description

Summarizes the criteria for the assessment of randomization procedures.

Details

Randomization in clinical trials is supposed to control certain properties in clinical trials. In the `randomizeR` package, these properties are called `issues`. It is crucial to decide which of the issues is relevant in the present clinical trial, because a randomization procedure that manages well one issue might behave very badly for another. The issues include

- **Selection bias** can occur if future treatment allocations are predictable due to restricted randomization and unmasking of past treatment assignments. The influence of selection bias on the test decision is represented by the `selBias` class. The measure for the predictability of a randomization procedure is implemented in the `corGuess` class representing the expected number of correct guesses.
- **Chronological bias** can occur if a time trend is present in the data. Time trends occur due to learning curves, relaxed inclusion/ exclusion criteria or new co-medication. Chronological bias is represented by the `chronBias` class.
- **Additive combination of chronological and selection bias** may occur if a time trend and selection bias are present in the data. The combined bias is represented by the `combineBias` class.
- **Balance** is important in order to ensure proper power estimation properties of the treatments. However, a high degree of balance favors selection bias. Depending on the clinical context, a randomization procedure should be chosen that admits a suitable imbalance. Imbalance bias is represented by the `imbal` class. The power loss due to imbalance can be assessed directly via the `setPower` class

See Also

Representation of randomization procedures: [randPar](#)

Generation of randomization sequences: [genSeq](#)

Assessment of randomization sequences: [assess](#)

Comparison of randomization sequences: [compare](#)

Other issues: [chronBias](#), [combineBias\(\)](#), [corGuess](#), [imbal](#), [selBias](#), [setPower\(\)](#)

K *Function returning the number of trial arms slot of an S4 object*

Description

Function returning the number of trial arms slot of an S4 object

Usage

`K(obj)`

Arguments

`obj` object of class `randPar`

Value

The number of trial arms

lambda *Method returning the rate parameter of an expEndp S4 object*

Description

Method returning the rate parameter of an `expEndp` S4 object

Usage

`lambda(obj)`

Arguments

`obj` object of class `expEndp`

method	<i>Function returning the method of an S4 object</i>
--------	--

Description

Function returning the method of an S4 object

Usage

```
method(obj)
```

Arguments

obj object inheriting from randPar

Value

The method of an S4 object

mpPar	<i>Representing Maximal Procedure</i>
-------	---------------------------------------

Description

Represents the Maximal Procedure.

Usage

```
mpPar(N, mti, ratio = c(1, 1), groups = LETTERS[1:2])
```

Arguments

N integer for the total sample size of the trial.
 mti maximum tolerated imbalance in patient numbers during the trial.
 ratio vector of length K. The total sample number N and all used block lengths (bc) have to be divisible by sum(ratio).
 groups character vector of labels for the different treatments.

Details

Fix the total sample size N and the mti. Afterwards, the patients are assigned to each treatment arm according to the ratio. All randomization sequences are equiprobable.

Value

S4 object of the class mpPar.

References

V.W. Berger, A. Ivanova and M.D. Knoll (2003) Minimizing predictability while retaining balance through the use of less restrictive randomization procedures. *Statistics in Medicine*, **19**, 3017-28.

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [bsdPar](#), [chenPar](#), [crPar](#), [createParam\(\)](#), [ebcPar](#), [gbsdPar](#), [hadaPar](#), [pbrPar](#), [rarPar](#), [rpbrPar](#), [rtbdPar](#), [tbdPar](#), [udPar](#)

mti	<i>Function returning the MTI slot of an S4 object</i>
-----	--

Description

Function returning the MTI slot of an S4 object

Usage

```
mti(obj)
```

Arguments

obj	object of class bsdPar or mpPar
-----	---------------------------------

mu	<i>Access the expectation value slot of a normEndp S4 object</i>
----	--

Description

Access the expectation value slot of a normEndp S4 object

Usage

```
mu(obj)
```

Arguments

obj	object of class normEndp
-----	--------------------------

N *Function returning the sample size slot of an S4 object*

Description

Function returning the sample size slot of an S4 object

Usage

N(obj)

Arguments

obj object inheriting from randPar

Value

the sample size slot of an S4 object

normEndp *Representation of normally distributed endpoints*

Description

Represents normally distributed endpoints in clinical trials.

Usage

normEndp(mu, sigma)

Arguments

mu vector of the expected responses of the treatment groups, should have length K (i.e. one entry for each treatment group).

sigma vector of the standard deviations in each treatment group, should have length K (i.e. one entry for each treatment group).

Details

The normEnd function is a constructor function for an S4 object of the class normEnd representing a normally distributed endpoint in a clinical trial. In conjunction with the assess function, normal endpoints admit the calculation of the exact type-I-error probability and power.

Value

A S4 object that represents a normally distributed endpoint in a clinical trial

See Also

Compute exact or simulated type-I-error: [assess](#).

Other endpoint types: [expEndp](#), [survEndp](#)

Examples

```
# set the parameters of two normally distributed endpoints
endp <- normEndp(mu = c(1, 2), sigma = c(1, 1))
```

 overview

Overview over the parameters used in the randomizeR package

Description

This list of parameters yields a comprehensive overview of the parameters used in the randomizeR package.

Arguments

<code>a</code>	nonnegative parameter which controls the degree of randomness: For decreasing <code>a</code> the allocations become deterministic, while for increasing <code>a</code> the randomization procedure tends to complete randomization.
<code>accrualTime</code>	duration of the accrual period in a survival study.
<code>add</code>	integer representing the number of balls that are added to the urn in each step.
<code>alpha</code>	the significance level of the test in each simulation.
<code>bc</code>	vector which contains the lengths k_1, \dots, k_l of each block. This means that the vector <code>bc</code> will have one entry for each block.
<code>b</code>	numeric vector of length at most 2 specifying the weight(s) for the punishment of deviations from the target value.
<code>cenRate</code>	exponential censoring rate in a survival study.
<code>cenTime</code>	total duration of a survival study (maximum length of followup).
<code>d</code>	effect size.
<code>df</code>	degrees of freedom (i.a. $N-2$).
<code>eta</code>	numeric specifying the magnitude of selection bias.
<code>file</code>	A connection, or a character string naming the file to write to.
<code>filledBlock</code>	logical whether the last block should be filled or not.
<code>FTI</code>	final tolerated imbalance. This is the difference in number of patients of groups A and B that is permitted at the end of a trial. Usually this is set to zero.
<code>groups</code>	character vector of labels for the different treatments.
<code>ini</code>	integer representing the initial urn composition.

