

# Package: grouprar (via r-universe)

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**Type** Package

**Title** Group Response Adaptive Randomization for Clinical Trials

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**Description** Implement group response-adaptive randomization procedures, which also integrates standard non-group response-adaptive randomization methods as specialized instances. It is also uniquely capable of managing complex scenarios, including those with delayed and missing responses, thereby expanding its utility in real-world applications. This package offers 16 functions for simulating a variety of response adaptive randomization procedures. These functions are essential for guiding the selection of statistical methods in clinical trials, providing a flexible and effective approach to trial design. Some of the detailed methodologies and algorithms used in this package, please refer to the following references:  
LJ Wei (1979) <[doi:10.1214/aos/1176344614](https://doi.org/10.1214/aos/1176344614)> L. J. WEI and S. DURHAM (1978) <[doi:10.1080/01621459.1978.10480109](https://doi.org/10.1080/01621459.1978.10480109)> Durham, S. D., FlournoY, N. AND LI, W. (1998) <[doi:10.2307/3315771](https://doi.org/10.2307/3315771)> Ivanova, A., Rosenberger, W. F., Durham, S. D. and Flournoy, N. (2000) <<https://www.jstor.org/stable/25053121>> Bai Z D, Hu F, Shen L. (2002) <[doi:10.1006/jmva.2001.1987](https://doi.org/10.1006/jmva.2001.1987)> Ivanova, A. (2003) <[doi:10.1007/s001840200220](https://doi.org/10.1007/s001840200220)> Hu, F., & Zhang, L. X. (2004) <[doi:10.1214/aos/1079120137](https://doi.org/10.1214/aos/1079120137)> Hu, F., & Rosenberger, W. F. (2006, ISBN:978-0-471-65396-7). Zhang, L. X., Chan, W. S., Cheung, S. H., & Hu, F. (2007) <<https://www.jstor.org/stable/26432528>> Zhang, L., & Rosenberger, W. F. (2006) <[doi:10.1111/j.1541-0420.2005.00496.x](https://doi.org/10.1111/j.1541-0420.2005.00496.x)> Hu, F., Zhang, L. X., Cheung, S. H., & Chan, W. S. (2008) <[doi:10.1002/cjs.5550360404](https://doi.org/10.1002/cjs.5550360404)>.

**License** GPL (>= 2)

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Bai Hu Shen's Urn      *Bai Hu Shen's Urn*

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## Description

Bai, Hu, and Shen *bai2002adaptive* proposed a new adaptive design for multi-arm clinical trials. The main idea behind this procedure is that the allocation probability adapts based on the performance of the most recent patients under their assigned treatment. Positive performance in that treatment increase the likelihood of the next patient being assigned to this group, whereas negative outcomes decrease it. This function is for simulating the Bai, Hu, and Shen's urn model under two-sided hypothesis testing in clinical trial context.

**Usage**

```
Bai.Hu.Shen.Urn(k, p, ssn, Y0 = NULL, nsim = 2000, alpha = 0.05)
```

**Arguments**

k	a positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k \geq 2$ )
p	a positive vector of length equals to k. The values specify the true success rates for the various treatments, and these rates are used to generate data for simulations.
ssn	a positive integer. The value specifies the total number of participants involved in each round of the simulation.
Y0	A vector of length k, specifying the initial probability of allocating a patient to each group. For instance, if $Y0 = c(1, 1, 1)$ , the initial probabilities are calculated as $Y0 / \text{sum}(Y0)$ . When Y0 is NULL, the initial urn will be set as If Y0 is NULL, then Y0 is set to a vector of length k, with all values equal to 1 by default.
nsim	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
alpha	A number between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

**Details**

Bai, Hu and Shen's urn can be describe as follows: An urn contains  $K$  types of balls initially. Balls of types  $1, 2, \dots, K$  represent treatments  $1, 2, \dots, K$ . A type  $k$  ball is drawn randomly from the urn, and then we assign the patient, who is waiting to be assigned, to the treatment  $k$ . After obtaining the response, we may adapt the composition of the urn. A success on treatment  $k$  adds a ball of type of  $k$  to the urn and a failure on treatment  $k$  adds  $\frac{p_i}{(M-p_k)}$  ball for each of the other  $K - 1$  types, where  $M = p_1 + \dots + p_K$ .

**Value**

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation propotion for each of treatment groups.
failRate	The propotion of individuals who do not achieve the expected outcome in each simulation, on average.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

**References**

Bai Z D, Hu F, Shen L. *An adaptive design for multi-arm clinical trials*[J]. Journal of Multivariate Analysis, 2002, 81(1): 1-18.

**Examples**

```
## a simple use
bhs.res = Bai.Hu.Shen.Urn(k = 3,
                          p = c(0.7, 0.8, 0.6),
                          ssn = 500,
                          Y0 = NULL,
                          nsim = 200,
                          alpha = 0.05)

## view the output
bhs.res

## view all simulation settings
bhs.res$name
bhs.res$parameter
bhs.res$k

## View the simulations results
bhs.res$propotion
bhs.res$failRate
bhs.res$pwCalc
bhs.res$assignment
```

---

 BirthDeathUrn

*Birth and Death Urn*


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**Description**

Simulating birth and death urn procedure (number of arms  $\geq 2$ ) with two-sided hypothesis testing in a clinical trial context.

**Usage**

```
BirthDeathUrn(k, p, ssn, Y0 = NULL, nsim = 2000, alpha = 0.05)
```

**Arguments**

k	a positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k \geq 2$ )
p	a positive vector of length equals to k. The values specify the true success rates for the various treatments, and these rates are used to generate data for simulations.
ssn	a positive integer. The value specifies the total number of participants involved in each round of the simulation.
Y0	A vector of length k, specifying the initial probability of allocating a patient to each group. For instance, if $Y0 = c(1, 1, 1)$ , the initial probabilities are calculated as $Y0 / \text{sum}(Y0)$ . When Y0 is NULL, the initial urn will be set as If Y0 is NULL, then Y0 is set to a vector of length k, with all values equal to 1 by default.

nsim	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
alpha	A number between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

### Details

The birth and death urn works as follows: Initially an urn contains balls of  $K$  types and  $a$  immigration balls. A ball is drawn randomly with replacement. If it is an immigration ball, one ball of each type is added to the urn, no patient is treated, and the next ball is drawn. The procedure is repeated until a type  $i$  ball ( $i = 1, \dots, K$ ) is drawn. Then the subject is assigned to treatment  $i$ . If a success, a type  $i$  ball is added in the urn; if a failure, a type  $i$  ball is removed. (Hu and Rosenberger (2006)). More detail could be found in paper *A birth and death urn for randomized clinical trials* written by Ivanova etl (2000).

