# Package 'survival'

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aareg

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Aalen's additive regression model for censored data

# Description

Returns an object of class "aareg" that represents an Aalen model.

# Usage

```
aareg(formula, data, weights, subset, na.action,
  qrtol=1e-07, nmin, dfbeta=FALSE, taper=1,
  test = c('aalen', 'variance', 'nrisk'),
  model=FALSE, x=FALSE, y=FALSE)
```

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#### **Arguments**

formula

a formula object, with the response on the left of a '~' operator and the terms, separated by + operators, on the right. The response must be a Surv object. Due to a particular computational approach that is used, the model MUST include an intercept term. If "-1" is used in the model formula the program will ignore it.

data

data frame in which to interpret the variables named in the formula, subset, and weights arguments. This may also be a single number to handle some speci al cases – see below for details. If data is missing, the variables in the model formula should be in the search path.

weights

vector of observation weights. If supplied, the fitting algorithm minimizes the sum of the weights multiplied by the squared residuals (see below for additional technical details). The length of weights must be the same as the number of observations. The weights must be nonnegative and it is recommended that they be strictly positive, since zero weights are ambiguous. To exclude particular observations from the model, use the subset argument instead of zero weights.

subset

expression specifying which subset of observations should be used in the fit. Th is can be a logical vector (which is replicated to have length equal to the numb er of observations), a numeric vector indicating the observation numbers to be included, or a character vector of the observation names that should be included. All observations are included by default.

na.action

a function to filter missing data. This is applied to the model.fr ame after any subset argument has be en applied. The default is na.fail, which returns a n error if any missing values are found. An alternative is na.excl ude, which deletes observations that contain one or more missing values.

qrtol

tolerance for detection of singularity in the QR decomposition

nmin

minimum number of observations for an estimate; defaults to 3 times the number of covariates. This essentially truncates the computations near the tail of the data set, when n is small and the calculations can become numerically unstable.

dfbeta

should the array of dfbeta residuals be computed. This implies computation of the sandwich variance estimate. The residuals will always be computed if there is a cluster term in the model formula.

taper

allows for a smoothed variance estimate. Var(x), where x is the set of covariates, is an important component of the calculations for the Aalen regression model. At any given time point t, it is computed over all subjects who are still at risk at time t. The tape argument allows smoothing these estimates, for example taper=(1:4)/4 would cause the variance estimate used at any event time to be a weighted average of the estimated variance matrices at the last 4 death times, with a weight of 1 for the current death time and decreasing to 1/4 for prior event times. The default value gives the standard Aalen model.

test

selects the weighting to be used, for computing an overall "average" coefficient vector over time and the subsequent test for equality to zero.

model, x, y

should copies of the model frame, the x matrix of predictors, or the response vector y be included in the saved result.

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#### **Details**

The Aalen model assumes that the cumulative hazard H(t) for a subject can be expressed as a(t) + X B(t), where a(t) is a time-dependent intercept term, X is the vector of covariates for the subject (possibly time-dependent), and B(t) is a time-dependent matrix of coefficients. The estimates are inherently non-parametric; a fit of the model will normally be followed by one or more plots of the estimates.

The estimates may become unstable near the tail of a data set, since the increment to B at time t is based on the subjects still at risk at time t. The tolerance and/or nmin parameters may act to truncate the estimate before the last death. The taper argument can also be used to smooth out the tail of the curve. In practice, the addition of a taper such as 1:10 appears to have little effect on death times when n is still reasonably large, but can considerably dampen wild occilations in the tail of the plot.

#### Value

an object of class "aareg" representing the fit, with the following components:

n vector containing the number of observations in the data set, the number of event

times, and the number of event times used in the computation

times vector of sorted event times, which may contain duplicates

nrisk vector containing the number of subjects at risk, of the same length as times coefficient matrix of coefficients, with one row per event and one column per covariate

test.statistic the value of the test statistic, a vector with one element per covariate

test.var variance-covariance matrix for the test

test the type of test; a copy of the test argument above

tweight matrix of weights used in the computation, one row per event

call a copy of the call that produced this result

# References

Aalen, O.O. (1989). A linear regression model for the analysis of life times. Statistics in Medicine, 8:907-925.

Aalen, O.O (1993). Further results on the non-parametric linear model in survival analysis. Statistics in Medicine. 12:1569-1588.

#### See Also

print.aareg, summary.aareg, plot.aareg

## **Examples**

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```
)
 n=227 (1 observations deleted due to missing values)
    138 out of 138 unique event times used
             slope
                        coef se(coef)
Intercept 5.26e-03 5.99e-03 4.74e-03 1.26 0.207000
     age 4.26e-05 7.02e-05 7.23e-05 0.97 0.332000
     sex -3.29e-03 -4.02e-03 1.22e-03 -3.30 0.000976
 ph.ecog 3.14e-03 3.80e-03 1.03e-03 3.70 0.000214
Chisq=26.73 on 3 df, p=6.7e-06; test weights=aalen
plot(lfit[4], ylim=c(-4,4)) # Draw a plot of the function for ph.ecog
## End(Not run)
lfit2 <- aareg(Surv(time, status) ~ age + sex + ph.ecog, data=lung,</pre>
                 nmin=1, taper=1:10)
## Not run: lines(lfit2[4], col=2) # Nearly the same, until the last point
# A fit to the mulitple-infection data set of children with
# Chronic Granuomatous Disease. See section 8.5 of Therneau and Grambsch.
fita2 <- aareg(Surv(tstart, tstop, status) ~ treat + age + inherit +</pre>
                        steroids + cluster(id), data=cgd)
## Not run:
 n = 203
   69 out of 70 unique event times used
                              coef se(coef) robust se
                    slope
Intercept
                 0.004670 0.017800 0.002780 0.003910 4.55 5.30e-06
treatrIFN-g
                -0.002520 -0.010100 0.002290 0.003020 -3.36 7.87e-04
                -0.000101 -0.000317 0.000115 0.000117 -2.70 6.84e-03
inheritautosomal 0.001330 0.003830 0.002800 0.002420 1.58 1.14e-01
steroids
                 Chisq=16.74 on 4 df, p=0.0022; test weights=aalen
## End(Not run)
```

aeqSurv

Adjudicate near ties in a Surv object

# Description

The check for tied survival times can fail due to floating point imprecision, which can make actual ties appear to be distinct values. Routines that depend on correct identification of ties pairs will then give incorrect results, e.g., a Cox model. This function rectifies these.

# Usage

```
aeqSurv(x, tolerance = sqrt(.Machine$double.eps))
```

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# Arguments

x a Surv object

tolerance the tolerance used to detect values that will be considered equal

#### **Details**

This routine is called by both survfit and coxph to deal with the issue of ties that get incorrectly broken due to floating point imprecision. See the short vignette on tied times for a simple example. Use the timefix argument of survfit or coxph.control to control the option if desired.

The rule for 'equality' is identical to that used by the all.equal routine. Pairs of values that are within round off error of each other are replaced by the smaller value. An error message is generated if this process causes a 0 length time interval to be created.

#### Value

a Surv object identical to the original, but with ties restored.

#### Author(s)

Terry Therneau

## See Also

```
survfit, coxph.control
```

aggregate.survfit

Average survival curves

## **Description**

For a survfit object containing multiple curves, create average curves over a grouping.

# Usage

```
## S3 method for class 'survfit'
aggregate(x, by = NULL, FUN = mean, ...)
```

# **Arguments**

x a survfit object which has a data dimension.

by an optional list or vector of grouping elements, each as long as dim(x)['data'].

FUN a function to compute the summary statistic of interest.

... optional further arguments to FUN.

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## **Details**

The primary use of this is to take an average over multiple survival curves that were created from a modeling function. That is, a marginal estimate of the survival. It is primarily used to average over multiple predicted curves from a Cox model.

#### Value

a survfit object of lower dimension.

#### See Also

survfit

# **Examples**

```
cfit <- coxph(Surv(futime, death) ~ sex + age*hgb, data=mgus2)
# marginal effect of sex, after adjusting for the others
dummy <- rbind(mgus2, mgus2)
dummy$sex <- rep(c("F", "M"), each=nrow(mgus2)) # population data set
dummy <- na.omit(dummy) # don't count missing hgb in our "population
csurv <- survfit(cfit, newdata=dummy)
dim(csurv) # 2 * 1384 survival curves
csurv2 <- aggregate(csurv, dummy$sex)</pre>
```

agreg.fit

Cox model fitting functions

# **Description**

These are the the functions called by coxph that do the actual computation. In certain situations, e.g. a simulation, it may be advantageous to call these directly rather than the usual coxph call using a model formula.

# Usage

```
agreg.fit(x, y, strata, offset, init, control, weights, method,
rownames, resid=TRUE, nocenter=NULL)
coxph.fit(x, y, strata, offset, init, control, weights, method,
rownames, resid=TRUE, nocenter=NULL)
```

# **Arguments**

X	Matix of predictors. This should <i>not</i> include an intercept.
у	a Surv object containing either 2 columns (coxph.fit) or 3 columns (agreg.fit).
strata	a vector containing the stratification, or NULL
offset	optional offset vector
init	initial values for the coefficients

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control the result of a call to coxph.control

weights optional vector of weights

method method for hanling ties, one of "breslow" or "efron"

rownames this is only needed for a NULL model, in which case it contains the rownames

(if any) of the original data.

resid compute and return residuals.

nocenter an optional list of values. Any column of the X matrix whose values lie strictly

within that set will not be recentered. Note that the coxph function has (-1, 0, 1)

as the default.

## **Details**

This routine does no checking that arguments are the proper length or type. Only use it if you know what you are doing!

The resid and concordance arguments will save some compute time for calling routines that only need the likelihood, the generation of a permutation distribution for instance.

#### Value

a list containing results of the fit

#### Author(s)

Terry Therneau

# See Also

coxph

aml

Acute Myelogenous Leukemia survival data

# **Description**

Survival in patients with Acute Myelogenous Leukemia. The question at the time was whether the standard course of chemotherapy should be extended ('maintainance') for additional cycles.

# Usage

```
aml
leukemia
data(cancer, package="survival")
```

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#### **Format**

time: survival or censoring time

status: censoring status

x: maintenance chemotherapy given? (factor)

## Source

Rupert G. Miller (1997), Survival Analysis. John Wiley & Sons. ISBN: 0-471-25218-2.

anova.coxph

Analysis of Deviance for a Cox model.

# **Description**

Compute an analysis of deviance table for one or more Cox model fits, based on the log partial likelihood.

# Usage

```
## S3 method for class 'coxph'
anova(object, ..., test = 'Chisq')
```

#### **Arguments**

object An object of class coxph
... Further coxph objects

test a character string. The appropriate test is a chisquare, all other choices result in

no test being done.

#### **Details**

Specifying a single object gives a sequential analysis of deviance table for that fit. That is, the reductions in the model Cox log-partial-likelihood as each term of the formula is added in turn are given in as the rows of a table, plus the log-likelihoods themselves. A robust variance estimate is normally used in situations where the model may be mis-specified, e.g., multiple events per subject. In this case a comparison of likelihood values does not make sense (differences no longer have a chi-square distribution), and anova will refuse to print results.

If more than one object is specified, the table has a row for the degrees of freedom and loglikelihood for each model. For all but the first model, the change in degrees of freedom and loglik is also given. (This only make statistical sense if the models are nested.) It is conventional to list the models from smallest to largest, but this is up to the user.

The table will optionally contain test statistics (and P values) comparing the reduction in loglik for each row.

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# Value

An object of class "anova" inheriting from class "data.frame".

## Warning

The comparison between two or more models by anova will only be valid if they are fitted to the same dataset. This may be a problem if there are missing values.

## See Also

```
coxph, anova.
```

# **Examples**

```
fit <- coxph(Surv(futime, fustat) ~ resid.ds *rx + ecog.ps, data = ovarian)
anova(fit)
fit2 <- coxph(Surv(futime, fustat) ~ resid.ds +rx + ecog.ps, data=ovarian)
anova(fit2,fit)</pre>
```

attrassign

Create new-style "assign" attribute

# **Description**

The "assign" attribute on model matrices describes which columns come from which terms in the model formula. It has two versions. R uses the original version, but the alternate version found in S-plus is sometimes useful.

# Usage

```
## Default S3 method:
attrassign(object, tt,...)
## S3 method for class 'lm'
attrassign(object,...)
```

# Arguments

```
object model matrix or linear model object
tt terms object
... ignored
```

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## **Details**

For instance consider the following

```
survreg(Surv(time, status) ~ age + sex + factor(ph.ecog), lung)
```

R gives the compact for for assign, a vector (0, 1, 2, 3, 3, 3); which can be read as "the first column of the X matrix (intercept) goes with none of the terms, the second column of X goes with term 1 of the model equation, the third column of X with term 2, and columns 4-6 with term 3".

The alternate (S-Plus default) form is a list

```
$(Intercept) 1
$age 2
$sex 3
$factor(ph.ecog) 4 5 6
```

#### Value

A list with names corresponding to the term names and elements that are vectors indicating which columns come from which terms

## See Also

```
terms, model. matrix
```

# **Examples**

```
formula <- Surv(time,status)~factor(ph.ecog)
tt <- terms(formula)
mf <- model.frame(tt,data=lung)
mm <- model.matrix(tt,mf)
## a few rows of data
mm[1:3,]
## old-style assign attribute
attr(mm,"assign")
## alternate style assign attribute
attrassign(mm,tt)</pre>
```

basehaz

Alias for the survfit function

# **Description**

Compute the predicted survival curve for a Cox model.

# Usage

```
basehaz(fit, centered=TRUE)
```

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## **Arguments**

fit a coxph fit

centered if TRUE return data from a predicted survival curve at the mean values of the

covariates fit\$mean, if FALSE return a prediction for all covariates equal to

zero.

#### **Details**

This function is simply an alias for survfit, which does the actual work and has a richer set of options. The alias exists only because some users look for predicted survival estimates under this name.

The function returns a data frame containing the time, cumhaz and optionally the strata (if the fitted Cox model used a strata statement), which are copied the survfit result. If there are factor variables in the model, then the default predictions at the "mean" are meaningless since they do not correspond to any possible subject; correct results require use of the newdata argument of survfit. Results for all covariates =0 are normally only of use as a building block for further calculations.

#### Value

a data frame with variable names of hazard, time and optionally strata. The first is actually the cumulative hazard.

#### See Also

survfit.coxph

bladder

Bladder Cancer Recurrences

# **Description**

Data on recurrences of bladder cancer, used by many people to demonstrate methodology for recurrent event modelling.

Bladder1 is the full data set from the study. It contains all three treatment arms and all recurrences for 118 subjects; the maximum observed number of recurrences is 9.

Bladder is the data set that appears most commonly in the literature. It uses only the 85 subjects with nonzero follow-up who were assigned to either thiotepa or placebo, and only the first four recurrences for any patient. The status variable is 1 for recurrence and 0 for everything else (including death for any reason). The data set is laid out in the competing risks format of the paper by Wei, Lin, and Weissfeld.

Bladder2 uses the same subset of subjects as bladder, but formatted in the (start, stop) or Anderson-Gill style. Note that in transforming from the WLW to the AG style data set there is a quite common programming mistake that leads to extra follow-up time for 12 subjects: all those with follow-up beyond their 4th recurrence. This "follow-up" is a side effect of throwing away all events after the fourth while retaining the last follow-up time variable from the original data. The bladder2 data set

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found here does not make this mistake, but some analyses in the literature have done so; it results in the addition of a small amount of immortal time bias and shrinks the fitted coefficients towards zero.

# Usage

bladder1
bladder2
data(cancer, package="survival")

## **Format**

#### bladder1

id: Patient id

treatment: Placebo, pyridoxine (vitamin B6), or thiotepa number: Initial number of tumours (8=8 or more) size: Size (cm) of largest initial tumour

recur: Number of recurrences

start,stop: The start and end time of each time interval status: End of interval code, 0=censored, 1=recurrence,

2=death from bladder disease, 3=death other/unknown cause

rtumor: Number of tumors found at the time of a recurrence

rsize: Size of largest tumor at a recurrence

enum: Event number (observation number within patient)

# bladder

id: Patient id

rx: Treatment 1=placebo 2=thiotepa number: Initial number of tumours (8=8 or more) size: size (cm) of largest initial tumour stop: recurrence or censoring time enum: which recurrence (up to 4)

# bladder2

id: Patient id

rx: Treatment 1=placebo 2=thiotepa number: Initial number of tumours (8=8 or more) size: size (cm) of largest initial tumour

start: start of interval (0 or previous recurrence time)

stop: recurrence or censoring time enum: which recurrence (up to 4)

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#### **Source**

Andrews DF, Hertzberg AM (1985), DATA: A Collection of Problems from Many Fields for the Student and Research Worker, New York: Springer-Verlag.

LJ Wei, DY Lin, L Weissfeld (1989), Regression analysis of multivariate incomplete failure time data by modeling marginal distributions. *Journal of the American Statistical Association*, **84**.

blogit

Bounded link functions

# **Description**

Alternate link functions that impose bounds on the input of their link function

# Usage

```
blogit(edge = 0.05)
bprobit(edge= 0.05)
bcloglog(edge=.05)
blog(edge=.05)
```

## **Arguments**

edge

input values less than the cutpoint are replaces with the cutpoint. For all be blog input values greater than (1-edge) are replaced with (1-edge)

# **Details**

When using survival psuedovalues for binomial regression, the raw data can be outside the range (0,1), yet we want to restrict the predicted values to lie within that range. A natural way to deal with this is to use glm with family = gaussian(link="logit"). But this will fail. The reason is that the family object has a component linkfun that does not accept values outside of (0,1).

This function is only used to create initial values for the iteration step, however. Mapping the offending input argument into the range of (egde, 1-edge) before computing the link results in starting estimates that are good enough. The final result of the fit will be no different than if explicit starting estimates were given using the etastart or mustart arguments. These functions create copies of the logit, probit, and complimentary log-log families that differ from the standard ones only in this use of a bounded input argument, and are called a "bounded logit" = blogit, etc.

The same argument hold when using RMST (area under the curve) pseudovalues along with a log link to ensure positive predictions, though in this case only the lower boundary needs to be mapped.

#### Value

```
a family object of the same form as make. family.
```

## See Also

```
stats{make.family}
```

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# **Examples**

```
py <- pseudo(survfit(Surv(time, status) ~1, lung), time=730) #2 year survival
range(py)
pfit <- glm(py ~ ph.ecog, data=lung, family=gaussian(link=blogit()))
# For each +1 change in performance score, the odds of 2 year survival
# are multiplied by 1/2 = exp of the coefficient.</pre>
```

cch

Fits proportional hazards regression model to case-cohort data

# **Description**

Returns estimates and standard errors from relative risk regression fit to data from case-cohort studies. A choice is available among the Prentice, Self-Prentice and Lin-Ying methods for unstratified data. For stratified data the choice is between Borgan I, a generalization of the Self-Prentice estimator for unstratified case-cohort data, and Borgan II, a generalization of the Lin-Ying estimator.

# Usage

```
cch(formula, data, subcoh, id, stratum=NULL, cohort.size,
  method =c("Prentice","SelfPrentice","LinYing","I.Borgan","II.Borgan"),
  robust=FALSE)
```

# **Arguments**

formula	A formula object that must have a Surv object as the response. The Surv object must be of type "right", or of type "counting".
subcoh	Vector of indicators for subjects sampled as part of the sub-cohort. Code 1 or TRUE for members of the sub-cohort, 0 or FALSE for others. If data is a data frame then subcoh may be a one-sided formula.
id	Vector of unique identifiers, or formula specifying such a vector.
stratum	A vector of stratum indicators or a formula specifying such a vector
cohort.size	Vector with size of each stratum original cohort from which subcohort was sampled
data	An optional data frame in which to interpret the variables occurring in the formula.
method	Three procedures are available. The default method is "Prentice", with options for "SelfPrentice" or "LinYing".
robust	For "LinYing" only, if robust=TRUE, use design-based standard errors even for phase $\boldsymbol{I}$

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#### **Details**

Implements methods for case-cohort data analysis described by Therneau and Li (1999). The three methods differ in the choice of "risk sets" used to compare the covariate values of the failure with those of others at risk at the time of failure. "Prentice" uses the sub-cohort members "at risk" plus the failure if that occurs outside the sub-cohort and is score unbiased. "SelfPren" (Self-Prentice) uses just the sub-cohort members "at risk". These two have the same asymptotic variance-covariance matrix. "LinYing" (Lin-Ying) uses the all members of the sub-cohort and all failures outside the sub-cohort who are "at risk". The methods also differ in the weights given to different score contributions.

The data argument must not have missing values for any variables in the model. There must not be any censored observations outside the subcohort.

#### Value

An object of class "cch" incorporating a list of estimated regression coefficients and two estimates of their asymptotic variance-covariance matrix.

coef regression coefficients.

naive.var Self-Prentice model based variance-covariance matrix.

var Lin-Ying empirical variance-covariance matrix.

#### Author(s)

Norman Breslow, modified by Thomas Lumley

#### References

Prentice, RL (1986). A case-cohort design for epidemiologic cohort studies and disease prevention trials. Biometrika 73: 1–11.

Self, S and Prentice, RL (1988). Asymptotic distribution theory and efficiency results for case-cohort studies. Annals of Statistics 16: 64–81.

Lin, DY and Ying, Z (1993). Cox regression with incomplete covariate measurements. Journal of the American Statistical Association 88: 1341–1349.

Barlow, WE (1994). Robust variance estimation for the case-cohort design. Biometrics 50: 1064–1072

Therneau, TM and Li, H (1999). Computing the Cox model for case-cohort designs. Lifetime Data Analysis 5: 99–112.

Borgan, O, Langholz, B, Samuelsen, SO, Goldstein, L and Pogoda, J (2000) Exposure stratified case-cohort designs. Lifetime Data Analysis 6, 39-58.

## See Also

twophase and svycoxph in the "survey" package for more general two-phase designs. http://faculty.washington.edu/tlumley/survey/

cgd 19

# **Examples**

```
## The complete Wilms Tumor Data
## (Breslow and Chatterjee, Applied Statistics, 1999)
## subcohort selected by simple random sampling.
subcoh <- nwtco$in.subcohort</pre>
selccoh <- with(nwtco, rel==1|subcoh==1)</pre>
ccoh.data <- nwtco[selccoh,]</pre>
ccoh.data$subcohort <- subcoh[selccoh]</pre>
## central-lab histology
ccoh.data$histol <- factor(ccoh.data$histol,labels=c("FH","UH"))</pre>
## tumour stage
ccoh.data$stage <- factor(ccoh.data$stage,labels=c("I","II","III","IV"))</pre>
ccoh.data$age <- ccoh.data$age/12 # Age in years</pre>
## Standard case-cohort analysis: simple random subcohort
fit.ccP <- cch(Surv(edrel, rel) ~ stage + histol + age, data =ccoh.data,</pre>
   subcoh = ~subcohort, id=~segno, cohort.size=4028)
fit.ccP
fit.ccSP <- cch(Surv(edrel, rel) ~ stage + histol + age, data =ccoh.data,</pre>
   subcoh = ~subcohort, id=~seqno, cohort.size=4028, method="SelfPren")
summary(fit.ccSP)
##
## (post-)stratified on instit
stratsizes<-table(nwtco$instit)</pre>
fit.BI<- cch(Surv(edrel, rel) ~ stage + histol + age, data =ccoh.data,</pre>
   subcoh = ~subcohort, id=~seqno, stratum=~instit, cohort.size=stratsizes,
   method="I.Borgan")
summary(fit.BI)
```

cgd

Chronic Granulotamous Disease data

# **Description**

Data are from a placebo controlled trial of gamma interferon in chronic granulotomous disease (CGD). Contains the data on time to serious infections observed through end of study for each patient.

cgd

## Usage