<code>k</code>	length of the block to be permuted. <code>k</code> should be divisible by the number of treatment arms.
<code>K</code>	number of treatment groups (e.g. $K=2$ if we compare one experimental against one control treatment).
<code>lb</code>	lower bound for the starting value of the poisson distribution.
<code>lambda</code>	vector of the exponential rate parameters in each treatment group.
<code>method</code>	method that is used to generate the (random) allocation sequence. It can take values PBR, RAR, HAD, PWR, EBC, BSD, CR, TBD, UD, and MP.
<code>mti</code>	maximum tolerated imbalance in patient numbers during the trial.
<code>mu</code>	vector of the expected responses of the treatment groups, should have length K (i.e. one entry for each treatment group).
<code>N</code>	integer for the total sample size of the trial.
<code>name</code>	name of a variable.
<code>obj</code>	object specifying the randomization procedure, see randPar or createParam .
<code>object</code>	any R object.
<code>p</code>	success probability of the biased coin (e.g. in Efron's Biased Coin Design).
<code>pr</code>	vector with patient responses, i.e. each patients resulting value after the treatment.
<code>q</code>	"cut-off" value in $[0.5, 1]$. This is the ratio of patients up from which the experimenter imposes selection bias on the data.
<code>r</code>	numeric indicating the number of random sequences to be generated at random, or missing.
<code>ratio</code>	vector of length K . The total sample number N and all used block lengths (<code>bc</code>) have to be divisible by <code>sum(ratio)</code> .
<code>rb</code>	block lengths of the blocks that can be selected equiprobable at random.
<code>rho</code>	nonnegative parameter which may be adjusted according to how strongly it is desired to balance the experiment. If $\rho = 1$, we have Wei's urn design with $\alpha = 0$. If $\rho = 0$, we have complete randomization.
<code>rsob</code>	randomization sequence (of one block).
<code>rs</code>	randomization sequence (of all blocks).
<code>S</code>	matrix for the computation of the probabilities in the maximal procedure.
<code>saltus</code>	integer or missing specifying the patient index (i.e. position) of the step in case of step time trend.
<code>seed</code>	a single value, interpreted as an integer, that specifies the seed for the random number generation.
<code>sigma</code>	vector of the standard deviations in each treatment group, should have length K (i.e. one entry for each treatment group).
<code>SLs</code>	numeric vector of length at most 2 specifying the lower and/or upper specified border.
<code>theta</code>	factor of the time trend for further details see <code>type</code> .

type	character vector indicating which biasing strategy the experimenter is using (selection bias) and which other bias is present in the clinical trial (e.g. time trend). All biases included in the vector are combined (i.e. added up) to form the total bias. Possible values are "none" (if no bias occurs), "CS" (resp. "DS") (if the experimenter uses the convergence (resp. divergence) strategy to invoke selection bias), LinT for linear time trend, LogT for log-linear time trend, StepT for step time trend, SigT for sigmoid time trend, PWR for knowledge of all up to the first observation in each block, MTI the next observation after reaching the maximal tolerated imbalance is reached will be known to the physician.
TV	numeric specifying the optimal desired value called the target value.
ub	upper bound for the last value of the poisson distribution.
varEq	logical parameter for the t.test: Shall the variances be treated as equal (TRUE=t.test) or different (FALSE= Welch.test).
x	a variable x.
allocRatio	numerical vector that represents the allocation ratio for the different strata in a clinical trial
strata	numeric specifying the number of strata in a clinical trial
maxcombo	logical specifying if the maxcombo test is used
weights	numeric specifying the weights used for the test. Unless specified an unweighted test is conducted.

pbrPar

Representing Permuted Block Randomization

Description

Represents the randomization procedure Permuted Block Randomization.

Usage

```
pbrPar(bc, K = 2, ratio = rep(1, K), groups = LETTERS[1:K])
```

Arguments

bc	vector which contains the lengths k_1, \dots, k_l of each block. This means that the vector bc will have one entry for each block.
K	number of treatment groups (e.g. $K=2$ if we compare one experimental against one control treatment).
ratio	vector of length K. The total sample number N and all used block lengths (bc) have to be divisible by $\text{sum}(\text{ratio})$.
groups	character vector of labels for the different treatments.

Details

Fix the block constellation bc , the number of treatment groups K , and the vector of the ratio. Afterwards, in each block the patients are assigned according to the ratio to the corresponding treatment groups. All generated randomization sequences are equiprobable.

Value

S4 object of the class `pbrPar`.

References

W. F. Rosenberger and J. M. Lachin (2002) *Randomization in Clinical Trials*. Wiley.

See Also

Other randomization procedures: `abcdPar`, `bbcdPar`, `bsdPar`, `chenPar`, `crPar`, `createParam()`, `ebcPar`, `gbcdPar`, `hadaPar`, `mpPar`, `rarPar`, `rpbrPar`, `rtbdPar`, `tbdPar`, `udPar`

plot	<i>Generic plotting of comparison objects</i>
------	---

Description

Generic plotting of comparison objects

Usage

```
plot(x, y, ...)
```

```
## S4 method for signature 'comparison,character'
```

```
plot(x, y)
```

```
## S4 method for signature 'comparison,missing'
```

```
plot(x, y)
```

Arguments

<code>x</code>	object of class <code>comparison</code> .
<code>y</code>	character <code>"boxplot"</code> , or <code>"violin"</code> , or <code>"missing"</code> .
<code>...</code>	<code>"missing"</code>

Details

Creates a box- or violinplot of an object `x` of the class `comparison`.

Value

A plot created with the additional package `ggplot2`.