### Value

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation propotion for each of treatment groups.
failRate	The proportion of individuals who do not achieve the expected outcome in each simulation, on average.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

### References

- Hu, F., & Rosenberger, W. F. (2006). *The theory of response-adaptive randomization in clinical trials*. John Wiley & Sons.
- Ivanova, A., Rosenberger, W. F., Durham, S. D. and Flournoy, N. (2000). *A birth and death urn for randomized clinical trials*. Sankhya B 62 104-118.

### Examples

```
## a simple use
bd.res = BirthDeathUrn(k = 3, p = c(0.6, 0.7, 0.6), ssn = 400, Y0 = NULL, nsim = 200, alpha = 0.05)

## view the output
bd.res

## view all simulation settings
bd.res$name
bd.res$parameter
bd.res$k
```

```
## View the simulations results
bd.res$propotion
bd.res$failRate
bd.res$pwCalc
bd.res$assignment
```

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CRDesign

*Complete Randomization*


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### Description

Simulating complete randomization with two-sided hypothesis testing in a clinical trial context.

### Usage

```
CRDesign(k, p, ssn, nsim = 2000, alpha = 0.05)
```

### Arguments

k	a positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k \geq 2$ )
p	a positive vector of length equals to k. The values specify the true success rates for the various treatments, and these rates are used to generate data for simulations.
ssn	a positive integer. The value specifies the total number of participants involved in each round of the simulation.
nsim	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
alpha	An integer between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

### Details

Complete randomization: Allocating participants or subjects to different treatment groups in a clinical trial in such a way that each participant has an equal and independent chance of being assigned to any of the treatment groups.

### Value

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation propotion for each of treatment groups.

failRate	The proportion of individuals who do not achieve the expected outcome in each simulation, on average.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

### Examples

```
## a simple use
CR.res = CRDesign(k=3, p = c(0.7, 0.8, 0.6), nsim = 500, ssn = 400)
## view the output
CR.res

## view all simulation settings
CR.res$name
CR.res$parameter
CR.res$k
## View the simulations results
CR.res$propotion
CR.res$failRate
CR.res$pwCalc
CR.res$assignment
```

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DBCD_Bin	<i>Hu and Zhang's Doubly Biased Coin Design with Binary Response Type</i>
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### Description

Simulating Hu and Zhang's doubly biased coin deisgn with binary response (number of arms  $\geq 2$ ) in a clinical trial context. (Inference: two-sided hypothesis testing t-test or chi-square test)

### Usage

```
DBCD_Bin(n0 = 20, p, k, ssn, theta0 = NULL, target.alloc = "RPW",
         r = 2, nsim = 2000, mRate = NULL, alpha = 0.05)
```

### Arguments

n0	A positive integer. n0 represents the initial patient population assigned through restricted randomization for initial parameter estimation.
p	A positive vector of length equals to k. The values specify the true success rates for the various treatments, and these rates are used to generate data for simulations.
k	A positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k \geq 2$ )

ssn	A positive integer. The value specifies the total number of participants involved in each round of the simulation.
theta0	A vector of length k. Each value in the vector represents a probability used for adjusting parameter estimates. If the argument is not provided, it defaults to a vector of length k, with all values set to 0.5.
target.alloc	Desired allocation proportion. The option for this argument could be one of "Neyman", "RSIHR", "RPW", "WeisUrn". The default is "RPW".
r	A positive number. Parameter for Hu and Zhang's doubly biased coin design and usually take values 2-4. The default value is 2.
nsim	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
mRate	a numerical value between 0 and 1, inclusive, representing the missing rate for the responses. This parameter pertains to missing-at-random data. The default value is NULL, indicating no missing values by default.
alpha	A number between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

### Details

The objective of Hu and Zhang's doubly biased coin design is to allocate patients sequentially while closely approximating the desired allocation proportion, which is a function of certain unknown parameters related to the response variable under each treatment.

The process begins by assigning  $n_0$  patients to treatment groups using restricted randomization and collecting their responses. Initial parameter estimates for the response variable are then obtained for each treatment group. Subsequently, based on these parameter estimates, the desired allocation proportion is calculated. Afterward, the Hu and Zhang's allocation function is applied to determine the probabilities for the next patient to be assigned to each treatment group, which force the allocation proportion close to the desired one. This process is repeated sequentially for each patient until the desired number of patients has been allocated, as predetermined.

This methodology was introduced by Hu and Zhang in their 2004 paper titled 'Asymptotic Properties of Doubly Adaptive Biased Coin Designs for Multitreatment Clinical Trials.

### Value

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation propotion for each of treatment groups.
failRate	The proportion of individuals who do not achieve the expected outcome in each simulation, on average.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.



## References

Hu, F., & Zhang, L. X. (2004). *Asymptotic properties of doubly adaptive biased coin designs for multi-treatment clinical trials*. The Annals of Statistics, 32(1), 268-301.

## Examples

```
DBCD_Bin(n0 = 20, p = c(0.7, 0.8), k = 2, ssn = 300, theta0 = NULL,
        target.alloc = "RPW", r = 2, nsim = 50, mRate = NULL, alpha = 0.05)
```

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DBCD_Cont	<i>Hu and Zhang's Doubly Biased Coin Design with Continuous Response Type</i>
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## Description

Simulating Hu and Zhang's doubly biased coin design with continuous response (number of arms  $\geq 2$ ) in a clinical trial context. (Inference: two-sided hypothesis testing t-test or chi-square test)

## Usage

```
DBCD_Cont(n0 = 20, theta, k, ssn, theta0 = NULL, target.alloc = "Neyman",
        r = 2, nsim = 2000, alpha = 0.05)
```

## Arguments

n0	A positive integer. n0 represents the initial patient population assigned through restricted randomization for initial parameter estimation.
theta	A numerical vector of length equal to 2k. These values specify the true parameters for each treatment and are used for generating data in simulations. For example, if k=2, you should provide two pairs of parameter values, each consisting of the mean and variance, like: theta = c(13, 4.0^2, 15, 2.5^2).
k	A positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k \geq 2$ )
ssn	A positive integer. The value specifies the total number of participants involved in each round of the simulation.
theta0	A vector of length 2k. Each value in the vector represents a probability used for adjusting parameter estimates. If the argument is not provided, it defaults to a vector of length 2k, with all parameter pair set to be (0, 1).
target.alloc	Desired allocation proportion. The option for this argument could be one of "Neyman", "ZR", "DaOptimal". The default is "Neyman". The details see Zhang L. and Rosenberger. W (2006).
r	A positive number. Parameter for Hu and Zhang's doubly biased coin design and usually take values 2-4. The default value is 2.
nsim	a positive integer. The value specifies the total number of simulations, with a default value of 2000.

alpha	A number between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.
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### Details

The objective of Hu and Zhang's doubly biased coin design is to allocate patients sequentially while closely approximating the desired allocation proportion, which is a function of certain unknown parameters related to the response variable under each treatment.