```
cgd
data(cgd)
```

#### **Format**

```
center enrolling center
random date of randomization
treatment placebo or gamma interferon
sex sex
age age in years, at study entry
height height in cm at study entry
weight weight in kg at study entry
inherit pattern of inheritance
steroids use of steroids at study entry,1=yes
propylac use of prophylactic antibiotics at study entry
hos.cat a categorization of the centers into 4 groups
tstart, tstop start and end of each time interval
status 1=the interval ends with an infection
enum observation number within subject
```

# **Details**

The cgd0 data set is in the form found in the references, with one line per patient and no recoding of the variables. The cgd data set (this one) has been cast into (start, stop) format with one line per event, and covariates such as center recoded as factors to include meaningful labels.

## **Source**

Fleming and Harrington, Counting Processes and Survival Analysis, appendix D.2.

## See Also

```
link{cgd0}
```

cgd0

cgd0

Chronic Granulotomous Disease data

## **Description**

Data are from a placebo controlled trial of gamma interferon in chronic granulotomous disease (CGD). Contains the data on time to serious infections observed through end of study for each patient.

# Usage

cgd0

## **Format**

id subject identification number

center enrolling center

random date of randomization

treatment placebo or gamma interferon

sex sex

age age in years, at study entry

height height in cm at study entry

weight weight in kg at study entry

inherit pattern of inheritance

steroids use of steroids at study entry,1=yes

propylac use of prophylactic antibiotics at study entry

**hos.cat** a categorization of the centers into 4 groups

futime days to last follow-up

etime1-etime7 up to 7 infection times for the subject

#### **Details**

The cgdraw data set (this one) is in the form found in the references, with one line per patient and no recoding of the variables.

The cgd data set has been further processed so as to have one line per event, with covariates such as center recoded as factors to include meaningful labels.

#### Source

Fleming and Harrington, Counting Processes and Survival Analysis, appendix D.2.

## See Also

cgd

22 cipoisson

cipoisson	Confidence limits for the Poisson	

## Description

Confidence interval calculation for Poisson rates.

# Usage

```
cipoisson(k, time = 1, p = 0.95, method = c("exact", "anscombe"))
```

# **Arguments**

k Number of successes time Total time on trial

p Probability level for the (two-sided) interval method The method for computing the interval.

#### **Details**

The likelihood method is based on equation 10.10 of Feller, which relates poisson probabilities to tail area of the gamma distribution. The Anscombe approximation is based on the fact that sqrt(k + 3/8) has a nearly constant variance of 1/4, along with a continuity correction.

There are many other proposed intervals: Patil and Kulkarni list and evaluate 19 different suggestions from the literature!. The exact intervals can be overly broad for very small values of k, many of the other approaches try to shrink the lengths, with varying success.

#### Value

a vector, matrix, or array. If both k and time are single values the result is a vector of length 2 containing the lower an upper limits. If either or both are vectors the result is a matrix with two columns. If k is a matrix or array, the result will be an array with one more dimension; in this case the dimensions and dimnames (if any) of k are preserved.

#### References

F.J. Anscombe (1949). Transformations of Poisson, binomial and negative-binomial data. Biometrika, 35:246-254.

W.F. Feller (1950). An Introduction to Probability Theory and its Applications, Volume 1, Chapter 6, Wiley.

V. V. Patil and H.F. Kulkarni (2012). Comparison of confidence intervals for the poisson mean: some new aspects. Revstat 10:211-227.

## See Also

```
ppois, qpois
```

clogit 23

# **Examples**

```
cipoisson(4) # 95\% confidence limit
# lower upper
# 1.089865 10.24153
ppois(4, 10.24153)  #chance of seeing 4 or fewer events with large rate
# [1] 0.02500096
1-ppois(3, 1.08986)  #chance of seeing 4 or more, with a small rate
# [1] 0.02499961
```

clogit

Conditional logistic regression

## **Description**

Estimates a logistic regression model by maximising the conditional likelihood. Uses a model formula of the form case.status~exposure+strata(matched.set). The default is to use the exact conditional likelihood, a commonly used approximate conditional likelihood is provided for compatibility with older software.

## Usage

```
clogit(formula, data, weights, subset, na.action,
method=c("exact", "approximate", "efron", "breslow"),
...)
```

# **Arguments**

formula	Model formula
data	data frame
weights	optional, names the variable containing case weights
subset	optional, subset the data
na.action	optional na.action argument. By default the global option na.action is used.
method	use the correct (exact) calculation in the conditional likelihood or one of the approximations
	optional arguments, which will be passed to coxph.control

## **Details**

It turns out that the loglikelihood for a conditional logistic regression model = loglik from a Cox model with a particular data structure. Proving this is a nice homework exercise for a PhD statistics class; not too hard, but the fact that it is true is surprising.

When a well tested Cox model routine is available many packages use this 'trick' rather than writing a new software routine from scratch, and this is what the clogit routine does. In detail, a stratified Cox model with each case/control group assigned to its own stratum, time set to a constant, status

24 clogit

of 1=case 0=control, and using the exact partial likelihood has the same likelihood formula as a conditional logistic regression. The clogit routine creates the necessary dummy variable of times (all 1) and the strata, then calls coxph.

The computation of the exact partial likelihood can be very slow, however. If a particular strata had say 10 events out of 20 subjects we have to add up a denominator that involves all possible ways of choosing 10 out of 20, which is  $20!/(10!\ 10!) = 184756$  terms. Gail et al describe a fast recursion method which partly ameliorates this; it was incorporated into version 2.36-11 of the survival package. The computation remains infeasible for very large groups of ties, say 100 ties out of 500 subjects, and may even lead to integer overflow for the subscripts – in this latter case the routine will refuse to undertake the task. The Efron approximation is normally a sufficiently accurate substitute.

Most of the time conditional logistic modeling is applied data with 1 case + k controls per set, in which case all of the approximations for ties lead to exactly the same result. The 'approximate' option maps to the Breslow approximation for the Cox model, for historical reasons.

Case weights are not allowed when the exact option is used, as the likelihood is not defined for fractional weights. Even with integer case weights it is not clear how they should be handled. For instance if there are two deaths in a strata, one with weight=1 and one with weight=2, should the likelihood calculation consider all subsets of size 2 or all subsets of size 3? Consequently, case weights are ignored by the routine in this case.

#### Value

An object of class "clogit", which is a wrapper for a "coxph" object.

#### References

Michell H Gail, Jay H Lubin and Lawrence V Rubinstein. Likelihood calculations for matched case-control studies and survival studies with tied death times. Biometrika 68:703-707, 1980.

John A. Logan. A multivariate model for mobility tables. Am J Sociology 89:324-349, 1983.

#### Author(s)

Thomas Lumley

# See Also

```
strata,coxph,glm
```

# **Examples**

```
## Not run: clogit(case ~ spontaneous + induced + strata(stratum), data=infert)

# A multinomial response recoded to use clogit

# The revised data set has one copy per possible outcome level, with new

# variable tocc = target occupation for this copy, and case = whether

# that is the actual outcome for each subject.

# See the reference below for the data.

resp <- levels(logan$occupation)

n <- nrow(logan)</pre>
```

cluster 25

cluster

Identify clusters.

# **Description**

This is a special function used in the context of survival models. It identifies correlated groups of observations, and is used on the right hand side of a formula. This style is now discouraged, use the cluster option instead.

## **Usage**

cluster(x)

## **Arguments**

Χ

A character, factor, or numeric variable.

#### Details

The function's only action is semantic, to mark a variable as the cluster indicator. The resulting variance is what is known as the "working independence" variance in a GEE model. Note that one cannot use both a frailty term and a cluster term in the same model, the first is a mixed-effects approach to correlation and the second a GEE approach, and these don't mix.

## Value

Χ

#### See Also

```
coxph, survreg
```

# **Examples**

26 colon

colon

Chemotherapy for Stage B/C colon cancer

## Description

These are data from one of the first successful trials of adjuvant chemotherapy for colon cancer. Levamisole is a low-toxicity compound previously used to treat worm infestations in animals; 5-FU is a moderately toxic (as these things go) chemotherapy agent. There are two records per person, one for recurrence and one for death

# Usage

colon

data(cancer, package="survival")

#### **Format**

id: id

study: 1 for all patients

rx: Treatment - Obs(ervation), Lev(amisole), Lev(amisole)+5-FU

sex: 1=male age: in years

obstruct: obstruction of colon by tumour

perfor: perforation of colon

adhere: adherence to nearby organs

nodes: number of lymph nodes with detectable cancer

time: days until event or censoring

status: censoring status

differ: differentiation of tumour (1=well, 2=moderate, 3=poor)

extent: Extent of local spread (1=submucosa, 2=muscle, 3=serosa, 4=contiguous structures)

surg: time from surgery to registration (0=short, 1=long)

node4: more than 4 positive lymph nodes etype: event type: 1=recurrence,2=death

#### Note

The study is originally described in Laurie (1989). The main report is found in Moertel (1990). This data set is closest to that of the final report in Moertel (1991). A version of the data with less follow-up time was used in the paper by Lin (1994).

Peter Higgins has pointed out a data inconsistency, revealed by table(colon\$nodes, colon\$node4). We don't know which of the two variables is actually correct so have elected not to 'fix' it. (Real data has warts, why not have some in the example data too?)

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#### References

JA Laurie, CG Moertel, TR Fleming, HS Wieand, JE Leigh, J Rubin, GW McCormack, JB Gerstner, JE Krook and J Malliard. Surgical adjuvant therapy of large-bowel carcinoma: An evaluation of levamisole and the combination of levamisole and fluorouracil: The North Central Cancer Treatment Group and the Mayo Clinic. J Clinical Oncology, 7:1447-1456, 1989.

DY Lin. Cox regression analysis of multivariate failure time data: the marginal approach. Statistics in Medicine, 13:2233-2247, 1994.

CG Moertel, TR Fleming, JS MacDonald, DG Haller, JA Laurie, PJ Goodman, JS Ungerleider, WA Emerson, DC Tormey, JH Glick, MH Veeder and JA Maillard. Levamisole and fluorouracil for adjuvant therapy of resected colon carcinoma. New England J of Medicine, 332:352-358, 1990.

CG Moertel, TR Fleming, JS MacDonald, DG Haller, JA Laurie, CM Tangen, JS Ungerleider, WA Emerson, DC Tormey, JH Glick, MH Veeder and JA Maillard, Fluorouracil plus Levamisole as an effective adjuvant therapy after resection of stage II colon carcinoma: a final report. Annals of Internal Med, 122:321-326, 1991.

concordance

Compute the concordance statistic for data or a model

# Description

The concordance statistic compute the agreement between an observed response and a predictor. It is closely related to Kendall's tau-a and tau-b, Goodman's gamma, and Somers' d, all of which can also be calculated from the results of this function.

## **Usage**

```
concordance(object, ...)
## S3 method for class 'formula'
concordance(object, data, weights, subset, na.action,
  cluster, ymin, ymax, timewt= c("n", "S", "S/G", "n/G", "n/G2", "I"),
  influence=0, ranks = FALSE, reverse=FALSE, timefix=TRUE, keepstrata=10, ...)
## S3 method for class 'lm'
concordance(object, ..., newdata, cluster, ymin, ymax,
  influence=0, ranks=FALSE, timefix=TRUE, keepstrata=10)
## S3 method for class 'coxph'
concordance(object, ..., newdata, cluster, ymin, ymax,
  timewt= c("n", "S", "S/G", "n/G", "n/G2", "I"), influence=0,
  ranks=FALSE, timefix=FALSE, keepstrata=10)
## S3 method for class 'survreg'
concordance(object, ..., newdata, cluster, ymin, ymax,
  timewt= c("n", "S", "S/G", "n/G", "n/G2", "I"), influence=0,
  ranks=FALSE, timefix=FALSE, keepstrata=10)
```

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#### **Arguments**

object a fitted model or a formula. The formula should be of the form y ~x or y ~ x + strata(z) with a single numeric or survival response and a single predictor. Counts of concordant, discordant and tied pairs are computed separately per stratum, and then added. data a data.frame in which to interpret the variables named in the formula, or in the subset and the weights argument. Only applicable if object is a formula. optional vector of case weights. Only applicable if object is a formula. weights expression indicating which subset of the rows of data should be used in the fit. subset Only applicable if object is a formula. na.action a missing-data filter function. This is applied to the model.frame after any subset argument has been used. Default is options()\$na.action. Only applicable if object is a formula. multiple fitted models are allowed. Only applicable if object is a model object. newdata optional, a new data frame in which to evaluate (but not refit) the models cluster optional grouping vector for calculating the robust variance compute the concordance over the restricted range ymin <= y <= ymax. (For ymin, ymax survival data this is a time range.) timewt the weighting to be applied. The overall statistic is a weighted mean over event influence 1= return the dfbeta vector, 2= return the full influence matrix, 3 = return both if TRUE, return a data frame containing the individual ranks that make up the ranks reverse if TRUE then assume that larger x values predict smaller response values y; a proportional hazards model is the common example of this. timefix if the response is a Surv object, correct for possible rounding error; otherwise this argument has no effect. See the vignette on tied times for more explanation. For the coxph and survreg methods this issue will have already been addressed in the parent routine, so should not be revisited. keepstrata either TRUE, FALSE, or an integer value. Computations are always done within stratum, then added. If the total number of strata greater than keepstrata, or keepstrata=FALSE, those subtotals are not kept in the output.

## **Details**

At each event time, compute the rank of the subject who had the event as compared to all others with a longer survival, where the rank is value between 0 and 1. The concordance is a weighted mean of these values, determined by the timewt option. For uncensored data each unique response value is compared to all those which are larger.

Using the default value for timewt gives the area under the receiver operating curve (AUC) for a binary response, and (d+1)/2 when y is continuous, where d is Somers' d. For a survival time, timewt of n gives Harrell's c-statistic, which is closely related to the Gehan-Wilcoxon test, S corresponds to the Peto-Wilcoxon, n/G2 is the weighted advocated by Umo, and S/G the weighting proposed by Schemper.

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When the number of strata is very large, such as in a conditional logistic regression for instance (clogit function), a much faster computation is available when the individual strata results are not retained. In the more general case the keepstrata = 10 default simply keeps the printout managable.

#### Value

An object of class concordance containing the following components:

concordance the estimated concordance value or values

count a vector containing the number of concordant pairs, discordant, tied on x but not

y, tied on y but not x, and tied on both x and y

n the number of observations

var a vector containing the estimated variance of the concordance based on the in-

finitesimal jackknife (IJ) method. If there are multiple models it contains the

estimtated variance/covariance matrix.

cvar a vector containing the estimated variance(s) of the concordance values, based

on the variance formula for the associated score test from a proportional hazards model. (This was the primary variance used in the survConcordance function.)

dfbeta optional, the vector of leverage estimates for the concordance

influence optional, the matrix of leverage values for each of the counts, one row per ob-

servation

ranks optional, a data frame containing the Somers' d rank at each event time, along

with the time weight, case weight of the observation with an event, and variance (contribution to the proportional hazards model information matrix). A

weighted mean of the ranks equals Somer's d.

# Author(s)

Terry Therneau

#### See Also

coxph

# **Examples**

```
fit1 <- coxph(Surv(ptime, pstat) ~ age + sex + mspike, mgus2)
concordance(fit1, timewt="n")

# logistic regression
fit2 <- glm(pstat ~ age + sex + mspike, binomial, data= mgus2)
concordance(fit2) # equal to the AUC</pre>
```

30 concordancefit

concord	lancef	it

Compute the concordance

## Description

This is the working routine behind the concordance function. It is not meant to be called by users, but is available for other packages to use. Input arguments, for instance, are assumed to all be the correct length and type, and missing values are not allowed: the calling routine is responsible for these things.

# Usage

```
concordancefit(y, x, strata, weights, ymin = NULL, ymax = NULL,
timewt = c("n", "S", "S/G", "n/G", "n/G2", "I"), cluster, influence =0,
ranks = FALSE, reverse = FALSE, timefix = TRUE, keepstrata=10)
```

# **Arguments**

y the response. It can be numeric, factor, or a Surv object

x the predictor, a numeric vector

strata optional numeric vector that stratifies the data

weights options vector of case weights

ymin, ymax restrict the comparison to response values in this range

timewt the time weighting to be used cluster, influence, ranks, reverse, timefix

see the help for the concordance function

keepstrata either TRUE, FALSE, or an integer value. Computations are always done within

stratum, then added. If the total number of strata greater than keepstrata, or

keepstrata=FALSE, those subtotals are not kept in the output.

# Value

a list containing the results

# Author(s)

Terry Therneau

# See Also

concordance

cox.zph 31

cox.zph Test the Proportional Hazards Assumption of a Cox Regression	cox.zph	Test the Proportional Hazards Assumption of a Cox Regression
--	---------	--

# **Description**

Test the proportional hazards assumption for a Cox regression model fit (coxph).

# Usage

```
cox.zph(fit, transform="km", terms=TRUE, singledf=FALSE, global=TRUE)
```

## **Arguments**

fit	the result of fitting a Cox regression model, using the coxph or coxme functions.
transform	a character string specifying how the survival times should be transformed before the test is performed. Possible values are "km", "rank", "identity" or a function of one argument.
terms	if TRUE, do a test for each term in the model rather than for each separate covariate. For a factor variable with k levels, for instance, this would lead to a k-1 degree of freedom test. The plot for such variables will be a single curve evaluating the linear predictor over time.
singledf	use a single degree of freedom test for terms that have multiple coefficients, i.e., the test that corresponds most closely to the plot. If terms=FALSE this argument has no effect.
global	should a global chi-square test be done, in addition to the per-variable or per-term tests tests.

## **Details**

The computations require the original x matrix of the Cox model fit. Thus it saves time if the x=TRUE option is used in coxph. This function would usually be followed by both a plot and a print of the result. The plot gives an estimate of the time-dependent coefficient  $\beta(t)$ . If the proportional hazards assumption holds then the true  $\beta(t)$  function would be a horizontal line. The table component provides the results of a formal score test for slope=0, a linear fit to the plot would approximate the test

Random effects terms such a frailty or random effects in a coxme model are not checked for proportional hazards, rather they are treated as a fixed offset in model.

If the model contains strata by covariate interactions, then the y matrix may contain structural zeros, i.e., deaths (rows) that had no role in estimation of a given coefficient (column). These are marked as NA. If an entire row is NA, for instance after subscripting a cox.zph object, that row is removed.

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# Value

an object of class "cox.zph", with components:

table	a matrix with one row for each variable, and optionally a last row for the global test. Columns of the matrix contain a score test of for addition of the time-dependent term, the degrees of freedom, and the two-sided p-value.
X	the transformed time axis.
time	the untransformed time values; there is one entry for each event time in the data
strata	for a stratified coxph model, the stratum of each of the events
у	the matrix of scaled Schoenfeld residuals. There will be one column per term or per variable (depending on the terms option above), and one row per event. The row labels are a rounded form of the original times.
var	a variance matrix for the covariates, used to create an approximate standard error band for plots
transform	the transform of time that was used

## Note

call

In versions of the package before survival 3.0 the function computed a fast approximation to the score test. Later versions compute the actual score test.

the calling sequence for the routine.

## References

P. Grambsch and T. Therneau (1994), Proportional hazards tests and diagnostics based on weighted residuals. *Biometrika*, **81**, 515-26.

## See Also

```
coxph, Surv.
```

# Examples

cox	n	h
CUA	ν.	

Fit Proportional Hazards Regression Model

# Description

Fits a Cox proportional hazards regression model. Time dependent variables, time dependent strata, multiple events per subject, and other extensions are incorporated using the counting process formulation of Andersen and Gill.

# Usage

```
coxph(formula, data=, weights, subset,
    na.action, init, control,
    ties=c("efron","breslow","exact"),
    singular.ok=TRUE, robust,
    model=FALSE, x=FALSE, y=TRUE, tt, method=ties,
    id, cluster, istate, statedata, nocenter=c(-1, 0, 1), ...)
```

# **Arguments**

formula	a formula object, with the response on the left of a $\sim$ operator, and the terms on the right. The response must be a survival object as returned by the Surv function.
data	a data.frame in which to interpret the variables named in the formula, or in the subset and the weights argument.
weights	vector of case weights, see the note below. For a thorough discussion of these see the book by Therneau and Grambsch.
subset	expression indicating which subset of the rows of data should be used in the fit. All observations are included by default.
na.action	a missing-data filter function. This is applied to the model.frame after any subset argument has been used. Default is options()\$na.action.
init	vector of initial values of the iteration. Default initial value is zero for all variables.
control	Object of class coxph.control specifying iteration limit and other control options. Default is coxph.control().
ties	a character string specifying the method for tie handling. If there are no tied death times all the methods are equivalent. Nearly all Cox regression programs use the Breslow method by default, but not this one. The Efron approximation is used as the default here, it is more accurate when dealing with tied death times, and is as efficient computationally. The "exact partial likelihood" is equivalent to a conditional logistic model, and is appropriate when the times are a small set of discrete values. See further below.

singular.ok	logical value indicating how to handle collinearity in the model matrix. If TRUE, the program will automatically skip over columns of the X matrix that are linear combinations of earlier columns. In this case the coefficients for such columns will be NA, and the variance matrix will contain zeros. For ancillary calculations, such as the linear predictor, the missing coefficients are treated as zeros.
robust	should a robust variance be computed. The default is TRUE if: there is a cluster argument, there are case weights that are not 0 or 1, or there are id values with more than one event.
id	optional variable name that identifies subjects. Only necessary when a subject can have multiple rows in the data, and there is more than one event type. This variable will normally be found in data.
cluster	optional variable which clusters the observations, for the purposes of a robust variance. If present, it implies robust. This variable will normally be found in data.
istate	optional variable giving the current state at the start each interval. This variable will normally be found in data.
statedata	optional data set used to describe multistate models.
model	logical value: if TRUE, the model frame is returned in component model.
x	logical value: if TRUE, the x matrix is returned in component x.
у	logical value: if TRUE, the response vector is returned in component y.
tt	optional list of time-transform functions.
method	alternate name for the ties argument.
nocenter	columns of the X matrix whose values lie strictly within this set are not recentered

## **Details**

The proportional hazards model is usually expressed in terms of a single survival time value for each person, with possible censoring. Andersen and Gill reformulated the same problem as a counting process; as time marches onward we observe the events for a subject, rather like watching a Geiger counter. The data for a subject is presented as multiple rows or "observations", each of which applies to an interval of observation (start, stop].

Other arguments will be passed to coxph.control

The routine internally scales and centers data to avoid overflow in the argument to the exponential function. These actions do not change the result, but lead to more numerical stability. Any column of the X matrix whose values lie within nocenter list are not recentered. The practical consequence of the default is to not recenter dummy variables corresponding to factors. However, arguments to offset are not scaled since there are situations where a large offset value is a purposefully used. In general, however, users should not avoid very large numeric values for an offset due to possible loss of precision in the estimates.

## Value

an object of class coxph representing the fit. See coxph. object for details.

#### **Side Effects**

Depending on the call, the predict, residuals, and survfit routines may need to reconstruct the x matrix created by coxph. It is possible for this to fail, as in the example below in which the predict function is unable to find tform.