See Also

[compare](#) for creating S4 objects of the class comparison

Examples

```
# compare Random Allocation Rule and Big Stick for N = 4 with respect to
# correct guesses
RAR <- getAllSeq(rarPar(4))
BSD <- getAllSeq(bsdPar(4, mti = 2))
corGuess <- corGuess("CS")
comp <- compare(corGuess, RAR, BSD)
plot(comp)
```

plotDes

desScore plotting

Description

Plot of an desScore object.

Usage

```
plotDes(desScore, labels, colAv = "red", quantiles = FALSE)
```

Arguments

desScore	object of type desScore.
labels	labels used in the plot. Can be NULL.
colAv	color of the line representing the average of the desirability scores in the plot.
quantiles	logical whether the quantiles should be depicted in the plot.

Value

a plot of an desScore object

See Also

Other desirability topics: [derFunc](#), [evaluate\(\)](#), [getDesScores\(\)](#), [plotEv\(\)](#), [probUnDes\(\)](#)

Examples

```
# compute the desirability scores of the full set of PBR(4)
sequences <- getAllSeq(rarPar(4))
issue1 <- corGuess("CS")
issue2 <- chronBias("linT", 1/4, "exact")
endp <- normEndp(mu = c(0,0), sigma = c(1,1))
A <- assess(sequences, issue1, issue2, endp = endp)
d1 <- derFunc(0.5, 0.75, 1)
d2 <- derFunc(0.05, 0.1, 1)

D <- getDesScores(A, d1, d2)
summary(D)
plotDes(D)
plotDes(D, quantiles = TRUE)
```

plotEv

Evaluation plotting

Description

Plot of an evaluation object.

Usage

```
plotEv(evaluation, labels, cols)
```

Arguments

evaluation object of type evaluation.
labels labels used in the plot. Can be NULL.
cols colors of the lines representing the desirability scores in the plot. Can be NULL.

Value

A plot of an evaluation object

See Also

Other desirability topics: [derFunc](#), [evaluate\(\)](#), [getDesScores\(\)](#), [plotDes\(\)](#), [probUnDes\(\)](#)

Examples

```
# compare Random Allocation Rule to Big Stick Design with respect to different issues
# and their corresponding desirability functions
issue1 <- corGuess("CS")
issue2 <- chronBias(type = "linT", theta = 1/4, method = "exact")
RAR <- getAllSeq(rarPar(4))
BSD <- getAllSeq(bsdPar(4, mti = 2))
```

```

A1 <- assess(RAR, issue1, issue2, endp = normEndp(c(0,0), c(1,1)))
A2 <- assess(BSD, issue1, issue2, endp = normEndp(c(0,0), c(1,1)))

d1 <- derFunc(TV = 0.5, 0.75, 2)
d2 <- derFunc(0.05, c(0, 0.1), c(1, 1))
DesScore <- getDesScores(A1, d1, d2, weights = c(5/6, 1/6))
DesScore2 <- getDesScores(A2, d1, d2, weights = c(5/6, 1/6))

E <- evaluate(DesScore, DesScore2)
plotEv(E)

```

plotSeq *Sequence plotting*

Description

Plot all randomization sequences of a randSeq object

Usage

```
plotSeq(sequences, plotAllSeq = FALSE, emph = NA, rs = NA)
```

Arguments

sequences	object of type randSeq
plotAllSeq	logical. If plotAllSeq=TRUE, the complete set of randomization sequences will be plotted in light gray.
emph	integer indicating which sequence should be highlighted in blue.
rs	vector of a randomization sequence that should be highlighted.

Value

A plot of all randomization sequences of a randSeq object.

probUnDes *Computing the probability of having desirability scores of zero*

Description

Computing the probability of having desirability scores of zero for each desirability function applied to an issue.

Usage

```
probUnDes(desScore)

## S4 method for signature 'desScores'
probUnDes(desScore)
```

Arguments

desScore an object of the class desScores, i.e. an object resulting from applying the function [getDesScores](#)

Details

The function probUnDes expects an object that results from the [getDesScores](#) function. For each issue it computes the probability that it achieves an undesirable score, i.e. a desirability score of 0. In doing so, it weights the zero desirability scores with the probability that the sequence occurs.

Value

S4 object of class probUnDesirable computing the probability of getting undesirable scores, i.e. desirability scores of 0.

See Also

Representation of randomization procedures: [randPar](#)

Generation of randomization sequences: [genSeq](#)

[issues](#) for the desirability of randomization sequences

Other desirability topics: [derFunc](#), [evaluate\(\)](#), [getDesScores\(\)](#), [plotDes\(\)](#), [plotEv\(\)](#)

Examples

```
# compare Random Allocation Rule to Big Stick Design with respect to different issues
# and their corresponding desirability functions
RAR <- getAllSeq(rarPar(4))
issue1 <- corGuess("CS")
issue2 <- corGuess("DS")
A1 <- assess(RAR, issue1, issue2)

d1 <- derFunc(TV = 0.1, 0.7, 2)
d2 <- derFunc(0.5, c(0.3, 0.8), c(1, 1))
DesScore <- getDesScores(A1, d1, d2, weights = c(5/6, 1/6))

probUnDes(DesScore)
```

randBlocks	<i>Function returning the block slot of an S4 object</i>
------------	--

Description

Function returning the block slot of an S4 object

Usage

```
randBlocks(obj)
```

Arguments

obj object of class pbrPar

Value

a vector with the lengths of each random block of a pbrPar object

randPar	<i>Settings for randomization procedures</i>
---------	--

Description

Randomization procedures in randomizeR are represented by objects that inherit from randPar. The representation can then be used in order to generate randomization sequences. In order generate a representation of a randomization procedure, call [createParam](#) or one of the following functions.

Supported randomization procedures

- Complete Randomization ([crPar](#))
- Random Allocation Rule ([rarPar](#))
- Permuted Block Randomization ([pbrPar](#))
- Permuted Block Randomization with random block length ([rpbrPar](#))
- Truncated Binomial Design ([tbdPar](#))
- Truncated Binomial Design with random block length ([rtbdPar](#))
- Efron's Biased Coin Design ([ebcPar](#))
- Big Stick Design ([bsdPar](#))
- Maximal Procedure ([mpPar](#))
- Wei's Urn Design ([udPar](#))
- Chen's Design ([chenPar](#))
- Generalized Biased Coin Design ([gbcdPar](#))
- Accelerated Biased Coin Design ([abcdPar](#))
- Bayesian Biased Coin Design ([bbcdPar](#))
- Hadamard Randomization ([hadaPar](#))

See Also

Generate randomization sequences [genSeq](#). Calculate the the complete set of randomization sequences of a randomization procedure. [getAllSeq](#).

 randSeq-class

An S4 Class for the representation of randomization sequences

Description

This set of classes provides functionality of storing randomization sequences of different randomization procedures along with the parameters representing the design.