The process begins by assigning  $n_0$  patients to treatment groups using restricted randomization and collecting their responses. Initial parameter estimates for the response variable are then obtained for each treatment group. Subsequently, based on these parameter estimates, the desired allocation proportion is calculated. Afterward, the Hu and Zhang's allocation function is applied to determine the probabilities for the next patient to be assigned to each treatment group, which force the allocation proportion close to the desired one. This process is repeated sequentially for each patient until the desired number of patients has been allocated, as predetermined.

This methodology was introduced by Hu and Zhang in their 2004 paper titled 'Asymptotic Properties of Doubly Adaptive Biased Coin Designs for Multitreatment Clinical Trials.

### Value

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation porpotion for each of treatment groups.
failRate	The average response value for the entire trial.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

### References

- Hu, F., Zhang, L. X., Cheung, S. H., & Chan, W. S. (2008). *Doubly adaptive biased coin designs with delayed responses*. Canadian Journal of Statistics, 36(4), 541-559.
- Zhang, L., & Rosenberger, W. F. (2006). *Response-adaptive randomization for clinical trials with continuous outcomes*. Biometrics, 62(2), 562-569.

### See Also

- See [DBCD\\_Bin](#) for simulations of Hu and Zhang's doubly biased coin deisgn with binary response.
- See [dylDBCD\\_Cont](#) for simulations of Hu and Zhang's doubly biased coin deisgn with delayed continuous response.

**Examples**

```

# A simple use!
# Define the arguments
## Arguments for generate the simulated data
theta = c(13, 4.0^2, 15, 2.5^2)
k = 2
ssn = 88
### Other arguments
target.alloc = "Neyman"

res = DBCD_Cont(n0 = 20, theta, k, ssn, theta0 = NULL,
               target.alloc = "Neyman", r = 2, nsim = 200, alpha = 0.05)

# View the output (A list of all results)
res

```

DLRule

*Drop the loser rule***Description**

Simulating drop the loser rule procedure with two-sided hypothesis testing in a clinical trial context.

**Usage**

```
DLRule(k, p, ssn, Y0 = NULL, nsim = 2000, alpha = 0.05)
```

**Arguments**

k	a positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k = 2$ )
p	a positive vector of length equals to k. The values specify the true success rates for the various treatments, and these rates are used to generate data for simulations.
ssn	a positive integer. The value specifies the total number of participants involved in each round of the simulation.
Y0	A vector of length k, specifying the initial probability of allocating a patient to each group. For instance, if $Y0 = c(1, 1, 1)$ , the initial probabilities are calculated as $Y0 / \text{sum}(Y0)$ . When Y0 is NULL, the initial urn will be set as If Y0 is NULL, then Y0 is set to a vector of length k, with all values equal to 1 by default.
nsim	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
alpha	A number between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

## Details

Drop the loser rule can be describe as follows: An urn contains three types of balls ( $A, B, 0$ ) initially. Balls of types  $A$  and  $B$  represent treatments  $A$  and  $B$ , balls of  $0$  type are immigration balls. If  $A$  (or  $B$ ) is drawn, then treatment  $A$  (or  $B$ ) is assigned to the subject and the response is observed. If the observed response is a failure, then the ball is not replaced, else replaced. If an immigration ball (type  $0$ ) is drawn, no treatment is assigned, and the ball is returned to the urn together with one  $A$  and one  $B$  ball.

## Value

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation propotion for each of treatment groups.
failRate	The proportion of individuals who do not achieve the expected outcome in each simulation, on average.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

## References

Ivanova, A. (2003). *A play-the-winner-type urn design with reduced variability*. *Metrika*, 58, 1-13.

## Examples

```
## a simple use
dl.res = DLRule(k = 2, p = c(0.7, 0.8), ssn = 400, Y0 = NULL, nsim = 200, alpha = 0.05)

## view the output
dl.res

## view all simulation settings
dl.res$name
dl.res$parameter
dl.res$k

## View the simulations results
dl.res$propotion
dl.res$failRate
dl.res$pwCalc
dl.res$assignment
```

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dyldBCD_Bin	<i>Hu and Zhang's Doubly Biased Coin Design with delayed Binary Response</i>
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### Description

Simulating Hu and Zhang's doubly biased coin design with delayed binary response (number of arms  $\geq 2$ ) in a clinical trial context. (Inference: two-sided hypothesis testing t-test or chi-square test)

### Usage

```
dyldBCD_Bin(n0 = 20, p, k, ssn, ent.param, rspT.dist,
            rspT.param, theta0 = NULL, target.alloc = "RPW", r = 2,
            nsim = 2000, mRate = NULL, alpha = 0.05)
```

### Arguments

n0	A positive integer. n0 represents the initial patient population assigned through restricted randomization for initial parameter estimation.
p	A positive vector of length equals to k. The values specify the true success rates for the various treatments, and these rates are used to generate data for simulations.
k	A positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k = 2$ )
ssn	A positive integer. The value specifies the total number of participants involved in each round of the simulation.
ent.param	A positive integer. The value specified the parameter for an exponential distribution which determine the time for each participant enter the trial.
rspT.dist	Distribution Type. Specifies the type of distribution that models the time spent for the availability of patient $i$ under treatment $k$ . Acceptable options for this argument include: "exponential", "normal", and "uniform".
rspT.param	A vector. Specifies the parameters required by the distribution that models the time spent for the availability under each treatment. (eg. If there are 3 treatments groups and each of them follows truncated normal distribution with parameter pair (3, 2), (2, 1), (4, 1), respectively. Then the <code>rspT.param = c(3, 2, 2, 1, 4, 1)</code> )
theta0	A vector of length k. Each value in the vector represents a probability used for adjusting parameter estimates. If the argument is not provided, it defaults to a vector of length k, with all values set to 0.5.
target.alloc	Desired allocation proportion. The option for this argument could be one of "Neyman", "RSIHR", "RPW", "WeisUrn". The default is "RPW".
r	A positive number. Parameter for Hu and Zhang's doubly biased coin design and usually take values 2-4. The default value is 2.

nsim	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
mRate	a numerical value between 0 and 1, inclusive, representing the missing rate for the responses. This parameter pertains to missing-at-random data. The default value is NULL, indicating no missing values by default.
alpha	a numerical value between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

### Details

Hu and Zhang's Doubly Biased Coin Design with delayed Binary Response employs the following treatment allocation scheme:

(a) Initially, due to limited information about treatment efficacy, the first  $n_0$  patients are assigned to  $K$  treatments using restricted randomization (as described by Rosenberger and Lachin, 2002).

(b) For  $m \geq n_0$ , patient  $(m + 1)$  is allocated to treatment  $k$  with a probability  $p_{m+1,k}$ , which depends on the available responses and estimated target allocation via  $g_k$ , as proposed by Hu and Zhang (2004).