```
tfun <- function(tform) coxph(tform, data=lung)
fit <- tfun(Surv(time, status) ~ age)
predict(fit)</pre>
```

In such a case add the model=TRUE option to the coxph call to obviate the need for reconstruction, at the expense of a larger fit object.

# Case weights

Case weights are treated as replication weights, i.e., a case weight of 2 is equivalent to having 2 copies of that subject's observation. When computers were much smaller grouping like subjects together was a common trick to used to conserve memory. Setting all weights to 2 for instance will give the same coefficient estimate but halve the variance. When the Efron approximation for ties (default) is employed replication of the data will not give exactly the same coefficients as the weights option, and in this case the weighted fit is arguably the correct one.

When the model includes a cluster term or the robust=TRUE option the computed variance treats any weights as sampling weights; setting all weights to 2 will in this case give the same variance as weights of 1.

# Special terms

as a term, in either survival or glm models.

There are three special terms that may be used in the model equation. A strata term identifies a stratified Cox model; separate baseline hazard functions are fit for each strata. The cluster term is used to compute a robust variance for the model. The term + cluster(id) where each value of id is unique is equivalent to specifying the robust=TRUE argument. If the id variable is not unique, it is assumed that it identifies clusters of correlated observations. The robust estimate arises from many different arguments and thus has had many labels. It is variously known as the Huber sandwich estimator, White's estimate (linear models/econometrics), the Horvitz-Thompson estimate (survey sampling), the working independence variance (generalized estimating equations), the infinitesimal jackknife, and the Wei, Lin, Weissfeld (WLW) estimate.

A time-transform term allows variables to vary dynamically in time. In this case the tt argument will be a function or a list of functions (if there are more than one tt() term in the model) giving the appropriate transform. See the examples below.

One user mistake that has recently arisen is to slavishly follow the advice of some coding guides and prepend survival:: onto everthing, including the special terms, e.g., survival::coxph(survival:Surv(time, status) ~ age + survival::cluster(inst),data=lung) First, this is unnecessary: arguments within the coxph call will be evaluated within the survival namespace, so another package's Surv or cluster function would not be noticed. (Full qualification of the coxph call itself may be protective, however.) Second, and more importantly, the call just above will not give the correct answer. The specials are recognized by their name, and survival::cluster is not the same as cluster; the above model would treat inst as an ordinary variable. A similar issue arises from using stats::offset

## Convergence

In certain data cases the actual MLE estimate of a coefficient is infinity, e.g., a dichotomous variable where one of the groups has no events. When this happens the associated coefficient grows at a steady pace and a race condition will exist in the fitting routine: either the log likelihood converges, the information matrix becomes effectively singular, an argument to exp becomes too large for the computer hardware, or the maximum number of interactions is exceeded. (Most often number 1 is the first to occur.) The routine attempts to detect when this has happened, not always successfully. The primary consequence for the user is that the Wald statistic = coefficient/se(coefficient) is not valid in this case and should be ignored; the likelihood ratio and score tests remain valid however.

#### **Ties**

There are three possible choices for handling tied event times. The Breslow approximation is the easiest to program and hence became the first option coded for almost all computer routines. It then ended up as the default option when other options were added in order to "maintain backwards compatability". The Efron option is more accurate if there are a large number of ties, and it is the default option here. In practice the number of ties is usually small, in which case all the methods are statistically indistinguishable.

Using the "exact partial likelihood" approach the Cox partial likelihood is equivalent to that for matched logistic regression. (The clogit function uses the coxph code to do the fit.) It is technically appropriate when the time scale is discrete and has only a few unique values, and some packages refer to this as the "discrete" option. There is also an "exact marginal likelihood" due to Prentice which is not implemented here.

The calculation of the exact partial likelihood is numerically intense. Say for instance 180 subjects are at risk on day 7 of which 15 had an event; then the code needs to compute sums over all 180-choose- $15 > 10^4$ 3 different possible subsets of size 15. There is an efficient recursive algorithm for this task, but even with this the computation can be insufferably long. With (start, stop) data it is much worse since the recursion needs to start anew for each unique start time.

A separate issue is that of artificial ties due to floating-point imprecision. See the vignette on this topic for a full explanation or the timefix option in coxph.control. Users may need to add timefix=FALSE for simulated data sets.

## Penalized regression

coxph can maximise a penalised partial likelihood with arbitrary user-defined penalty. Supplied penalty functions include ridge regression (ridge), smoothing splines (pspline), and frailty models (frailty).

## References

Andersen, P. and Gill, R. (1982). Cox's regression model for counting processes, a large sample study. *Annals of Statistics* **10**, 1100-1120.

Therneau, T., Grambsch, P., Modeling Survival Data: Extending the Cox Model. Springer-Verlag, 2000.

#### See Also

coxph.object, coxph.control, cluster, strata, Surv, survfit, pspline, ridge.

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### **Examples**

```
# Create the simplest test data set
test1 <- list(time=c(4,3,1,1,2,2,3),
              status=c(1,1,1,0,1,1,0),
              x=c(0,2,1,1,1,0,0),
              sex=c(0,0,0,0,1,1,1)
# Fit a stratified model
coxph(Surv(time, status) \sim x + strata(sex), test1)
# Create a simple data set for a time-dependent model
test2 <- list(start=c(1,2,5,2,1,7,3,4,8,8),
              stop=c(2,3,6,7,8,9,9,9,14,17),
              event=c(1,1,1,1,1,1,1,0,0,0),
              x=c(1,0,0,1,0,1,1,1,0,0))
summary(coxph(Surv(start, stop, event) ~ x, test2))
# Create a simple data set for a time-dependent model
test2 <- list(start=c(1, 2, 5, 2, 1, 7, 3, 4, 8, 8),
                stop =c(2, 3, 6, 7, 8, 9, 9, 9, 14, 17),
                event=c(1, 1, 1, 1, 1, 1, 1, 0, 0, 0),
                x = c(1, 0, 0, 1, 0, 1, 1, 1, 0, 0)
summary( coxph( Surv(start, stop, event) ~ x, test2))
# Fit a stratified model, clustered on patients
bladder1 <- bladder[bladder$enum < 5, ]</pre>
coxph(Surv(stop, event) ~ (rx + size + number) * strata(enum),
      cluster = id, bladder1)
# Fit a time transform model using current age
coxph(Surv(time, status) ~ ph.ecog + tt(age), data=lung,
     tt=function(x,t,...) pspline(x + t/365.25))
```

coxph.control

Ancillary arguments for controlling coxph fits

### **Description**

This is used to set various numeric parameters controlling a Cox model fit. Typically it would only be used in a call to coxph.

## Usage

```
coxph.control(eps = 1e-09, toler.chol = .Machine$double.eps^0.75,
iter.max = 20, toler.inf = sqrt(eps), outer.max = 10, timefix=TRUE)
```

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## Arguments

eps	Iteration continues until the relative change in the log partial likelihood is less than eps, or the absolute change is less than sqrt(eps). Must be positive.
toler.chol	Tolerance for detection of singularity during a Cholesky decomposition of the variance matrix, i.e., for detecting a redundant predictor variable.
iter.max	Maximum number of iterations to attempt for convergence.
toler.inf	Tolerance criteria for the warning message about a possible infinite coefficient value.
outer.max	For a penalized coxph model, e.g. with pspline terms, there is an outer loop of iteration to determine the penalty parameters; maximum number of iterations for this outer loop.
timefix	Resolve any near ties in the time variables.

#### **Details**

The convergence tolerances are a balance. Users think they want THE maximum point of the likelihood surface, and for well behaved data sets where this is quadratic near the max a high accuracy is fairly inexpensive: the number of correct digits approximately doubles with each iteration. Conversely, a drop of .0001 from the maximum in any given direction will be correspond to only about 1/20 of a standard error change in the coefficient. Statistically, more precision than this is straining at a gnat. Based on this the author originally had set the tolerance to 1e-5, but relented in the face of multiple "why is the answer different than package X" queries.

Asking for results that are too close to machine precision (double.eps) is a fool's errand; a reasonable critera is often the square root of that precision. The Cholesky decompostion needs to be held to a higher standard than the overall convergence criterion, however. The tolerance.inf value controls a warning message; if it is too small incorrect warnings can appear, if too large some actual cases of an infinite coefficient will not be detected.

The most difficult cases are data sets where the MLE coefficient is infinite; an example is a data set where at each death time, it was the subject with the largest covariate value who perished. In that situation the coefficient increases at each iteration while the log-likelihood asymptotes to a maximum. As iteration proceeds there is a race condition condition for three endpoint: exp(coef) overflows, the Hessian matrix become singular, or the change in loglik is small enough to satisfy the convergence criterion. The first two are difficult to anticipate and lead to numeric diffculties, which is another argument for moderation in the choice of eps.

See the vignette "Roundoff error and tied times" for a more detailed explanation of the timefix option. In short, when time intervals are created via subtraction then two time intervals that are actually identical can appear to be different due to floating point round off error, which in turn can make coxph and survfit results dependent on things such as the order in which operations were done or the particular computer that they were run on. Such cases are unfortunatedly not rare in practice. The timefix=TRUE option adds logic similar to all.equal to ensure reliable results. In analysis of simulated data sets, however, where often by defintion there can be no duplicates, the option will often need to be set to FALSE to avoid spurious merging of close numeric values.

## Value

a list containing the values of each of the above constants

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### See Also

coxph

coxph.detail Details of a Cox Model Fit

## Description

Returns the individual contributions to the first and second derivative matrix, at each unique event time.

# Usage

```
coxph.detail(object, riskmat=FALSE, rorder=c("data", "time"))
```

### **Arguments**

object a Cox model object, i.e., the result of coxph.
riskmat include the at-risk indicator matrix in the output?

rorder if riskmat=TRUE, the rows of riskmat will be in the original data order, other-

wise sorted by time within strata.

### **Details**

This function may be useful for those who wish to investigate new methods or extensions to the Cox model. The example below shows one way to calculate the Schoenfeld residuals.

### Value

a list with components

time the vector of unique event times

nevent the number of events at each of these time points.

means a matrix with one row for each event time and one column for each variable in

the Cox model, containing the weighted mean of the variable at that time, over all subjects still at risk at that time. The weights are the risk weights exp(x %\*%

fit\$coef).

nrisk number of subjects at risk.

score the contribution to the score vector (first derivative of the log partial likelihood)

at each time point.

imat the contribution to the information matrix (second derivative of the log partial

likelihood) at each time point.

hazard the hazard increment. Note that the hazard and variance of the hazard are always

for some particular future subject. This routine uses object\$mean as the future

subject.

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varhaz the variance of the hazard increment.

x,y copies of the input data.

strata only present for a stratified Cox model, this is a table giving the number of time points of component time that were contributed by each of the strata.

riskmat a matrix with one row for each observation and one colum for each unique event time, containing a 0/1 value to indicate whether that observation was (1) or was not (0) at risk at the given time point. Rows are in the order of the original data (after removal of any missings by coxph), or in time order.

# See Also

```
coxph, residuals.coxph
```

## **Examples**

```
fit <- coxph(Surv(futime,fustat) ~ age + rx + ecog.ps, ovarian, x=TRUE)
fitd <- coxph.detail(fit)
# There is one Schoenfeld residual for each unique death. It is a
# vector (covariates for the subject who died) - (weighted mean covariate
# vector at that time). The weighted mean is defined over the subjects
# still at risk, with exp(X beta) as the weight.

events <- fit$y[,2]==1
etime <- fit$y[events,1] #the event times --- may have duplicates
indx <- match(etime, fitd$time)
schoen <- fit$x[events,] - fitd$means[indx,]</pre>
```

coxph.object

Proportional Hazards Regression Object

### **Description**

This class of objects is returned by the coxph class of functions to represent a fitted proportional hazards model. Objects of this class have methods for the functions print, summary, residuals, predict and survfit.

## **Arguments**

coefficients	the vector of coefficients. If the model is over-determined there will be missing values in the vector corresponding to the redundant columns in the model matrix.
var	the variance matrix of the coefficients. Rows and columns corresponding to any missing coefficients are set to zero.
naive.var	this component will be present only if the robust option was true. If so, the var component will contain the robust estimate of variance, and this component will contain the ordinary estimate.
loglik	a vector of length 2 containing the log-likelihood with the initial values and with the final values of the coefficients.

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score value of the efficient score test, at the initial value of the coefficients.

rscore the robust log-rank statistic, if a robust variance was requested.

wald.test the Wald test of whether the final coefficients differ from the initial values.

iter number of iterations used.

linear.predictors

the vector of linear predictors, one per subject. Note that this vector has been

centered, see predict.coxph for more details.

residuals the martingale residuals.

means vector of column means of the X matrix. Subsequent survival curves are ad-

justed to this value.

n the number of observations used in the fit.

nevent the number of events (usually deaths) used in the fit.

concordance a vector of length 6, containing the number of pairs that are concordant, discor-

dant, tied on x, tied on y, and tied on both, followed by the standard error of the

concordance statistic.

first the first derivative vector at the solution.

weights the vector of case weights, if one was used.

method the method used for handling tied survival times.

na.action the na.action attribute, if any, that was returned by the na.action routine.

timefix the value of the timefix option used in the fit

cmap the coefficient map, present for multi-state coxph fits. There a column for each

transition and a row for each coefficient, the value maps that transition/coefficient pair to a position in the coefficient vector. If a particular covariate is not used by the transition the matrix will contain a zero, if two transitions share a coefficient

the matrix will contain repeats.

stratum\_map stratum mapping, present for multi-state coxph fits. The row labeled '(Baseline)'

identifies transitions that do or do not share a baseline stratum. Further rows correspond to strata() terms in the model, each of which may apply to some

transitions and not others.

... The object will also contain the following, for documentation see the 1m object:

terms, assign, formula, call, and, optionally, x, y, and/or frame.

### **Components**

The following components must be included in a legitimate coxph object.

### See Also

coxph, coxph.detail, cox.zph, residuals.coxph, survfit, survreg.

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coxph.wtest

Compute a quadratic form

# **Description**

This function is used internally by several survival routines. It computes a simple quadratic form, while properly dealing with missings.

# Usage

```
coxph.wtest(var, b, toler.chol = 1e-09)
```

# **Arguments**

var variance matrix

b vector

toler.chol tolerance for the internal cholesky decomposition

### **Details**

Compute b' V-inverse b. Equivalent to sum(b \* solve(V,b)), except for the case of redundant covariates in the original model, which lead to NA values in V and b.

# Value

a real number

## Author(s)

Terry Therneau

coxsurv.fit

A direct interface to the 'computational engine' of survfit.coxph

# Description

This program is mainly supplied to allow other packages to invoke the survfit.coxph function at a 'data' level rather than a 'user' level. It does no checks on the input data that is provided, which can lead to unexpected errors if that data is wrong.

# Usage

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### **Arguments**

stype	survival curve computation: 1=direct, 2=exp(-cumulative hazard)
ctype	cumulative hazard computation: 1=Breslow, 2=Efron
se.fit	if TRUE, compute standard errors
varmat	the variance matrix of the coefficients
cluster	vector to control robust variance
У	the response variable used in the Cox model. (Missing values removed of course.)
х	covariate matrix used in the Cox model
wt	weight vector for the Cox model. If the model was unweighted use a vector of 1s.
risk	the risk score $exp(X beta + offset)$ from the fitted Cox model.
position	optional argument controlling what is counted as 'censored'. Due to time dependent covariates, for instance, a subject might have start, stop times of $(1,5)(5,30)(30,100)$ . Times 5 and 30 are not 'real' censorings. Position is 1 for a real start, 2 for an actual end, 3 for both, 0 for neither.
strata	strata variable used in the Cox model. This will be a factor.
oldid	identifier for subjects with multiple rows in the original data.
y2, x2, risk2, s	trata2
	variables for the hypothetical subjects, for which prediction is desired
id2	optional; if present and not NULL this should be a vector of identifiers of length nrow(x2). A non-null value signifies that x2 contains time dependent covariates, in which case this identifies which rows of x2 go with each subject.
unlist	if FALSE the result will be a list with one element for each strata. Otherwise the

### Value

a list containing nearly all the components of a survfit object. All that is missing is to add the confidence intervals, the type of the original model's response (as in a coxph object), and the class.

strata are "unpacked" into the form found in a survfit object.

# Note

The source code for for both this function and survfit.coxph is written using noweb. For complete documentation see the inst/sourcecode.pdf file.

# Author(s)

Terry Therneau

### See Also

survfit.coxph

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diabetic

Ddiabetic retinopathy

### **Description**

Partial results from a trial of laser coagulation for the treatment of diabetic retinopathy.

## Usage

```
diabetic
data(diabetic, package="survival")
```

#### **Format**

A data frame with 394 observations on the following 8 variables.

```
id subject id
laser laser type: xenon or argon
age age at diagnosis
eye a factor with levels of left right
trt treatment: 0 = no treatment, 1= laser
risk risk group of 6-12
time time to event or last follow-up
status status of 0= censored or 1 = visual loss
```

### **Details**

The 197 patients in this dataset were a 50% random sample of the patients with "high-risk" diabetic retinopathy as defined by the Diabetic Retinopathy Study (DRS). Each patient had one eye randomized to laser treatment and the other eye received no treatment. For each eye, the event of interest was the time from initiation of treatment to the time when visual acuity dropped below 5/200 two visits in a row. Thus there is a built-in lag time of approximately 6 months (visits were every 3 months). Survival times in this dataset are therefore the actual time to blindness in months, minus the minimum possible time to event (6.5 months). Censoring was caused by death, dropout, or end of the study.

## References

```
Huster, Brookmeyer and Self, Biometrics, 1989.
American Journal of Ophthalmology, 1976, 81:4, pp 383-396
```

### **Examples**

dsurvreg 45

dsurvreg	Distributions available in survreg.	

### Description

Density, cumulative distribution function, quantile function and random generation for the set of distributions supported by the survreg function.

## Usage

```
dsurvreg(x, mean, scale=1, distribution='weibull', parms)
psurvreg(q, mean, scale=1, distribution='weibull', parms)
qsurvreg(p, mean, scale=1, distribution='weibull', parms)
rsurvreg(n, mean, scale=1, distribution='weibull', parms)
```

### **Arguments**

V	vector of quantiles. Missing values (NAs) are allowed.
Х	vector of quantities. Wissing values (WAS) are anowed.
q	vector of quantiles. Missing values (NAs) are allowed.
р	vector of probabilities. Missing values (NAs) are allowed.
n	number of random deviates to produce
mean	vector of linear predictors for the model. This is replicated to be the same length as p, q or n.
scale	vector of (positive) scale factors. This is replicated to be the same length as $p$ , $q$ or $n$ .
distribution	character string giving the name of the distribution. This must be one of the elements of survreg.distributions
parms	optional parameters, if any, of the distribution. For the t-distribution this is the degrees of freedom.

### **Details**

Elements of q or p that are missing will cause the corresponding elements of the result to be missing. The location and scale values are as they would be for survreg. The label "mean" was an unfortunate choice (made in mimicry of qnorm); since almost none of these distributions are symmetric it will not actually be a mean, but corresponds instead to the linear predictor of a fitted model. Translation to the usual parameterization found in a textbook is not always obvious. For example, the Weibull distribution is fit using the Extreme value distribution along with a log transformation. Letting  $F(t) = 1 - \exp[-(at)^p]$  be the cumulative distribution of the Weibull using a standard parameterization in terms of a and p, the survreg location corresponds to  $-\log(a)$  and the scale to 1/p (Kalbfleisch and Prentice, section 2.2.2).

#### Value

density (dsurvreg), probability (psurvreg), quantile (qsurvreg), or for the requested distribution with mean and scale parameters mean and sd.

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### References

Kalbfleisch, J. D. and Prentice, R. L. (1970). *The Statistical Analysis of Failure Time Data* Wiley, New York

### See Also

```
survreg, Normal
```

## **Examples**

```
# List of distributions available
names(survreg.distributions)
## Not run:
[1] "extreme"
                   "logistic"
                                 "gaussian"
                                                "weibull"
                                                              "exponential"
[6] "rayleigh"
                   "loggaussian" "lognormal"
                                                "loglogistic" "t"
## End(Not run)
# Compare results
all.equal(dsurvreg(1:10, 2, 5, dist='lognormal'), dlnorm(1:10, 2, 5))
# Hazard function for a Weibull distribution
   <- seq(.1, 3, length=30)
haz <- dsurvreg(x, 2, 3)/ (1-psurvreg(x, 2, 3))
## Not run:
plot(x, haz, log='xy', ylab="Hazard") #line with slope (1/scale -1)
## End(Not run)
```

finegray

Create data for a Fine-Gray model

# Description

The Fine-Gray model can be fit by first creating a special data set, and then fitting a weighted Cox model to the result. This routine creates the data set.

# Usage

```
finegray(formula, data, weights, subset, na.action= na.pass, etype,
    prefix="fg", count, id, timefix=TRUE)
```

### **Arguments**

formula a standard model formula, with survival on the left and covariates on the right.

data an optional data frame, list or environment (or object coercible by as.data.frame to a data frame) containing the variables in the model.

weights optional vector of observation weights

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subset	an optional vector specifying a subset of observations to be used in the fitting process.
na.action	a function which indicates what should happen when the data contain NAs. The default is set by the na.action setting of options.
etype	the event type for which a data set will be generated. The default is to use whichever is listed first in the multi-state survival object.
prefix	the routine will add 4 variables to the data set: a start and end time for each interval, status, and a weight for the interval. The default names of these are "fgstart", "fgstop", "fgstatus", and "fgwt"; the prefix argument determines the initial portion of the new names.
count	a variable name in the output data set for an optional variable that will contain the the replication count for each row of the input data. If a row is expanded into multiple lines it will contain 1, 2, etc.
id	optional, the variable name in the data set which identifies subjects.
timefix	process times through the aeqSurv function to eliminate potential roundoff issues.

### **Details**

The function expects a multi-state survival expression or variable as the left hand side of the formula, e.g. Surv(atime, astat) where astat is a factor whose first level represents censoring and remaining levels are states. The output data set will contain simple survival data (status = 0 or 1) for a single endpoint of interest. In the output data set subjects who did not experience the event of interest become censored subjects whose times are artificially extended over multiple intervals, with a decreasing case weight from interval to interval. The output data set will normally contain many more rows than the input.

Time dependent covariates are allowed, but not (currently) delayed entry. If there are time dependent covariates, e.g.., the input data set had Surv(entry,exit,stat) as the left hand side, then an id statement is required. The program does data checks in this case, and needs to know which rows belong to each subject.

The output data set will often have gaps. Say that there were events at time 50 and 100 (and none between) and censoring at 60, 70, and 80. Formally, a non event subjects at risk from 50 to 100 will have different weights in each of the 3 intervals 50-60, 60-70, and 80-100, but because the middle interval does not span any event times the subsequent Cox model will never use that row. The finegray output omits such rows.

See the competing risks vignette for more details.

## Value

a data frame

### Author(s)

Terry Therneau

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### References

Fine JP and Gray RJ (1999) A proportional hazards model for the subdistribution of a competing risk. JASA 94:496-509.

Geskus RB (2011). Cause-Specific Cumulative Incidence Estimation and the Fine and Gray Model Under Both Left Truncation and Right Censoring. Biometrics 67, 39-49.

### See Also

```
coxph, aeqSurv
```

### **Examples**

flchain

Assay of serum free light chain for 7874 subjects.

## Description

This is a stratified random sample containing 1/2 of the subjects from a study of the relationship between serum free light chain (FLC) and mortality. The original sample contains samples on approximately 2/3 of the residents of Olmsted County aged 50 or greater.

### Usage

```
flchain
data(flchain, package="survival")
```

# Format

A data frame with 7874 persons containing the following variables.