Slots

N total number of patients included in the trial

M matrix containing randomization sequences of length N in its rows.

K number of treatment groups

groups character string of length K defining the names of the treatment groups

ratio ratio of patients between the different groups

 rarPar

Representing Random Allocation Rule

Description

Represents the randomization procedure Random Allocation Rule.

Usage

```
rarPar(N, K = 2, ratio = rep(1, K), groups = LETTERS[1:K])
```

Arguments

N integer for the total sample size of the trial.

K number of treatment groups (e.g. K=2 if we compare one experimental against one control treatment).

ratio vector of length K. The total sample number N and all used block lengths (bc) have to be divisible by `sum(ratio)`.

groups character vector of labels for the different treatments.

Details

Fix a total sample size N the number of treatment groups K , and the vector of the `ratio`. Afterwards, all patients are assigned according to the `ratio` to the corresponding treatment groups. All randomization sequences are equiprobable.

Value

S4 object of the class `rarPar`.

References

W. F. Rosenberger and J. M. Lachin (2002) *Randomization in Clinical Trials*. Wiley.

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [bsdPar](#), [chenPar](#), [crPar](#), [createParam\(\)](#), [ebcPar](#), [gbcdPar](#), [hadaPar](#), [mpPar](#), [pbrPar](#), [rpbrPar](#), [rtbdPar](#), [tbdPar](#), [udPar](#)

`ratio`

Function returning the allocation ratio slot of an S4 object

Description

Function returning the allocation ratio slot of an S4 object

Usage

```
ratio(obj)
```

Arguments

`obj` object of class `randPar`

Value

A vector containng the allocation ratio of an S4 object

rho	<i>Function returning the adjusting parameter rho slot of an S4 object</i>
-----	--

Description

Function returning the adjusting parameter rho slot of an S4 object

Usage

```
rho(obj)
```

Arguments

obj	object of class randPar
-----	-------------------------

Value

the value of the rho parameter of an S4 object

rpbrPar	<i>Representing Randomized Permuted Block Randomization</i>
---------	---

Description

Represents the randomization procedure Randomized Permuted Block Randomization.

Usage

```
rpbrPar(
  N,
  rb,
  K = 2,
  ratio = rep(1, K),
  groups = LETTERS[1:K],
  filledBlock = FALSE
)
```

Arguments

N	integer for the total sample size of the trial.
rb	block lengths of the blocks that can be selected equiprobable at random.
K	number of treatment groups (e.g. K=2 if we compare one experimental against one control treatment).
ratio	vector of length K. The total sample number N and all used block lengths (bc) have to be divisible by sum(ratio).
groups	character vector of labels for the different treatments.
filledBlock	logical whether the last block should be filled or not.

Details

Fix the possible random block lengths `rb`, the number of treatment groups `K`, the sample size `N` and the vector of the `ratio`. Afterwards, one block length is randomly selected of the random block lengths. The patients are assigned according to the ratio to the corresponding treatment groups. This procedure is repeated until `N` patients are assigned. Within each block all possible randomization sequences are equiprobable.

Value

S4 object of the class `rpbrPar`.

References

W. F. Rosenberger and J. M. Lachin (2002) *Randomization in Clinical Trials*. Wiley.

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [bsdPar](#), [chenPar](#), [crPar](#), [createParam\(\)](#), [ebcPar](#), [gbcdPar](#), [hadaPar](#), [mpPar](#), [pbrPar](#), [rarPar](#), [rtbdPar](#), [tbdPar](#), [udPar](#)

 rtbdPar

Representing Randomized Truncated Binomial Design

Description

Represents the randomization procedure Randomized Truncated Binomial Design.

Usage

```
rtbdPar(N, rb = N, groups = LETTERS[1:2], filledBlock = FALSE)
```

Arguments

<code>N</code>	integer for the total sample size of the trial.
<code>rb</code>	block lengths of the blocks that can be selected equiprobable at random.
<code>groups</code>	character vector of labels for the different treatments.
<code>filledBlock</code>	logical whether the last block should be filled or not.

Details

Fix the possible random block lengths `rb` and the sample size of the trial `N`. Afterwards, one block length is randomly selected of the random block lengths. In this block a fair coin is tossed for the patient assignments until half of the patients have been assigned to one of the treatment arms. Afterwards, the block is filled with the other treatment. This procedure is repeated until `N` patients are assigned.

Value

S4 object of the class `rtbdPar`.

References

W. F. Rosenberger and J. M. Lachin (2002) *Randomization in Clinical Trials*. Wiley.

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [bsdPar](#), [chenPar](#), [crPar](#), [createParam\(\)](#), [ebcPar](#), [gbcdPar](#), [hadaPar](#), [mpPar](#), [pbrPar](#), [rarPar](#), [rpbrPar](#), [tbdPar](#), [udPar](#)

saveAssess

Saving an assess object

Description

Saves the full information of an assess object in a .csv data sheet.

Usage

```
saveAssess(obj, file = "assessObject.csv")
```

Arguments

`obj` object of class `assessment`, e.g. the output of the [assess](#) function.
`file` A connection, or a character string naming the file to write to.

Value

Creates a .csv data in the home folder.

See Also

Other saving functions: [saveRand\(\)](#)

saveRand	<i>Saving a randomization lists</i>
----------	-------------------------------------

Description

Saves the parameters of a randSeq object in a .csv data sheet.

Usage

```
saveRand(obj, file = "randList.csv")
```

Arguments

obj	object of class randSeq containing a single randomization sequence.
file	A connection, or a character string naming the file to write to.

Value

Creates a .csv data in the home folder and saves the randomization list as a column vector.

