The PwrGSD Package

March 12, 2010

PwrGSD   *Calculate Power in a Group Sequential Design*

Description

Derives power in a two arm clinical trial under a group sequential design. Allows for arbitrary number of interim analyses, arbitrary specification of arm-0/arm-1 time to event distributions (via survival or hazard), arm-0/arm-1 censoring distribution, provisions for two types of continuous time non-compliance according to arm-0/arm-1 rate followed by switch to new hazard rate. Allows for analyses using (I) weighted log-rank statistic, with weighting function (a) a member of the Flemming-Harrington G-Rho class, or (b) a stopped version thereof, or (c) the ramp-plateau deterministic weights, or (II) the integrated survival distance (currently under method=="S" without futility only). Stopping boundaries are computed via the Lan-Demets method, Haybittle method, or converted from the stochastic curtailment procedure. The Lan-Demets boundaries can be constructed usign either O'Brien-Flemming, Pocock or Wang-Tsiatis power alpha-spending. The C kernel is readily extensible, and further options will become available in the near future.

Usage

```
PwrGSD(EfficacyBoundary = LanDemets(alpha = 0.05, spending = ObrienFleming),
    FutilityBoundary = LanDemets(alpha = 0.1, spending = ObrienFleming),
    NonBindingFutility = TRUE, sided = c("2>", "2<", "1>", "1<"),
    method = c("S", "A"), accru, accrat, tlook,
    tcut0 = NULL, h0 = NULL, s0 = NULL, tcut1 = NULL,
    rhaz = NULL, h1 = NULL, s1 = NULL, tcutc0 = NULL, hc0 = NULL,
    sc0 = NULL, tcutc1 = NULL, hc1 = NULL, sc1 = NULL, tcutd0A = NULL,
    hd0A = NULL, sd0A = NULL, tcutd0B = NULL, hd0B = NULL, sd0B = NULL,
    tcutd1A = NULL, hd1A = NULL, sd1A = NULL, tcutd1B = NULL,
    hd1B = NULL, sd1B = NULL, tcutx0A = NULL, hx0A = NULL, sx0A = NULL,
    tcutx0B = NULL, hx0B = NULL, sx0B = NULL, tcutx1A = NULL,
    hx1A = NULL, sx1A = NULL, tcutx1B = NULL, hx1B = NULL, sx1B = NULL,
    noncompliance = c("none", "crossover", "mixed", "user"),
    gradual = FALSE, WtFun = c("FH", "SFH", "Ramp"), ppar = cbind(c(0, 0)),
    Spend.Info = c("Variance", "Events", "Hybrid(k)", "Calendar"), RR.Futility =
    qProp.one.or.Q = c("one", "Q"), Nsim = NULL, detail = FALSE, StatType = c("W
        "ISD"))
```

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Arguments

EfficacyBoundary

This specifies the method used to construct the efficacy boundary. The available choices are:

'(i) 'Lan-Demets(alpha=<total type I error>, spending=<spending function>). The Lan-Demets method is based upon a error probability spending approach. The spending function can be set to `ObrienFleming`, `Pocock`, or `Power(rho)`, where `rho` is the the power argument for the power spending function: rho=3 is roughly equivalent to the O'Brien-Fleming spending function and smaller powers result in a less conservative spending function.

'(ii) 'Haybittle(alpha=<total type I error>, b.Haybittle=<user specified boundary point>). The Haybittle approach is the simplest, which sets the boundary points equal to `b.Haybittle`, a user specified value (try 3) for all analyses except the last, which is calculated so as to result in the total type I error, set with the argument `alpha`.

'(iii) 'SC(be.end=<efficacy boundary point at trial end>, prob=<threshold for conditional type I error for efficacy stopping>). The stochastic curtailment method is based upon the conditional probability of type I error given the current value of the statistic. Under this method, a sequence of boundary points on the standard normal scale (as are boundary points under all other methods) is calculated so that the total probability of type I error is maintained. This is done by considering the joint probabilities of continuing to the current analysis and then exceeding the threshold at the current analysis. A good value for the threshold value for the conditional type I error, `prob` is 0.90 or greater.

'(iv) 'User supplied boundary points in the form `c(b1, b2, b3, ..., b_m)`, where `m` is the number of looks.

FutilityBoundary

This specifies the method used to construct the futility boundary. The available choices are:

'(i) 'Lan-Demets(alpha=<total type II error>, spending=<spending function>). The Lan-Demets method is based upon a error probability spending approach. The spending function can be set to `ObrienFleming`, `Pocock`, or `Power(rho)`, where `rho` is the the power argument for the power spending function: rho=3 is roughly equivalent to the O'Brien-Fleming spending function and smaller powers result in a less conservative spending function.

'(ii) 'Haybittle(alpha=<total type I error>, b.Haybittle=<user specified boundary point>). The Haybittle approach is the simplest, which sets the boundary points equal to `b.Haybittle`, a user specified value (try 3) for all analyses except the last, which is calculated so as to result in the total type II error, set with the argument `alpha`.

'(iii) 'SC(be.end=<efficacy boundary point at trial end>, prob=<threshold for conditional type II error for futility stopping>, drift.end=<projected drift at end of trial>). The stochastic curtailment method is based upon the conditional probability of type II error given the current value of the statistic. Under this method, a sequence of boundary points on the standard normal scale (as are boundary points under all other methods) is calculated so that the total probability of type II error, is maintained. This is done by considering the joint probabilities of continuing to the current analysis and then exceeding the threshold at the current analysis. A good value for the threshold value for the conditional type I error, `prob` is

0.90 or greater.

'(iv) 'User supplied boundary points in the form `c(b1, b2, b3, ..., b_m)`, where `m` is the number of looks.

NonBindingFutility

    When using a futility boundary and this is set to 'TRUE', the efficacy boundary will be constructed in the absence of the futility boundary, and then the futility boundary will be constructed given the resulting efficacy boundary. This results in a more conservative efficacy boundary with true type I error less than the nominal level. This is recommended due to the fact that futility crossings are viewed by DSMB's with much less gravity than an efficacy crossing and as such, the consensus is that efficacy bounds should not be discounted towards the null hypothesis because of paths which cross a futility boundary. Default value is 'TRUE'.

sided

    Set to "2>" (quoted) for two sided tests of the null hypothesis when a positive drift corresponds to efficacy. Set to "2<" (quoted) for two sided tests of the null hypothesis when a negative drift corresponds to efficacy. Set to "1>" or "1<" for one sided tests of H0 when efficacy corresponds to a positive or negative drift, respectively. If `method`=="S" then this must be of the same length as `StatType` because the interpretation of `sided` is different depending upon whether `StatType`=="WLR" (negative is benefit) or `StatType`=="ISD" (positive is benefit)

method

    Determines how to calculate the power. Set to "A" (Asymptotic method, the default) or "S" (Simulation method)

accru

    The upper endpoint of the accrual period beginning with time 0.

accrat

    The rate of accrual per unit of time.

tlook

    The times of planned interim analyses.

tcut0

    Left hand endpoints for intervals upon which the arm-0 specific mortality is constant. The last given component is the left hand endpoint of the interval having right hand endpoint infinity.

h0

    A vector of the same length as `tcut0` which specifies the piecewise constant arm-0 mortality rate.

s0

    Alternatively, the arm-0 mortality distribution can be supplied via this argument, in terms of of the corresponding survival function values at the times given in the vector `tcut0`. If `s0` is supplied, then `h0`is derived internally, assuming the piecewise exponential distrubiton. If you specify `s0`, the first element must be 1, and correspondingly, the first component of `tcut0` will be the lower support point of the distribution. You must supply either `h0` or `s0` but not both.

tcut1

    Left hand endpoints for intervals upon which the arm-1 specific mortality is constant. The last given component is the left hand endpoint of the interval having right hand endpoint infinity.

rhaz

    A vector of piecewise constant arm-1 versus arm-0 mortality rate ratios. If `tcut1` and `tcut0` are not identical, then `tcut1`, `h0`, and `rhaz` are internally rederived at the union of the sequences `tcut0` and `tcut1`. In all cases the arm-1 mortality rate is then derived at the time cutpoints `tcut1` as `rhaz` times`h0`.

h1

    Alternatively, the arm-1 mortality distribution can be supplied via this argument by specifying the piecewise constant arm-1 mortality rate. See the comments above.

s1	Alternatively, the arm-1 mortality distribution can be supplied via this argument, in terms of of the corresponding survival function values at the times given in the vector `tcut1`. Comments regarding `s0` above apply here as well. You must supply exactly one of the following: `h1`, `rhaz`, or `s1`.
tcutc0	Left hand endpoints for intervals upon which the arm-0 specific censoring distribution hazard function is constant. The last given component is the left hand endpoint of the interval having right hand endpoint infinity.
hc0	A vector of the same length as `tcutc0` which specifies the arm-0 censoring distribution in terms of a piecewise constant hazard function.
sc0	Alternatively, the arm-0 censoring distribution can be supplied via this argument, in terms of of the corresponding survival function values at the times given in the vector `tcutc0`. See comments above. You must supply either `hc0` or `sc0` but not both.
tcutc1	Left hand endpoints for intervals upon which the arm-1 specific censoring distribution hazard function is constant. The last given component is the left hand endpoint of the interval having right hand endpoint infinity.
hc1	A vector of the same length as `tcutc1` which specifies the arm-1 censoring distribution in terms of a piecewise constant hazard function.
sc1	Alternatively, the arm-1 censoring distribution can be supplied via this argument, in terms of of the corresponding survival function values at the times given in the vector `tcutc1`. See comments above. You must supply either `hc1` or `sc1` but not both.
noncompliance	
	(i) Seting `noncompliance` to "none" for no non-compliance will automatically set the non-compliance arguments, below, to appropriate values for no compliance. This requires no additional user specification of non-compliance parameters. (ii) Setting `noncompliance` to "crossover" will automatically set crossover values in the arm 0/1 specific *post-cause-B-delay-mortality* for cross-over, i.e. simple interchange of the arm 0 and arm 1 mortalities. The user is required to specify all parameters corresponding to the arm 0/1 specific *cause-B-delay* distributions. The *cause-A-delay* and *post-cause-A-delay-mortality* are automatically set so as not to influence the calculations. Setting `noncompliance` to "mixed" will set the arm 0/1 specific *post-cause-B-delay-mortality* distributions for crossover as defined above. The user specifies the arm 0/1 specific *cause-B-delay* distribution as above, and in addition, all parameters related to the arm 0/1 specific *cause-A-delay* distributions and corresponding arm 0/1 specific *post-cause-A-delay-mortality* distributions. (iii) Setting `noncompliance` to "user" requires the user to specify all non-compliance distribution parameters.
tcutd0A	Left hand endpoints for intervals upon which the arm-0 specific *cause-A delay* distribution hazard function is constant. The last given component is the left hand endpoint of the interval having right hand endpoint infinity. Required only when `noncompliance` is set to "mixed" or "user".
hd0A	A vector of the same length as `tcutd0A` containing peicewise constant hazard rates for the arm-0 *cause-A delay* distribution. Required only when `noncompliance` is set to "mixed" or "user".
sd0A	When required, the arm-0 *cause-A-delay* distribution is alternately specified via a survival function. A vector of the same length as `tcutd0A`.
tcutd0B	Left hand endpoints for intervals upon which the arm-0 specific *cause-B delay* distribution hazard function is constant. The last given component is the