```
age age in years
sex F=female, M=male
sample.yr the calendar year in which a blood sample was obtained
```

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kappa serum free light chain, kappa portion

lambda serum free light chain, lambda portion

flc.grp the FLC group for the subject, as used in the original analysis

creatinine serum creatinine

mgus 1 if the subject had been diagnosed with monoclonal gammapothy (MGUS)

futime days from enrollment until death. Note that there are 3 subjects whose sample was obtained on their death date.

death 0=alive at last contact date, 1=dead

chapter for those who died, a grouping of their primary cause of death by chapter headings of the International Code of Diseases ICD-9

#### **Details**

In 1995 Dr. Robert Kyle embarked on a study to determine the prevalence of monoclonal gammopathy of undetermined significance (MGUS) in Olmsted County, Minnesota, a condition which is normally only found by chance from a test (serum electrophoresis) which is ordered for other causes. Later work suggested that one component of immunoglobulin production, the serum free light chain, might be a possible marker for immune disregulation. In 2010 Dr. Angela Dispenzieri and colleagues assayed FLC levels on those samples from the original study for which they had patient permission and from which sufficient material remained for further testing. They found that elevated FLC levels were indeed associated with higher death rates.

Patients were recruited when they came to the clinic for other appointments, with a final random sample of those who had not yet had a visit since the study began. An interesting side question is whether there are differences between early, mid, and late recruits.

This data set contains an age and sex stratified random sample that includes 7874 of the original 15759 subjects. The original subject identifiers and dates have been removed to protect patient identity. Subsampling was done to further protect this information.

# Source

The primary investigator (A Dispenzieri) and statistician (T Therneau) for the study.

### References

A Dispenzieri, J Katzmann, R Kyle, D Larson, T Therneau, C Colby, R Clark, G Mead, S Kumar, LJ Melton III and SV Rajkumar (2012). Use of monclonal serum immunoglobulin free light chains to predict overall survival in the general population, Mayo Clinic Proceedings 87:512-523.

R Kyle, T Therneau, SV Rajkumar, D Larson, M Plevak, J Offord, A Dispenzieri, J Katzmann, and LJ Melton, III, 2006, Prevalence of monoclonal gammopathy of undetermined significance, New England J Medicine 354:1362-1369.

# **Examples**

50 frailty

frailty	Random effects terms	

# Description

The frailty function allows one to add a simple random effects term to a Cox model.

# Usage

# **Arguments**

eps

x	the variable to be entered as a random effect. It is always treated as a factor.
distribution	either the gamma, gaussian or t distribution may be specified. The routines frailty.gamma, frailty.gaussian and frailty.t do the actual work.
	Arguments for specific distribution, including (but not limited to)
sparse	cutoff for using a sparse coding of the data matrix. If the total number of levels of x is larger than this value, then a sparse matrix approximation is used. The correct cutoff is still a matter of exploration: if the number of levels is very large (thousands) then the non-sparse calculation may not be feasible in terms of both memory and compute time. Likewise, the accuracy of the sparse approximation appears to be related to the maximum proportion of subjects in any one class, being best when no one class has a large membership.
theta	if specified, this fixes the variance of the random effect. If not, the variance is a parameter, and a best solution is sought. Specifying this implies method='fixed'.
df	if specified, this fixes the degrees of freedom for the random effect. Specifying this implies method='df'. Only one of theta or df should be specified.
method	the method used to select a solution for theta, the variance of the random effect. The fixed corresponds to a user-specified value, and no iteration is done. The df selects the variance such that the degrees of freedom for the random effect matches a user specified value. The aic method seeks to maximize Akaike's information criteria 2*(partial likelihood - df). The em and reml methods are specific to Cox models with gamma and gaussian random effects, respectively. Please see further discussion below.
tdf	the degrees of freedom for the t-distribution.

convergence criteria for the iteration on theta.

frailty 51

### **Details**

The frailty plugs into the general penalized modeling framework provided by the coxph and survreg routines. This framework deals with likelihood, penalties, and degrees of freedom; these aspects work well with either parent routine.

Therneau, Grambsch, and Pankratz show how maximum likelihood estimation for the Cox model with a gamma frailty can be accomplished using a general penalized routine, and Ripatti and Palmgren work through a similar argument for the Cox model with a gaussian frailty. Both of these are specific to the Cox model. Use of gamma/ml or gaussian/reml with survreg does not lead to valid results.

The extensible structure of the penalized methods is such that the penalty function, such as frailty or pspine, is completely separate from the modeling routine. The strength of this is that a user can plug in any penalization routine they choose. A weakness is that it is very difficult for the modeling routine to know whether a sensible penalty routine has been supplied.

Note that use of a frailty term implies a mixed effects model and use of a cluster term implies a GEE approach; these cannot be mixed.

The coxme package has superseded this method. It is faster, more stable, and more flexible.

#### Value

this function is used in the model statement of either coxph or survreg. It's results are used internally.

### References

S Ripatti and J Palmgren, Estimation of multivariate frailty models using penalized partial likelihood, Biometrics, 56:1016-1022, 2000.

T Therneau, P Grambsch and VS Pankratz, Penalized survival models and frailty, J Computational and Graphical Statistics, 12:156-175, 2003.

### See Also

coxph, survreg

### **Examples**

52 gbsg

gbsg

Breast cancer data sets used in Royston and Altman (2013)

## **Description**

The gbsg data set contains patient records from a 1984-1989 trial conducted by the German Breast Cancer Study Group (GBSG) of 720 patients with node positive breast cancer; it retains the 686 patients with complete data for the prognostic variables.

### Usage

```
gbsg
data(cancer, package="survival")
```

### **Format**

A data set with 686 observations and 11 variables.

```
pid patient identifier
age age, years
meno menopausal status (0= premenopausal, 1= postmenopausal)
size tumor size, mm
grade tumor grade
nodes number of positive lymph nodes
pgr progesterone receptors (fmol/l)
er estrogen receptors (fmol/l)
hormon hormonal therapy, 0= no, 1= yes
rfstime recurrence free survival time; days to first of reccurence, death or last follow-up
status 0= alive without recurrence, 1= recurrence or death
```

### **Details**

These data sets are used in the paper by Royston and Altman. The Rotterdam data is used to create a fitted model, and the GBSG data for validation of the model. The paper gives references for the data source.

# References

Patrick Royston and Douglas Altman, External validation of a Cox prognostic model: principles and methods. BMC Medical Research Methodology 2013, 13:33

### See Also

rotterdam

heart 53

heart

Stanford Heart Transplant data

## **Description**

Survival of patients on the waiting list for the Stanford heart transplant program.

# Usage

```
heart
data(heart, package="survival")
```

### **Format**

jasa: original data

birth.dt: birth date

accept.dt: acceptance into program

tx.date: transplant date fu.date: end of followup fustat: dead or alive

surgery: prior bypass surgery age: age (in years) futime: followup time

wait.time: time before transplant transplant: transplant indicator mismatch: mismatch score

hla.a2: particular type of mismatch mscore: another mismatch score reject: rejection occurred

# jasa1, heart: processed data

start, stop, event: Entry and exit time and status for this interval of time

age: age-48 years

year: year of acceptance (in years after 1 Nov 1967)

surgery: prior bypass surgery 1=yes transplant: received transplant 1=yes

id: patient id

### **Source**

J Crowley and M Hu (1977), Covariance analysis of heart transplant survival data. *Journal of the American Statistical Association*, **72**, 27–36.

54 is.ratetable

## See Also

stanford2

is.ratetable

Verify that an object is of class ratetable.

# Description

The function verifies not only the class attribute, but the structure of the object.

# Usage

```
is.ratetable(x, verbose=FALSE)
```

# **Arguments**

x the object to be verified.

verbose

if TRUE and the object is not a ratetable, then return a character string describing the way(s) in which x fails to be a proper ratetable object.

# **Details**

Rate tables are used by the pyears and survexp functions, and normally contain death rates for some population, categorized by age, sex, or other variables. They have a fairly rigid structure, and the verbose option can help in creating a new rate table.

## Value

returns TRUE if x is a ratetable, and FALSE or a description if it is not.

## See Also

```
pyears, survexp.
```

# **Examples**

```
is.ratetable(survexp.us) # True
is.ratetable(lung) # False
```

kidney 55

kidney

Kidney catheter data

## Description

Data on the recurrence times to infection, at the point of insertion of the catheter, for kidney patients using portable dialysis equipment. Catheters may be removed for reasons other than infection, in which case the observation is censored. Each patient has exactly 2 observations.

This data has often been used to illustrate the use of random effects (frailty) in a survival model. However, one of the males (id 21) is a large outlier, with much longer survival than his peers. If this observation is removed no evidence remains for a random subject effect.

# Usage

```
kidney
data(cancer, package="survival")
```

#### **Format**

patient: id
time: time
status: event status
age: in years
sex: 1=male, 2=female

disease: disease type (0=GN, 1=AN, 2=PKD, 3=Other)

frail: frailty estimate from original paper

### Note

The original paper ignored the issue of tied times and so is not exactly reproduced by the survival package.

## Source

CA McGilchrist, CW Aisbett (1991), Regression with frailty in survival analysis. *Biometrics* **47**, 461–66.

# Examples

```
kfit <- coxph(Surv(time, status)~ age + sex + disease + frailty(id), kidney)
kfit0 <- coxph(Surv(time, status)~ age + sex + disease, kidney)
kfitm1 <- coxph(Surv(time, status) ~ age + sex + disease +
frailty(id, dist='gauss'), kidney)</pre>
```

56 lines.survfit

levels.Surv

Return the states of a multi-state Surv object

## **Description**

For a multi-state Surv object, this will return the names of the states.

# Usage

```
## S3 method for class 'Surv'
levels(x)
```

## **Arguments**

Х

a Surv object

#### Value

for a multi-state Surv object, the vector of state names (excluding censoring); or NULL for an ordinary Surv object

# **Examples**

lines.survfit

Add Lines or Points to a Survival Plot

# Description

Often used to add the expected survival curve(s) to a Kaplan-Meier plot generated with plot.survfit.

# Usage

lines.survfit 57

### **Arguments**

x a survival object, generated from the survfit or survexp functions.

type the line type, as described in lines. The default is a step function for survfit

objects, and a connected line for survexp objects. All other arguments for

lines.survexp are identical to those for lines.survfit.

col, lty, lwd, cex

vectors giving the mark symbol, color, line type, line width and character size

for the added curves. Of this set only color is applicable to points.

pch plotting characters for points, in the style of matplot, i.e., either a single string

of characters of which the first will be used for the first curve, etc; or a vector of

characters or integers, one element per curve.

mark a historical alias for pch

censor should censoring times be displayed for the points function?

mark.time controls the labeling of the curves. If FALSE, no labeling is done. If TRUE, then

curves are marked at each censoring time. If mark.time is a numeric vector,

then curves are marked at the specified time points.

xmax optional cutoff for the right hand of the curves.

fun an arbitrary function defining a transformation of the survival curve. For ex-

ample fun=log is an alternative way to draw a log-survival curve (but with the axis labeled with  $\log(S)$  values). Four often used transformations can be specified with a character argument instead: "log" is the same as using the log=T option, "event" plots cumulative events (f(y) = 1-y), "cumhaz" plots the cumulative hazard function  $(f(y) = -\log(y))$  and "cloglog" creates a complimentary

log-log survival plot  $(f(y) = \log(-\log(y)))$  along with log scale for the x-axis.

conf.int if TRUE, confidence bands for the curves are also plotted. If set to "only", then

only the CI bands are plotted, and the curve itself is left off. This can be useful for fine control over the colors or line types of a plot. A numeric value, e.g.

conf.int = .90, can be used to

conf. times optional vector of times at which to place a confidence bar on the curve(s). If

present, these will be used instead of confidence bands.

conf. cap width of the horizontal cap on top of the confidence bars; only used if conf.times

is used. A value of 1 is the width of the plot region.

conf.offset the offset for confidence bars, when there are multiple curves on the plot. A

value of 1 is the width of the plot region. If this is a single number then each curve's bars are offset by this amount from the prior curve's bars, if it is a vector

the values are used directly.

58 lines.survfit

conf.type	One of "plain", "log" (the default), "log-log", "logit", or "none". Only enough of the string to uniquely identify it is necessary. The first option causes
	confidence intervals not to be generated. The second causes the standard inter-
	vals curve +-k *se(curve), where k is determined from conf.int. The log
	option calculates intervals based on the cumulative hazard or log(survival). The
	log-log option bases the intervals on the log hazard or log(-log(survival)), and
	the logit option on log(survival/(1-survival)).
noplot	for multi-state models, curves with this label will not be plotted. The default corresponds to an unspecified state.
cumhaz	plot the cumulative hazard, rather than the survival or probability in state.
	other graphical parameters

### **Details**

When the survfit function creates a multi-state survival curve the resulting object has class 'survfitms'. The only difference in the plots is that that it defaults to a curve that goes from lower left to upper right (starting at 0), where survival curves default to starting at 1 and going down. All other options are identical.

If the user set an explicit range in an earlier plot.survfit call, e.g. via xlim or xmax, subsequent calls to this function remember the right hand cutoff. This memory can be erased by options(plot.survfit) <-NULL.

### Value

a list with components x and y, containing the coordinates of the last point on each of the curves (but not of the confidence limits). This may be useful for labeling.

### **Side Effects**

one or more curves are added to the current plot.

## See Also

```
lines, par, plot. survfit, survfit, survexp.
```

### **Examples**

logan 59

```
text(temp, c("Male", "Female"), adj= -.1) #labels just past the ends
title(main="Primary Biliary Cirrhosis, Observed and Expected")
```

logan

Data from the 1972-78 GSS data used by Logan

# Description

Intergenerational occupational mobility data with covariates.

# Usage

```
logan
data(logan, package="survival")
```

### **Format**

A data frame with 838 observations on the following 4 variables.

**occupation** subject's occupation, a factor with levels farm, operatives, craftsmen, sales, and professional

focc father's occupation

education total years of schooling, 0 to 20

race levels of non-black and black

# Source

General Social Survey data, see the web site for detailed information on the variables. https://gss.norc.org/.

# References

Logan, John A. (1983). A Multivariate Model for Mobility Tables. *American Journal of Sociology* 89: 324-349.

60 logLik.coxph

logLik.coxph

logLik method for a Cox model

# Description

The logLik function for survival models

# Usage

```
## S3 method for class 'coxph'
logLik(object, ...)
## S3 method for class 'survreg'
logLik(object, ...)
```

## Arguments

object the result of a coxph or survreg fit

... optional arguments for other instances of the method

### **Details**

The logLik function is used by summary functions in R such as AIC. For a Cox model, this method returns the partial likelihood. The number of degrees of freedom (df) used by the fit and the effective number of observations (nobs) are added as attributes. Per Raftery and others, the effective number of observations is the taken to be the number of events in the data set.

For a survreg model the proper value for the effective number of observations is still an open question (at least to this author). For right censored data the approach of logLik.coxph is the possible the most sensible, but for interval censored observations the result is unclear. The code currently does not add a *nobs* attribute.

## Value

an object of class logLik

# Author(s)

Terry Therneau

### References

Robert E. Kass and Adrian E. Raftery (1995). "Bayes Factors". J. American Statistical Assoc. 90 (430): 791.

Raftery A.E. (1995), "Bayesian Model Selection in Social Research", Sociological methodology, 111-196.

### See Also

logLik

lung 61

lung NCCTG Lung Cancer Data

# **Description**

Survival in patients with advanced lung cancer from the North Central Cancer Treatment Group. Performance scores rate how well the patient can perform usual daily activities.

# Usage

```
lung
data(cancer, package="survival")
```

### **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 as rated by the physician. 0=asymptomatic, 1= symptomatic but completely ambulator

ph.karno: Karnofsky performance score (bad=0-good=100) rated by physician

pat.karno: Karnofsky performance score as rated by patient

meal.cal: Calories consumed at meals wt.loss: Weight loss in last six months

# Note

The use of 1/2 for alive/dead instead of the usual 0/1 is a historical footnote. For data contained on punch cards, IBM 360 Fortran treated blank as a zero, which led to a policy within the section of Biostatistics to never use "0" as a data value since one could not distinguish it from a missing value. The policy became a habit, as is often the case; and the 1/2 coding endured long beyond the demise of punch cards and Fortran.

# Source

Terry Therneau

### References

Loprinzi CL. Laurie JA. Wieand HS. Krook JE. Novotny PJ. Kugler JW. Bartel J. Law M. Bateman M. Klatt NE. et al. Prospective evaluation of prognostic variables from patient-completed questionnaires. North Central Cancer Treatment Group. Journal of Clinical Oncology. 12(3):601-7, 1994.

62 mgus

mgus	Monoclonal gammopathy data	

## Description

Natural history of 241 subjects with monoclonal gammopathy of undetermined significance (MGUS).

## Usage

```
mgus
mgus1
data(cancer, package="survival")
```

### **Format**

mgus: A data frame with 241 observations on the following 12 variables.

id: subject id

age: age in years at the detection of MGUS

sex: male or female dxyr: year of diagnosis

pcdx: for subjects who progress to a plasma cell malignancy

the subtype of malignancy: multiple myeloma (MM) is the

most common, followed by amyloidosis (AM), macroglobulinemia (MA),

and other lymphprolifative disorders (LP)

pctime: days from MGUS until diagnosis of a plasma cell malignancy

futime: days from diagnosis to last follow-up

death: 1= follow-up is until death
alb: albumin level at MGUS diagnosis
creat: creatinine at MGUS diagnosis
hgb: hemoglobin at MGUS diagnosis

mspike: size of the monoclonal protein spike at diagnosis

mgus1: The same data set in start,stop format. Contains the id, age, sex, and laboratory variable described above along with

start, stop: sequential intervals of time for each subject

status: =1 if the interval ends in an event

event: a factor containing the event type: censor, death, or plasma cell malignancy

enum: event number for each subject: 1 or 2

## **Details**

Plasma cells are responsible for manufacturing immunoglobulins, an important part of the immune defense. At any given time there are estimated to be about  $10^6$  different immunoglobulins in the circulation at any one time. When a patient has a plasma cell malignancy the distribution will

mgus2 63

become dominated by a single isotype, the product of the malignant clone, visible as a spike on a serum protein electrophoresis. Monoclonal gammopathy of undertermined significance (MGUS) is the presence of such a spike, but in a patient with no evidence of overt malignancy. This data set of 241 sequential subjects at Mayo Clinic was the groundbreaking study defining the natural history of such subjects. Due to the diligence of the principle investigator 0 subjects have been lost to follow-up.

Three subjects had MGUS detected on the day of death. In data set mgus1 these subjects have the time to MGUS coded as .5 day before the death in order to avoid tied times.

These data sets were updated in Jan 2015 to correct some small errors.

### **Source**

Mayo Clinic data courtesy of Dr. Robert Kyle.

#### References

R Kyle, Benign monoclonal gammopathy – after 20 to 35 years of follow-up, Mayo Clinic Proc 1993; 68:26-36.

# **Examples**

mgus2

Monoclonal gammopathy data

## **Description**

Natural history of 1341 sequential patients with monoclonal gammopathy of undetermined significance (MGUS). This is a superset of the mgus data, at a later point in the accrual process

### Usage

```
mgus2
data(cancer, package="survival")
```

64 model.frame.coxph

### **Format**

```
A data frame with 1384 observations on the following 10 variables.
```

```
id subject identifier
age age at diagnosis, in years
sex a factor with levels F M
dxyr year of diagnosis
hgb hemoglobin
creat creatinine
mspike size of the monoclonal serum splike
ptime time until progression to a plasma cell malignancy (PCM) or last contact, in months
pstat occurrence of PCM: 0=no, 1=yes
futime time until death or last contact, in months
death occurrence of death: 0=no, 1=yes
```

### **Details**

This is an extension of the study found in the mgus data set, containing enrollment through 1994 and follow-up through 1999.

### **Source**

Mayo Clinic data courtesy of Dr. Robert Kyle. All patient identifiers have been removed, age rounded to the nearest year, and follow-up times rounded to the nearest month.

### References

R. Kyle, T. Therneau, V. Rajkumar, J. Offord, D. Larson, M. Plevak, and L. J. Melton III, A long-terms study of prognosis in monoclonal gammopathy of undertermined significance. New Engl J Med, 346:564-569 (2002).

model.frame.coxph

Model.frame method for coxph objects

## **Description**

Recreate the model frame of a coxph fit.

### Usage

```
## S3 method for class 'coxph'
model.frame(formula, ...)
```

model.matrix.coxph 65

## **Arguments**

```
formula the result of a coxph fit
... other arguments to model.frame
```

### **Details**

For details, see the manual page for the generic function. This function would rarely be called by a user, it is mostly used inside functions like residual that need to recreate the data set from a model in order to do further calculations.

### Value

the model frame used in the original fit, or a parallel one for new data.

### Author(s)

Terry Therneau

#### See Also

```
model.frame
```

model.matrix.coxph

Model.matrix method for coxph models

# **Description**

Reconstruct the model matrix for a cox model.

# Usage

```
## S3 method for class 'coxph'
model.matrix(object, data=NULL, contrast.arg =
  object$contrasts, ...)
```

# **Arguments**

object the result of a coxph model

data optional, a data frame from which to obtain the data

contrast.arg optional, a contrasts object describing how factors should be coded

... other possible argument to model.frame

### **Details**

When there is a data argument this function differs from most of the other model.matrix methods in that the response variable for the original formula is *not* required to be in the data.

If the data frame contains a terms attribute then it is assumed to be the result of a call to model.frame, otherwise a call to model.frame is applied with the data as an argument.

66 myeloid

# Value

The model matrix for the fit

### Author(s)

Terry Therneau

### See Also

```
model.matrix
```

# **Examples**

myeloid

Acute myeloid leukemia

# **Description**

This simulated data set is based on a trial in acute myeloid leukemia.

# Usage

```
myeloid
data(cancer, package="survival")
```

## **Format**

A data frame with 646 observations on the following 9 variables.