For a more comprehensive description of the procedure, please refer to the paper titled 'Doubly adaptive biased coin designs with delayed responses' authored by Hu et al. in 2008.

### Value

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation propotion for each of treatment groups.
failRate	The proportion of individuals who do not achieve the expected outcome in each simulation, on average.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

### References

Hu, F., Zhang, L. X., Cheung, S. H., & Chan, W. S. (2008). *Doubly adaptive biased coin designs with delayed responses*. Canadian Journal of Statistics, 36(4), 541-559.

### Examples

```
# a simple use
# Define the arguments
## Arguments for generate the simulated data
### For response simulation
p = c(0.6, 0.8)
k = 2
```

```

ssn = 100
### for enter time and response time simulation
ent.param = 0.7
rspT.dist = "exponential"
rspT.param = c(1, 1, 3, 1)

## Arguments for the deisgn
n0 = 20
target.alloc = "RSIHR"

dyldBCD_Bin(n0 = n0, p = p, k = k, ssn = ssn, ent.param, rspT.dist, rspT.param, theta0 = NULL,
            target.alloc, r = 2, nsim = 150, mRate = NULL, alpha = 0.05)

```

---

dyldBCD_Cont	<i>Hu and Zhang's Doubly Biased Coin Deisgn with delayed Continuous Response</i>
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---

### Description

Simulating Hu and Zhang's doubly biased coin deisgn with delayed continuous response (number of arms  $\geq 2$ ) in a clinical trial context. (Inference: two-sided hypothesis testing t-test or chi-square test)

### Usage

```

dyldBCD_Cont(n0 = 20, theta, k, ssn, ent.param, rspT.dist, rspT.param,
            target.alloc = "Neyman", r = 2, nsim = 2000, mRate = NULL, alpha = 0.05)

```

### Arguments

n0	A positive integer. n0 represents the initial patient population assigned through restricted randomization for initial parameter estimation.
theta	A numerical vector of length equal to 2k. These values specify the true parameters for each treatment and are used for generating data in simulations. For example, if k=2, you should provide two pairs of parameter values, each consisting of the mean and variance, like: theta = c(13, 4.0^2, 15, 2.5^2).
k	A positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k \geq 2$ )
ssn	A positive integer. The value specifies the total number of participants involved in each round of the simulation.
ent.param	A positive integer. The value specified the parameter for an exponential distribution which determine the time for each participant enter the trial.
rspT.dist	Distribution Type. Specifies the type of distribution that models the time spent for the availability of patient $i$ under treatment $k$ . Acceptable options for this argument include: "exponential", "normal", and "uniform".

<code>rspT.param</code>	A vector. Specifies the parameters required by the distribution that models the time spent for the availability under each treatment. (eg. If there are 3 treatment groups and each of them follows truncated normal distribution with parameter pair (3, 2), (2, 1), (4, 1), respectively. Then the <code>rspT.param = c(3, 2, 2, 1, 4, 1)</code> )
<code>target.alloc</code>	Desired allocation proportion. The option for this argument could be one of "Neyman", "ZR", "DaOptimal". The default is "Neyman". The details see Zhang L. and Rosenberger. W (2006).
<code>r</code>	A positive number. Parameter for Hu and Zhang's doubly biased coin design and usually take values 2-4. The default value is 2.
<code>nsim</code>	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
<code>mRate</code>	a numerical value between 0 and 1, inclusive, representing the missing rate for the responses. This parameter pertains to missing-at-random data. The default value is NULL, indicating no missing values by default.
<code>alpha</code>	a numerical value between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

### Details

Hu and Zhang's Doubly Biased Coin Design with delayed Continuous Response employs the following treatment allocation scheme:

(a) Initially, due to limited information about treatment efficacy, the first  $n_0$  patients are assigned to  $K$  treatments using restricted randomization (as described by Rosenberger and Lachin, 2002).

(b) For  $m \geq n_0$ , patient  $(m + 1)$  is allocated to treatment  $k$  with a probability  $p_{m+1,k}$ , which depends on the available responses and estimated target allocation via  $g_k$ , as proposed by Hu and Zhang (2004).

For a more comprehensive description of the procedure, please refer to the paper titled 'Doubly adaptive biased coin designs with delayed responses' authored by Hu et al. in 2008.

### Value

<code>name</code>	The name of procedure.
<code>parameter</code>	The true parameters used to do the simulations.
<code>assignment</code>	The randomization sequence.
<code>propotion</code>	Average allocation propotion for each of treatment groups.
<code>failRate</code>	The average response value for the entire trial.
<code>pwClac</code>	The probability of the study to detect a significant difference or effect if it truly exists.
<code>k</code>	Number of arms involved in the trial.



## References

Hu, F., & Zhang, L. X. (2004). *Asymptotic properties of doubly adaptive biased coin designs for multi-treatment clinical trials*. The Annals of Statistics, 32(1), 268-301.

Hu, F., Zhang, L. X., Cheung, S. H., & Chan, W. S. (2008). *Doubly adaptive biased coin designs with delayed responses*. Canadian Journal of Statistics, 36(4), 541-559.

## Examples

```
# a simple use
# Define the arguments
## Arguments for generate the simulated data
### For response simulation
theta = c(13, 4.0^2, 15, 2.5^2)
k = 2
ssn = 88

### for enter time and response time simulation
ent.param = 5
rspT.param = rep(10, 2)
rspT.dist = "exponential"

## Arguments for the deisgn
target.alloc = "Neyman"

res = dyldBCD_Cont(n0 = 10, theta, k, ssn, ent.param, rspT.dist,
                  rspT.param, target.alloc, r = 2, nsim = 200,
                  mRate = 0.2, alpha = 0.05)

# View the output (A list of all results)
res
```

---

GDLRule

*Generalized drop-the-loser rule*

---

## Description

Simulating generalized drop-the-loser rule procedure (number of arms > 2) with two-sided hypothesis testing in a clinical trial context.