	left hand endpoint of the interval having right hand endpoint infinity. Always required when `noncompliance` is set to any value other than "none".
hd0B	A vector of the same length as `tcutd0B` containing peicewise constant hazard rates for the arm-0 *cause-B delay* distribution. Always required when `noncompliance` is set to any value other than "none".
sd0B	When required, the arm-0 *cause-B-delay* distribution is alternately specified via a survival function. A vector of the same length as `tcutd0B`.
tcutd1A	Left hand endpoints for intervals upon which the arm-1 specific *cause-A delay* distribution hazard function is constant. The last given component is the left hand endpoint of the interval having right hand endpoint infinity. Required only when `noncompliance` is set to "mixed" or "user".
hd1A	A vector of the same length as `tcutd1A` containing peicewise constant hazard rates for the arm-1 *cause-A delay* distribution. Required only when `noncompliance` is set to "mixed" or "user".
sd1A	When required, the arm-1 *cause-A-delay* distribution is alternately specified via a survival function. A vector of the same length as `tcutd1A`.
tcutd1B	Left hand endpoints for intervals upon which the arm-1 specific *cause-B delay* distribution hazard function is constant. The last given component is the left hand endpoint of the interval having right hand endpoint infinity. Always required when `noncompliance` is set to any value other than "none".
hd1B	A vector of the same length as `tcutd1B` containing peicewise constant hazard rates for the arm-1 *cause-B delay* distribution. Always required when `noncompliance` is set to any value other than "none".
sd1B	When required, the arm-1 *cause-A-delay* distribution is alternately specified via a survival function. A vector of the same length as `tcutd1A`.
tcutx0A	Left hand endpoints for intervals upon which the arm-0 specific *post-cause-A-delay-mortality* rate is constant. The last given component is the left hand endpoint of the interval having right hand endpoint infinity. Required only when `noncompliance` is set to "mixed" or "user".
hx0A	A vector of the same length as `tcutx0A` containing the arm-0 *post-cause-A-delay mortality* rates. Required only when `noncompliance` is set to "mixed" or "user".
sx0A	When required, the arm-0 *post-cause-A-delay mortality* distribution is alternately specified via a survival function. A vector of the same length as `tcutx0A`.
tcutx0B	Left hand endpoints for intervals upon which the arm-0 specific *post-cause-B-delay-mortality* rate is constant. The last given component is the left hand endpoint of the interval having right hand endpoint infinity. Always required when `noncompliance` is set to any value other than "none".
hx0B	A vector of the same length as `tcutx0B` containing the arm-0 *post-cause-B-delay mortality* rates. Always required when `noncompliance` is set to any value other than "none".
sx0B	When required, the arm-0 *post-cause-B-delay mortality* distribution is alternately specified via a survival function. A vector of the same length as `tcutx0B`.
tcutx1A	Left hand endpoints for intervals upon which the arm-1 specific *post-cause-A-delay-mortality* rate is constant. The last given component is the left hand endpoint of the interval having right hand endpoint infinity. Required only when `noncompliance` is set to "mixed" or "user".

hx1A   A vector of the same length as `tcutx1A` containing the arm-1 *post-cause-A-delay mortality* rates. Required only when `noncompliance` is set to "mixed" or "user".

sx1A   When required, the arm-1 *post-cause-A-delay mortality* distribution is alternately specified via a survival function. A vector of the same length as `tcutx1A`.

tcutx1B   Left hand endpoints for intervals upon which the arm-1 specific *post-cause-B-delay-mortality* rate is constant. The last given component is the left hand endpoint of the interval having right hand endpoint infinity. Always required when `noncompliance` is set to any value other than "none".

hx1B   A vector of the same length as `tcutx1B` containing the arm-1 *post-cause-B-delay mortality* rates. Always required when `noncompliance` is set to any value other than "none".

sx1B   When required, the arm-1 *post-cause-B-delay mortality* distribution is alternately specified via a survival function. A vector of the same length as `tcutx1B`.

gradual   Should the conversion to post-noncompliance mortality be gradual. Under the default behavior, `gradual=FALSE`, there is an immediate conversion to the post-noncompliance mortality rate function. If `gradual` is set to `TRUE` then this conversion is done "gradually". In truth, at the individual level what is done is that the new mortality rate function is a convex combination of the pre-noncompliance mortality and the post-noncompliance mortality, with the weighting in proportion to the time spent in compliance with the study arm protocal.

WtFun   Specifies the name of a weighting function (of time) for assigning relative weights to events according to the times at which they occur. The default, "FH", a two parameter weight function, specifies the 'Fleming-Harrington' `g-rho` family of weighting functions defined as the pooled arm survival function (Kaplan-Meier estimate) raised to the `g` times its complement raised to the `rho`. Note that `g=rho=0` corresponds to the unweighted log-rank statistic. A second choice is the "SFH" function, (for 'Stopped Fleming-Harrington'), meaning that the "FH" weights are capped at their value at a user specified time, which has a total of 3 parameters. A third choice is `Ramp(tcut)`. Under this choice, weights are assigned in a linearly manner from time 0 until a user specified cut-off time, `tcut`, after which events are weighted equally. It is possible to conduct computations on `nstat` candidate statistics within a single run. In this case, `WtFun` should be a character vector of length `nstat` having components set from among the available choices.

ppar   A vector containing all the weight function parameters, in the order determined by that of "WtFun". For example, if `WtFun` is set to `c("FH","SFH","Ramp")` then `ppar` should be a vector of length six, with the "FH" parameters in the first two elements, "SFH" parameters in the next 3 elements, and "Ramp" parameter in the last element.

RR.Futility   The relative risk corresponding to the alternative alternative hypothesis that is required in the construction of the futility boundary. Required if `Boundary.Futility` is set to a non-null value.

Spend.Info   When the test statistic is something other than the unweighted log-rank statistic, the variance information, i.e. the ratio of variance at interim analysis to variance at the end of trial, is something other than the ratio of events at interim analysis to the events at trial end. The problem is that in practice one doesn't necessarily have a good idea what the end of trial variance should be. In this

case the user may wish to spend the type I and type II error probabilities according to a different time scale. Possible choices are "Variance", (default), which just uses the variance ratio scale, "Events", which uses the events ratio scale, "Hybrid(k)", which makes a linear transition from the "Variance" scale to the "Events" scale beginning with analysis number `k`. The last choice, "Calendar", uses the calendar time scale

qProp.one.or.Q

If a futility boundary is specified, what assumption should be made about the drift function (the mean value of the weighted log-rank statistic at analysis `j` normalized by the square root of the variance function at analysis `k`). In practice we don't presume to know the shape of the drift function. Set to "one" or "Q". The choice "one" results in a more conservative boundary.

Nsim   If you specify `method=="S"`, then you must specify the number of simulations. 1000 should be sufficient.

detail   If you specify `method=="S"`, and want to see the full level of detail regarding arguments returned from the C level code, specify `detail==TRUE`

StatType   If you specify `method=="S"`, then the available choices are "WLR" (weighted log-rank) and "ISD" (integrated survival difference).

Value

Returns a value of class `PwrGSD` which has components listed below. Note that the print method will display a summary table of estimated powers and type I errors as a `nstat` by 2 matrix. The summary method returns the same object invisibly, but after computing the summary table mentioned above, and it is included in the returned value as a commponent `TBL`. See examples below.

dPower   A `length(tlook)` by `nstat` matrix containing in each column, an increment in power that resulted at that analysis time for the given statistic.

dErrorI   A `length(tlook)` by `nstat` matrix containing in each column, an increment in type I error that resulted at that analysis time for the given statistic. Always sums to the total alpha specified in `alphatot`

detail   A list with components equal to the arguments of the C-call, which correspond in a natural way to the arguments specified in the R call, along with the computed results in `palpha0vec`, `palpha1vec`, `inffrac`, and `mu`. The first two are identical to `dErrorI` and `dPower`, explained above. The last two are `length(tlook)` by `nstat` matrices. For each statistic specified in `par`, the corresponding columns of `pinffrac` and `mu` contain the information fraction and drift at each of the analysis times.

call   the call

Author(s)

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References

Gu, M.-G. and Lai, T.-L. "Determination of power and sample size in the design of clinical trials with failure-time endpoints and interim analyses." Controlled Clinical Trials 20 (5): 423-438. 1999

Izmirlian, G. "The PwrGSD package." NCI Div. of Cancer Prevention Technical Report. 2004

Jennison, C. and Turnbull, B.W. (1999) Group Sequential Methods: Applications to Clinical Trials Chapman & Hall/Crc, Boca Raton FL

Proschan, M.A., Lan, K.K.G., Wittes, J.T. (2006), corr 2nd printing (2008) Statistical Monitoring of Clinical Trials A Unified Approach Springer Verlag, New York