```
id subject identifier, 1-646

trt treatment arm A or B

sex f=female, m=male

futime time to death or last follow-up

death 1 if futime is a death, 0 for censoring

txtime time to hematropetic stem cell transplant

crtime time to complete response

rltime time to relapse of disease
```

nafld 67

### **Details**

This data set is used to illustrate multi-state survival curves. The correlation between within-subject event times strongly resembles that from an actual trial, but none of the actual data values are from that source.

### References

Le-Rademacher JG, Peterson RA, Therneau TM, Sanford BL, Stone RM, Mandrekar SJ. Application of multi-state models in cancer clinical trials. Clin Trials. 2018 Oct; 15 (5):489-498

### **Examples**

```
coxph(Surv(futime, death) ~ trt, data=myeloid)
# See the mstate vignette for a more complete analysis
```

nafld

Non-alcohol fatty liver disease

## **Description**

Data sets containing the data from a population study of non-alcoholic fatty liver disease (NAFLD). Subjects with the condition and a set of matched control subjects were followed forward for metabolic conditions, cardiac endpoints, and death.

## Usage

## **Format**

nafld1 is a data frame with 17549 observations on the following 10 variables.

```
id subject identifier
age age at entry to the study
male 0=female, 1=male
weight weight in kg
height height in cm
bmi body mass index
case.id the id of the NAFLD case to whom this subject is matched
futime time to death or last follow-up
status 0= alive at last follow-up, 1=dead
```

68 nafld

naf1d2 is a data frame with 400123 observations and 4 variables containing laboratory data

id subject identifier

days days since index date

test the type of value recorded

value the numeric value

nafld3 is a data frame with 34340 observations and 3 variables containing outcomes

id subject identifier

days days since index date

event the endpoint that occurred

### **Details**

The primary reference for the NAFLD study is Allen (2018). The incidence of non-alcoholic fatty liver disease (NAFLD) has been rising rapidly in the last decade and it is now one of the main drivers of hepatology practice *Tapper2018*. It is essentially the presence of excess fat in the liver, and parallels the ongoing obesity epidemic. Approximately 20-25% of NAFLD patients will develop the inflammatory state of non-alcoholic steatohepatitis (NASH), leading to fibrosis and eventual end-stage liver disease. NAFLD can be accurately diagnosed by MRI methods, but NASH diagnosis currently requires a biopsy.

The current study constructed a population cohort of all adult NAFLD subjects from 1997 to 2014 along with 4 potential controls for each case. To protect patient confidentiality all time intervals are in days since the index date; none of the dates from the original data were retained. Subject age is their integer age at the index date, and the subject identifier is an arbitrary integer. As a final protection, we include only a 90% random sample of the data. As a consequence analyses results will not exactly match the original paper.

There are 3 data sets: nafld1 contains baseline data and has one observation per subject, nafld2 has one observation for each (time dependent) continuous measurement, and nafld3 has one observation for each yes/no outcome that occured.

## Source

Data obtained from the author.

### References

AM Allen, TM Therneau, JJ Larson, A Coward, VK Somers and PS Kamath, Nonalcoholic Fatty Liver Disease Incidence and Impact on Metabolic Burden and Death: A 20 Year Community Study, Hepatology 67:1726-1736, 2018.

neardate 69

neardate	Find the index of the closest value in data set 2, for each entry in data set one.

# **Description**

A common task in medical work is to find the closest lab value to some index date, for each subject.

# Usage

```
neardate(id1, id2, y1, y2, best = c("after", "prior"),
nomatch = NA_integer_)
```

## **Arguments**

id1	vector of subject identifiers for the index group
id2	vector of identifiers for the reference group
y1	normally a vector of dates for the index group, but any orderable data type is allowed
y2	reference set of dates
best	if best='prior' find the index of the first y2 value less than or equal to the target y1 value, for each subject. If best='after' find the first y2 value which is greater than or equal to the target y1 value, for each subject.
nomatch	the value to return for items without a match

## **Details**

This routine is closely related to match and to findInterval, the first of which finds exact matches and the second closest matches. This finds the closest matching date within sets of exactly matching identifiers. Closest date matching is often needed in clinical studies. For example data set 1 might contain the subject identifier and the date of some procedure and data set set 2 has the dates and values for laboratory tests, and the query is to find the first test value after the intervention but no closer than 7 days.

The id1 and id2 arguments are similar to match in that we are searching for instances of id1 that will be found in id2, and the result is the same length as id1. However, instead of returning the first match with id2 this routine returns the one that best matches with respect to y1.

The y1 and y2 arguments need not be dates, the function works for any data type such that the expression c(y1,y2) gives a sensible, sortable result. Be careful about matching Date and DateTime values and the impact of time zones, however, see as.POSIXct. If y1 and y2 are not of the same class the user is on their own. Since there exist pairs of unmatched data types where the result could be sensible, the routine will in this case proceed under the assumption that "the user knows what they are doing". Caveat emptor.

70 nsk

### Value

the index of the matching observations in the second data set, or the nomatch value for no successful match

### Author(s)

Terry Therneau

#### See Also

match, findInterval

### **Examples**

```
data1 <- data.frame(id = 1:10,</pre>
                    entry.dt = as.Date(paste("2011", 1:10, "5", sep='-')))
temp1 <- c(1,4,5,1,3,6,9, 2,7,8,12,4,6,7,10,12,3)
data2 <- data.frame(id = c(1,1,1,2,2,4,4,5,5,5,6,8,8,9,10,10,12),
                     lab.dt = as.Date(paste("2011", temp1, "1", sep='-')),
                    chol = round(runif(17, 130, 280)))
#first cholesterol on or after enrollment
indx1 <- neardate(data1$id, data2$id, data1$entry.dt, data2$lab.dt)</pre>
data2[indx1, "chol"]
# Closest one, either before or after.
indx2 <- neardate(data1$id, data2$id, data1$entry.dt, data2$lab.dt,</pre>
                   best="prior")
ifelse(is.na(indx1), indx2, # none after, take before
       ifelse(is.na(indx2), indx1, #none before
       ifelse(abs(data2$lab.dt[indx2]- data1$entry.dt) <</pre>
              abs(data2$lab.dt[indx1]- data1$entry.dt), indx2, indx1)))
# closest date before or after, but no more than 21 days prior to index
indx2 <- ifelse((data1$entry.dt - data2$lab.dt[indx2]) >21, NA, indx2)
ifelse(is.na(indx1), indx2, # none after, take before
       ifelse(is.na(indx2), indx1, #none before
       ifelse(abs(data2$lab.dt[indx2]- data1$entry.dt) <</pre>
              abs(data2$lab.dt[indx1]- data1$entry.dt), indx2, indx1)))
```

nsk

Natural splines with knot heights as the basis.

### **Description**

Create the design matrix for a natural spline, such that the coefficient of the resulting fit are the values of the function at the knots.

nsk 71

### Usage

```
nsk(x, df = NULL, knots = NULL, intercept = FALSE, b = 0.05, Boundary.knots = quantile(x, c(b, 1 - b), na.rm = TRUE))
```

### **Arguments**

x the predictor variable. Missing values are allowed.

df degrees of freedom. One can supply df rather than knots; ns() then chooses df - 1

- intercept knots at suitably chosen quantiles of x (which will ignore missing values). The default, df = NULL, sets the number of inner knots as length(knots).

knots breakpoints that define the spline. The default is no knots; together with the

natural boundary conditions this results in a basis for linear regression on x. Typical values are the mean or median for one knot, quantiles for more knots.

See also Boundary.knots.

intercept if TRUE, an intercept is included in the basis; default is FALSE

b default placement of the boundary knots. A value of bs=0 will replicate the

default behavior of ns.

Boundary.knots boundary points at which to impose the natural boundary conditions and anchor

the B-spline basis. Beyond these points the function is assumed to be linear. If both knots and Boundary.knots are supplied, the basis parameters do not depend

on x. Data can extend beyond Boundary.knots

#### **Details**

The nsk function behaves identically to the ns function, with two exceptions. The primary one is that the returned basis is such that coefficients correspond to the value of the fitted function at the knot points. If intercept = FALSE, there will be k-1 coefficients corresponding to the k knots, and they will be the difference in predicted value between knots 2-k and knot 1. The primary advantage to the basis is that the coefficients are directly interpretable. A second is that tests for the linear and non-linear components are simple contrasts.

The second difference with ns is one of opinion with respect to the default position for the boundary knots. The default here is closer to that found in the rms::rcs function.

### Value

A matrix of dimension length(x) \* df where either df was supplied or, if knots were supplied, df = length(knots) + 1 + intercept. Attributes are returned that correspond to the arguments to kns, and explicitly give the knots, Boundary.knots etc for use by predict.kns().

#### Note

A thin flexible metal or wooden strip is called a spline, and is the traditional method for laying out a smooth curve, e.g., for a ship's hull or an airplane wing. Pins are put into a board and the strip is passed through them, each pin is a 'knot'.

A mathematical spline is a piecewise function between each knot. A linear spline will be a set of connected line segments, a quadratic spline is a set of connected local quadratic functions, constrained to have a continuous first derivative, a cubic spline is cubic between each knot, constrained

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to have continuous first and second derivatives, and etc. Mathematical splines are not an exact representation of natural splines: being a physical object the wood or metal strip will have continuous derivatives of all orders. Cubic splines are commonly used because they are sufficiently smooth to look natural to the human eye.

If the mathematical spline is further constrained to be linear beyond the end knots, this is often called a 'natural spline', due to the fact that a wooden or metal spline will also be linear beyond the last knots. Another name for the same object is a 'restricted cubic spline', since it is achieved in code by adding a further constraint. Given a vector of data points and a set of knots, it is possible to create a basis matrix X with one column per knot, such that ordinary regression of X on y will fit the cubic spline function, hence these are also called 'regression splines'. One label is no better than another.

Given a basis matrix X, the matrix Z= XT for any X by X nonsingular matrix X is also a basis matrix, and will result in identical predicted values, but a new set of coefficients gamma = (X-inverse) beta in place of beta. One can choose the basis function so that X is easy to construct, to make the regression numerically stable, to make tests easier, or based on other considerations. It seems as though every spline function returns a different basis set, which unfortunately makes fits difficult to compare; and this is yet one more, chosen to make the coefficients more interpretable.

### See Also

ns

### **Examples**

```
# make some dummy data
tdata <- data.frame(x= lung$age, y = 10*log(lung$age-35) + rnorm(228, 0, 2))
fit1 <- lm(y ~ -1 + nsk(x, df=4, intercept=TRUE) , data=tdata)
fit2 <- lm(y ~ nsk(x, df=3), data=tdata)

# the knots (same for both fits)
knots <- unlist(attributes(fit1$model[[2]])[c('Boundary.knots', 'knots')])
knots
unname(coef(fit1)) # predictions at the knot points

unname(coef(fit1)[-1] - coef(fit1)[1]) # differences: yhat[2:4] - yhat[1]
unname(coef(fit2))

## Not run:
plot(y ~ x, data=tdata)
points(sort(knots), coef(fit1), col=2, pch=19)
coef(fit)[1] + c(0, coef(fit)[-1])

## End(Not run)</pre>
```

Data from the National Wilm's Tumor Study

ovarian 73

## **Description**

Measurement error example. Tumor histology predicts survival, but prediction is stronger with central lab histology than with the local institution determination.

## Usage

```
nwtco
data(nwtco, package="survival")
```

### **Format**

A data frame with 4028 observations on the following 9 variables.

```
seqno id number
instit Histology from local institution
histol Histology from central lab
stage Disease stage
study study
rel indicator for relapse
edrel time to relapse
age age in months
in.subcohort Included in the subcohort for the example in the paper
```

### References

NE Breslow and N Chatterjee (1999), Design and analysis of two-phase studies with binary outcome applied to Wilms tumour prognosis. *Applied Statistics* **48**, 457–68.

## **Examples**

```
with(nwtco, table(instit,histol))
anova(coxph(Surv(edrel,rel)~histol+instit,data=nwtco))
anova(coxph(Surv(edrel,rel)~instit+histol,data=nwtco))
```

ovarian

Ovarian Cancer Survival Data

### **Description**

Survival in a randomised trial comparing two treatments for ovarian cancer

# Usage

```
ovarian
data(cancer, package="survival")
```

## **Format**

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futime: survival or censoring time

fustat: censoring status

age: in years

resid.ds: residual disease present (1=no,2=yes)

rx: treatment group

ecog.ps: ECOG performance status (1 is better, see reference)

#### Source

Terry Therneau

#### References

Edmunson, J.H., Fleming, T.R., Decker, D.G., Malkasian, G.D., Jefferies, J.A., Webb, M.J., and Kvols, L.K., Different Chemotherapeutic Sensitivities and Host Factors Affecting Prognosis in Advanced Ovarian Carcinoma vs. Minimal Residual Disease. Cancer Treatment Reports, 63:241-47, 1979.

pbc

Mayo Clinic Primary Biliary Cholangitis Data

### **Description**

Primary sclerosing cholangitis is an autoimmune disease leading to destruction of the small bile ducts in the liver. Progression is slow but inexhortable, eventually leading to cirrhosis and liver decompensation. The condition has been recognised since at least 1851 and was named "primary biliary cirrhosis" in 1949. Because cirrhosis is a feature only of advanced disease, a change of its name to "primary biliary cholangitis" was proposed by patient advocacy groups in 2014.

This data is from the Mayo Clinic trial in PBC conducted between 1974 and 1984. A total of 424 PBC patients, referred to Mayo Clinic during that ten-year interval, met eligibility criteria for the randomized placebo controlled trial of the drug D-penicillamine. The first 312 cases in the data set participated in the randomized trial and contain largely complete data. The additional 112 cases did not participate in the clinical trial, but consented to have basic measurements recorded and to be followed for survival. Six of those cases were lost to follow-up shortly after diagnosis, so the data here are on an additional 106 cases as well as the 312 randomized participants.

A nearly identical data set found in appendix D of Fleming and Harrington; this version has fewer missing values.

## Usage

```
pbc
data(pbc, package="survival")
```

## Format

pbcseq 75

age: in years

albumin: serum albumin (g/dl)

alk.phos: alkaline phosphotase (U/liter)

ascites: presence of ascites

ast: aspartate aminotransferase, once called SGOT (U/ml)

bili: serum bilirunbin (mg/dl) chol: serum cholesterol (mg/dl) copper: urine copper (ug/day)

edema: 0 no edema, 0.5 untreated or successfully treated

1 edema despite diuretic therapy

hepato: presence of hepatomegaly or enlarged liver

id: case number platelet: platelet count

protime: standardised blood clotting time

sex: m/f

spiders: blood vessel malformations in the skin stage: histologic stage of disease (needs biopsy)

status: status at endpoint, 0/1/2 for censored, transplant, dead time: number of days between registration and the earlier of death,

transplantion, or study analysis in July, 1986

trt: 1/2/NA for D-penicillmain, placebo, not randomised

trig: triglycerides (mg/dl)

#### **Source**

T Therneau and P Grambsch (2000), *Modeling Survival Data: Extending the Cox Model*, Springer-Verlag, New York. ISBN: 0-387-98784-3.

## See Also

pbcseq

pbcseq

Mayo Clinic Primary Biliary Cirrhosis, sequential data

### Description

This data is a continuation of the PBC data set, and contains the follow-up laboratory data for each study patient. An analysis based on the data can be found in Murtagh, et. al.

The primary PBC data set contains only baseline measurements of the laboratory parameters. This data set contains multiple laboratory results, but only on the 312 randomized patients. Some baseline data values in this file differ from the original PBC file, for instance, the data errors in prothrombin time and age which were discovered after the original analysis (see Fleming and Harrington, figure 4.6.7).

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One "feature" of the data deserves special comment. The last observation before death or liver transplant often has many more missing covariates than other data rows. The original clinical protocol for these patients specified visits at 6 months, 1 year, and annually thereafter. At these protocol visits lab values were obtained for a large pre-specified battery of tests. "Extra" visits, often undertaken because of worsening medical condition, did not necessarily have all this lab work. The missing values are thus potentially informative.

## Usage

```
pbcseq
data(pbc, package="survival")
```

#### **Format**

id: case number age: in years sex: m/f

trt: 1/2/NA for D-penicillmain, placebo, not randomised

time: number of days between registration and the earlier of death,

transplantion, or study analysis in July, 1986

status: status at endpoint, 0/1/2 for censored, transplant, dead day: number of days between enrollment and this visit date

all measurements below refer to this date

albumin: serum albumin (mg/dl) alk.phos: alkaline phosphotase (U/liter)

ascites: presence of ascites

ast: aspartate aminotransferase, once called SGOT (U/ml)

bili: serum bilirunbin (mg/dl) chol: serum cholesterol (mg/dl) copper: urine copper (ug/day)

edema: 0 no edema, 0.5 untreated or successfully treated

1 edema despite diuretic therapy

hepato: presence of hepatomegaly or enlarged liver

platelet: platelet count

protime: standardised blood clotting time spiders: blood vessel malformations in the skin stage: histologic stage of disease (needs biopsy)

trig: triglycerides (mg/dl)

### Source

T Therneau and P Grambsch, "Modeling Survival Data: Extending the Cox Model", Springer-Verlag, New York, 2000. ISBN: 0-387-98784-3.

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### References

Murtaugh PA. Dickson ER. Van Dam GM. Malinchoc M. Grambsch PM. Langworthy AL. Gips CH. "Primary biliary cirrhosis: prediction of short-term survival based on repeated patient visits." Hepatology. 20(1.1):126-34, 1994.

Fleming T and Harrington D., "Counting Processes and Survival Analysis", Wiley, New York, 1991.

### See Also

pbc

## **Examples**

```
# Create the start-stop-event triplet needed for coxph
first <- with(pbcseq, c(TRUE, diff(id) !=0)) #first id for each subject
last <- c(first[-1], TRUE) #last id

time1 <- with(pbcseq, ifelse(first, 0, day))
time2 <- with(pbcseq, ifelse(last, futime, c(day[-1], 0)))
event <- with(pbcseq, ifelse(last, status, 0))

fit1 <- coxph(Surv(time1, time2, event) ~ age + sex + log(bili), pbcseq)</pre>
```

plot.aareg

Plot an aareg object.

# Description

Plot the estimated coefficient function(s) from a fit of Aalen's additive regression model.

### Usage

```
## S3 method for class 'aareg'
plot(x, se=TRUE, maxtime, type='s', ...)
```

### **Arguments**

x	the result of a call to the aareg function
se	if TRUE, standard error bands are included on the plot
maxtime	upper limit for the x-axis.
type	graphical parameter for the type of line, default is "steps".
	other graphical parameters such as line type, color, or axis labels.

### **Side Effects**

A plot is produced on the current graphical device.

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## References

Aalen, O.O. (1989). A linear regression model for the analysis of life times. Statistics in Medicine, 8:907-925.

## See Also

aareg

plot.cox.zph

Graphical Test of Proportional Hazards

# Description

Displays a graph of the scaled Schoenfeld residuals, along with a smooth curve.

# Usage

# Arguments

X	result of the cox.zph function.
resid	a logical value, if TRUE the residuals are included on the plot, as well as the smooth fit. $$
se	a logical value, if TRUE, confidence bands at two standard errors will be added.
df	the degrees of freedom for the fitted natural spline, df=2 leads to a linear fit.
nsmo	number of points to use for the lines
var	the set of variables for which plots are desired. By default, plots are produced in turn for each variable of a model. Selection of a single variable allows other features to be added to the plot, e.g., a horizontal line at zero or a main title. This has been superseded by a subscripting method; see the example below.
hr	if TRUE, label the y-axis using the estimated hazard ratio rather than the estimated coefficient. (The plot does not change, only the axis label.)
xlab	label for the x-axis of the plot
ylab	optional label for the y-axis of the plot. If missing a default label is provided. This can be a vector of labels.
lty, col, lwd	line type, color, and line width for the overlaid curve. Each of these can be vector of length 2, in which case the second element is used for the confidence interval.
	additional graphical arguments passed to the plot function.

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#### **Side Effects**

a plot is produced on the current graphics device.

#### See Also

```
coxph, cox.zph.
```

#### **Examples**

plot.survfit

Plot method for survfit objects

## **Description**

A plot of survival curves is produced, one curve for each strata. The log=T option does extra work to avoid log(0), and to try to create a pleasing result. If there are zeros, they are plotted by default at 0.8 times the smallest non-zero value on the curve(s).

Curves are plotted in the same order as they are listed by print (which gives a 1 line summary of each). This will be the order in which col, 1ty, etc are used.

## Usage