See Also

Other saving functions: [saveAssess\(\)](#)

scale	<i>Method returning the scale parameter of an survEndp S4 object</i>
-------	--

Description

Method returning the scale parameter of an survEndp S4 object

Usage

```
scale(obj)
```

Arguments

obj	object of class survEndp
-----	--------------------------

seed	<i>Function returning the allocation seed slot of an object</i>
------	---

Description

Returns the seed that was either generated at random or user specified. The seed can be specified for any random operation e.g. `genSeq`.

Usage

```
seed(obj)
```

Arguments

`obj` object specifying the randomization procedure, see [randPar](#) or [createParam](#).

selBias	<i>Representing selection bias</i>
---------	------------------------------------

Description

Represents the issue of selection bias in a clinical trial.

Usage

```
selBias(type, eta, method, alpha = 0.05, delta = 0)
```

Arguments

<code>type</code>	character string, should be one of "CS", "CS2" or "DS", see Details.
<code>eta</code>	numeric specifying the magnitude of selection bias.
<code>method</code>	character string, should be one of "sim" or "exact", see Details.
<code>alpha</code>	significance level.
<code>delta</code>	parameter of selection bias used for calculating shape and scale of the Weibull distribution with exponential endpoints

Details

Selection bias can be an issue in the design of a clinical trial. The `selBias` function is a constructor function for an S4 object of the class `selBias` representing the issue of third order selection bias in a clinical trial. It supports two possible modes, `method="sim"` and `method="exact"`. This representation is particularly useful in interaction with the [assess](#) function.

method="sim" Represents the simulated type-I-error rate given the level alpha, the selection effect eta and the biasing strategy type. When calling assess for a selBias object with method="sim", one test decision is computed for each sequence of randSeq. The type-I-error rate (power) is the proportion of falsely (correctly) rejected null hypotheses.

method="exact" Represents the exact type-I-error probability given the level alpha, the selection effect eta and the biasing strategy type. When calling assess for a selBias object with method="exact", the p -value of each randomization sequence is computed. For normal endpoints and two treatment groups these p -values are exact values which can be calculated from the sum of the corresponding quantiles of the doubly noncentral t-distribution. For more than two treatment groups, exact p -values are computed using a doubly noncentral F distribution. For exponential endpoints the p -values are obtained using an approximation formula.

It also supports three types of selection bias:

type="DS" Refers to the divergence strategy according to Blackwell and Hodges (1957). Under this guessing strategy, the investigator guesses that the upcoming treatment is the one that has so far been allocated *more* frequently.

type="CS" Refers to the convergence strategy according to Blackwell and Hodges (1957). Under this guessing strategy, the investigator guesses that the upcoming treatment is the one that has so far been allocated *less* frequently. In multi-arm trials, type="CS" refers to the first generalization of the convergence strategy according to Uschner et al (2018). The investigator guesses the treatment that had been allocated less frequently whenever all the treatments of the opposite group are larger than the smallest of the present group.

type="CS2" In trials with two treatment arms, type="CS2" is equivalent to type="CS". In multi-arm trials, type="CS2" refers to the second generalization of convergence strategy according to Uschner et al (2018). The investigator guesses the treatment that had been allocated less frequently whenever all the treatments of the opposite group are larger than the smallest of the present group.

Value

S4 object of class selBias, a formal representation of the issue of selection bias in a clinical trial.

References

- D. Blackwell and J.L. Hodges Jr. (1957) Design for the control of selection bias. *Annals of Mathematical Statistics*, **25**, 449-60.
- M. Proschan (1994) Influence of selection bias on the type-I-error rate under random permuted block designs. *Statistica Sinica*, **4**, 219-31.
- D. Uschner, R.-D. Hilgers, N. Heussen (2018) The impact of selection bias in randomized multi-arm parallel group clinical trials *PLOS ONE*, **13**(1), 1-18.

See Also

Compute exact or simulated rejection probability: [assess](#).

Other issues: [chronBias](#), [combineBias\(\)](#), [corGuess](#), [imbal](#), [issue](#), [setPower\(\)](#)

Examples

```
# create a selection bias of the convergency strategy type with eta = 0.25 for which
# the exact rejection probabilities are calculated
sbias <- selBias("CS", 0.25, "exact")
```

setPower	<i>Representing the power</i>
----------	-------------------------------

Description

Represents the expected power of the individual randomization sequences.

Usage

```
setPower(d, method, alpha = 0.05)
```

Arguments

d	effect size.
method	character string, should be one of "sim" or "exact", see Description.
alpha	significance level.

Details

The attained power of an individual randomization sequence can be an issue in the design of a clinical trial. The power of a randomization sequence is computed depending on the effect size d and the difference in group sizes.

If `method="sim"`, the object represents the simulated power of an individual randomization sequence. When calling `assess` for a power object with `method="sim"`, one test decision is computed for each randomization sequence of `randSeq`. The power is the proportion of falsely (correctly) rejected null hypotheses.

If `method="exact"`, the object represents the exact power of an individual randomization sequence. When calling `assess` for a power object with `method="exact"`, the exact p -value of each randomization sequence is computed. So far, this is only supported for normal endpoints. Then the power is the sum of the corresponding quantiles of the noncentral t -distribution.

Value

S4 object of class `power`, a formal representation of the issue of power in a clinical trial.

See Also

Other issues: [chronBias](#), [combineBias\(\)](#), [corGuess](#), [imbal](#), [issue](#), [selBias](#)

shape	<i>Method returning the shape parameter of an survEndp S4 object</i>
-------	--

Description

Method returning the shape parameter of an survEndp S4 object

Usage

shape(obj)

Arguments

obj	object of class survEndp
-----	--------------------------

sigma	<i>Function returning the standard deviation slot of a normEndp S4 object</i>
-------	---

Description

Function returning the standard deviation slot of a normEndp S4 object

Usage

sigma(obj)

Arguments

obj	object of class normEndp
-----	--------------------------

summary	<i>Summary of assessments of a randomization procedure</i>
---------	--

Description

Summary of assessments of a randomization procedure

Summary of desirability scores of a randomization procedure

Usage

```
summary(object, ...)

## S4 method for signature 'assessment'
summary(object)

summary(object, ...)

## S4 method for signature 'desScores'
summary(object)
```

Arguments

```
object      assessment object.
...         additional arguments affecting the summary that will be produced.
```

Details

For each issue the assessment of the sequences is summarized to permit a design-based assessment of the randomization procedure. This approach uses the sequence-wise values of the assessment and the probabilities in order to give an overall summary.