See Also

[cpd.PwrGSD](#)

Examples

```
library(PwrGSD)

test.example <-
  PwrGSD(EfficacyBoundary = LanDemets(alpha = 0.05, spending = ObrienFleming),
         FutilityBoundary = LanDemets(alpha = 0.1, spending = ObrienFleming),
 RR.Futility = 0.82, sided="1<",method="A",accru =7.73, accrat =9818.65,
         tlook =c(7.14, 8.14, 9.14, 10.14, 10.64, 11.15, 12.14, 13.14,
                  14.14, 15.14, 16.14, 17.14, 18.14, 19.14, 20.14),
         tcut0 =0:19, h0 =c(rep(3.73e-04, 2), rep(7.45e-04, 3),
                            rep(1.49e-03, 15)),
         tcut1 =0:19, rhaz =c(1, 0.9125, 0.8688, 0.7814, 0.6941,
                              0.6943, 0.6072, 0.5202, 0.4332, 0.6520,
                              0.6524, 0.6527, 0.6530, 0.6534, 0.6537,
                              0.6541, 0.6544, 0.6547, 0.6551, 0.6554),
         tcutc0 =0:19, hc0 =c(rep(1.05e-02, 2), rep(2.09e-02, 3),
                              rep(4.19e-02, 15)),
         tcutc1 =0:19, hc1 =c(rep(1.05e-02, 2), rep(2.09e-02, 3),
                              rep(4.19e-02, 15)),
         tcutd0B =c(0, 13), hd0B =c(0.04777, 0),
         tcutd1B =0:6, hd1B =c(0.1109, 0.1381, 0.1485, 0.1637, 0.2446,
                               0.2497, 0),
         noncompliance =crossover, gradual =TRUE,
         WtFun =c("FH", "SFH", "Ramp"),
         ppar =c(0, 1, 0, 1, 10, 10))
```

| LanDemets | *The Lan-Demets method of Boundary Construction* |

Description

The function `LanDemets` is used in calls to the functions `GrpSeqBnds` and `PwrGSD` as a possible setting for the arguments `EfficacyBoundary` and `FutilityBoundary`, in specification of the method whereby efficacy and or futility boundaries are to be constructed. The Lan-Demets method is one of four currently availiable choices, the others being `SC` (stochastic curtailment), `Haybittle` and user specified.

Usage

```
LanDemets(alpha, spending, from = NULL, to = NULL)
```

Arguments

alpha	If `LanDemets` is used to specify the `EfficacyBoundary` then the argument `alpha` is the total probability of type I error. If `LanDemets` is used to specify the `FutilityBoundary` then the argument `alpha` is the total probability of type II error.
spending	Specify the alpha spending function. Set this to `ObrienFleming, Pow(rho=<x>)`, or `Pocock`. See help files for these spending functions.
from	WARNING EXPERIMENTAL: you can actually construct boundaries via a hybrid of the 3 boundary construction methods, `LanDemets`, `SC`, and 'user specified'. When using a hybrid boundry, set the argument `EfficacyBoundary` or `FutilityBoundary` respectively, to a list with components `LanDemets`, `SC`, or user specified numbers. In the former two cases, `from` and `to` are used in `LanDemets` and also in `SC` to stipulate how many interim analyses they are in effect. See the help for `GrpSeqBnds` and `PwrGSD`
to	See above.

Details

The cornerstone of the Lan-Demets method is that the amount of alpha (type I or II error probability) that is "spent" at a given interim analysis is determined via a user specified "spending function". A spending function is a monotone increasing mapping on (0,1) with range (0,alpha). The 'alpha' spent at a given analysis is determined by the increment in the values of the spending function at the current and at the most recent information fractions.

Value

An object of class `boundary.construction.method` which is really a list with the following components. The print method displays the original call.

type	Gives the boundary construction method type, which is the character string "LanDemets"
alpha	The numeric value passed to the argument 'alpha' which is the total probability of type I (efficacy) or type II (futility) error.
spending	The spending function that was passed to the argument 'spending'. Note that this will be of class 'name' for 'ObrienFleming' and 'Pocock', but will be of class 'function' for 'Pow'
from	The numeric value passed to the argument 'from'. See above.
to	The numeric value passed to the argument 'to'. See above.
call	returns the call

Note

The print method returns the call by default

Author(s)

Grant Izmirlian

References

see references under PwrGSD

See Also

SC, ObrienFleming, Pow, Pocock, GrpSeqBnds, and PwrGSD

Examples

```
## example 1: what is the result of calling a Boundary Construction Method function
    ## A call to 'LanDemets' just returns the call
    LanDemets(alpha=0.05, spending=ObrienFleming)

    ## It does arguement checking...this results in an error
    ## Not run:
      LanDemets(alpha=0.05)

## End(Not run)

    ## but really its value is a list with the a component containing
    ## the boundary method type, "LanDemts", and components for each
    ## of the arguments.
    names(LanDemets(alpha=0.05, spending=ObrienFleming))

    LanDemets(alpha=0.05, spending=ObrienFleming)$type
    LanDemets(alpha=0.05, spending=ObrienFleming)$alpha
    LanDemets(alpha=0.05, spending=ObrienFleming)$spending
    class(LanDemets(alpha=0.05, spending=ObrienFleming)$spending)
    LanDemets(alpha=0.05, spending=Pow(2))$spending
    class(LanDemets(alpha=0.05, spending=Pow(2))$spending)
    LanDemets(alpha=0.05, spending=ObrienFleming)$call

## example 2: ...But the intended purpose of the spending functions its
    ## intended purpose is in constructing calls to 'GrpSeqBnds' and to 'PwrGSD':


    frac <- c(0.07614902,0.1135391,0.168252,0.2336901,0.3186155,
              0.4164776,0.5352199,0.670739,0.8246061,1)
    drift <- c(0.3836636,0.5117394,0.6918584,0.8657705,1.091984,
               1.311094,1.538582,1.818346,2.081775,2.345386)

    test <- GrpSeqBnds(frac=frac, EfficacyBoundary=LanDemets(alpha=0.05, spending=ObrienF
                       FutilityBoundary=LanDemets(alpha=0.10, spending=Pocock),
                       drift=drift)
```

ObrienFleming   *The O'Brien-Fleming Alpha Spending Function*

Description

Stipulates alpha spending according to the O'Brien-Fleming spending function in the Lan-Demets boundary construction method. Its intended purpose is in constructing calls to GrpSeqBnds and PwrGSD.

Usage

```
ObrienFleming()
```

Value

An object of class `spending.function` which is really a list with the following components. The print method displays the original call.

type   Gives the spending function type, which is the character string "ObrienFleming"

call   returns the call

Note

The print method returns the call by default

Author(s)

Grant Izmirlian

References

see references under PwrGSD

See Also

LanDemets, Pow, Pocock, GrpSeqBnds, PwrGSD

Examples

```
## example 1: what is the result of calling a spending function
    ## A call to 'ObrienFleming' just returns the call
    ObrienFleming()

    ## but really its value is a list with a component named
    ## 'type' equal to "ObrienFleming" and a component named
    ## 'call' equal to the call.
    names(ObrienFleming)

    ObrienFleming()$type

    ObrienFleming()$call

## example 2: ...But the intended purpose of the spending functions its
    ## intended purpose is in constructing calls to 'GrpSeqBnds' and to 'PwrGSD':


    frac <- c(0.07614902,0.1135391,0.168252,0.2336901,0.3186155,
              0.4164776,0.5352199,0.670739,0.8246061,1)
    drift <- c(0.3836636,0.5117394,0.6918584,0.8657705,1.091984,
               1.311094,1.538582,1.818346,2.081775,2.345386)

    test <- GrpSeqBnds(frac=frac, EfficacyBoundary=LanDemets(alpha=0.05, spending=ObrienF
                       FutilityBoundary=LanDemets(alpha=0.10, spending=Pocock),
                       drift=drift)
```

Pow *The Wang-Tsiatis Power Alpha Spending Function*

Description

Stipulates alpha spending according to the Wang-Tsiatis Power function in the Lan-Demets boundary construction method. Its intended purpose is in constructing calls to `GrpSeqBnds` and `PwrGSD`.

Usage

```
Pow(rho)
```

Arguments

rho   The exponent for the Wang-Tsiatis power spending function

Details

Larger `rho` results in more conservative boundaries. `rho=3` is roughly equivalent to Obrien-Fleming spending. `rho=1` spends `alpha` linearly in the information fraction

Value

An object of class `spending.function` which is really a list with the following components. The print method displays the original call.

type   Gives the spending function type, which is the character string "Pow"

rho   the numeric value passed to the single argument, `rho`

call   returns the call

Note

The print method returns the call by default

Author(s)

Grant Izmirlian

References

see references under PwrGSD

See Also

LanDemets, ObrienFleming, Pocock, GrpSeqBnds, PwrGSD

Examples

```
## example 1: what is the result of calling a spending function
    ## A call to 'Pow' just returns the call
    Pow(rho=2)

    ## It does argument checking...the following results in an error:
    ## Not run:
      Pow()

## End(Not run)

    ## it doesn't matter whether the argument is named or not,
    ## either produces the same result
    Pow(2)

    ## but really its value is a list with a component named
    ## 'type' equal to "Pow", a component named 'rho' equal
    ## to the numeric value passed to the single argument 'rho'
    ## and a component  named 'call' equal to the call.
    names(Pow(rho=2))

    names(Pow(2))

    Pow(rho=2)$type
    Pow(rho=2)$rho
    Pow(rho=2)$call

## example 2: ...But the intended purpose of the spending functions its
    ## intended purpose is in constructing calls to 'GrpSeqBnds' and to 'PwrGSD':


    frac <- c(0.07614902,0.1135391,0.168252,0.2336901,0.3186155,
              0.4164776,0.5352199,0.670739,0.8246061,1)
    drift <- c(0.3836636,0.5117394,0.6918584,0.8657705,1.091984,
               1.311094,1.538582,1.818346,2.081775,2.345386)

    test <- GrpSeqBnds(frac=frac, EfficacyBoundary=LanDemets(alpha=0.05, spending=Pow(2)),
                       FutilityBoundary=LanDemets(alpha=0.10, spending=ObrienFleming),
                       drift=drift)
```

Pocock	*The Pocock Alpha Spending Function*

Description

Stipulates alpha spending according to the Pocock spending function in the Lan-Demets boundary construction method. Its intended purpose is in constructing calls to `GrpSeqBnds` and `PwrGSD`.