```
## S3 method for class 'survfit'
plot(x, conf.int=, mark.time=FALSE,
  pch=3, col=1, lty=1, lwd=1, cex=1, log=FALSE, xscale=1, yscale=1,
  xlim, ylim, xmax, fun,
  xlab="", ylab="", xaxs="r", conf.times, conf.cap=.005,
  conf.offset=.012,
  conf.type = c("log", "log-log", "plain", "logit", "arcsin"),
  mark, noplot="(s0)", cumhaz=FALSE,
  firstx, ymin, ...)
```

#### **Arguments**

x an object of class survfit, usually returned by the survfit function.

conf.int determines whether pointwise confidence intervals will be plotted. The

determines whether pointwise confidence intervals will be plotted. The default is to do so if there is only 1 curve, i.e., no strata, using 95% confidence intervals Alternatively, this can be a numeric value giving the desired confidence level.

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mark.time controls the labeling of the curves. If set to FALSE, no labeling is done. If TRUE,

then curves are marked at each censoring time. If mark is a numeric vector then

curves are marked at the specified time points.

pch vector of characters which will be used to label the curves. The points help

file contains examples of the possible marks. A single string such as "abcd" is treated as a vector c("a","b","c","d"). The vector is reused cyclically if it is shorter than the number of curves. If it is present this implies mark.time =

TRUE.

col a vector of integers specifying colors for each curve. The default value is 1.

1ty a vector of integers specifying line types for each curve. The default value is 1.

lwd a vector of numeric values for line widths. The default value is 1.

cex a numeric value specifying the size of the marks. This is not treated as a vector;

all marks have the same size.

log a logical value, if TRUE the y axis wll be on a log scale. Alternately, one of the

standard character strings "x", "y", or "xy" can be given to specific logarithmic

horizontal and/or vertical axes.

xscale a numeric value used like yscale for labels on the x axis. A value of 365.25

will give labels in years instead of the original days.

yscale a numeric value used to multiply the labels on the y axis. A value of 100, for

instance, would be used to give a percent scale. Only the labels are changed, not the actual plot coordinates, so that adding a curve with "lines(surv.exp(...))",

say, will perform as it did without the yscale argument.

xlim, ylim optional limits for the plotting region.

xmax the maximum horizontal plot coordinate. This can be used to shrink the range

of a plot. It shortens the curve before plotting it, so that unlike using the xlim graphical parameter, warning messages about out of bounds points are not gen-

erated.

fun an arbitrary function defining a transformation of the survival (or probability

in state, or cumulative hazard) curves. For example fun=log is an alternative way to draw a log-survival curve (but with the axis labeled with log(S) values), and fun=sqrt would generate a curve on square root scale. Four often used transformations can be specified with a character argument instead: "S" gives the usual survival curve, "log" is the same as using the log=T option, "event" or "F" plots the empirical CDF F(t) = 1 - S(t) (f(y) = 1-y), and "cloglog" creates a complimentary log-log survival plot (f(y) = log(-log(y)) along with log scale for the x-axis). The terms "identity" and "surv" are allowed as synonyms for type="S". The argument "cumhaz" causes the cumulative hazard

function to be plotted.

xlab label given to the x-axis.

ylab label given to the y-axis.

xaxs either "S" for a survival curve or a standard x axis style as listed in par; "r"

(regular) is the R default. Survival curves have historically been displayed with the curve touching the y-axis, but not touching the bounding box of the plot on the other 3 sides, Type "S" accomplishes this by manipulating the plot range and then using the "i" style internally. The "S" style is becoming increasingly

less common, however.

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conf.times optional vector of times at which to place a confidence bar on the curve(s). If present, these will be used instead of confidence bands. conf.cap width of the horizontal cap on top of the confidence bars; only used if conf.times is used. A value of 1 is the width of the plot region. conf.offset the offset for confidence bars, when there are multiple curves on the plot. A value of 1 is the width of the plot region. If this is a single number then each curve's bars are offset by this amount from the prior curve's bars, if it is a vector the values are used directly. One of "plain", "log" (the default), "log-log" or "logit". Only enough of conf.type the string to uniquely identify it is necessary. The first option causes confidence intervals not to be generated. The second causes the standard intervals curve +-k \*se(curve), where k is determined from conf.int. The log option calculates intervals based on the cumulative hazard or log(survival). The log-log option bases the intervals on the log hazard or log(-log(survival)), and the logit option on log(survival/(1-survival)). a historical alias for pch mark noplot for multi-state models, curves with this label will not be plotted. (Also see the istate0 argument in survcheck.) plot the cumulative hazard rather than the probability in state or survival. Opcumhaz tionally, this can be a numeric vector specifying which columns of the cumhaz component to plot. this will normally be given as part of the ylim argument ymin firstx this will normally be given as part of the xlim argument. other arguments that will be passed forward to the underlying plot method, such as xlab or ylab.

## Details

If the object contains a cumulative hazard curve, then fun='cumhaz' will plot that curve, otherwise it will plot  $-\log(S)$  as an approximation. Theoretically,  $S = \exp(-\Lambda)$  where S is the survival and  $\Lambda$  is the cumulative hazard. The same relationship holds for estimates of S and  $\Lambda$  only in special cases, but the approximation is often close.

When the survfit function creates a multi-state survival curve the resulting object also has class 'survfitms'. Competing risk curves are a common case. In this situation the fun argument is ignored.

When the conf.times argument is used, the confidence bars are offset by conf.offset units to avoid overlap. The bar on each curve are the confidence interval for the time point at which the bar is drawn, i.e., different time points for each curve. If curves are steep at that point, the visual impact can sometimes substantially differ for positive and negative values of conf.offset.

## Value

a list with components x and y, containing the coordinates of the last point on each of the curves (but not the confidence limits). This may be useful for labeling.

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#### Note

In prior versions the behavior of xscale and yscale differed: the first changed the scale both for the plot and for all subsequent actions such as adding a legend, whereas yscale affected only the axis label. This was normalized in version 2-36.4, and both parameters now only affect the labeling.

In versions prior to approximately 2.36 a survfit object did not contain the cumulative hazard as a separate result, and the use of fun="cumhaz" would plot the approximation -log(surv) to the cumulative hazard. When cumulative hazards were added to the object, the cumhaz=TRUE argument to the plotting function was added. In version 2.3-8 the use of fun="cumhaz" became a synonym for cumhaz=TRUE.

### See Also

```
points.survfit, lines.survfit, par, survfit
```

### **Examples**

```
leukemia.surv <- survfit(Surv(time, status) ~ x, data = aml)
plot(leukemia.surv, lty = 2:3)
legend(100, .9, c("Maintenance", "No Maintenance"), lty = 2:3)
title("Kaplan-Meier Curves\nfor AML Maintenance Study")
lsurv2 <- survfit(Surv(time, status) ~ x, aml, type='fleming')
plot(lsurv2, lty=2:3, fun="cumhaz",
xlab="Months", ylab="Cumulative Hazard")</pre>
```

predict.coxph

Predictions for a Cox model

## Description

Compute fitted values and regression terms for a model fitted by coxph

#### **Usage**

```
## S3 method for class 'coxph'
predict(object, newdata,
type=c("lp", "risk", "expected", "terms", "survival"),
se.fit=FALSE, na.action=na.pass, terms=names(object$assign), collapse,
reference=c("strata", "sample", "zero"), ...)
```

### **Arguments**

object

the results of a coxph fit.

newdata

Optional new data at which to do predictions. If absent predictions are for the data frame used in the original fit. When coxph has been called with a formula argument created in another context, i.e., coxph has been called within another function and the formula was passed as an argument to that function, there can be problems finding the data set. See the note below.

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type the type of predicted value. Choices are the linear predictor ("lp"), the risk score exp(lp) ("risk"), the expected number of events given the covariates and follow-up time ("expected"), and the terms of the linear predictor ("terms"). The survival probability for a subject is equal to exp(-expected). se.fit if TRUE, pointwise standard errors are produced for the predictions. na.action applies only when the newdata argument is present, and defines the missing value action for the new data. The default is to include all observations. When there is no newdata, then the behavior of missing is dictated by the na.action option of the original fit. terms if type="terms", this argument can be used to specify which terms should be included; the default is all. optional vector of subject identifiers. If specified, the output will contain one collapse entry per subject rather than one entry per observation. reference for centering predictions, see details below reference For future methods

#### **Details**

The Cox model is a *relative* risk model; predictions of type "linear predictor", "risk", and "terms" are all relative to the sample from which they came. By default, the reference value for each of these is the mean covariate within strata. The underlying reason is both statistical and practial. First, a Cox model only predicts relative risks between pairs of subjects within the same strata, and hence the addition of a constant to any covariate, either overall or only within a particular stratum, has no effect on the fitted results. Second, downstream calculations depend on the risk score exp(linear predictor), which will fall prey to numeric overflow for a linear predictor greater than .Machine\$double.max.exp. The coxph routines try to approximately center the predictors out of self protection. Using the reference="strata" option is the safest centering, since strata occassionally have different means. When the results of predict are used in further calculations it may be desirable to use a single reference level for all observations. Use of reference="sample" will use the overall means, and agrees with the linear.predictors component of the coxph object (which uses the overall mean for backwards compatability with older code). Predictions of type="terms" are almost invariably passed forward to further calculation, so for these we default to using the sample as the reference. A reference of "zero" causes no centering to be done.

Predictions of type "expected" incorporate the baseline hazard and are thus absolute instead of relative; the reference option has no effect on these. These values depend on the follow-up time for the future subjects as well as covariates so the newdata argument needs to include both the right and *left* hand side variables from the formula. (The status variable will not be used, but is required since the underlying code needs to reconstruct the entire formula.)

Models that contain a frailty term are a special case: due to the technical difficulty, when there is a newdata argument the predictions will always be for a random effect of zero.

### Value

a vector or matrix of predictions, or a list containing the predictions (element "fit") and their standard errors (element "se.fit") if the se.fit option is TRUE.

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#### Note

Some predictions can be obtained directly from the coxph object, and for others it is necessary for the routine to have the entirety of the original data set, e.g., for type = terms or if standard errors are requested. This extra information is saved in the coxph object if model=TRUE, if not the original data is reconstructed. If it is known that such residuals will be required overall execution will be slightly faster if the model information is saved.

In some cases the reconstruction can fail. The most common is when coxph has been called inside another function and the formula was passed as one of the arguments to that enclosing function. Another is when the data set has changed between the original call and the time of the prediction call. In each of these the simple solution is to add model=TRUE to the original coxph call.

#### See Also

```
predict,coxph,termplot
```

## **Examples**

```
options(na.action=na.exclude) # retain NA in predictions
fit <- coxph(Surv(time, status) ~ age + ph.ecog + strata(inst), lung)
#lung data set has status coded as 1/2
mresid <- (lung$status-1) - predict(fit, type='expected') #Martingale resid
predict(fit,type="lp")
predict(fit,type="expected")
predict(fit,type="risk",se.fit=TRUE)
predict(fit,type="terms",se.fit=TRUE)

# For someone who demands reference='zero'
pzero <- function(fit)
    predict(fit, reference="sample") + sum(coef(fit) * fit$means, na.rm=TRUE)</pre>
```

predict.survreg

Predicted Values for a 'survreg' Object

#### **Description**

Predicted values for a survreg object

## Usage

```
## S3 method for class 'survreg'
predict(object, newdata,
type=c("response", "link", "lp", "linear", "terms", "quantile",
   "uquantile"),
   se.fit=FALSE, terms=NULL, p=c(0.1, 0.9), na.action=na.pass, ...)
```

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### **Arguments**

object result of a model fit using the survreg function. newdata data for prediction. If absent predictions are for the subjects used in the original fit. the type of predicted value. This can be on the original scale of the data (retype sponse), the linear predictor ("linear", with "lp" as an allowed abbreviation), a predicted quantile on the original scale of the data ("quantile"), a quantile on the linear predictor scale ("uquantile"), or the matrix of terms for the linear predictor ("terms"). At this time "link" and linear predictor ("lp") are identical. se.fit if TRUE, include the standard errors of the prediction in the result. subset of terms. The default for residual type "terms" is a matrix with one terms column for every term (excluding the intercept) in the model. р vector of percentiles. This is used only for quantile predictions. na.action applies only when the newdata argument is present, and defines the missing value action for the new data. The default is to include all observations.

## Value

a vector or matrix of predicted values.

for future methods

#### References

Escobar and Meeker (1992). Assessing influence in regression analysis with censored data. *Biometrics*, 48, 507-528.

## See Also

```
survreg, residuals.survreg
```

## **Examples**

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print.aareg

Print an aareg object

## **Description**

Print out a fit of Aalen's additive regression model

## Usage

```
## S3 method for class 'aareg'
print(x, maxtime, test=c("aalen", "nrisk"),scale=1,...)
```

# **Arguments**

x	the result of a call to the aareg function
maxtime	the upper time point to be used in the test for non-zero slope
test	the weighting to be used in the test for non-zero slope. The default weights are based on the variance of each coefficient, as a function of time. The alternative weight is proportional to the number of subjects still at risk at each time point.
scale	scales the coefficients. For some data sets, the coefficients of the Aalen model will be very small (10-4); this simply multiplies the printed values by a constant, say 1e6, to make the printout easier to read.
	for future methods

### **Details**

The estimated increments in the coefficient estimates can become quite unstable near the end of follow-up, due to the small number of observations still at risk in a data set. Thus, the test for slope will sometimes be more powerful if this last 'tail' is excluded.

## Value

the calling argument is returned.

### **Side Effects**

the results of the fit are displayed.

# References

Aalen, O.O. (1989). A linear regression model for the analysis of life times. Statistics in Medicine, 8:907-925.

print.summary.coxph 87

### See Also

aareg

```
print.summary.coxph Print method for summary.coxph objects
```

### **Description**

Produces a printed summary of a fitted coxph model

## Usage

```
## S3 method for class 'summary.coxph'
print(x, digits=max(getOption("digits") - 3, 3),
    signif.stars = getOption("show.signif.stars"), expand=FALSE, ...)
```

### **Arguments**

x the result of a call to summary.coxph

digits significant digits to print

signif.stars Show stars to highlight small p-values

expand if the summary is for a multi-state coxph fit, print the results in an expanded

format.

... For future methods

```
print.summary.survexp Print Survexp Summary
```

## **Description**

Prints the results of summary.survexp

### Usage

```
## S3 method for class 'summary.survexp'
print(x, digits = max(options()$digits - 4, 3), ...)
```

### **Arguments**

```
x an object of class summary.survexp.
```

digits the number of digits to use in printing the result.

... for future methods

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## Value

x, with the invisible flag set to prevent further printing.

## Author(s)

Terry Therneau

## See Also

```
link{summary.survexp}, survexp
```

```
print.summary.survfit Print Survfit Summary
```

# Description

Prints the result of summary.survfit.

## Usage

```
## S3 method for class 'summary.survfit'
print(x, digits = max(options() $digits-4, 3), ...)
```

## **Arguments**

x an object of class "summary.survfit", which is the result of the summary.survfit function.

digits the number of digits to use in printing the numbers.

... for future methods

# Value

x, with the invisible flag set to prevent printing.

## **Side Effects**

prints the summary created by summary.survfit.

## See Also

```
options, print, summary.survfit.
```

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print.survfit

Print a Short Summary of a Survival Curve

### **Description**

Print number of observations, number of events, the restricted mean survival and its standard error, and the median survival with confidence limits for the median.

### **Usage**

```
## S3 method for class 'survfit'
print(x, scale=1, digits = max(options()$digits - 4,3),
    print.rmean=getOption("survfit.print.rmean"),
    rmean = getOption('survfit.rmean'),...)
```

### **Arguments**

x the result of a call to the survfit function.

scale a numeric value to rescale the survival time, e.g., if the input data to survfit were

in days, scale=365 would scale the printout to years.

digits Number of digits to print

print.rmean,rmean

Options for computation and display of the restricted mean.

.. for future results

## **Details**

The mean and its variance are based on a truncated estimator. That is, if the last observation(s) is not a death, then the survival curve estimate does not go to zero and the mean is undefined. There are four possible approaches to resolve this, which are selected by the rmean option. The first is to set the upper limit to a constant, e.g.,rmean=365. In this case the reported mean would be the expected number of days, out of the first 365, that would be experienced by each group. This is useful if interest focuses on a fixed period. Other options are "none" (no estimate), "common" and "individual". The "common" option uses the maximum time for all curves in the object as a common upper limit for the auc calculation. For the "individual" options the mean is computed as the area under each curve, over the range from 0 to the maximum observed time for that curve. Since the end point is random, values for different curves are not comparable and the printed standard errors are an underestimate as they do not take into account this random variation. This option is provided mainly for backwards compatability, as this estimate was the default (only) one in earlier releases of the code. Note that SAS (as of version 9.3) uses the integral up to the last *event* time of each individual curve; we consider this the worst of the choices and do not provide an option for that calculation.

The median and its confidence interval are defined by drawing a horizontal line at 0.5 on the plot of the survival curve and its confidence bands. The intersection of the line with the lower CI band defines the lower limit for the median's interval, and similarly for the upper band. If any of the intersections is not a point the we use the center of the intersection interval, e.g., if the survival

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curve were exactly equal to 0.5 over an interval. When data is uncensored this agrees with the usual definition of a median.

#### Value

x, with the invisible flag set to prevent printing. (The default for all print functions in R is to return the object passed to them; print.survfit complies with this pattern. If you want to capture these printed results for further processing, see the table component of summary.survfit.)

## **Side Effects**

The number of observations, the number of events, the median survival with its confidence interval, and optionally the restricted mean survival (rmean) and its standard error, are printed. If there are multiple curves, there is one line of output for each.

### References

Miller, Rupert G., Jr. (1981). Survival Analysis. New York: Wiley, p 71.

Pseudo values for survival.

#### See Also

summary.survfit, quantile.survfit

## **Description**

pseudo

Produce pseudo values from a survival curve.

# Usage

```
pseudo(fit, times, type, addNA=TRUE, data.frame=FALSE, minus1=FALSE, ...)
```

## **Arguments**

fit	a survfit object, or one that inherits that class.
times	a vector of time points, at which to evaluate the pseudo values.
type	the type of value, either the probabilty in state pstate, the cumulative hazard cumhaz or the expected sojourn time in the state sojourn.
addNA	If any observations were removed due to missing values in the fit object, add those rows (as NA) into the return. This causes the result of pseudo to match the original dataframe.
data.frame	if TRUE, return the data in "long" form as a data.frame with id, time, and pseudo as variables.
minus1	use n-1 as the multiplier rather than n.
•••	other arguments to the residuals.survfit function, which does the majority of the work, e.g., collapse and weighted.

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#### **Details**

This function computes pseudo values based on a first order Taylor series, also known as the "infinitesimal jackknife" (IJ) or "dfbeta" residuals. To be completely correct these results could perhaps be called 'IJ pseudo values' or even pseudo psuedo-values. For moderate to large data, however, the resulting values will be almost identical, numerically, to the ordinary jackknife.

A primary advantage of this approach is computational speed. Other features, neither good nor bad, are that they will agree with robust standard errors of other survival package estimates, which are based on the IJ, and that the mean of the estimates, over subjects, is exactly the underlying survival estimate.

For the type variable, surv is an acceptable synonym for pstate, and rmst, rmts are equivalent to sojourn. All of these are case insensitive.

#### Value

A vector, matrix, or array. The first dimension is always the number of observations in fit object, in the same order as the original data set (less any missing values that were removed when creating the survfit object); the second, if applicable, corresponds to fit\$states, e.g., multi-state survival, and the last dimension to the selected time points.

For the data frame option, a data frame containing values for id, time, and pseudo. If the original survfit call contained an id statement, then the values in the id column will be taken from that variable. If the id statement has a simple form, e.g., id = patno, then the name of the id column will be 'patno', otherwise it will be named '(id)'.

### References

PK Andersen and M Pohar-Perme, Pseudo-observations in surivival analysis, Stat Methods Medical Res, 2010; 19:71-99

#### See Also

```
residuals.survfit
```

## **Examples**

```
fit1 <- survfit(Surv(time, status) ~ 1, data=lung)
yhat <- pseudo(fit1, times=c(365, 730))
dim(yhat)
lfit <- lm(yhat[,1] ~ ph.ecog + age + sex, data=lung)</pre>
```

pspline

Smoothing splines using a pspline basis

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### **Description**

Specifies a penalised spline basis for the predictor. This is done by fitting a comparatively small set of splines and penalising the integrated second derivative. Traditional smoothing splines use one basis per observation, but several authors have pointed out that the final results of the fit are indistinguishable for any number of basis functions greater than about 2-3 times the degrees of freedom. Eilers and Marx point out that if the basis functions are evenly spaced, this leads to significant computational simplification, they refer to the result as a p-spline.

### Usage

```
pspline(x, df=4, theta, nterm=2.5 * df, degree=3, eps=0.1, method,
    Boundary.knots=range(x), intercept=FALSE, penalty=TRUE, combine, ...)
psplineinverse(x)
```

### **Arguments**

x for psline: a covariate vector. The function does not apply to factor variables.
--

For psplineinverse x will be the result of a pspline call.

df the desired degrees of freedom. One of the arguments df or theta' must be

given, but not both. If df=0, then the AIC = (loglik -df) is used to choose an "optimal" degrees of freedom. If AIC is chosen, then an optional argument

'caic=T' can be used to specify the corrected AIC of Hurvich et. al.

theta roughness penalty for the fit. It is a monotone function of the degrees of freedom,

with theta=1 corresponding to a linear fit and theta=0 to an unconstrained fit of

nterm degrees of freedom.

nterm number of splines in the basis

degree degree of splines
eps accuracy for df

method the method for choosing the tuning parameter theta. If theta is given, then

'fixed' is assumed. If the degrees of freedom is given, then 'df' is assumed. If method='aic' then the degrees of freedom is chosen automatically using Akaike's

information criterion.

... optional arguments to the control function

Boundary knots the spline is linear beyond the boundary knots. These default to the range of the

data.

intercept if TRUE, the basis functions include the intercept.

penalty if FALSE a large number of attributes having to do with penalized fits are ex-

cluded. This is useful to create a pspline basis matrix for other uses.

combine an optional vector of increasing integers. If two adjacent values of combine are

equal, then the corresponding coefficients of the fit are forced to be equal. This

is useful for monotone fits, see the vignette for more details.

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### Value

Object of class pspline, coxph.penalty containing the spline basis, with the appropriate attributes to be recognized as a penalized term by the coxph or survreg functions.

For psplineinverse the original x vector is reconstructed.

#### References

Eilers, Paul H. and Marx, Brian D. (1996). Flexible smoothing with B-splines and penalties. Statistical Science, 11, 89-121.

Hurvich, C.M. and Simonoff, J.S. and Tsai, Chih-Ling (1998). Smoothing parameter selection in nonparametric regression using an improved Akaike information criterion, JRSSB, volume 60, 271–293.

#### See Also