For each issue the desirability score of the sequences is summarized to permit a design-based desirability score of the randomization procedure. This approach uses the sequence-wise values of the desirability and the probabilities in order to give an overall summary.

Value

Data frame with a summary of the assessment object.
Data frame with a summary of the desirability scores object.

Examples

```
# assess the full set of PBR(4)
seq <- getAllSeq(pbrPar(4))
issue <- corGuess("CS")
A <- assess(seq, issue)
summary(A)

# compute the desirability scores of the full set of PBR(4)
seq <- getAllSeq(pbrPar(4))
issue1 <- corGuess("CS")
issue2 <- corGuess("DS")
A <- assess(seq, issue1, issue2)
d1 <- derFunc(0.5, c(0.1, 0.8), c(1, 1))
d2 <- derFunc(0.1, 0.7, 2)
D <- getDesScores(A, d1, d2, weights = c(5/6, 1/6))
summary(D)
```

survEndp	<i>Representation of survival endpoints</i>
----------	---

Description

Represents survival endpoints in clinical trials.

Usage

```
survEndp(  
  cenRate,  
  accrualTime = 0,  
  cenTime,  
  shape,  
  scale,  
  weights = c(0, 0),  
  maxcombo = FALSE  
)
```

Arguments

cenRate	exponential censoring rate in a survival study.
accrualTime	duration of the accrual period in a survival study.
cenTime	total duration of a survival study (maximum length of followup).
shape	parameter of the Weibull distribution (must be positive)
scale	parameter of the Weibull distribution (must be positive)
weights	numeric specifying the weights used for the test. Unless specified an unweighted test is conducted.
maxcombo	logical specifying if the maxcombo test is used

Details

The `survEnd` function is a constructor function for an S4 object of the class `survEnd` representing a survival endpoint in a clinical trial.

Value

A S4 object representing a survival endpoint in a clinical trial.

See Also

Compute exact or simulated type-I-error: [assess](#).

Other endpoint types: [expEndp](#), [normEndp](#)

tbdPar	<i>Representing Truncated Binomial Design</i>
--------	---

Description

Represents the Truncated Binomial Design.

Usage

```
tbdPar(bc = N, groups = LETTERS[1:2])
```

Arguments

bc	vector which contains the lengths k_1, \dots, k_l of each block. This means that the vector bc will have one entry for each block.
groups	character vector of labels for the different treatments.

Details

A fair coin is tossed until half of the patients have been assigned to one of the treatment arms. Afterwards, the randomization list is filled with the other treatment.

Value

S4 object of the class tbdPar.

References

W. F. Rosenberger and J. M. Lachin (2002) *Randomization in Clinical Trials*. Wiley.

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [bsdPar](#), [chenPar](#), [crPar](#), [createParam\(\)](#), [ebcPar](#), [gbcdPar](#), [hadaPar](#), [mpPar](#), [pbrPar](#), [rarPar](#), [rpbrPar](#), [rtbdPar](#), [udPar](#)

TV	<i>Function returning the target value slot of an S4 object</i>
----	---

Description

Function returning the target value slot of an S4 object

Usage

```
TV(obj)
```

Arguments

obj object inheriting from derFunc

type *Get type of an object*

Description

Accesses the type slot of an S4 object

Usage

type(obj)

Arguments

obj a bias object (i.e. S4 object inheriting from bias)

Value

Character string specifying the type of bias obj represents, e.g. "linT" in case of chronological bias.

udPar *Representing Wei's Urn Design*

Description

Represents Wei's Urn Design.

Usage

udPar(N, ini, add, groups = LETTERS[1:2])

Arguments

N integer for the total sample size of the trial.
 ini integer representing the initial urn composition.
 add integer representing the number of balls that are added to the urn in each step.
 groups character vector of labels for the different treatments.

Details

An urn is filled with a number of ini balls of both of the treatments. Afterwards, a ball is drawn randomly from the urn. Finally, add balls are added to the urn from the opposite treatment. This procedure is repeated until N patients are assigned.

Value

S4 object of the class udPar.

References

L.J. Wei (1977) A Class of Designs for Sequential Clinical Trials. *Journal of the American Statistical Association*, **72**, 382-6.

See Also

Other randomization procedures: [abcdPar](#), [bbcdPar](#), [bsdPar](#), [chenPar](#), [crPar](#), [createParam\(\)](#), [ebcPar](#), [gbcdPar](#), [hadaPar](#), [mpPar](#), [pbrPar](#), [rarPar](#), [rpbrPar](#), [rtbdPar](#), [tbdPar](#)

Index

* desirability topics

derFunc, 19
evaluate, 22
getDesScores, 37
plotDes, 56
plotEv, 57
probUnDes, 58

* endpoint types

expEndp, 24
normEndp, 51
survEndp, 72

* issues

chronBias, 12
combineBias, 14
corGuess, 16
imbal, 46
issue, 47
selBias, 67
setPower, 69

* randomization procedures

abcdPar, 5
bbcdPar, 8
bsdPar, 10
chenPar, 11
createParam, 17
crPar, 18
ebcPar, 21
gbcdPar, 25
hadaPar, 45
mpPar, 49
pbrPar, 54
rarPar, 61
rpbrPar, 63
rtbdPar, 64
tbdPar, 73
udPar, 74