Usage

```
Pocock()
```

Value

An object of class `spending.function`

`type`	Gives the spending function type, which is the character string "Pocock"
`call`	returns the call

Note

The print method returns the call by default

Author(s)

Grant Izmirlian

References

see references under `PwrGSD`

See Also

`LanDemets`, `ObrienFleming`, `Pow`, `GrpSeqBnds`, `PwrGSD`

Examples

```
## example 1: what is the result of calling a spending function

    ## A call to 'Pocock' just returns the call
    Pocock()

    ## but really its value is a list with a component named
    ## 'type' equal to "Pocock" and a component named
    ## 'call' equal to the call.
    names(Pocock)

    Pocock()$type

    Pocock()$call

## example 2: ...But the intended purpose of the spending functions its
    ## intended purpose is in constructing calls to 'GrpSeqBnds' and to 'PwrGSD':


    frac <- c(0.07614902,0.1135391,0.168252,0.2336901,0.3186155,
              0.4164776,0.5352199,0.670739,0.8246061,1)
    drift <- c(0.3836636,0.5117394,0.6918584,0.8657705,1.091984,
               1.311094,1.538582,1.818346,2.081775,2.345386)

    test <- GrpSeqBnds(frac=frac, EfficacyBoundary=LanDemets(alpha=0.05, spending=Pocock)
                       FutilityBoundary=LanDemets(alpha=0.10, spending=ObrienFleming),
                       drift=drift)
```

SC   *The Stochastic Curtailment method of Boundary Construction*

Description

The function `SC` is used in calls to the functions `GrpSeqBnds` and `PwrGSD` as a possible setting for the arguments `EfficacyBoundary` and `FutilityBoundary`, in specification of the method whereby efficacy and or futility boundaries are to be constructed. The Stochastic Curtailment method is one of four currently availiable choices, the others being `LanDemets`, `Haybittle` and user specified.

Usage

```
SC(be.end, prob, drift.end = NULL, from = NULL, to = NULL)
```

Arguments

`be.end`	The value of the efficacy criterion in the scale of a standardized normal. This should be set to something further from the null than the single test Z_{alpha}. For example if the total type I error probability is 0.05 in a two sided test of the null than set be.end to 2.10 or larger (instead of 1.96).
`prob`	The criterion, a probability to be exceeded in order to stop. 0.90 or above is a good choice. See detail below.
`drift.end`	Required only if you are using `SC` to set the `FutilityBoundary`. In this case, set `drift.end` to the value of the drift function anticipated at the end of the trial. See detail below.
`from`	WARNING EXPERIMENTAL: you can actually construct boundaries via a hybrid of the 3 boundary construction methods, `LanDemets`, `SC`, and 'user specified'. When using a hybrid boundry, set the argument `EfficacyBoundary` or `FutilityBoundary` respectively, to a list with components `LanDemets`, `SC`, or user specified numbers. In the former two cases, `from` and `to` are used in `LanDemets` and also in `SC` to stipulate how many interim analyses they are in effect. See the help for `GrpSeqBnds` and `PwrGSD`
`to`	See above.

Details

When the stochastic curtailment procedure is used to construct the efficacy boundary, i.e. `EfficacyBoundary=SC(...`, the efficacy criterion is reached when the conditional probability, under the null hypothesis, that the last analysis results in statistical significance, given the present value of the statistic, exceeds 'prob'. In of itself, this doesn't produce a boundary on the scale of a standard normal, but it is easily converted to one as is done here. When this is used to construct a futility boundary, i.e. `FutilityBoundary=SC(...)`, the futility criterion is reached when the conditional probability, under the design alternative hypothesis, that the last analysis does not result in statistical significance, given the present value of the statistic, exceeds 'prob'. The design alternative corresponds to a drift function, which is the expected value of the statistic normalized to have variance equal to the information fraction at each interim analysis. For the unweighted log-rank statistic, the drift function is $(V_T)^{(1/2)} B f$, where B is the logged relative risk, V_T is the variance at the end of the trial and f is the information fraction. If the two trial arms are balanced and the number at risk is roughly constant throughout the trial then $V_T = pi (1-pi) N_T$, where pi is the constant proportion at risk in one of the trial arms and N_T is the anticipated number of events.

Value

An object of class `boundary.construction.method` which is really a list with the following components. The print method displays the original call.

`type`	Gives the boundary construction method type, which is the character string "SC"
`be.end`	The numeric value passed to the argument 'be.end', which is the value of the efficacy criterion in the scale of a standardized normal.
`prob`	The numeric value passed to the argument 'prob', which is the probability to be exceeded in order to stop.
`drift.end`	The numeric value passed to the argument 'drift.end', which is the value of the drift function at the end of the trial. See details.
`from`	The numeric value passed to the argument 'from'. See above.
`to`	The numeric value passed to the argument 'to'. See above.
`call`	returns the call

Note

The print method returns the call by default

Author(s)

Grant Izmirlian

References

see references under PwrGSD

See Also

LanDemets, GrpSeqBnds, PwrGSD

Examples

```
## example 1: what is the result of calling a Boundary Construction Method function
    ## A call to 'SC' just returns the call
    SC(be.end=2.10, prob=0.90)

    ## It does arguement checking...this results in an error
    ## Not run:
      SC(be.end=2.10)

## End(Not run)

    ## but really its value is a list with the a component containing
    ## the boundary method type, "LanDemts", and components for each
    ## of the arguments.
    names(SC(be.end=2.10, prob=0.90))

    SC(be.end=2.10, prob=0.90, drift.end=2.34)$type
    SC(be.end=2.10, prob=0.90, drift.end=2.34)$be.end
    SC(be.end=2.10, prob=0.90, drift.end=2.34)$prob
    SC(be.end=2.10, prob=0.90, drift.end=2.34)$drift.end
```

```
## example 2: ...But the intended purpose of the spending functions its
    ## intended purpose is in constructing calls to 'GrpSeqBnds' and to 'PwrGSD':


    frac <- c(0.07614902,0.1135391,0.168252,0.2336901,0.3186155,
              0.4164776,0.5352199,0.670739,0.8246061,1)
    drift <- c(0.3836636,0.5117394,0.6918584,0.8657705,1.091984,
       1.311094,1.538582,1.818346,2.081775,2.345386)


    test <- GrpSeqBnds(frac=frac, EfficacyBoundary=LanDemets(alpha=0.05, spending=ObrienF
                       FutilityBoundary=SC(be.end=2.10, prob=0.90, drift.end=drift[10]),
                       drift=drift)
```

Haybittle	*The Haybittle method of Boundary Construction*

Description

The function `Haybittle` is used in calls to the functions `GrpSeqBnds` and `PwrGSD` as a possible setting for the argument `EfficacyBoundary`. NOTE: the Haybittle method is not used for construction a futility boundary (does this even make sense?). The Haybittle method is one of four currently availiable choices, the others being `LanDemets`, `SC` (stochastic curtailment), and user specified.

Usage

```
Haybittle(alpha, b.Haybittle, from = NULL, to = NULL)
```

Arguments

`alpha`	The total probability of type I error.
`b.Haybittle`	User specified efficacy boundary at all but the last analysis.
`from`	WARNING EXPERIMENTAL: See the documentation under `LanDemets` or `SC`. I'm not quite sure if this works or even makes sense. Don't use it, ok?
`to`	See above.

Details

The Haybittle method is the simplest of all methods for efficacy boundary construction. You just specify a value such as 3 which is so large that multiple testing with standard normal criterion 3 (upper quantile is 0.00135) means that after 20 analyses you've spent only 0.027. The value of the criterion for the final analysis is determined so that the total type I error is obtained.

Value

An object of class `boundary.construction.method` which is really a list with the following components. The print method displays the original call.

`type`	Gives the boundary construction method type, which is the character string "Haybittle"
`alpha`	The numeric value passed to the argument 'alpha' which is the total probability of type I error.

`b.Haybittle`	The numeric value passed to the argument 'b.Haybittle' which is the user specified efficacy boundary at all but the last analysis.
`from`	Description of 'comp2'
`to`	You're not using this, right?
`call`	see above.

Note

The print method returns the call by default

Author(s)

Grant Izmirlian

References

see references under PwrGSD

See Also

LanDemets, SC, GrpSeqBnds, and PwrGSD

Examples

```
## example 1: what is the result of calling a Boundary Construction Method function
    ## A call to 'Haybittle' just returns the call
    Haybittle(alpha=0.05, b.Haybittle=3)

    ## It does arguement checking...this results in an error
    ## Not run:
      Haybittle(alpha=0.05)

## End(Not run)

    ## but really its value is a list with the a component containing
    ## the boundary method type, "LanDemts", and components for each
    ## of the arguments.
    names(Haybittle(alpha=0.05, b.Haybittle=3))

    Haybittle(alpha=0.05, b.Haybittle=3)$type
    Haybittle(alpha=0.05, b.Haybittle=3)$alpha
    Haybittle(alpha=0.05, b.Haybittle=3)$b.Haybittle
    Haybittle(alpha=0.05, b.Haybittle=3)$call

## example 2: ...But the intended purpose of the spending functions its
    ## intended purpose is in constructing calls to 'GrpSeqBnds' and to 'PwrGSD':


    frac <- c(0.07614902,0.1135391,0.168252,0.2336901,0.3186155,
              0.4164776,0.5352199,0.670739,0.8246061,1)


    test <- GrpSeqBnds(frac=frac, EfficacyBoundary=Haybittle(alpha=0.025, b.Haybittle=3))
```

---

cpd.PwrGSD *Create a skeleton compound PwrGSD object*

Description

Given a user defined indexing dataframe as its only argument, creates a skeleton compound PwrGSD object having a component `Elements`, a list of `PwrGSD` objects, of length equal to the number of rows in the indexing dataframe

Usage

```
cpd.PwrGSD(descr)
```

Arguments

descr   A dataframe of a number of rows equal to the length of the resulting list, `Elements`, of `PwrGSD` objects. The user defines the mapping between rows of `descr` and components of `Elements` and uses it to set up a loop over scenarios. There are several S3 classes and methods for example `plot.cpd.PwrGSD`, which exploit this mapping between characteristics of a run and the rows of `desr` for subsetting and constructing conditioned plots. See the example below.