```
coxph,survreg,ridge, frailty
```

## **Examples**

```
lfit6 <- survreg(Surv(time, status)~pspline(age, df=2), lung)
plot(lung$age, predict(lfit6), xlab='Age', ylab="Spline prediction")
title("Cancer Data")
fit0 <- coxph(Surv(time, status) ~ ph.ecog + age, lung)
fit1 <- coxph(Surv(time, status) ~ ph.ecog + pspline(age,3), lung)
fit3 <- coxph(Surv(time, status) ~ ph.ecog + pspline(age,8), lung)
fit0
fit1
fit3</pre>
```

pyears

Person Years

#### **Description**

This function computes the person-years of follow-up time contributed by a cohort of subjects, stratified into subgroups. It also computes the number of subjects who contribute to each cell of the output table, and optionally the number of events and/or expected number of events in each cell.

## Usage

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## **Arguments**

formula	a formula object. The response variable will be a vector of follow-up times for each subject, or a Surv object containing the survival time and an event indicator. The predictors consist of optional grouping variables separated by + operators (exactly as in survfit), time-dependent grouping variables such as age (specified with tcut), and optionally a ratetable term. This latter matches each subject to his/her expected cohort.
data	a data frame in which to interpret the variables named in the formula, or in the subset and the weights argument.
weights	case weights.
subset	expression saying that only a subset of the rows of the data should be used in the fit.
na.action	a missing-data filter function, applied to the model.frame, after any subset argument has been used. Default is options()\$na.action.
rmap	an optional list that maps data set names to the ratetable names. See the details section below.
ratetable	a table of event rates, such as survexp.uswhite.
scale	a scaling for the results. As most rate tables are in units/day, the default value of 365.25 causes the output to be reported in years.
expect	should the output table include the expected number of events, or the expected number of person-years of observation. This is only valid with a rate table.
data.frame	return a data frame rather than a set of arrays.
model, x, y	If any of these is true, then the model frame, the model matrix, and/or the vector of response times will be returned as components of the final result.

## Details

Because pyears may have several time variables, it is necessary that all of them be in the same units. For instance, in the call

the natural unit of the ratetable is hazard per day, it is important that futime, age and entry.dt all be in days. Given the wide range of possible inputs, it is difficult for the routine to do sanity checks of this aspect.

The ratetable being used may have different variable names than the user's data set, this is dealt with by the rmap argument. The rate table for the above calculation was survexp.us, a call to summary{survexp.us} reveals that it expects to have variables age = age in days, sex, and year = the date of study entry, we create them in the rmap line. The sex variable is not mapped, therefore the code assumes that it exists in mydata in the correct format. (Note: for factors such as sex, the program will match on any unique abbreviation, ignoring case.)

A special function tcut is needed to specify time-dependent cutpoints. For instance, assume that age is in years, and that the desired final arrays have as one of their margins the age groups 0-2, 2-10,

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10-25, and 25+. A subject who enters the study at age 4 and remains under observation for 10 years will contribute follow-up time to both the 2-10 and 10-25 subsets. If cut(age,c(0,2,10,25,100)) were used in the formula, the subject would be classified according to his starting age only. The tcut function has the same arguments as cut, but produces a different output object which allows the pyears function to correctly track the subject.

The results of pyears are normally used as input to further calculations. The print routine, therefore, is designed to give only a summary of the table.

### Value

a list with components:

pyears	an array containing the person-years of exposure. (Or other units, depending on the rate table and the scale). The dimension and dimnames of the array correspond to the variables on the right hand side of the model equation.
n	an array containing the number of subjects who contribute time to each cell of the pyears array.
event	an array containing the observed number of events. This will be present only if the response variable is a Surv object.
expected	an array containing the expected number of events (or person years if expect ="pyears"). This will be present only if there was a ratetable term.
data	if the data. frame option was set, a data frame containing the variables n, event, pyears and event that supplants the four arrays listed above, along with variables corresponding to each dimension. There will be one row for each cell in the arrays.
offtable	the number of person-years of exposure in the cohort that was not part of any cell in the pyears array. This is often useful as an error check; if there is a mismatch of units between two variables, nearly all the person years may be off table.
tcut	whether the call included any time-dependent cutpoints.
summary	a summary of the rate-table matching. This is also useful as an error check.
call	an image of the call to the function.
observations	the number of observations in the input data set, after any missings were removed.
na.action	the na.action attribute contributed by an na.action routine, if any.

## See Also

```
ratetable, survexp, Surv.
```

# **Examples**

```
# Look at progression rates jointly by calendar date and age
#
temp.yr <- tcut(mgus$dxyr, 55:92, labels=as.character(55:91))
temp.age <- tcut(mgus$age, 34:101, labels=as.character(34:100))</pre>
```

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```
ptime <- ifelse(is.na(mgus$pctime), mgus$futime, mgus$pctime)</pre>
pstat <- ifelse(is.na(mgus$pctime), 0, 1)</pre>
pfit <- pyears(Surv(ptime/365.25, pstat) ~ temp.yr + temp.age + sex, mgus,</pre>
     data.frame=TRUE)
# Turn the factor back into numerics for regression
tdata <- pfit$data
tdata$age <- as.numeric(as.character(tdata$temp.age))</pre>
tdata$year<- as.numeric(as.character(tdata$temp.yr))</pre>
fit1 <- glm(event ~ year + age+ sex +offset(log(pyears)),</pre>
             data=tdata, family=poisson)
## Not run:
# fit a gam model
gfit.m <- gam(y ~ s(age) + s(year) + offset(log(time)),
                         family = poisson, data = tdata)
## End(Not run)
# Example #2 Create the hearta data frame:
hearta <- by(heart, heart$id,
             function(x)x[x$stop == max(x$stop),])
hearta <- do.call("rbind", hearta)</pre>
# Produce pyears table of death rates on the surgical arm
# The first is by age at randomization, the second by current age
fit1 <- pyears(Surv(stop/365.25, event) \sim cut(age + 48, c(0,50,60,70,100)) +
       surgery, data = hearta, scale = 1)
fit2 <- pyears(Surv(stop/365.25, event) ~ tcut(age + 48, c(0,50,60,70,100)) +
       surgery, data = hearta, scale = 1)
fit1$event/fit1$pyears #death rates on the surgery and non-surg arm
fit2$event/fit2$pyears #death rates on the surgery and non-surg arm
```

quantile.survfit

Quantiles from a survfit object

### **Description**

Retrieve quantiles and confidence intervals for them from a survfit object.

## Usage

```
## S3 method for class 'survfit'
quantile(x, probs = c(0.25, 0.5, 0.75), conf.int = TRUE,
    scale, tolerance= sqrt(.Machine$double.eps), ...)
## S3 method for class 'survfitms'
quantile(x, probs = c(0.25, 0.5, 0.75), conf.int = TRUE,
    scale, tolerance= sqrt(.Machine$double.eps), ...)
```

quantile.survfit 97

#### **Arguments**

x a result of the survfit function
probs numeric vector of probabilities with values in [0,1]
conf.int should lower and upper confidence limits be returned?

scale optional scale factor, e.g., scale=365.25 would return results in years if the fit

object were in days.

tolerance tolerance for checking that the survival curve exactly equals one of the quantiles

... optional arguments for other methods

#### **Details**

The kth quantile for a survival curve S(t) is the location at which a horizontal line at height p= 1-k intersects the plot of S(t). Since S(t) is a step function, it is possible for the curve to have a horizontal segment at exactly 1-k, in which case the midpoint of the horizontal segment is returned. This mirrors the standard behavior of the median when data is uncensored. If the survival curve does not fall to 1-k, then that quantile is undefined.

In order to be consistent with other quantile functions, the argument prob of this function applies to the cumulative distribution function F(t) = 1-S(t).

Confidence limits for the values are based on the intersection of the horizontal line at 1-k with the upper and lower limits for the survival curve. Hence confidence limits use the same p-value as was in effect when the curve was created, and will differ depending on the conf.type option of survfit. If the survival curves have no confidence bands, confidence limits for the quantiles are not available.

When a horizontal segment of the survival curve exactly matches one of the requested quantiles the returned value will be the midpoint of the horizontal segment; this agrees with the usual definition of a median for uncensored data. Since the survival curve is computed as a series of products, however, there may be round off error. Assume for instance a sample of size 20 with no tied times and no censoring. The survival curve after the 10th death is (19/20)(18/19)(17/18) ... (10/11) = 10/20, but the computed result will not be exactly 0.5. Any horizontal segment whose absolute difference with a requested percentile is less than tolerance is considered to be an exact match.

### Value

The quantiles will be a vector if the survfit object contains only a single curve, otherwise it will be a matrix or array. In this case the last dimension will index the quantiles.

If confidence limits are requested, then result will be a list with components quantile, lower, and upper, otherwise it is the vector or matrix of quantiles.

# Author(s)

Terry Therneau

#### See Also

survfit, print.survfit, qsurvreg

98 ratetable

### **Examples**

ratetable

Rate table structure

### **Description**

Description of the rate tables used by expected survival routines.

#### **Details**

A rate table contains event rates per unit time for some particular endpoint. Death rates are the most common use, the survexp.us table, for instance, contains death rates for the United States by year of age, sex, and calendar year.

A rate table is structured as a multi-way array with the following attributes:

dim the dimensions of the array

dimnames a named list of dimnames. The names are used to match user data to the dimensions, e.g., see the rmap argument in the pyears example. If a dimension is categorical, such as sex in survexp.us, then the dimname itself is matched against user's data values. The matching ignores case and allows abbreviations, e.g., "M", "Male", and "m" all successfully match the survexp.us dimname of sex=c("male", "female").

type a vector giving the type of each dimension, which will be 1= categorical, 2= continuous, 3= date, 4= calendar year of a US rate table. If type is 3 or 4, then the corresponding cutpoints must be one of the calendar date types: Date, POSIXt, date, or chron. This allows the code to properly match user data to the ratetable. (The published US decennial rate tables' definition is that a subject does not begin to experience a new years' death rate on Jan 1, but rather on their next birthday. The actual impact of this delay on any given subjects' calculation is neglible, but the code has always tried to be correct.)

cutpoints a list with one elment per dimension. If type=1 then the corresponding list element should be NULL, otherwise it should be a vector of length dim[i] containing the starting point of the interval to which the corresponding row/col of the array applies. Cutpoints must be in the same units as the underlying table, e.g., the survexp.us table contains death rates per day, so the age cutpoint vector contains age in days while year contains a vector of Dates. Cutpoints do not need to be evenly spaced: the survexp.us table, for instance, originally had age divided up as 0-1 days, 1-7 days, 7-28 days, 28 days - 1 year, 2, 3, ... 119 years. (Changes in the source of the tables made it difficult to continue splitting out the first year.)

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summary an optional summarization function. If present, it will be called with a numeric matrix that has one column per dimension and one row per observation. The function returns a character string giving a summary of the data. This is used by some routines to print an informative message, and provides one way to inform users of a data mistake, e.g., if the printout states that all subjects are between 0.14 and 0.23 years old it is likely that the user's age variable was in years when it should have been in days.

**dimid** optional attribute containing the names of the dimnames. If the dimnames list itself has names, this attribute will be ignored.

## See Also

survexp, pyears, survexp.us

ratetableDate

Convert date objects to ratetable form

### **Description**

This method converts dates from various forms into the internal form used in ratetable objects.

## Usage

ratetableDate(x)

### **Arguments**

Х

a date. The function currently has methods for Date, date, POSIXt, timeDate, and chron objects.

# Details

This function is useful for those who create new ratetables, but is normally invisible to users. It is used internally by the survexp and pyears functions to map the various date formats; if a new method is added then those routines will automatically be adapted to the new date type.

### Value

a numeric vector, the number of days since 1/1/1960.

# Author(s)

Terry Therneau

#### See Also

pyears, survexp

100 ratetables

ratetables	Census Data Sets for the Expected Survival and Person Years Functions

### Description

Census data sets for the expected survival and person years functions.

### Usage

```
data(survexp, package="survival")
```

### **Details**

survexp.us total United States population, by age and sex, 1940 to 2012.

**survexp.usr** United States population, by age, sex and race, 1940 to 2014. Race is white, nonwhite, or black. For 1960 and 1970 the black population values were not reported separately, so the nonwhite values were used.

**survexp.mn** total Minnesota population, by age and sex, 1970 to 2013.

Each of these tables contains the daily hazard rate for a matched subject from the population, defined as  $-\log(1-q)/365.25$  where q is the 1 year probability of death as reported in the original tables from the US Census. For age 25 in 1970, for instance, p=1-q is is the probability that a subject who becomes 25 years of age in 1970 will achieve his/her 26th birthday. The tables are recast in terms of hazard per day for computational convenience.

Each table is stored as an array, with additional attributes, and can be subset and manipulated as standard R arrays. See the help page for ratetable for details.

All numeric dimensions of a rate table must be in the same units. The survexp.us rate table contains daily hazard rates, the age cutpoints are in days, and the calendar year cutpoints are a Date.

## See Also

```
ratetable, survexp, pyears
```

### **Examples**

```
survexp.uswhite <- survexp.usr[,,"white",]</pre>
```

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rats

Rat treatment data from Mantel et al

### Description

Rat treatment data from Mantel et al. Three rats were chosen from each of 100 litters, one of which was treated with a drug, and then all followed for tumor incidence.

### Usage

```
rats
data(cancer, package="survival")
```

### **Format**

litter: litter number from 1 to 100 rx: treatment,(1=drug, 0=control) time: time to tumor or last follow-up

status: event status, 1=tumor and 0=censored

sex: male or female

#### Note

Since only 2/150 of the male rats have a tumor, most analyses use only females (odd numbered litters), e.g. Lee et al.

### Source

N. Mantel, N. R. Bohidar and J. L. Ciminera. Mantel-Haenszel analyses of litter-matched time to response data, with modifications for recovery of interlitter information. Cancer Research, 37:3863-3868, 1977.

### References

E. W. Lee, L. J. Wei, and D. Amato, Cox-type regression analysis for large number of small groups of correlated failure time observations, in "Survival Analysis, State of the Art", Kluwer, 1992.

rats2

Rat data from Gail et al.

### **Description**

48 rats were injected with a carcinogen, and then randomized to either drug or placebo. The number of tumors ranges from 0 to 13; all rats were censored at 6 months after randomization.

102 reliability

### Usage

```
rats2
data(cancer, package="survival")
```

#### **Format**

rat: id

trt: treatment,(1=drug, 0=control)

observation: within rat start: entry time stop: exit time

status: event status, 1=tumor, 0=censored

#### Source

MH Gail, TJ Santner, and CC Brown (1980), An analysis of comparative carcinogenesis experiments based on multiple times to tumor. *Biometrics* **36**, 255–266.

reliability

Reliability data sets

### **Description**

A set of data for simple reliablility analyses, taken from the book by Meeker and Escobar.

## Usage

```
data(reliability, package="survival")
```

#### **Details**

- capacitor: Data from a factorial experiment on the life of glass capacitors as a function of voltage and operating temperature. There were 8 capacitors at each combination of temperature and voltage. Testing at each combination was terminated after the fourth failure.
  - temperature: temperature in degrees celcius
  - voltage: applied voltage
  - time: time to failure
  - status: 1=failed, 0=censored
- cracks: Data on the time until the development of cracks in a set of 167 identical turbine parts. The parts were inspected at 8 selected times.
  - day: time of inspection
  - fail: number of fans found to have cracks, at this inspection

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- Data set genfan: Time to failure of 70 diesel engine fans.
  - hours: hours of service
  - status: 1=failure, 0=censored

Data set ifluid: A data frame with two variables describing the time to electrical breakdown of an insulating fluid.

- time: hours to breakdown
- voltage: test voltage in kV
- Data set imotor: Breakdown of motor insulation as a function of temperature.
  - temp: temperature of the test
  - time: time to failure or censoring
  - status: 0=censored, 1=failed
- Data set turbine: Each of 432 turbine wheels was inspected once to determine whether a crack had developed in the wheel or not.
  - hours: time of inspection (100s of hours)
  - inspected: number that were inspected
  - failed: number that failed

Data set valveSeat: Time to replacement of valve seats for 41 diesel engines. More than one seat may be replaced at a particular service, leading to duplicate times in the data set. The final inspection time for each engine will have status=0.

- id: engine identifier
- time: time of the inspection, in days
- status: 1=replacement occured, 0= not

#### References

Meeker and Escobar, Statistical Methods for Reliability Data, 1998.

### **Examples**

```
survreg(Surv(time, status) ~ temperature + voltage, capacitor)
```

residuals.coxph

Calculate Residuals for a 'coxph' Fit

### **Description**

Calculates martingale, deviance, score or Schoenfeld residuals for a Cox proportional hazards model.

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## **Usage**

## **Arguments**

object	an object inheriting from class coxph, representing a fitted Cox regression model. Typically this is the output from the coxph function.
type	character string indicating the type of residual desired. Possible values are "martingale", "deviance", "score", "schoenfeld", "dfbeta", "dfbetas", "scaledsch" and "partial". Only enough of the string to determine a unique match is required.
collapse	vector indicating which rows to collapse (sum) over. In time-dependent models more than one row data can pertain to a single individual. If there were 4 individuals represented by 3, 1, 2 and 4 rows of data respectively, then collapse=c(1,1,1,2,3,3,4,4,4,4) could be used to obtain per subject rather than per observation residuals.
weighted	if TRUE and the model was fit with case weights, then the weighted residuals are returned.
	other unused arguments

#### Value

For martingale and deviance residuals, the returned object is a vector with one element for each subject (without collapse). For score residuals it is a matrix with one row per subject and one column per variable. The row order will match the input data for the original fit. For Schoenfeld residuals, the returned object is a matrix with one row for each event and one column per variable. The rows are ordered by time within strata, and an attribute strata is attached that contains the number of observations in each strata. The scaled Schoenfeld residuals are used in the cox.zph function.

The score residuals are each individual's contribution to the score vector. Two transformations of this are often more useful: dfbeta is the approximate change in the coefficient vector if that observation were dropped, and dfbetas is the approximate change in the coefficients, scaled by the standard error for the coefficients.

residuals.survfit 105

## NOTE

For deviance residuals, the status variable may need to be reconstructed. For score and Schoenfeld residuals, the X matrix will need to be reconstructed.

#### References

T. Therneau, P. Grambsch, and T. Fleming. "Martingale based residuals for survival models", *Biometrika*, March 1990.

### See Also

coxph

## **Examples**

residuals.survfit

IJ residuals from a survfit object.

# Description

Return infinitesimal jackknife residuals from a survfit object, for the survival, cumulative hazard, or restricted mean time in state (RMTS).

### Usage

```
## S3 method for class 'survfit'
residuals(object, times,
    type="pstate", collapse, weighted=FALSE,
    method=1, ...)
```

## **Arguments**

object a survfit object

times a vector of times at which the residuals are desired

type the type of residual, see below

collapse add the residuals for all subjects in a cluster

weight the residuals by each observation's weight

method controls a choice of algorithm. Current an internal debugging option.

... arguments for other methods

106 residuals.survreg

#### **Details**

This function is designed to efficiently compute the leverage residuals at a small number of time points; a primary use is the creation of pseudo-values. If the residuals at all time points are needed, e.g. to compute a robust pointwise confidence interval for the survival curve, then this can be done more efficiently using the influence argument of the underlying survfit function. But be aware that such matrices can get very large.

The residuals are the impact of each observation or cluster on the resulting probability in state curves at the given time points, the cumulative hazard curvsurv at those time points, or the expected sojourn time in each state up to the given time points. For a simple Kaplan-Meier the survfit object contains only the probability in the "initial" state, i.e., the survival fraction. For the KM case the sojourn time, the expected amount of time spent in the initial state, up to the specified endpoint, is more commonly known as the restricted mean survival time (RMST). For a multistate model this same quantity is also referred to as the restricted mean time in state (RMTS). It can be computed as the area under the respective probability in state curve. The program allows any of pstate, surv, cumhaz, chaz, sojourn, rmst, rmts or auc for the type argument, ignoring upper/lowercase, so users can choose whichever abbreviation they like best.

When collapse=TRUE the result has the cluster identifier (which defaults to the id variable) as the dimname for the first dimension. If the fit object contains more than one curve, and the same identifier is reused in two different curves this approach does not work and the routine will stop with an error. In principle this is not necessary, e.g., the result could contain two rows with the same label, showing the separate effect on each curve, but this was deemed too confusing.

#### Value

A matrix or array with one row per observation or cluster, and one column for each value in times. For a multi-state model the three dimensions are observation, time and state. For cumulative hazard, the last dimension is the set of transitions. (A competing risks model for instance has 3 states and 2 transitions.)

#### See Also

```
survfit, survfit.formula
```

### **Examples**

```
fit <- survfit(Surv(time, status) ~ x, aml)
resid(fit, times=c(24, 48), type="RMTS")</pre>
```

residuals.survreg

Compute Residuals for 'survreg' Objects

### Description

This is a method for the function residuals for objects inheriting from class survreg.

residuals.survreg 107

### Usage

```
## S3 method for class 'survreg'
residuals(object, type=c("response", "deviance","dfbeta","dfbetas",
"working","ldcase","ldresp","ldshape", "matrix"), rsigma=TRUE,
collapse=FALSE, weighted=FALSE, ...)
```

## **Arguments**

object	an object inheriting from class survreg.
type	type of residuals, with choices of "response", "deviance", "dfbeta", "dfbetas", "working", "ldcase", "lsresp", "ldshape", and "matrix".
rsigma	include the scale parameters in the variance matrix, when doing computations. (I can think of no good reason not to).
collapse	optional vector of subject groups. If given, this must be of the same length as the residuals, and causes the result to be per group residuals.
weighted	give weighted residuals? Normally residuals are unweighted.
	other unused arguments

### Value

A vector or matrix of residuals is returned. Response residuals are on the scale of the original data, working residuals are on the scale of the linear predictor, and deviance residuals are on log-likelihood scale. The dfbeta residuals are a matrix, where the ith row gives the approximate change in the coefficients due to the addition of subject i. The dfbetas matrix contains the dfbeta residuals, with each column scaled by the standard deviation of that coefficient.

The matrix type produces a matrix based on derivatives of the log-likelihood function. Let L be the log-likelihood, p be the linear predictor  $X\beta$ , and s be  $\log(\sigma)$ . Then the 6 columns of the matrix are L, dL/dp,  $\partial^2 L/\partial p^2$ , dL/ds,  $\partial^2 L/\partial s^2$  and  $\partial^2 L/\partial p\partial s$ . Diagnostics based on these quantities are discussed in the book and article by Escobar and Meeker. The main ones are the likelihood displacement residuals for perturbation of a case weight (ldcase), the response value (ldresp), and the shape.

For a transformed distribution such as the log-normal or Weibull, matrix residuals are based on the log-likelihood of the transformed data log(y). For a monotone function f the density of f(X) is the density of X divided by the derivative of f (the Jacobian), so subtract log(derivative) from each uncensored observation's loglik value in order to match the loglik component of the result. The other colums of the matrix residual are unchanged by the transformation.

### References

Escobar, L. A. and Meeker, W. Q. (1992). Assessing influence in regression analysis with censored data. *Biometrics* **48**, 507-528.

Escobar, L. A. and Meeker, W. Q. (1998). Statistical Methods for Reliablity Data. Wiley.

### See Also

```
predict.survreg
```

108 retinopathy

## **Examples**

```
fit <- survreg(Surv(futime, death) ~ age + sex, mgus2)
summary(fit)  # age and sex are both important

rr <- residuals(fit, type='matrix')
sum(rr[,1]) - with(mgus2, sum(log(futime[death==1]))) # loglik
plot(mgus2$age, rr[,2], col= (1+mgus2$death)) # ldresp</pre>
```

retinopathy

Diabetic Retinopathy

## Description

A trial of laser coagulation as a treatment to delay diabetic retinopathy.

## Usage

```
retinopathy
data(retinopathy, package="survival")
```

#### **Format**

A data frame with 394 observations on the following 9 variables.

```
id numeric subject id
laser type of laser used: xenon argon
eye which eye was treated: right left
age age at diagnosis of diabetes
type type of diabetes: juvenile adult, (diagnosis before age 20)
trt 0 = control eye, 1 = treated eye
futime time to loss of vision or last follow-up
status 0 = censored, 1 = loss of vision in this eye
```

risk a risk score for the eye. This high risk subset is defined as a score of 6 or greater in at least one eye.