* saving functions

saveAssess, 65
saveRand, 66

a, 5

abcdPar, 5, 9–11, 18, 19, 22, 25, 46, 50, 55,
60, 62, 64, 65, 73, 75

analyse, 6

assess, 4, 7, 24, 37, 48, 52, 65, 67, 68, 72

assess, randSeq, endpoint-method
(assess), 7

assess, randSeq, missing-method (assess),
7

bbcdPar, 6, 8, 10, 11, 18, 19, 22, 25, 46, 50,
55, 60, 62, 64, 65, 73, 75

blocks, 9

bsdPar, 6, 9, 10, 11, 18, 19, 22, 25, 46, 50, 55,
60, 62, 64, 65, 73, 75

calcProb (getProbabilities), 41

calculateProbabilities
(getProbabilities), 41

chenPar, 6, 9, 10, 11, 18, 19, 22, 25, 46, 50,
55, 60, 62, 64, 65, 73, 75

chronBias, 12, 14, 17, 40, 47, 48, 68, 69

coin, 13

combineBias, 13, 14, 17, 47, 48, 68, 69

compare, 4, 15, 48, 56

compare, issue, endpoint-method
(compare), 15

compare, issue, missing-method (compare),
15

corGuess, 13, 14, 16, 47, 48, 68, 69

createParam, 6, 9–11, 17, 19, 22, 25, 27, 32,
43, 46, 50, 53, 55, 60, 62, 64, 65, 67,
73, 75

createSeq, 18

crPar, 6, 9–11, 18, 18, 22, 25, 46, 50, 55, 60,
62, 64, 65, 73, 75

derFunc, 19, 20, 23, 37, 56, 57, 59

desirability, 20

- ebcPar, [6](#), [9–11](#), [18](#), [19](#), [21](#), [25](#), [46](#), [50](#), [55](#), [60](#), [62](#), [64](#), [65](#), [73](#), [75](#)
- evaluate, [20](#), [22](#), [37](#), [56](#), [57](#), [59](#)
- evaluate, character-method (evaluate), [22](#)
- evaluate, missing-method (evaluate), [22](#)
- expEndp, [24](#), [52](#), [72](#)

- gbcdPar, [6](#), [9–11](#), [18](#), [19](#), [22](#), [25](#), [46](#), [50](#), [55](#), [60](#), [62](#), [64](#), [65](#), [73](#), [75](#)
- generateAllSequences, [26](#)
- generateRandomSequences, [28](#)
- genNcps_new, [33](#)
- genSeq, [4](#), [7](#), [15](#), [23](#), [37](#), [48](#), [59](#), [61](#)
- genSeq (generateRandomSequences), [28](#)
- genSeq, abcdPar, missing, missing-method (generateRandomSequences), [28](#)
- genSeq, abcdPar, missing, numeric-method (generateRandomSequences), [28](#)
- genSeq, abcdPar, numeric, missing-method (generateRandomSequences), [28](#)
- genSeq, abcdPar, numeric, numeric-method (generateRandomSequences), [28](#)
- genSeq, bbcdPar, missing, missing-method (generateRandomSequences), [28](#)
- genSeq, bbcdPar, missing, numeric-method (generateRandomSequences), [28](#)
- genSeq, bbcdPar, numeric, missing-method (generateRandomSequences), [28](#)
- genSeq, bbcdPar, numeric, numeric-method (generateRandomSequences), [28](#)
- genSeq, bsdPar, missing, missing-method (generateRandomSequences), [28](#)
- genSeq, bsdPar, missing, numeric-method (generateRandomSequences), [28](#)
- genSeq, bsdPar, numeric, missing-method (generateRandomSequences), [28](#)
- genSeq, bsdPar, numeric, numeric-method (generateRandomSequences), [28](#)
- genSeq, chenPar, missing, missing-method (generateRandomSequences), [28](#)
- genSeq, chenPar, missing, numeric-method (generateRandomSequences), [28](#)
- genSeq, chenPar, numeric, missing-method (generateRandomSequences), [28](#)
- genSeq, chenPar, numeric, numeric-method (generateRandomSequences), [28](#)
- genSeq, crPar, missing, missing-method (generateRandomSequences), [28](#)
- genSeq, crPar, missing, numeric-method (generateRandomSequences), [28](#)
- genSeq, crPar, numeric, missing-method (generateRandomSequences), [28](#)
- genSeq, crPar, numeric, numeric-method (generateRandomSequences), [28](#)
- genSeq, ebcPar, missing, missing-method (generateRandomSequences), [28](#)
- genSeq, ebcPar, missing, numeric-method (generateRandomSequences), [28](#)
- genSeq, ebcPar, numeric, missing-method (generateRandomSequences), [28](#)
- genSeq, ebcPar, numeric, numeric-method (generateRandomSequences), [28](#)
- genSeq, gbcdPar, missing, missing-method (generateRandomSequences), [28](#)
- genSeq, gbcdPar, missing, numeric-method (generateRandomSequences), [28](#)
- genSeq, gbcdPar, numeric, missing-method (generateRandomSequences), [28](#)
- genSeq, gbcdPar, numeric, numeric-method (generateRandomSequences), [28](#)
- genSeq, hadaPar, missing, missing-method (generateRandomSequences), [28](#)
- genSeq, hadaPar, missing, numeric-method (generateRandomSequences), [28](#)
- genSeq, hadaPar, numeric, missing-method (generateRandomSequences), [28](#)
- genSeq, hadaPar, numeric, numeric-method (generateRandomSequences), [28](#)
- genSeq, mpPar, missing, missing-method (generateRandomSequences), [28](#)
- genSeq, mpPar, missing, numeric-method (generateRandomSequences), [28](#)
- genSeq, mpPar, numeric, missing-method (generateRandomSequences), [28](#)
- genSeq, mpPar, numeric, numeric-method (generateRandomSequences), [28](#)
- genSeq, pbrPar, missing, missing-method (generateRandomSequences), [28](#)
- genSeq, pbrPar, missing, numeric-method (generateRandomSequences), [28](#)
- genSeq, pbrPar, numeric, missing-method (generateRandomSequences), [28](#)
- genSeq, pbrPar, numeric, numeric-method (generateRandomSequences), [28](#)
- genSeq, rarPar, missing, missing-method (generateRandomSequences), [28](#)