Value

An object of class `cpd.PwrGSD` containing elements:

date   the POSIX date that the object was created–its quite useful

Elements   a list of length equal to the number of rows of `descr` which will later contain objects of class `PwrGSD`

descr   a copy of the indexing dataframe argument for use in navigating the compound object in subsequent calls to other functions such as the related `plot` method, and the subset extractor, `Elements`

Note

A `cpd.PwrGSD` object essentially a list of `PwrGSD` objects that a user may set up in order to investigate the space of possible trial scenarios, test statistics, and boundary construction options. One could store a list of results without appealing at all to these internal indexing capabilities. The advantage of setting up a `cpd.PwrGSD` object is the nice summarization functionality provided, for example the plot method for the `cpd.PwrGSD` class.

The key ingredient to (i) the construction of the empty object, (ii) and summarizing the results in tabular or plotted form via its manipulation in subsequent function calls, is the indexing dataset, `descr` (for description). The correspondence between rows of `descr` and elements in the list of `PwrGSD` objects is purposely left very loose. In the example outlined below, the user creates a "base case" call to `PwrGSD` and then decides which quantities in this "base case" call to vary in order to navigate the space of possible trial scenarios, monitoring statistics and boundary construction methods. Next, for each one of these settings being varied, a variable with levels that determine each possible setting is created. The dataset `descr` is created with one line corresponding to each combination of the selection variables so created. In order to ensure that there is 1-1 correspondence between the order of the rows in `descr` and the order in the list `Elements` of `PwrGSD` objects, the user carries out the computation in a loop over rows of `descr` in which the values of the

selection variables in each given row of `descr` are used to create the corresponding component of `Elements` via an update the "base case" call.

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

See Also

`Elements`, `plot.cpd.PwrGSD` and `Power`

Examples

```
## don't worry--these examples are guaranteed to work,
## its just inconvenient for the package checker
## Not run:
  library(PwrGSD)

## In order to set up a compound object of class `cpd.PwrGSD'
## we first construct a base case: a two arm trial randomized in just
## under eight years with a maximum of 20 years of follow-up.
## We compute power at a specific alternative, `rhaz', under
## an interim analysis plan with roughly one annual analysis, some
## crossover between intervention and control arms, with Efficacy
## and futility boundaries constructed via the Lan-Demets procedure
## with O'Brien-Fleming spending on the hybrid scale. Investigate
## the behavior of three weighted log-rank statistics.

test.example <-
  PwrGSD(EfficacyBoundary = LanDemets(alpha = 0.05, spending = ObrienFleming),
         FutilityBoundary = LanDemets(alpha = 0.1, spending = ObrienFleming),
 RR.Futility = 0.82, sided="1<",method="A",accru =7.73, accrat =9818.65,
         tlook =c(7.14, 8.14, 9.14, 10.14, 10.64, 11.15, 12.14, 13.14,
                  14.14, 15.14, 16.14, 17.14, 18.14, 19.14, 20.14),
         tcut0 =0:19, h0 =c(rep(3.73e-04, 2), rep(7.45e-04, 3),
                           rep(1.49e-03, 15)),
         tcut1 =0:19, rhaz =c(1, 0.9125, 0.8688, 0.7814, 0.6941,
                              0.6943, 0.6072, 0.5202, 0.4332, 0.6520,
                              0.6524, 0.6527, 0.6530, 0.6534, 0.6537,
                              0.6541, 0.6544, 0.6547, 0.6551, 0.6554),
         tcutc0 =0:19, hc0 =c(rep(1.05e-02, 2), rep(2.09e-02, 3),
                             rep(4.19e-02, 15)),
         tcutc1 =0:19, hc1 =c(rep(1.05e-02, 2), rep(2.09e-02, 3),
                             rep(4.19e-02, 15)),
         tcutd0B =c(0, 13), hd0B =c(0.04777, 0),
         tcutd1B =0:6, hd1B =c(0.1109, 0.1381, 0.1485, 0.1637, 0.2446,
                              0.2497, 0),
         noncompliance =crossover, gradual =TRUE,
         WtFun =c("FH", "SFH", "Ramp"),
         ppar =c(0, 1, 0, 1, 10, 10))

## we will construct a variety of alternate hypotheses relative to the
## base case specified above

  rhaz <-
    c(1, 0.9125, 0.8688, 0.7814, 0.6941, 0.6943, 0.6072, 0.5202, 0.4332,
    0.652, 0.6524, 0.6527, 0.653, 0.6534, 0.6537, 0.6541, 0.6544,
```

```
     0.6547, 0.6551, 0.6554)

  max.effect <- 0.80 + 0.05*(0:8)
  n.me <- length(max.effect)

## we will also vary extent of censoring relative to the base case
## specified above

  hc <- c(rep(0.0105, 2), rep(0.0209, 3), rep(0.0419, 15))

  cens.amt <- 0.75 + 0.25*(0:2)
  n.ca <- length(cens.amt)

## we may also wish to compare the Lan-Demets/O'Brien-Fleming efficacy
## boundary with a Lan-Demets/linear spending boundary

  Eff.bound.choice <- 1:2
  ebc.nms <- c("LanDemets(alpha=0.05, spending=ObrienFleming)",
               "LanDemets(alpha=0.05, spending=Pow(1))")
  n.ec <- length(Eff.bound.choice)

## The following line creates the indexing dataframe, `descr', with one
## line for each possible combination of the selection variables we've
## created.


  descr <- as.data.frame(
              cbind(Eff.bound.choice=rep(Eff.bound.choice, each=n.ca*n.me),
                    cens.amt=rep(rep(cens.amt, each=n.me), n.ec),
                    max.effect=rep(max.effect, n.ec*n.ca)))

  descr$Eff.bound.choice <- ebc.nms[descr$Eff.bound.choice]

## Now descr contains one row for each combination of the levels of
## the user defined selection variables, `Eff.bound.choice',
## `max.effect' and `cens.amt'. Keep in mind that the names and number
## of these variables is arbitrary. Next we create a skeleton
## `cpd.PwrGSD' object with a call to the function `cpd.PwrGSD' with
## argument `descr'

  test.example.set <- cpd.PwrGSD(descr)

## Now, the newly created object, of class `cpd.PwrGSD', contains
## an element `descr', a component `date', the date created
## and a component `Elements', an empty list of length equal
## to the number of rows in `descr'.  Next we do the computation in
## a loop over the rows of `descr'.

  n.descr <- nrow(descr)

  for(k in 1:n.descr){

    ## First, we copy the original call to the current call,
    ## `Elements[[k]]$call'

    test.example.set$Elements[[k]]$call <- test.example$call
```

```
## Use the efficacy boundary choice in the kth row of `descr'
## to set the efficacy boundary choice in the current call

test.example.set$Elements[[k]]$call$EfficacyBoundary <-
parse(text=as.character(descr[k,"Eff.bound.choice"]))[[1]]

## Derive the `rhaz' defined by the selection variable "max.effect"
## in the kth row of `descr' and use this to set the `rhaz'
## components of the current call

test.example.set$Elements[[k]]$call$rhaz <-
                          exp(descr[k,"max.effect"] * log(rhaz))

## Derive the censoring components from the selection variable
## "cens.amt" in the kth row of `descr' and place that result
## into the current call

test.example.set$Elements[[k]]$call$hc0 <-
test.example.set$Elements[[k]]$call$hc1 <- descr[k, "cens.amt"] * hc

## Now the current call corresponds exactly to the selection
## variable values in row `k' of `descr'. The computation is
## done by calling `update'

test.example.set$Elements[[k]] <- update(test.example.set$Elements[[k]])
cat(k/n.descr, "\r")
}

## We can create a new `cpd.PwrGSD' object by subsetting on
## the selection variables in `descr':

Elements(test.example.set,
         subset=(substring(Eff.bound.choice, 32, 34)=="Obr" &
                          max.effect >= 1))


## or we can plot the results -- see the help under `plot.cpd.PwrGSD'

plot(test.example.set, formula = ~ max.effect | stat * cens.amt,
     subset=(substring(Eff.bound.choice, 32, 34)=="Obr"))

plot(test.example.set, formula = ~ max.effect | stat * cens.amt,
     subset=(substring(Eff.bound.choice, 32, 34)=="Pow"))

## Notice the appearance of the selection variable `stat' which was
## not defined in the dataset `descr'.

## Recall that each single "PwrGSD" object can contain results
## for a list of test statistics, as in the example shown here where
## we have results on three statistics per component of `Elements'.
## For this reason the variable `stat' can be also be referenced in
## the `subset' or `formula' arguments of calls to this `plot' method,
## and in the `subset' argument of the function `Power' shown below.

## The function `Power' is used to convert the `cpd.PwrGSD' object
## into  a dataframe, stacked by rows of `descr' and by `stat'
## (there are three statistics being profiled per each component of
```

```
## `Elements'), for generating tables or performing other
## computations.

Power(test.example.set,
      subset=(substring(Eff.bound.choice, 32, 34)=="Pow" & stat %in% c(1,3)))


## End(Not run)
```

Power   *Extract the Power results*

Description

The function 'Power' is used to summarize the 'cpd.PwrGSD' object into a dataframe containing power and type II error, summed over analysis times. The data frame is stacked by rows of 'descr' and by 'stat' (if there are multiple statistics being profiled per each component of 'Elements'), for generating tables or performing other computations.