## Usage

```
GDLRule(k, p, ssn, aK, Y0 = NULL, nsim = 2000, alpha = 0.05)
```

## Arguments

**k** a positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k \geq 2$ )

p	a positive vector of length equals to k. The values specify the true success rates for the various treatments, and these rates are used to generate data for simulations.
ssn	a positive integer. The value specifies the total number of participants involved in each round of the simulation.
aK	a positive vector of length equals to k. The values specifies when the immigration ball is drawn, the number of each treatment ball added to the urn.
Y0	A vector of length k, specifying the initial probability of allocating a patient to each group. For instance, if $Y0 = c(1, 1, 1)$ , the initial probabilities are calculated as $Y0 / \text{sum}(Y0)$ . When Y0 is NULL, the initial urn will be set as If Y0 is NULL, then Y0 is set to a vector of length k, with all values equal to 1 by default.
nsim	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
alpha	A number between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

### Details

Consider an urn containing balls of  $K + 1$  types. Balls of types \$1, \dots, K\$ represent treatments, balls of type 0 will be called immigration balls. When the subject arrived for randomization, a ball is drawn at random. If the ball is of type 0 (i.e, an immigration ball), no subject is treated, and the ball is returned to the urn together with  $A = a_1 + \dots + a_K$  additional balls,  $a_k$  of treatment type  $k, k = 1, \dots, K$ . If a treatment ball is drawn (say, of type  $k$ , for some  $k = 1, \dots, K$ ) the next subject is given treatment  $k$  and the ball is not replaced. If the observed response of this subject is a success, then the ball is replaced, otherwise not replaced.

### Value

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation propotion for each of treatment groups.
failRate	The proportion of individuals who do not achieve the expected outcome in each simulation, on average.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

### References

Zhang, L. X., Chan, W. S., Cheung, S. H., & Hu, F. (2007). *A generalized drop-the-loser urn for clinical trials with delayed responses*. *Statistica Sinica*, 17(1), 387-409.

**Examples**

```
## a simple use
gdl.res = GDLRule(k = 3, p = c(0.6, 0.7, 0.6),
                 ssn = 400, aK = c(1, 1, 1), Y0 = NULL, nsim = 200, alpha = 0.05)

## view the output
gdl.res

## view all simulation settings
gdl.res$name
gdl.res$parameter
gdl.res$k

## View the simulations results
gdl.res$propotion
gdl.res$failRate
gdl.res$pwCalc
gdl.res$assignment
```

---

Group.DBCD\_Bin

*Group Doubly Biased Coin Design with Binary Response Type*


---

**Description**

Simulating group doubly biased coin design with binary response (number of arms  $\geq 2$ ) in a clinical trial context. (Inference: two-sided hypothesis testing t-test or chi-square test)

**Usage**

```
Group.DBCD_Bin(n0 = 20, p, k, gsize.param, ssn, theta0 = NULL,
               target.alloc = "RPW", r = 2, nsim = 2000,
               mRate = NULL, alpha = 0.05)
```

**Arguments**

<code>n0</code>	A positive integer. <code>n0</code> represents the initial patient population assigned through restricted randomization for initial parameter estimation.
<code>p</code>	A positive vector of length equals to <code>k</code> . The values specify the true success rates for the various treatments, and these rates are used to generate data for simulations.
<code>k</code>	A positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k \geq 2$ )
<code>gsize.param</code>	A positive integer. It represents the expected number of people enrolling in a specified interval, assuming that the enrollment rate per unit time follows a Poisson distribution.

ssn	A positive integer. The value specifies the total number of participants involved in each round of the simulation.
theta0	A vector of length k. Each value in the vector represents a probability used for adjusting parameter estimates. If the argument is not provided, it defaults to a vector of length k, with all values set to 0.5.
target.alloc	Desired allocation proportion. The option for this argument could be one of "Neyman", "RSIHR", "RPW", "WeisUrn". The default is "RPW".
r	A positive number. Parameter for Hu and Zhang's doubly biased coin design and usually take values 2-4. The default value is 2.
nsim	A positive integer. The value specifies the total number of simulations, with a default value of 2000.
mRate	A numerical value between 0 and 1, inclusive, representing the missing rate for the responses. This parameter pertains to missing-at-random data. The default value is NULL, indicating no missing values by default.
alpha	A number between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

## Details

Hu and Zhang's Doubly Biased Coin Design (DBCD) adjusts the probability of assigning each patient to a specific treatment group in a clinical trial, based on the responses of all previous patients. The Group DBCD is an enhanced version of this approach, offering a more practical perspective. It dynamically updates the allocation probabilities for patients in each group based on the responses of all preceding groups, either when the data available or at fixed time intervals (weekly or biweekly).

The process begins by assigning  $n_0$  (perhaps the first few groups of) patients to treatment groups using restricted randomization and collecting their responses. Initial parameter estimates for the response variable are then obtained for each treatment group. Subsequently, based on these parameter estimates, the estimated desired allocation proportion is calculated. Afterward, the Hu and Zhang's allocation function is applied to determine the probabilities for the next group of patients to be assigned to each treatment group, which force the allocation proportion close to the desired one. This process is repeated sequentially for each group until the desired number of patients has been allocated, as predetermined.

This methodology was introduced by Zhai, Li, Zhang and Hu in their 2023 paper titled 'Group Response-Adaptive Randomization with Delayed and Missing Responses'.

## Value

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation propotion for each of treatment groups.
failRate	The proportion of individuals who do not achieve the expected outcome in each simulation, on average.

pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

## References

Zhai, G., Li, Y., Zhang, L. X. & Hu, F. (2023). *Group Response-Adaptive Randomization with Delayed and Missing Responses*.

## Examples

```
Group.DBCD_Bin(n0 = 20, p = c(0.65, 0.8), k = 2, gsize.param = 5,
              ssn = 300, theta0 = NULL, target.alloc = "RPW",
              r = 2, nsim = 400, mRate = NULL, alpha = 0.05)
```

---

Group.DBCD_Cont	<i>Group Doubly Biased Coin Design with Continuous Response Type</i>
-----------------	--

---

## Description

Simulating group doubly biased coin design with continuous response (number of arms  $\geq 2$ ) in a clinical trial context. (Inference: two-sided hypothesis testing t-test or chi-square test)

## Usage

```
Group.DBCD_Cont(n0 = 20, theta, k, gsize.param, ssn,
               theta0 = NULL, target.alloc = "Neyman",
               r = 2, nsim = 2000, mRate = NULL, alpha = 0.05)
```

## Arguments

n0	A positive integer. n0 represents the initial patient population assigned through restricted randomization for initial parameter estimation.
theta	A numerical vector of length equal to 2k. These values specify the true parameters for each treatment and are used for generating data in simulations. For example, if k=2, you should provide two pairs of parameter values, each consisting of the mean and variance, like: theta = c(13, 4.0^2, 15, 2.5^2).
k	A positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k = 2$ )
ssn	A positive integer. The value specifies the total number of participants involved in each round of the simulation.
gsize.param	A positive integer. It represents the expected number of people enrolling in a specified interval, assuming that the enrollment rate per unit time follows a Poisson distribution.
theta0	A vector of length 2k. Each value in the vector represents a probability used for adjusting parameter estimates. If the argument is not provided, it defaults to a vector of length 2k, with all parameter pair set to be (0, 1).