- genSeq, rarPar, missing, numeric-method
(generateRandomSequences), 28
- genSeq, rarPar, numeric, missing-method
(generateRandomSequences), 28
- genSeq, rarPar, numeric, numeric-method
(generateRandomSequences), 28
- genSeq, rpbrPar, missing, missing-method
(generateRandomSequences), 28
- genSeq, rpbrPar, missing, numeric-method
(generateRandomSequences), 28
- genSeq, rpbrPar, numeric, missing-method
(generateRandomSequences), 28
- genSeq, rpbrPar, numeric, numeric-method
(generateRandomSequences), 28
- genSeq, rtbdPar, missing, missing-method
(generateRandomSequences), 28
- genSeq, rtbdPar, missing, numeric-method
(generateRandomSequences), 28
- genSeq, rtbdPar, numeric, missing-method
(generateRandomSequences), 28
- genSeq, rtbdPar, numeric, numeric-method
(generateRandomSequences), 28
- genSeq, tbdPar, missing, missing-method
(generateRandomSequences), 28
- genSeq, tbdPar, missing, numeric-method
(generateRandomSequences), 28
- genSeq, tbdPar, numeric, missing-method
(generateRandomSequences), 28
- genSeq, tbdPar, numeric, numeric-method
(generateRandomSequences), 28
- genSeq, udPar, missing, missing-method
(generateRandomSequences), 28
- genSeq, udPar, missing, numeric-method
(generateRandomSequences), 28
- genSeq, udPar, numeric, missing-method
(generateRandomSequences), 28
- genSeq, udPar, numeric, numeric-method
(generateRandomSequences), 28
- get_p_values_new, 43
- getAllSeq, 7, 61
- getAllSeq (generateAllSequences), 26
- getAllSeq, abcdPar-method
(generateAllSequences), 26
- getAllSeq, bbcdPar-method
(generateAllSequences), 26
- getAllSeq, bsdPar-method
(generateAllSequences), 26
- getAllSeq, chenPar-method
(generateAllSequences), 26
- getAllSeq, crPar-method
(generateAllSequences), 26
- getAllSeq, ebcPar-method
(generateAllSequences), 26
- getAllSeq, gbcdPar-method
(generateAllSequences), 26
- getAllSeq, hadaPar-method
(generateAllSequences), 26
- getAllSeq, mpPar-method
(generateAllSequences), 26
- getAllSeq, pbrPar-method
(generateAllSequences), 26
- getAllSeq, rarPar-method
(generateAllSequences), 26
- getAllSeq, tbdPar-method
(generateAllSequences), 26
- getAllSeq, udPar-method
(generateAllSequences), 26
- getCorGuesses, 33
- getDesFunc, 34
- getDesFunc, derringerLs-method
(getDesFunc), 34
- getDesFunc, derringerRs-method
(getDesFunc), 34
- getDesFunc, derringerTs-method
(getDesFunc), 34
- getDesign, 34
- getDesign, abcdPar-method (getDesign), 34
- getDesign, abcdSeq-method (getDesign), 34
- getDesign, bbcdPar-method (getDesign), 34
- getDesign, bbcdSeq-method (getDesign), 34
- getDesign, bsdPar-method (getDesign), 34
- getDesign, bsdSeq-method (getDesign), 34
- getDesign, chenPar-method (getDesign), 34
- getDesign, chenSeq-method (getDesign), 34
- getDesign, crPar-method (getDesign), 34
- getDesign, crSeq-method (getDesign), 34
- getDesign, ebcPar-method (getDesign), 34
- getDesign, ebcSeq-method (getDesign), 34
- getDesign, gbcdPar-method (getDesign), 34
- getDesign, gbcdSeq-method (getDesign), 34
- getDesign, hadaPar-method (getDesign), 34
- getDesign, hadaSeq-method (getDesign), 34
- getDesign, mpPar-method (getDesign), 34
- getDesign, mpSeq-method (getDesign), 34
- getDesign, pbrPar-method (getDesign), 34
- getDesign, pbrSeq-method (getDesign), 34

- GSD_POC, 44
- hadaPar, 6, 9–11, 18, 19, 22, 25, 45, 50, 55, 60, 62, 64, 65, 73, 75
- imbal, 13, 14, 17, 46, 47, 48, 68, 69
- issue, 13, 14, 17, 47, 47, 68, 69
- issues, 4, 7, 15, 23, 37, 59
- issues (issue), 47
- K, 48
- lambda, 48
- method, 49
- mpPar, 6, 9–11, 18, 19, 22, 25, 46, 49, 55, 60, 62, 64, 65, 73, 75
- mti, 50
- mu, 50
- N, 51
- normEndp, 24, 40, 51, 72
- overview, 52
- pbrPar, 6, 9–11, 18, 19, 22, 25, 46, 50, 54, 60, 62, 64, 65, 73, 75
- plot, 55
- plot, comparison, character-method (plot), 55
- plot, comparison, missing-method (plot), 55
- plotDes, 20, 23, 37, 56, 57, 59
- plotEv, 20, 23, 37, 56, 57, 59
- plotSeq, 58
- probUnDes, 20, 23, 37, 56, 57, 58
- probUnDes, desScores-method (probUnDes), 58
- randBlocks, 60
- randomizeR (randomizeR-package), 4
- randomizeR-package, 4
- randPar, 4, 7, 15, 17, 23, 27, 32, 37, 43, 48, 53, 59, 60, 67
- randSeq, 27, 32
- randSeq-class, 61
- rarPar, 6, 9–11, 18, 19, 22, 25, 46, 50, 55, 60, 61, 64, 65, 73, 75
- ratio, 62
- rho, 63
- rpbrPar, 6, 9–11, 18, 19, 22, 25, 46, 50, 55, 60, 62, 63, 65, 73, 75
- rtbdPar, 6, 9–11, 18, 19, 22, 25, 46, 50, 55, 60, 62, 64, 64, 73, 75
- saveAssess, 65, 66
- saveRand, 65, 66
- scale, 66
- seed, 67
- selBias, 13, 14, 17, 40, 47, 48, 67, 69
- setPower, 13, 14, 17, 47, 48, 68, 69
- shape, 70
- sigma, 70
- summary, 70
- summary, assessment-method (summary), 70
- summary, desScores-method (summary), 70
- survEndp, 24, 52, 72
- tbdPar, 6, 9–11, 18, 19, 22, 25, 46, 50, 55, 60, 62, 64, 65, 73, 75
- TV, 73
- type, 74
- udPar, 6, 9–11, 18, 19, 22, 25, 46, 50, 55, 60, 62, 64, 65, 73, 74