Usage

```
Power(object, subset, nlook.ind = NULL)
```

Arguments

object	an object of class cpd.PwrGSD
subset	you may extract a subset via a logical expression in the variables of the index dataframe, descr
nlook.ind	(optional) a vector containing a subset of the indices of analysis times over which the sum is formed. Use this for example if you want to know the probability of stopping by the kth analysis under an unfavorable alternative. Set nlook.ind to 1:k

Value

a dataframe, stacked by rows of 'descr' and then by choices of 'stat'

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

See Also

cpd.PwrGSD and PwrGSD

Examples

```
## See the `cpd.PwrGSD' example
```

---

as.boundaries   *Convert a "PwrGSD" object to a "boundaries" object*

Description

Convert a `PwrGSD` object to a `boundaries` object

Usage

```
as.boundaries(object, ...)
```

Arguments

object   an object of class `PwrGSD`

...   if `object` is of class `PwrGSD` and there are more than one statistic under in-
                vestigation, then you may specify an argument `stat`. The default value is 1,
                meaning the first one.

Value

an object of class `boundaries`. See the documentation for `GrpSeqBnds`

Author(s)

Grant Izmirlian <izmirlian@nih.gov

See Also

`GrpSeqBnds`

Examples

```
## none as yet
```

Elements   *Create a subset of a "cpd.PwrGSD" object*

Description

Create a subset of a `cpd.PwrGSD` object

Usage

```
Elements(object, subset, na.action = na.pass)
```

Arguments

object   an object of class `cpd.PwrGSD`

subset   you may extract a subset via a logical expression in the variables of the index
                dataframe, `descr`

na.action   a method for handling `NA` values in the variables in the subset expression.

Value

an object of class `cpd.PwrGSD`. See help on that topic for details.

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

See Also

`cpd.PwrGSD` and `PwrGSD`

Examples

```
## See the `cpd.PwrGSD' example
```

`plot.cpd.PwrGSD`   *Plot Method for cpd.PwrGSD objects*

Description

Creates a trellis plot of type II error probability and power at each interim analysis, stacked, versus an effect size variable, conditioned upon levels of up to two factors.

Usage

```
## S3 method for class 'cpd.PwrGSD':
plot(x, formula, subset, na.action,...)
```

Arguments

x	an object of class `cpd.PwrGSD`
formula	a one sided formula of the form ~ `effect	f1` or ~ `effect	f1 *` `f2` where `effect`, `f1`, and `f2` are variables in the indexing dataframe `descr`, or the special variable `stat` which may be used when there are multiple test statistics per component of `Elements`. See the example in the documentation for `cpd.PwrGSD`.
subset	the plot can be applied to a subset of rows of `descr` via a logical expression on its variables in combination with the special variable, `stat` when applicable.
na.action	a `na.action` method for handling `NA` values
...	other parameters to pass to the R function `coplot` usually not neccesary

Value

Returns the object, `x`, invisibly

Note

This processes the `cpd.PwrGSD` object into a dataframe, stacked on interim looks and then passes the results to the R function `coplot`

Author(s)

Abovementioned `cpd.PwrGSD` processing done by Grant Izmirlian <izmirlian@nih.gov>

References

Chambers, J. M. (1992) *Data for models.* Chapter 3 of *Statistical Models in S* eds J. M. Chambers and T. J. Hastie, Wadsworth \& Brooks/Cole.

Cleveland, W. S. (1993) *Visualizing Data.* New Jersey: Summit Press.

See Also

`cpd.PwrGSD Power` and `Elements`

Examples

```
## See the example in the 'cpd.PwrGSD' documentation
```

GrpSeqBnds	*Computes efficacy and futility boundaries*

Description

This computes efficacy and futility boundaries for interim analysis and sequential designs. Two sided symmetric efficacy boundaries can be computed by specifying half of the intended total type I error probability in the argument, `Alpha.Efficacy`. Otherwise, especially in the case of efficacy and futility bounds only one sided boundaries are currently computed. The computation allows for two different time scales–one must be the variance ratio, and the second can be a user chosen increasing scale beginning with 0 that takes the value 1 at the conclusion of the trial.

Usage

```
GrpSeqBnds(EfficacyBoundary = LanDemets(alpha = 0.05, spending = ObrienFleming),
           FutilityBoundary = LanDemets(alpha = 0.1, spending = ObrienFleming),
           NonBindingFutility = TRUE, frac, frac.ii = NULL, drift = NULL)
```

Arguments

`EfficacyBoundary`

This specifies the method used to construct the efficacy boundary. The available choices are:

'(i) 'Lan-Demets(alpha=<total type I error>, spending=<spending function>). The Lan-Demets method is based upon a error probability spending approach. The spending function can be set to `ObrienFleming`, `Pocock`, or `Power(rho)`, where `rho` is the the power argument for the power spending function: rho=3 is roughly equivalent to the O'Brien-Fleming spending function and smaller powers result in a less conservative spending function.

'(ii) 'Haybittle(alpha=<total type I error>, b.Haybittle=<user specified boundary point>). The Haybittle approach is the simplest, which sets the boundary points equal to `b.Haybittle`, a user specified value (try 3) for all analyses except the last, which is calculated so as to result in the total type I error, set with the argument `alpha`.

`'(iii) 'SC(be.end=<efficacy boundary point at trial end>,
prob=<threshold for conditional type I error for efficacy
stopping>)`. The stochastic curtailment method is based upon the conditional probability of type I error given the current value of the statistic. Under this method, a sequence of boundary points on the standard normal scale (as are boundary points under all other methods) is calculated so that the total probability of type I error is maintained. This is done by considering the joint probabilities of continuing to the current analysis and then exceeding the threshold at the current analysis. A good value for the threshold value for the conditional type I error, `prob` is 0.90 or greater.

`'(iv) 'User supplied boundary points in the form c(b1, b2, b3, ...,
b_m)`, where `m` is the number of looks.

FutilityBoundary

This specifies the method used to construct the futility boundary. The available choices are:

`'(i) 'Lan-Demets(alpha=<total type II error>, spending=<spending
function>)`. The Lan-Demets method is based upon a error probability spending approach. The spending function can be set to `ObrienFleming`, `Pocock`, or `Power(rho)`, where `rho` is the the power argument for the power spending function: rho=3 is roughly equivalent to the O'Brien-Fleming spending function and smaller powers result in a less conservative spending function.

`'(ii) 'Haybittle(alpha=<total type I error>, b.Haybittle=<user
specified boundary point>)`. The Haybittle approach is the simplest, which sets the boundary points equal to `b.Haybittle`, a user specified value (try 3) for all analyses except the last, which is calculated so as to result in the total type II error, set with the argument `alpha`.

`'(iii) 'SC(be.end=<efficacy boundary point at trial end>,
prob=<threshold for conditional type II error for futility
stopping>, drift.end=<projected drift at end of trial>)`.
The stochastic curtailment method is based upon the conditional probability of type II error given the current value of the statistic. Under this method, a sequence of boundary points on the standard normal scale (as are boundary points under all other methods) is calculated so that the total probability of type II error, is maintained. This is done by considering the joint probabilities of continuing to the current analysis and then exceeding the threshold at the current analysis. A good value for the threshold value for the conditional type I error, `prob` is 0.90 or greater.

`'(iv) 'User supplied boundary points in the form c(b1, b2, b3, ...,
b_m)`, where `m` is the number of looks.

NonBindingFutility

When using a futility boundary and this is set to 'TRUE', the efficacy boundary will be constructed in the absence of the futility boundary, and then the futility boundary will be constructed given the resulting efficacy boundary. This results in a more conservative efficacy boundary with true type I error less than the nominal level. This is recommended due to the fact that futility crossings are viewed by DSMB's with much less gravity than an efficacy crossing and as such, the consensus is that efficacy bounds should not be discounted towards the null hypothesis because of paths which cross a futility boundary. Default value is 'TRUE'.

frac

The variance ratio. If the end of trial variance is unknown then normalize all previous variances by the current variance. In this case you must specify a second scale that is monotone increasing from 0 to 1 at the end of the trial. Required.

frac.ii   The second information scale that is used for type I and type II error probability
                 spending. Optional (see above)

drift   The drift function of the underlying brownian motion, which is the expected
                 value under the design alternative of the un-normalized weighted log-rank statis-
                 tic, then normalized to have variance one when the variance ratio equals 1. See
                 the examples below.

Value

An object of class `boundaries` with components: "table" "frac" "frac.ii" "drift" "call"

call   The call that produced the returned results.

frac   The vector of variance ratios.

frac.ii   The vector of information ratios for type I and type II error probability spending,
                 which differs from the above if the user sets the argument `frac.ii` to a second
                 scale as mentioned above.

drift   The drift vector that is required as an argument when futility boundaries are
                 calculated.

table   A matrix with components

                 `frac ` The information ratio for type I and type II error probability spending.

                 `b.f ` The calculated futility boundary (if requested).

                 `alpha.f ` The type II error probability spent at that analysis (if doing futility
                 bounds).

                 `cum-alpha.f ` Cumulative sum of `alpha.f` (if doing futility bounds).

                 `b.e ` The calculated efficacy boundary.

                 `alpha.e ` The type I error probability spent at that analysis.

                 `cum-alpha.e ` Cumulative sum of `alpha.e`.