target.alloc	Desired allocation proportion. The option for this argument could be one of "Neyman", "ZR", "DaOptimal". The default is "Neyman". The details see Zhang L. and Rosenberger. W (2006).
r	A positive number. Parameter for Hu and Zhang's doubly biased coin design and usually take values 2-4. The default value is 2.
nsim	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
mRate	a numerical value between 0 and 1, inclusive, representing the missing rate for the responses. This parameter pertains to missing-at-random data. The default value is NULL, indicating no missing values by default.
alpha	a numerical value between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

### Details

Hu and Zhang's Doubly Biased Coin Design (DBCD) adjusts the probability of assigning each patient to a specific treatment group in a clinical trial, based on the responses of all previous patients. The Group DBCD is an enhanced version of this approach, offering a more practical perspective. It dynamically updates the allocation probabilities for patients in each group based on the responses of all preceding groups, either when the data available or at fixed time intervals (weekly or biweekly). The process begins by assigning  $n_0$  (perhaps the first few groups of) patients to treatment groups using restricted randomization and collecting their responses. Initial parameter estimates for the response variable are then obtained for each treatment group. Subsequently, based on these parameter estimates, the estimated desired allocation proportion is calculated. Afterward, the Hu and Zhang's allocation function is applied to determine the probabilities for the next group of patients to be assigned to each treatment group, which force the allocation proportion close to the desired one. This process is repeated sequentially for each group until the desired number of patients has been allocated, as predetermined.

This methodology was introduced by Zhai, Li, Zhang and Hu in their 2023 paper titled 'Group Response-Adaptive Randomization with Delayed and Missing Responses'.

### Value

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation porpotion for each of treatment groups.
failRate	The average response value for the entire trial.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

### References

Zhai, G., Li, Y., Zhang, L. X. & Hu, F. (2023). *Group Response-Adaptive Randomization with Delayed and Missing Responses*.

**Examples**

```
theta = c(13, 4.0^2, 15, 2.5^2)
k = 2
gsize.param = 5
ssn = 120
Group.DBCD_Cont(n0 = 20, theta, k, gsize.param, ssn,
               target.alloc = "Neyman", r = 2, nsim = 500,
               mRate = NULL, alpha = 0.05)
```

---

Group.dyldBCD\_Bin      *Group Doubly Biased Coin Design with Delayed Discrete Response*

---

**Description**

Simulating group doubly biased coin design with delayed discrete response (number of arms  $\geq 2$ ) in a clinical trial context. (Inference: two-sided hypothesis testing t-test or chi-square test)

**Usage**

```
Group.dyldBCD_Bin(n0 = 20, p, k, ssn, gsize.param,
                 rspT.dist, rspT.param, theta0 = NULL,
                 target.alloc = "RPW", r = 2, nsim = 2000,
                 eTime = 7, mRate = NULL, alpha = 0.05)
```

**Arguments**

<code>n0</code>	A positive integer. <code>n0</code> represents the initial patient population assigned through restricted randomization for initial parameter estimation.
<code>p</code>	A positive vector of length equals to <code>k</code> . The values specify the true success rates for the various treatments, and these rates are used to generate data for simulations.
<code>k</code>	A positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k = 2$ )
<code>ssn</code>	A positive integer. The value specifies the total number of participants involved in each round of the simulation.
<code>gsize.param</code>	A positive integer. It represents the expected number of people enrolling in a specified interval, assuming that the enrollment rate per unit time follows a Poisson distribution.
<code>rspT.dist</code>	Distribution Type. Specifies the type of distribution that models the time spent for the availability of patient $i$ under treatment $k$ . Acceptable options for this argument include: "exponential", "normal", and "uniform".
<code>rspT.param</code>	A vector with length $2k$ . Specifies the parameters required by the distribution that models the time spent for the availability under each treatment and each response. (eg. If there are 3 treatments groups with 0 or 1 as response and each of them follows exponential distribution with parameter (3, 2, 3, 3, 1, 2), respectively. Then the <code>rspT.param = c(3, 2, 2, 1, 4, 1)</code> )

<code>theta0</code>	A vector of length $k$ . Each value in the vector represents a probability used for adjusting parameter estimates. If the argument is not provided, it defaults to a vector of length $k$ , with all values set to 0.5.
<code>target.alloc</code>	Desired allocation proportion. The option for this argument could be one of "Neyman", "RSIHR", "RPW", "WeisUrn". The default is "RPW".
<code>r</code>	A positive number. Parameter for Hu and Zhang's doubly biased coin design and usually take values 2-4. The default value is 2.
<code>nsim</code>	A positive integer. The value specifies the total number of simulations, with a default value of 2000.
<code>eTime</code>	A positive number. The interval time between enrollment of participants in each group. The default is 7.
<code>mRate</code>	a numerical value between 0 and 1, inclusive, representing the missing rate for the responses. This parameter pertains to missing-at-random data. The default value is NULL, indicating no missing values by default.
<code>alpha</code>	a numerical value between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

## Details

Hu and Zhang's Doubly Biased Coin Design (DBCD) adjusts the probability of assigning each patient to a specific treatment group in a clinical trial, based on the responses of all previous patients. The Group DBCD is an enhanced version of this approach, offering a more practical perspective. It dynamically updates the allocation probabilities for patients in each group based on the available responses of all preceding groups, either when the data available or at fixed time intervals (weekly or biweekly). Here, the function focuses on implementing the group doubly biased coin design, tailored to delayed binary responses.

The process begins by assigning  $n_0$  (perhaps the first few groups of) patients to treatment groups using restricted randomization and collecting their responses. Initial parameter estimates for the response variable are then obtained for each treatment group. Subsequently, based on these parameter estimates, the estimated desired allocation proportion is calculated. Afterward, the Hu and Zhang's allocation function is applied to determine the probabilities for the next group of patients to be assigned to each treatment group, which force the allocation proportion close to the desired one. This process is repeated sequentially for each group until the desired number of patients has been allocated, as predetermined.

This methodology was introduced by Zhai, Li, Zhang and Hu in their 2023 paper titled 'Group Response-Adaptive Randomization with Delayed and Missing Responses'.

## Value

<code>name</code>	The name of procedure.
<code>parameter</code>	The true parameters used to do the simulations.
<code>assignment</code>	The randomization sequence.
<code>propotion</code>	Average allocation propotion for each of treatment groups.
<code>failRate</code>	The proportion of individuals who do not achieve the expected outcome in each simulation, on average.



pwClac            The probability of the study to detect a significant difference or effect if it truly exists.

k                 Number of arms involved in the trial.

## References

Zhai, G., Li, Y., Zhang, L. X. & Hu, F. (2023). *Group Response-Adaptive Randomization with Delayed and Missing Responses*.