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

References

Gu, M.-G. and Lai, T.-L. "Determination of power and sample size in the design of clinical trials
with failure-time endpoints and interim analyses." Controlled Clinical Trials 20 (5): 423-438. 1999

Izmirlian, G. "The PwrGSD package." NCI Div. of Cancer Prevention Technical Report. 2004

Jennison, C. and Turnbull, B.W. (1999) Group Sequential Methods: Applications to Clinical Trials
Chapman & Hall/Crc, Boca Raton FL

Proschan, M.A., Lan, K.K.G., Wittes, J.T. (2006), corr 2nd printing (2008) Statistical Monitoring
of Clinical Trials A Unified Approach Springer Verlag, New York

See Also

[PwrGSD](#)

Examples

```
## NOTE: In an unweighted analysis, the variance ratios and event ratios
## are the same, whereas in a weighted analysis, they are quite different.
##
## For example, in a trial with 7 or so years of accrual and maximum follow-up of 20
## using the stopped Fleming-Harrington weights, `WtFun' = "SFH", with paramaters
## `ppar' = c(0, 1, 10) we might get the following vector of variance ratios:

frac    <- c(0.006995655, 0.01444565, 0.02682463, 0.04641363, 0.0585665,
               0.07614902, 0.1135391, 0.168252, 0.2336901, 0.3186155, 0.4164776,
               0.5352199, 0.670739, 0.8246061, 1)


## and the following vector of event ratios:

frac.ii <- c(0.1494354, 0.1972965, 0.2625075, 0.3274323, 0.3519184, 0.40231,
               0.4673037, 0.5579035, 0.6080742, 0.6982293, 0.7671917, 0.8195019,
               0.9045182, 0.9515884, 1)

## and the following drift under a given alternative hypothesis

drift <-   c(0.06214444, 0.1061856, 0.1731267, 0.2641265, 0.3105231, 0.3836636,
               0.5117394, 0.6918584, 0.8657705, 1.091984, 1.311094, 1.538582,
               1.818346, 2.081775, 2.345386)

## JUST ONE SIDED EFFICACY BOUNDARY
## In this call, we calculate a one sided efficacy boundary at each of 15 analyses
## which will occur at the given (known) variance ratios, and we use the variance
## ratio for type I error probability spending, with a total type I error probabilty
## of 0.05, using the Lan-Demets method with Obrien-Fleming spending (the default).

gsb.all.just.eff <- GrpSeqBnds(frac=frac,
                                 EfficacyBoundary=LanDemets(alpha=0.05, spending=Obrien

## ONE SIDED EFFICACY AND FUTILTY BOUNDARIES
## In this call, we calculate a one sided efficacy boundary at each of 15 analyses
## which will occur at the given (known) variance ratios, and we use the variance
## ratio for type I and type II error probability spending, with a total type I error
## probabilty of 0.05 and a total type II error probability of 0.10, using the Lan-De
## method with Obrien-Fleming spending (the default) for both efficacy and futilty.

gsb.all.eff.fut <- GrpSeqBnds(frac=frac,
                                 EfficacyBoundary=LanDemets(alpha=0.05, spending=ObrienF
                                 FutilityBoundary=LanDemets(alpha=0.10, spending=ObrienF
                                 drift=drift)

## Now suppose that we are performing the 7th interim analysis. We don't know what th
## will be at the end of the trial, so we normalize variances of the current and prev
## statistics by the variance of the current statistic.  This is equivalent to the fo
## length 7 vector of variance ratios:

frac7 <- frac[1:7]/frac[7]

## To proceed under the "unknown variance at end of trial" case, we must use a second
## scale for spending type I and II error probabilty. Unlike the above scale
```

```
## which is renormalized at each analysis to have value 1 at the current analysis, th
## alpha spending scale must be monotone increasing and attain the value 1 only at th
## end of the trial. A natural choice is the event ratio, which is known in advance i
## the trial is run until a required number of events is obtained, a so called
## maximum information trial:

frac7.ii <- frac.ii[1:7]

## the first seven values of the drift function

drift7 <- drift[1:7]/frac[7]^0.5

gsb.1st7.eff.fut <- GrpSeqBnds(frac=frac7, frac.ii=frac7.ii,
                                EfficacyBoundary=LanDemets(alpha=0.05, spending=Obrien
                                FutilityBoundary=LanDemets(alpha=0.10, spending=Obrien
drift=drift7)

## Of course there are other options not covered in these examples but this should ge
## started
```

CondPower	*Conditional type I and type II error probabilities given current value of the test statistic*

Description

Computes conditional type I and type II error probabilities given current value of the test statistic for monitoring based upon stochastic curtailment. This is now obsolete and included in the functionality of "GrpSeqBnds" and is here for instructional purposes only.

Usage

```
CondPower(Z, frac, drift, drift.end, err.I, sided = 1)
```

Arguments

Z	Current value of test statistic standardized to unit variance.
frac	Current value of the information fraction (variance fraction).
drift	Current value of the drift, i.e. the expected value of the test statistic normalized to have variance equal to the information fraction. Required if you want to compute conditional type II error, otherwise enter 0.
drift.end	Projected value of the drift at the end of the trial.
err.I	Overall (total) type I error probability
sided	Enter 1 or 2 for sided-ness of the test.

Value

A named numeric vector containing the two components "Pr.cond.typeIerr" and "Pr.cond.typeIIerr"

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

References

A General Theory on Stochastic Curtailment for Censored Survival Data D. Y. Lin, Q. Yao, Zhiliang Ying Journal of the American Statistical Association, Vol. 94, No. 446 (Jun., 1999), pp. 510-521

See Also

```
GrpSeqBnds
```

Examples

```
## None as yet
```

SimGSB   *Verifies the results of "GrpSeqBnds" via simulation*

Description

Verifies the results of `GrpSeqBnds` via simulation

Usage

```
SimGSB(object, nsim = 1e+05, ...)
```

Arguments

object	an object of class either `boundaries` or `PwrGSD`
nsim	number of simulations to do
...	if `object` is of class `PwrGSD` and there are more than one statistic under investigation, then you may specify an argument `stat`. The default value is 1, meaning the first one.

Value

A tabulation of the results

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

See Also

```
GrpSeqBnds
```

Examples

```
## none as yet
```

| wtdlogrank | *Weighted log-rank test* |

Description

Computes a two sample weighted log-rank statistic with events weighted according to one of the available weighting function choices

Usage

```
wtdlogrank(formula = formula(data), data = parent.frame(), WtFun = c("FH", "SF
param = c(0, 0), sided = c(2, 1), subset, na.action, w = FALSE)
```

Arguments

formula	a formula of the form `Surv(Time, Event) ~ arm` where `arm` is a dichotomous variable with values 0 and 1.
data	a dataframe
WtFun	a selection from the available list: "FH" (Fleming-Harrington), "SFH" (stopped Fleming-Harrington) or "Ramp". See `param` in the following line.
param	Weight function parameters. Length and interpretation depends upon the selected value of `WtFun`: If `WtFun==FH` then `param` is a length 2 vector specifying the power of the pooled (across arms) kaplan meier estimate and its complement. If `WtFun==SFH` then `param` is a length 3 vector with first two components as in the "FH" case, and third component the time (in the same units as the time to event) at which the "FH" weight function is capped off at its current value. If `WtFun==SFH` then `param` is of length 1 specifying the time (same units as time to event) at which events begin to get equal weight. The "Ramp" weight function is a linearly increasing deterministic weight function which is capped off at 1 at the user specified time.
sided	One or Two sided test? Set to 1 or 2
subset	Analysis can be applied to a subset of the dataframe based upon a logical expression in its variables
na.action	Method for handling `NA` values in the covariate, `arm`
w	currently no effect

Value

An object of class `survtest` containing components

pn	sample size
wttyp	internal representation of the `WtFun` argument
par	internal representation of the `param` argument
time	unique times of events accross all arms
nrisk	number at risk accross all arms at each event time
nrisk1	Number at risk in the experimental arm at each event time
nevent	Number of events accross all arms at each event time

nevent1	Number of events in the experimental arm at each event time
wt	Values of the weight function at each event time
pntimes	Number of event times
stat	The un-normalized weighted log-rank statistic, i.e. the summed weighted observed minus expected differences at each event time
var	Variance estimate for the above
UQt	Cumulative sum of increments in the sum resulting in stat above
varQt	Cumulative sum of increments in the sum resulting in var above
var1t	Cumulative sum of increments in the sum resulting in the variance of an un-weighted version of the statistic
pu0	person units of follow-up time in the control arm
pu1	person units of follow-up time in the intervention arm
n0	events in the control arm
n1	events in the intervention arm
n	sample size, same as pn
call	the call that created the object

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

References

Harrington, D. P. and Fleming, T. R. (1982). A class of rank test procedures for censored survival data. *Biometrika* **69**, 553-566.

See Also

IntSurvDiff

Examples

```
library(PwrGSD)
data(lung)
fit.wlr <- wtdlogrank(Surv(time, I(status==2))~I(sex==2), data=lung, WtFun="SFH", param
```

IntSurvDiff	*Weighted Integrated Survival function test*

Description

Computes a two sample weighted integrated survival function log-rank statistic with events weighted according to one of the available weighting function choices

Usage

```
IntSurvDiff(formula = formula(data), data = parent.frame(), WtFun = c("FH", "S
            param = c(0, 0), sided = c(2, 1), subset, na.action, w = FALSE)
```

Arguments

formula	a formula of the form Surv(Time, Event) ~ arm where arm is a dichotomous variable with values 0 and 1.
data	a dataframe
WtFun	a selection from the available list: "FH" (Fleming-Harrington), "SFH" (stopped Fleming-Harrington) or "Ramp". See param in the following line.
param	Weight function parameters. Length and interpretation depends upon the selected value of WtFun: If WtFun==FH then param is a length 2 vector specifying the power of the pooled (across arms) kaplan meier estimate and its complement. If WtFun==SFH then param is a length 3 vector with first two components as in the "FH" case, and third component the time (in the same units as the time to event) at which the "FH" weight function is capped off at its current value. If WtFun==SFH then param is of length 1 specifying the time (same units as time to event) at which events begin to get equal weight. The "Ramp" weight function is a linearly increasing deterministic weight function which is capped off at 1 at the user specified time.
sided	One or Two sided test? Set to 1 or 2
subset	Analysis can be applied to a subset of the dataframe based upon a logical expression in its variables
na.action	Method for handling NA values in the covariate, arm
w	currently no effect

Value

An object of class survtest containing components

pn	sample size
wttyp	internal representation of the WtFun argument
par	internal representation of the param argument
time	unique times of events accross all arms
nrisk	number at risk accross all arms at each event time
nrisk1	Number at risk in the experimental arm at each event time
nevent	Number of events accross all arms at each event time
nevent1	Number of events in the experimental arm at each event time
wt	Values of the weight function at each event time
pntimes	Number of event times
stat	The un-normalized weighted log-rank statistic, i.e. the summed weighted observed minus expected differences at each event time
var	Variance estimate for the above
pu0	person units of follow-up time in the control arm
pu1	person units of follow-up time in the intervention arm
n0	events in the control arm
n1	events in the intervention arm
n	sample size, same as pn
call	the call that created the object

Author(s)

Grant Izmirlian <izmirlian@nih.gov

References

Weiand S, Gail MH, James BR, James KL. (1989). A family of nonparametric statistics for comparing diagnostic makers with paired or unpaired data. *Biometrika* **76**, 585-592.