## Examples

```
# a simple use
# Define the arguments
## Arguments for generate the simulated data
### For response simulation
p = c(0.7, 0.5)
k = 2
ssn = 200

### for enter time and response time simulation
eTime = 7
rspT.param = rep(10, 4)
rspT.dist = "exponential"
gsize.param = 5

## Arguments for the deisgn
n0 = 10
target.alloc = "RPW"

res = Group.dyldDBCD_Bin(n0, p, k, ssn, gsize.param,
                        rspT.dist, rspT.param, theta0 = NULL,
                        target.alloc = "RPW", r = 2,
                        nsim = 120, eTime = 7, mRate = NULL, alpha = 0.05)

# View the output (A list of all results)
res
```

---

Group.dyldDBCD\_Cont    *Group Doubly Biased Coin Design with Delayed Continuous Response*

---

## Description

Simulating group doubly biased coin deisgn with delayed continuous response (number of arms  $\geq 2$ ) in a clinical trial context. (Inference: two-sided hypothesis testing t-test or chi-square test)

**Usage**

```
Group.dyldBCD_Cont(n0 = 20, theta, k, ssn,
                  gsize.param, rspT.dist, rspT.param,
                  target.alloc = "Neyman", r = 2,
                  nsim = 2000, eTime = 7,
                  mRate = NULL, alpha = 0.05)
```

**Arguments**

<code>n0</code>	A positive integer. <code>n0</code> represents the initial patient population assigned through restricted randomization for initial parameter estimation.
<code>theta</code>	A numerical vector of length equal to $2k$ . These values specify the true parameters for each treatment and are used for generating data in simulations. For example, if $k=2$ , you should provide two pairs of parameter values, each consisting of the mean and variance, like: <code>theta = c(13, 4.0^2, 15, 2.5^2)</code> .
<code>k</code>	A positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k = 2$ )
<code>ssn</code>	A positive integer. The value specifies the total number of participants involved in each round of the simulation.
<code>gsize.param</code>	A positive integer. It represents the expected number of people enrolling in a specified interval, assuming that the enrollment rate per unit time follows a Poisson distribution.
<code>rspT.dist</code>	Distribution Type. Specifies the type of distribution that models the time spent for the availability of patient $i$ under treatment $k$ . Acceptable options for this argument include: "exponential", "normal", and "uniform".
<code>rspT.param</code>	A vector. Specifies the parameters required by the distribution that models the time spent for the availability under each treatment. (eg. If there are 3 treatments groups and each of them follows truncated normal distribution with parameter pair (3, 2), (2, 1), (4, 1), respectively. Then the <code>rspT.param = c(3, 2, 2, 1, 4, 1)</code> )
<code>target.alloc</code>	Desired allocation proportion. The option for this argument could be one of "Neyman", "ZR", "DaOptimal". The default is "Neyman". The details see Zhang L. and Rosenberger. W (2006).
<code>r</code>	A positive number. Parameter for Hu and Zhang's doubly biased coin design and usually take values 2-4. The default value is 2.
<code>nsim</code>	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
<code>eTime</code>	A positive number. The interval time between enrollment of participants in each group. The default is 7.
<code>mRate</code>	a numerical value between 0 and 1, inclusive, representing the missing rate for the responses. This parameter pertains to missing-at-random data. The default value is NULL, indicating no missing values by default.
<code>alpha</code>	a numerical value between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

## Details

Hu and Zhang's Doubly Biased Coin Design (DBCD) adjusts the probability of assigning each patient to a specific treatment group in a clinical trial, based on the responses of all previous patients. The Group DBCD is an enhanced version of this approach, offering a more practical perspective. It dynamically updates the allocation probabilities for patients in each group based on the available responses of all preceding groups, either when the data available or at fixed time intervals (weekly or biweekly). Here, the function focuses on implementing the group doubly biased coin design, tailored to delayed continuous responses.

The process begins by assigning  $n_0$  (perhaps the first few groups of) patients to treatment groups using restricted randomization and collecting their responses. Initial parameter estimates for the response variable are then obtained for each treatment group. Subsequently, based on these parameter estimates, the estimated desired allocation proportion is calculated. Afterward, the Hu and Zhang's allocation function is applied to determine the probabilities for the next group of patients to be assigned to each treatment group, which force the allocation proportion close to the desired one. This process is repeated sequentially for each group until the desired number of patients has been allocated, as predetermined.

This methodology was introduced by Zhai, Li, Zhang and Hu in their 2023 paper titled 'Group Response-Adaptive Randomization with Delayed and Missing Responses'.

## Value

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation porpotion for each of treatment groups.
failRate	The average response value for the entire trial.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

## References

Zhai, G., Li, Y., Zhang, L. X. & Hu, F. (2023). *Group Response-Adaptive Randomization with Delayed and Missing Responses*.

## Examples

```
# a simple use
# Define the arguments
## Arguments for generate the simulated data
### For response simulation
theta = c(13, 4.0^2, 15, 2.5^2)
k = 2
ssn = 120

### for enter time and response time simulation
eTime = 7
```

```

gsize.param = 5
rspT.param = rep(10, 2)
rspT.dist = "exponential"

## Arguments for the deisgn
n0 = 10
target.alloc = "Neyman"

res = Group.dylDBCD_Cont(n0, theta, k, ssn, gsize.param,
                        rspT.dist, rspT.param, target.alloc = "Neyman",
                        r = 2, nsim = 200, eTime = 7, mRate = 0.2, alpha = 0.05)

# View the output (A list of all results)
res

```

---

grouprar

*grouprar-package: Group Response-Adaptive Randomization for Clinical Trials*


---

## Description

Implement group response-adaptive randomization procedures, which also integrates standard non-group response-adaptive randomization methods as specialized instances. It is also uniquely capable of managing complex scenarios, including those with delayed and missing responses, thereby expanding its utility in real-world applications. This package offers 16 functions for simulating a variety of response adaptive randomization procedures. These functions are essential for guiding the selection of statistical methods in clinical trials, providing a flexible and effective approach to trial design. Some of the detailed methodologies and algorithms used in this package, please refer to the following references: LJ Wei (1979) <doi:10.1214/aos/1176344614> L. J. WEI and S. DURHAM (1978) <doi:10.1080/01621459.1978.10480109> Durham, S. D., Flournoy, N. AND LI, W. (1998) <doi:10.2307/3315771> Ivanova, A., Rosenberger, W. F., Durham, S. D. and Flournoy, N. (2000) <https://www.jstor.org/stable/25053121> Bai Z D, Hu F, Shen L. (2002) <doi:10.1006/jmva.2001.1987> Ivanova, A. (2003) <doi:10.1007/s001840200220> Hu, F., & Zhang, L. X. (2004) <doi:10.1214/aos/1079120137> Hu, F., & Rosenberger, W. F. (2006, ISBN:978-0-471-65396-7). Zhang, L. X., Chan, W. S., Cheung, S. H., & Hu, F. (2007) <https://www.jstor.org/stable/26432528> Zhang, L., & Rosenberger, W. F. (2006) <doi:10.1111/j.1541-0420.2005.00496.x> Hu, F., Zhang, L. X., Cheung, S. H., & Chan, W. S. (2008) <doi:10.1002/cjs.5550360404>.