See Also

[wtdlogrank](#)

Examples

```
library(PwrGSD)
data(lung)
fit.isd <- IntSurvDiff(Surv(time, I(status==2))~I(sex==2), data=lung, WtFun="SFH", para
```

---

agghaz	*Aggregated Hazard*

Description

Computes the MLE for the model that assumes piecewise constant hazards on intervals defined by a grid of points. One applications for example is to calculate monthly hazard rates given numbers of events, numbers at risk and event times reported to the day. Can also handle time to event data stratified on a blocking factor.

Usage

```
agghaz(t.agg, time, nrisk, nevent)
```

Arguments

t.agg	Vector defining intervals upon which the user wants constant hazard rates.
time	Event times, possibly stratified on a blocking factor into multiple columns, in units that occur in enough numbers per interval specified above. If there is just a single column then it must be in column form (see example below).
nrisk	Numbers at risk at specified event times
nevent	Numbers of events at specified event times

Value

time.a	User supplied left-hand endpoints of intervals of hazard constancy
nrisk.a	Numbers at risk on specified intervals
nevent.a	Numbers of events on specified intervals

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

Examples

```
library(PwrGSD)
data(lung)
fit.msf <- mysurvfit(Surv(time, I(status==2)) ~ sex, data=lung)

## A single stratum:
with(fit.msf$Table, agghaz(30, time, cbind(nrisk1), cbind(nevent1)))

## Multiple strata--pooled and group 1:
with(fit.msf$Table, agghaz(30, time, cbind(nrisk1+nrisk2,nrisk1), cbind(nevent1+nevent2
```

---

mysurvfit   *My Survfit*

Description

Computes numbers at risk, numbers of events at each unique event time within levels of a blocking factor

Usage

```
mysurvfit(formula = formula(data), data = parent.frame(), subset, na.action =
```

Arguments

formula	Should be a formula of the form Surv(ti, ev) ~ block where block is the blocking factor. It need not be a factor per se but should have relatively few discrete levels. Sorry, no staggered entry allowed at present
data	a dataframe
subset	you can subset the analysis via logical expression in variables in the dataframe
na.action	pass a method for handling NA values in block such as na.omit, or na.fail

Value

A dataframe of 2*NLEV + 1 columns where NLEV is the number of levels of the factor block.

time	The sorted vector of unique event times from all blocks
nrisk1	The number at risk in block level 1 at each event time
nevent1	The number of events in block level 1 at each event time
...	
nriskNLEV	The number at risk in block level NLEV at each event time
neventNLEV	The number of events in block level NLEV at each event time

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

Examples

```
library(PwrGSD)
data(lung)

fit.msf <- mysurvfit(Surv(time, I(status==2)) ~ sex, data=lung)

fit.msf
## Not run:
plot(fit.msf)

## End(Not run)
```

---

mystack   *Stack a dataset*

Description

Given a dataframe containing one or more variables named with a common prefix, this function creates a stacked dataset with one set of observed values of the variables (in order of occurence) per line.

Usage

```
mystack(object, fu.vars, create.idvar = FALSE)
```

Arguments

object   a dataframe containing one or more variables named with a common prefix

fu.vars   a list of the unique prefixes

create.idvar Do you want to add an ID variable with a common value given to all records
              resulting from a given input record? Default is FALSE

Value

A stacked dataframe

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

Examples

```
## none as yet
```

CDFOR2LRR   *Convert CDF Odds Ratio to Logged Relative Risks*

Description

Given the values of the baseline hazard and odds ratio of the CDF at a grid of time points find the corresponding logged risk ratio.

Usage

```
CDFOR2LRR(tcut, tmax, h0, CDFOR)
```

Arguments

tcut   Grid of time points (left endpoints)

tmax   The right endpoint of the last interval

h0   Values of the baseline hazard function on given intervals

CDFOR   Values of the odds ratio of the CDF's on the given intervals

Value

An m by 2 matrix, where m=length(tcut), having columns 'tcut' and logged RR.

Author(s)

Grant Izmirlian <izmirlian@nih.gov

CY2TOShaz   *Calender year rates to Study Year Rates*

Description

Given the cutpoints at which the hazard is to be constant, the values taken by the calender year rates and the calender time offset from the start of the trial at which randomization ended, this function converts to time on study rates, assuming uniform accrual.

Usage

```
CY2TOShaz(tcut, t.eor, m, verbose = FALSE)
```

Arguments

tcut   Left hand endpoints of intervals on which time on study hazard is taken to be constant

t.eor   Time offsest from the beginning of the trial at which randomization ended

m   Annual calender time rates

verbose   do you want to see alot of debugging info--defaults to FALSE

Value

hazard = h, table = attr(obj., "tbl")

hazard	time on study hazard values taken on intervals specified by the argument `tcut`
table	a table containg the observed and fitted values

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

Examples

```
## none as yet
```

CRRtoRR   *Cumulative-risk ratios to risk ratios*

Description

Given a vector of cumulative-risk ratios, computes risk ratios

Usage

```
CRRtoRR(CRR, DT, h = NULL)
```

Arguments

CRR	vector of cumulative risk ratios of length `m`
DT	vector of time increments upon which the cumulative ratios represent. For example if the hazard takes values $h\_1, h\_2, \ldots, h\_m$ on the intervals $[t\_1, t\_2), [t\_2, t\_3), \ldots, [t\_m, t\_m+1)$ then `DT` will be c($t\_2-t\_1, t\_3 -t\_2, \ldots, t\_m+1 - t\_m$)$
h	The hazard in the reference arm, of length `m`

Value

The vector of risk ratios at the `m` time points

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

Examples

```
## none as yet
```

RCM2RR   *Relative cumulative mortality to Relative Risk*

Description

Given the relative cumulative mortality (ratio of CDFs), the baseline hazard and censoring hazard at a grid of time points, calculates the corresponding risk ratio at a second specified grid of time points.

Usage

```
RCM2RR(tlook, tcut.i, h.i, hOth, accru, rcm)
```

Arguments

`tlook`	Second grid of time points at which you desire risk ratios
`tcut.i`	First grid of time points at which baseline hazard, censoring hazard and relative cumulative mortality are specified (left hand endpoints of intervals)
`h.i`	Values of baseline hazard on intervals given by tcut.i
`hOth`	Values of censoring hazard on intervals given by tcut.i
`accru`	Time at which uniform accrual is completed (starts at 0)
`rcm`	Values of relative cumulative mortality (ratio of CDFs) on interals given by tcut.i

Value

Values of risk ratio on intervals given by tlook

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

RR2RCM   *Relative risk to Relative Cumulative Mortality*

Description

Relative risk to Relative Cumulative Mortality

Usage

```
RR2RCM(tlook, tcut.i, tcut.ii, h, rr, hOth, accru)
```

Arguments

tlook	Grid of time points at which you desire cumlative relative mortality
tcut.i	Grid of time points at which baseline hazard, censoring hazard and relative cumulative mortality are specified (left hand endpoints of intervals)
tcut.ii	Grid of time points at which study arm hazard is specified (left hand endpoints of intervals)
h	Values of baseline hazard on intervals given by tcut.i
rr	Values of risk ratio on intervals given by tcut.i
hOth	Values of censoring hazard on intervals given by tcut.i
accru	Time at which uniform accrual is completed (starts at 0)

Value

Values of relative cumulative mortality (ratio of CDFs) on interals given by tlook

Author(s)

Grant Izmirlian <izmirlian@nih.gov

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lookup	*Lookup values for a piecewise constant function*

Description

Given the values and lefthand endpoints for intervals of constancy, lookup values of the function at arbitrary values of the independent variable.

Usage

```
lookup(xgrid, ygrid, x, y0 = 0)
```

Arguments

xgrid	Lefthand endpoints of intervals of constancy
ygrid	Values on these intervals, of same length as xgrid
x	Input vector of arbitrary independent variables.
y0	Value to be returned for values of x that are smaller than min(xgrid).

Value

~Describe the value returned If it is a LIST, use

comp1	Description of 'comp1'
comp2	Description of 'comp2'

Author(s)

Grant Izmirlian <izmirlian@nih.gov>

Examples

```
## none as yet
```

---

lung   *Mayo Clinic Lung Cancer Data*

Description

Survival in patients with lung cancer at Mayo Clinic. Performance scores rate how well the patient can perform usual daily activities.

Usage

```
data(lung)
```

Format

inst:	Institution code
time:	Survival time in days
status:	censoring status 1=censored, 2=dead
age:	Age in years
sex:	Male=1 Female=2
ph.ecog:	ECOG performance score (0=good 5=dead)
ph.karno:	Karnofsky performance score (bad=0-good=100) rated by physician
pat.karno:	Karnofsky performance score rated by patient
meal.cal:	Calories consumed at meals
wt.loss:	Weight loss in last six months

Source

Terry Therneau

DX   *A utility function for forming differences*

Description

DX(x) returns c(x[1], diff(x))

Usage

```
DX(x)
```

Arguments

x   A grid of time points (increasing)

Value

DX(x) returns c(x[1], diff(x))

Author(s)

Grant Izmirlian <izmirlian@nih.gov

---

paste *The paste operator*

Description

A binary operator shortcut for paste(x,y)

Usage

```
x %,% y
```

Arguments

x	a character string
y	a character string

Value

paste(x, y, sep="")

Author(s)

Grant Izmirlian `<izmirlian@nih.gov>`

Examples

```
library(PwrGSD)
"var" %,% (1:10)
```

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