## Author(s)

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---

 PolyaUrn

 Randomized Pólya urn procedure
 

---

### Description

Simulating randomized Pólya urn procedure with two-sided hypothesis testing in a clinical trial context.

### Usage

PolyaUrn(k, p, ssn, Y0 = NULL, nsim = 2000, alpha = 0.05)

### Arguments

- |     |  |
|-----|--|
| k   | a positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k \geq 2$ )  |
| p   | a positive vector of length equals to k. The values specify the true success rates for the various treatments, and these rates are used to generate data for simulations.  |
| ssn | a positive integer. The value specifies the total number of participants involved in each round of the simulation.   |
| Y0  | A vector of length k, specifying the initial probability of allocating a patient to each group. For instance, if $Y0 = c(1, 1, 1)$ , the initial probabilities are calculated as $Y0 / \text{sum}(Y0)$ . When Y0 is NULL, the initial urn will be set as If Y0 is NULL, then Y0 is set to a vector of length k, with all values equal to 1 by default. |

nsim	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
alpha	A number between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

### Details

The randomized Pólya urn (RPU) procedure can be describe as follows: An urn contains at least one ball of each treatment type (totally  $K$  treatments) initially. A ball is drawn from the urn with replacement. If a type  $i$  ball is drawn,  $i = 1, \dots, K$ , then treatment  $i$  is assigned to the next patient. If the response is a success, a ball of type  $i$  is added to the urn. Otherwise the urn remains unchanged.

### Value

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation propotion for each of treatment groups.
failRate	The proportion of individuals who do not achieve the expected outcome in each simulation, on average.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

### References

Durham, S. D., FlournoY, N. AND LI, W. (1998). *Sequential designs for maximizing the probability of a favorable response*. Canadian Journal of Statistics, 3, 479-495.

### Examples

```
## a simple use
Polya.res = PolyaUrn(k = 3, p = c(0.6, 0.7, 0.6), ssn = 400, Y0 = NULL, nsim = 200, alpha = 0.05)

## view the output
Polya.res

## view all simulation settings
Polya.res$name
Polya.res$parameter
Polya.res$k

## View the simulations results
Polya.res$propotion
Polya.res$failRate
Polya.res$pwCalc
Polya.res$assignment
```

---

RPWRule

*Randomized Play-the-winner Rule*


---

### Description

Simulating randomized play-the-winner rule with two-sided hypothesis testing in a clinical trial context.

### Usage

```
RPWRule(k, p, ssn, Y0 = NULL, nsim = 2000, alpha = 0.05)
```

### Arguments

k	a positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k = 2$ )
p	a positive vector of length equals to k. The values specify the true success rates for the various treatments, and these rates are used to generate data for simulations.
ssn	a positive integer. The value specifies the total number of participants involved in each round of the simulation.
Y0	A vector of length k, specifying the initial probability of allocating a patient to each group. For instance, if $Y0 = c(1, 1, \dots)$ , the initial probabilities are calculated as $Y0 / \text{sum}(Y0)$ . When Y0 is NULL, the initial urn will be set as If Y0 is NULL, then Y0 is set to a vector of length k, with all values equal to 1 by default.
nsim	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
alpha	An integer between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

### Details

The Randomized Play-the-Winner Rule allocates future subjects in a clinical trial to treatment groups based on the performance of previously treated subjects. This rule increases the likelihood of future patients being assigned to the better-performing treatment, as determined by the outcomes of previously treated subjects.

### Value

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.

propotion	Average allocation porpotion for each of treatment groups.
failRate	The proportion of individuals who do not achieve the expected outcome in each simulation, on average.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

## References

L. J. WEI and S. DURHAM (1978) *The Randomized Play-the-Winner Rule in Medical Trials*. Journal of the American Statistical Association, 73, 364, 840–843.

## Examples

```
## a simple use
RPW.res = RPWRule(k = 2, p = c(0.7, 0.8), ssn = 400, Y0 = NULL, nsim = 200, alpha = 0.05)
## view the output
RPW.res

## view all simulation settings
RPW.res$name
RPW.res$parameter
RPW.res$k

## View the simulations results
RPW.res$propotion
RPW.res$failRate
RPW.res$pwCalc
RPW.res$assignment
```

---

WeiUrn

*Randomized Play-the-winner rule with multiple arms ( $k > 2$ )*

---

## Description

Simulating randomized play-the-winner rule (multiple arms) with two-sided hypothesis testing in a clinical trial context.

## Usage

```
WeiUrn(k, p, ssn, Y0 = NULL, nsim = 2000, alpha = 0.05)
```



**Arguments**

k	a positive integer. The value specifies the number of treatment groups involved in a clinical trial. ( $k > 2$ )
p	a positive vector of length equals to k. The values specify the true success rates for the various treatments, and these rates are used to generate data for simulations.
ssn	a positive integer. The value specifies the total number of participants involved in each round of the simulation.
Y0	A vector of length k, specifying the initial probability of allocating a patient to each group. For instance, if $Y0 = c(1, 1, 1)$ , the initial probabilities are calculated as $Y0 / \text{sum}(Y0)$ . When Y0 is NULL, the initial urn will be set as If Y0 is NULL, then Y0 is set to a vector of length k, with all values equal to 1 by default.
nsim	a positive integer. The value specifies the total number of simulations, with a default value of 2000.
alpha	A number between 0 and 1. The value represents the predetermined level of significance that defines the probability threshold for rejecting the null hypothesis, with a default value of 0.05.

**Details**

Wei's urn procedure is obtained by extending the randomized play the winner rule (Wei1978) from the case  $k = 2$  to  $k > 2$ . Hence, It enables to conduct multi-arm clinical trials, and offers a greater range of applications.

**Value**

name	The name of procedure.
parameter	The true parameters used to do the simulations.
assignment	The randomization sequence.
propotion	Average allocation porpotion for each of treatment groups.
failRate	The proportion of individuals who do not achieve the expected outcome in each simulation, on average.
pwClac	The probability of the study to detect a significant difference or effect if it truly exists.
k	Number of arms involved in the trial.

**References**

LJ Wei (1979). *The generalized polya's urn design for sequential medical trials*. The Annals of Statistics, 7(2):291–296, 19

**Examples**

```
## a simple use
wei.res = WeiUrn(k = 3, p = c(0.7, 0.8, 0.7), ssn = 400, Y0 = NULL, nsim = 200, alpha = 0.05)

## view the output
wei.res

## view all simulation settings
wei.res$name
wei.res$parameter
wei.res$k

## View the simulations results
wei.res$propotion
wei.res$failRate
wei.res$pWCalc
wei.res$assignment
```

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