## **Details**

The 197 patients in this dataset were a 50% random sample of the patients with "high-risk" diabetic retinopathy as defined by the Diabetic Retinopathy Study (DRS). Each patient had one eye randomized to laser treatment and the other eye received no treatment, and has two observations in the data set. For each eye, the event of interest was the time from initiation of treatment to the time when visual acuity dropped below 5/200 two visits in a row. Thus there is a built-in lag time of approximately 6 months (visits were every 3 months). Survival times in this dataset are the actual time to vision loss in months, minus the minimum possible time to event (6.5 months). Censoring was caused by death, dropout, or end of the study.

rhDNase 109

#### References

W. J. Huster, R. Brookmeyer and S. G. Self (1989). Modelling paired survival data with covariates, Biometrics 45:145-156.

A. L. Blair, D. R. Hadden, J. A. Weaver, D. B. Archer, P. B. Johnston and C. J. Maguire (1976). The 5-year prognosis for vision in diabetes, American Journal of Ophthalmology, 81:383-396.

### **Examples**

```
coxph(Surv(futime, status) ~ type + trt, cluster= id, retinopathy)
```

rhDNase

rhDNASE data set

## **Description**

Results of a randomized trial of rhDNase for the treatment of cystic fibrosis.

### Usage

```
rhDNase
data(rhDNase, package="survival")
```

### **Format**

A data frame with 767 observations on the following 8 variables.

id subject id
inst enrolling institution
trt treatment arm: 0=placebo, 1= rhDNase
entry.dt date of entry into the study
end.dt date of last follow-up

fev forced expriatory volume at enrollment, a measure of lung capacity

ivstart days from enrollment to the start of IV antibiotics

ivstop days from enrollment to the cessation of IV antibiotics

#### **Details**

In patients with cystic fibrosis, extracellular DNA is released by leukocytes that accumulate in the airways in response to chronic bacterial infection. This excess DNA thickens the mucus, which then cannot be cleared from the lung by the cilia. The accumulation leads to exacerbations of respiratory symptoms and progressive deterioration of lung function. At the time of this study more than 90% of cystic fibrosis patients eventually died of lung disease.

Deoxyribonuclease I (DNase I) is a human enzyme normally present in the mucus of human lungs that digests extracellular DNA. Genentech, Inc. cloned a highly purified recombinant DNase I (rhD-Nase or Pulmozyme) which when delivered to the lungs in an aerosolized form cuts extracellular

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DNA, reducing the viscoelasticity of airway secretions and improving clearance. In 1992 the company conducted a randomized double-blind trial comparing rhDNase to placebo. Patients were then monitored for pulmonary exacerbations, along with measures of lung volume and flow. The primary endpoint was the time until first pulmonary exacerbation; however, data on all exacerbations were collected for 169 days.

The definition of an exacerbation was an infection that required the use of intravenous (IV) antibiotics. Subjects had 0–5 such episodes during the trial, those with more than one have multiple rows in the data set, those with none have NA for the IV start and end times. A few subjects were infected at the time of enrollment, subject 173 for instance has a first infection interval of -21 to 7. We do not count this first infection as an "event", and the subject first enters the risk set at day 7. Subjects who have an event are not considered to be at risk for another event during the course of antibiotics, nor for an additional 6 days after they end. (If the symptoms reappear immediately after cessation then from a medical standpoint this would not be a new infection.)

This data set reproduces the data in Therneau and Grambsch, is does not exactly reproduce those in Therneau and Hamilton due to data set updates.

### References

T. M. Therneau and P. M. Grambsch, Modeling Survival Data: Extending the Cox Model, Springer, 2000.

T. M. Therneau and S.A. Hamilton, rhDNase as an example of recurrent event analysis, Statistics in Medicine, 16:2029-2047, 1997.

### **Examples**

ridge 111

## **Description**

When used in a coxph or survreg model formula, specifies a ridge regression term. The likelihood is penalised by theta/2 time the sum of squared coefficients. If scale=T the penalty is calculated for coefficients based on rescaling the predictors to have unit variance. If df is specified then theta is chosen based on an approximate degrees of freedom.

# Usage

```
ridge(..., theta, df=nvar/2, eps=0.1, scale=TRUE)
```

## **Arguments**

	predictors to be ridged
theta	penalty is theta/2 time sum of squared coefficients
df	Approximate degrees of freedom
eps	Accuracy required for df
scale	Scale variables before applying penalty?

## Value

An object of class coxph. penalty containing the data and control functions.

### Note

If the expression ridge(x1,x2,x3,...) is too many characters long then the internal terms() function will add newlines to the variable name and then the coxph routine simply gets lost. (Some labels will have the newline and some won't.) One solution is to bundle all of the variables into a single matrix and use that matrix as the argument to ridge so as to shorten the call, e.g. mdata\$many <-as.matrix(mydata[,5:53]).

# References

Gray (1992) "Flexible methods of analysing survival data using splines, with applications to breast cancer prognosis" JASA 87:942–951

### See Also

```
coxph,survreg,pspline,frailty
```

## **Examples**

112 rotterdam

rotterdam

Breast cancer data set used in Royston and Altman (2013)

## Description

The rotterdam data set includes 2982 primary breast cancers patients whose data whose records were included in the Rotterdam tumor bank.

# Usage

```
rotterdam
data(cancer, package="survival")
```

#### **Format**

A data frame with 2982 observations on the following 15 variables.

```
pid patient identifier
year year of surgery
age age at surgery
meno menopausal status (0= premenopausal, 1= postmenopausal)
size tumor size, a factor with levels <=20 20-50 >50
grade differentiation grade
nodes number of positive lymph nodes
pgr progesterone receptors (fmol/l)
er estrogen receptors (fmol/l)
hormon hormonal treatment (0=no, 1=yes)
chemo chemotherapy
rtime days to relapse or last follow-up
recur 0= no relapse, 1= relapse
dtime days to death or last follow-up
death 0= alive, 1= dead
```

## **Details**

These data sets are used in the paper by Royston and Altman. The Rotterdam data is used to create a fitted model, and the GBSG data for validation of the model. The paper gives references for the data source.

## References

Patrick Royston and Douglas Altman, External validation of a Cox prognostic model: principles and methods. BMC Medical Research Methodology 2013, 13:33

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## See Also

gbsg

## **Examples**

```
rfstime <- pmin(rotterdam$rtime, rotterdam$dtime)</pre>
status <- pmax(rotterdam$recur, rotterdam$death)</pre>
fit1 <- coxph(Surv(rfstime, status) ~ pspline(age) + meno + size +</pre>
        pspline(nodes) + er,
     data=rotterdam, subset = (nodes > 0))
# Royston and Altman used fractional polynomials for the nonlinear terms
```

royston

Compute Royston's D for a Cox model

# **Description**

Compute the D statistic and R^2 for a coxph model, proposed by Royston and Sauerbrei

## Usage

```
royston(fit, newdata, ties = TRUE, adjust = FALSE)
```

## **Arguments**

adjust

fit a coxph fit optional validation data set newdata ties make a correction for ties in the risk score adjust for possible overfitting

### **Details**

The adjustment is based on the ratio r= (number of events)/(number of coefficients). For models which have sufficient sample size (r>20) the adjustment will be small.

## Value

a vector containing the value of D, R-squared and the estimated standard error of D.

# References

P. Royston and W. Sauerbrei, A new measure of prognostic separation in survival data. Statistics in Medicine 23:723-748, 2004.

114 rttright

## **Examples**

rttright

Compute redistribute-to-the-right weights

# Description

For many survival estimands, one approach is to redistribute each censored observation's weight to those other observations with a longer survival time (think of distributing an estate to the heirs). Then compute on the remaining, uncensored data.

# Usage

```
rttright(formula, data, weights, subset, na.action, times, id, timefix = TRUE)
```

## **Arguments**

formula	a formula object, which must have a Surv object as the response on the left of the ~ operator and, if desired, terms separated by + operators on the right. Each unique combination of predictors will define a separate strata.
data	$\boldsymbol{a}$ data frame in which to interpret the variables named in the formula, subset and weights arguments.
weights	The weights must be nonnegative and it is strongly recommended that they be strictly positive, since zero weights are ambiguous, compared to use of the subset argument.
subset	expression saying that only a subset of the rows of the data should be used in the fit.
na.action	a missing-data filter function, applied to the model frame, after any subset argument has been used. Default is options()\$na.action.
times	a vector of time points, for which to return updated weights. If missing, a time after the largest time in the data is assumed.
id	optional: if the data set has multiple rows per subject, a a variable containing the subect identifier of each row.
timefix	correct for possible round-off error

solder 115

### **Details**

The formula argument is treated exactly the same as in the survfit function.

Redistribution is recursive: redistribute the weight of the smallest censored observation to all those with longer time, which may include other censored observations. Then redistribute the next smallest and etc. up to the specified time value. After re-distributing the weight for a censored observation to other observations that are not censored, ordinary non-censored methods can often be applied. For example, redistribution of the weights, followed by computation of the weighted cumulative distribution function, reprises the Kaplan-Meier estimator.

A primary use of this routine is illustration of methods or exploration of new methods. Methods that use RTTR directly, such as the Brier score, will normally do these computations internally.

### Value

a vector or matrix of weights, with one column for each requested time

## See Also

```
survfit
```

## **Examples**

```
afit <- survfit(Surv(time, status) ~1, data=aml)
rwt <- rttright(Surv(time, status) ~1, data=aml)

index <- order(aml$time)
cdf <- cumsum(rwt[index]) # weighted CDF
cdf <- cdf[!duplicated(aml$time[index], fromLast=TRUE)] # remove duplicates
cbind(time=afit$time, KM= afit$surv, RTTR= 1-cdf)</pre>
```

solder

Data from a soldering experiment

## Description

In 1988 an experiment was designed and implemented at one of AT&T's factories to investigate alternatives in the "wave soldering" procedure for mounting electronic componentes to printed circuit boards. The experiment varied a number of factors relevant to the process. The response, measured by eye, is the number of visible solder skips.

# Usage

```
solder
data(solder, package="survival")
```

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### **Format**

A data frame with 900 observations on the following 6 variables.

Opening the amount of clearance around the mounting pad (3 levels)

Solder the amount of solder (Thick or Thin)

Mask type and thickness of the material used for the solder mask (A1.5, A3, A6, B3, B6)

PadType the geometry and size of the mounting pad (10 levels)

Panel each board was divided into 3 panels

skips the number of skips

### **Details**

This data set is used as a detailed example in chapter 1 of Chambers and Hastie. Observations 1-360 and 541-900 form a balanced design of 3\*2\*10\*3= 180 observations for four of the pad types (A1.5, A3, B3, B6), while rows 361-540 match 3 of the 6 Solder\*Opening combinations with pad type A6 and the other 3 with pad type A3.

## References

J Chambers and T Hastie, Statistical models in S. Chapman and Hall, 1993.

## **Examples**

```
# The balanced subset used by Chambers and Hastie
# contains the first 180 of each mask and deletes mask A6.
index <- 1 + (1:nrow(solder)) - match(solder$Mask, solder$Mask)
solder.balance <- droplevels(subset(solder, Mask != "A6" & index <= 180))</pre>
```

stanford2

More Stanford Heart Transplant data

## **Description**

This contains the Stanford Heart Transplant data in a different format. The main data set is in heart.

## Usage

stanford2

## **Format**

id: ID number

time: survival or censoring time

status: censoring status

age: in years

t5: T5 mismatch score

statefig 117

### Source

LA Escobar and WQ Meeker Jr (1992), Assessing influence in regression analysis with censored data. *Biometrics* **48**, 507–528. Page 519.

#### See Also

```
predict.survreg, heart
```

statefig

Draw a state space figure.

## **Description**

For multi-state survival models it is useful to have a figure that shows the states and the possible transitions between them. This function creates a simple "box and arrows" figure. It's goal was simplicity.

## Usage

```
statefig(layout, connect, margin = 0.03, box = TRUE, cex = 1, col = 1,
lwd=1, lty=1, bcol=col, acol=col, alwd=lwd, alty=lty, offset=0)
```

### **Arguments**

layout describes the layout of the boxes on the page. See the detailed description below.

connect a square matrix with one row for each state. If connect[i,j]!=0 then an arrow

is drawn from state i to state j. The row names of the matrix are used as the labels

for the states.

margin the fraction of white space between the label and the surrounding box, and be-

tween the box and the arrows, as a function of the plot region size.

box should boxes be drawn? TRUE or FALSE.

cex, col, lty, lwd

default graphical parameters used for the text and boxes. The last 3 can be a

vector of values.

bcol color for the box, if it differs from that used for the text.

acol, alwd, alty

color, line type and line width for the arrows.

offset

used to slight offset the arrows between two boxes x and y if there is a transition in both directions. The default of 0 leads to a double headed arrow in this case – to arrows are drawn but they coincide. A positive value causes each arrow to shift to the left, from the view of someone standing at the foot of a arrow and looking towards the arrowhead, a negative offset shifts to the right. A value of 1 corresponds to the size of the plotting region.

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### **Details**

The arguments for color, line type and line width can all be vectors, in which case they are recycled as needed. Boxes and text are drawn in the order of the rownames of connect, and arrows are drawn in the usual R matrix order.

The layout argument is normally a vector of integers, e.g., the vector (1, 3, 2) describes a layout with 3 columns. The first has a single state, the second column has 3 states and the third has 2. The coordinates of the plotting region are 0 to 1 for both x and y. Within a column the centers of the boxes are evenly spaced, with 1/2 a space between the boxes and the margin, e.g., 4 boxes would be at 1/8, 3/8, 5/8 and 7/8. If layout were a 1 column matrix with values of (1, 3, 2) then the layout will have three rows with 1, 3, and 2 boxes per row, respectively. Alternatively, the user can supply a 2 column matrix that directly gives the centers.

The values of the connect matrix should be 0 for pairs of states that do not have a transition and values between 0 and 2 for those that do. States are connected by an arc that passes through the centers of the two boxes and a third point that is between them. Specifically, consider a line segment joining the two centers and erect a second segment at right angles to the midpoint of length d times the distance from center to midpoint. The arc passes through this point. A value of d=0 gives a straight line, d=1 a right hand half circle centered on the midpoint and d=-1 a left hand half circle. The connect matrix contains values of d+1 with -1 < d < 1.

The connecting arrow are drawn from (center of box 1 + offset) to (center of box 2 + offset), where the amount of offset (white space) is determined by the box and margin parameters. If a pair of states are too close together this can result in an arrow that points the wrong way.

## Value

a matrix containing the centers of the boxes, with the invisible attribute set.

## Note

The goal of this function is to make "good enough" figures as simply as possible, and thereby to encourage users to draw them. The layout argument was inspired by the diagram package, which can draw more complex and well decorated figures, e.g., many different shapes, shading, multiple types of connecting lines, etc., but at the price of greater complexity.

Because curved lines are drawn as a set of short line segments, line types have almost no effect for that case.

# Author(s)

Terry Therneau

### **Examples**

```
# Draw a simple competing risks figure
states <- c("Entry", "Complete response", "Relapse", "Death")
connect <- matrix(0, 4, 4, dimnames=list(states, states))
connect[1, -1] <- c(1.1, 1, 0.9)
statefig(c(1, 3), connect)</pre>
```

strata 119

strata	Identify Stratification Variables	

## **Description**

This is a special function used in the context of the Cox survival model. It identifies stratification variables when they appear on the right hand side of a formula.

## Usage

```
strata(..., na.group=FALSE, shortlabel, sep=', ')
```

# Arguments

	any number of variables. All must be the same length.
na.group	a logical variable, if TRUE, then missing values are treated as a distinct level of each variable.
shortlabel	if TRUE omit variable names from resulting factor labels. The default action is to omit the names if all of the arguments are factors, and none of them was named.
sep	the character used to separate groups, in the created label

## **Details**

When used outside of a coxph formula the result of the function is essentially identical to the interaction function, though the labels from strata are often more verbose.

# Value

a new factor, whose levels are all possible combinations of the factors supplied as arguments.

## See Also

```
coxph, interaction
```

## **Examples**

```
a <- factor(rep(1:3,4), labels=c("low", "medium", "high"))
b <- factor(rep(1:4,3))
levels(strata(b))
levels(strata(a,b,shortlabel=TRUE))

coxph(Surv(futime, fustat) ~ age + strata(rx), data=ovarian)</pre>
```

120 summary.aareg

summary.aareg	Summarize an aareg fit

### **Description**

Creates the overall test statistics for an Aalen additive regression model

## Usage

```
## S3 method for class 'aareg'
summary(object, maxtime, test=c("aalen", "nrisk"), scale=1,...)
```

## **Arguments**

object the result of a call to the aareg function

maxtime truncate the input to the model at time "maxtime"

test the relative time weights that will be used to compute the test

scale scales the coefficients. For some data sets, the coefficients of the Aalen model

will be very small (10-4); this simply multiplies the printed values by a constant,

say 1e6, to make the printout easier to read.

for future methods

### **Details**

It is not uncommon for the very right-hand tail of the plot to have large outlying values, particularly for the standard error. The maxtime parameter can then be used to truncate the range so as to avoid these. This gives an updated value for the test statistics, without refitting the model.

The slope is based on a weighted linear regression to the cumulative coefficient plot, and may be a useful measure of the overall size of the effect. For instance when two models include a common variable, "age" for instance, this may help to assess how much the fit changed due to the other variables, in leiu of overlaying the two plots. (Of course the plots are often highly non-linear, so it is only a rough substitute). The slope is not directly related to the test statistic, as the latter is invariant to any monotone transformation of time.

## Value

a list is returned with the following components

table a matrix with rows for the intercept and each covariate, and columns giving a

slope estimate, the test statistic, it's standard error, the z-score and a p-value

test the time weighting used for computing the test statistics

test.statistic

the vector of test statistics

test.var the model based variance matrix for the test statistic test.var2 optionally, a robust variance matrix for the test statistic

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chisq n the overall test (ignoring the intercept term) for significance of any variable a vector containing the number of observations, the number of unique death times used in the computation, and the total number of unique death times

### See Also

aareg, plot.aareg

# Examples

```
afit <- aareg(Surv(time, status) ~ age + sex + ph.ecog, data=lung,
    dfbeta=TRUE)
summary(afit)
## Not run:
                   test se(test) robust se
             slope
                                               Z
Intercept 5.05e-03
                   1.9 1.54
                                     1.55 1.23 0.219000
     age 4.01e-05 108.0 109.00
                                    106.00 1.02 0.307000
                          5.90
     sex -3.16e-03 -19.5
                                     5.95 -3.28 0.001030
 ph.ecog 3.01e-03 33.2
                             9.18
                                      9.17 3.62 0.000299
Chisq=22.84 on 3 df, p=4.4e-05; test weights=aalen
## End(Not run)
summary(afit, maxtime=600)
## Not run:
                    test se(test) robust se
             slope
                                                Z
Intercept 4.16e-03
                   2.13
                           1.48
                                     1.47 1.450 0.146000
                           106.00
     age 2.82e-05 85.80
                                    100.00 0.857 0.392000
                          5.61
     sex -2.54e-03 -20.60
                                    5.63 -3.660 0.000256
                             8.91
 ph.ecog 2.47e-03 31.60
                                      8.67 3.640 0.000271
Chisq=27.08 on 3 df, p=5.7e-06; test weights=aalen
## End(Not run)
```

summary.coxph

Summary method for Cox models

## **Description**

Produces a summary of a fitted coxph model

## Usage

```
## S3 method for class 'coxph'
summary(object, conf.int=0.95, scale=1,...)
```

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## **Arguments**

object the result of a coxph fit

conf.int level for computation of the confidence intervals. If set to FALSE no confidence

intervals are printed

scale vector of scale factors for the coefficients, defaults to 1. The printed coefficients,

se, and confidence intervals will be associated with one scale unit.

... for future methods

## Value

An object of class summary.coxph, with components:

n, nevent number of observations and number of events, respectively, in the fit

loglik the log partial likelihood at the initial and final values

coefficients a matrix with one row for each coefficient, and columns containing the coeffi-

cient, the hazard ratio exp(coef), standard error, Wald statistic, and P value.

conf.int a matrix with one row for each coefficient, containing the confidence limits for

exp(coef)

logtest, sctest, waldtest

the overall likelihood ratio, score, and Wald test statistics for the model

concordance the concordance statistic and its standard error

used.robust whether an asymptotic or robust variance was used

rsq an approximate R^2 based on Nagelkirke (Biometrika 1991).

fail a message, if the underlying coxph call failed

call a copy of the call

na.action information on missing values

## Note

The pseudo r-squared of Nagelkirke is attractive because it is simple, but further work has shown that it has poor properties and it is now depricated. The value is no longer printed by default, and will eventually be removed from the object.

### See Also

```
coxph, print.coxph
```

### **Examples**

```
fit <- coxph(Surv(time, status) ~ age + sex, lung)
summary(fit)</pre>
```

summary.pyears 123

|--|

# Description

Create a printable table of a person-years result.

# Usage

```
## S3 method for class 'pyears'
summary(object, header = TRUE, call = header, n = TRUE,
event = TRUE, pyears = TRUE, expected = TRUE, rate = FALSE, rr =expected,
ci.r = FALSE, ci.rr = FALSE, totals=FALSE, legend = TRUE, vline = FALSE,
vertical= TRUE, nastring=".", conf.level = 0.95,
scale = 1, ...)
```

# Arguments

object	a pyears object
header	print out a header giving the total number of observations, events, person-years, and total time (if any) omitted from the table
call	print out a copy of the call
n, event, pyear	s, expected
	logical arguments: should these elements be printed in the table?
rate, ci.r	logical arguments: should the incidence rate and/or its confidence interval be given in the table?
rr, ci.rr	logical arguments: should the hazard ratio and/or its confidence interval be given in the table?
totals	should row and column totals be added?
legend	should a legend be included in the printout?
vline	should vertical lines be included in the printed tables?
vertical	when there is only a single predictor, should the table be printed with the predictor on the left (vertical=TRUE) or across the top (vertical=FALSE)?
nastring	what to use for missing values in the table. Some of these are structural, e.g., risk ratios for a cell with no follow-up time.
conf.level	confidence level for any confidence intervals
scale	a scaling factor for printed rates
•••	optional arguments which will be passed to the format function; common choices would be digits= $2$ or nsmall= $1$ .

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### **Details**

The pyears function is often used to create initial descriptions of a survival or time-to-event variable; the type of material that is often found in "table 1" of a paper. The summary routine prints this information out using one of pandoc table styles. A primary reason for choosing this style is that Rstudio is then able to automatically render the results in multiple formats: html, rtf, latex, etc.

If the pyears call has only a single covariate then the table will have that covariate as one margin and the statistics of interest as the other. If the pyears call has two predictors then those two predictors are used as margins of the table, while each cell of the table contains the statistics of interest as multiple rows within the cell. If there are more than two predictors then multiple tables are produced, in the same order as the standard R printout for an array.

The "N" entry of a pyears object is the number of observations which contributed to a particular cell. When the original call includes tcut objects then a single observation may contribute to multiple cells.

## Value

a copy of the object

### Notes

The pandoc system has four table types: with or without vertical bars, and with single or multiple rows of data in each cell. This routine produces all 4 styles depending on options, but currently not all of them are recognized by the Rstudio-pandoc pipeline. (And we don't yet see why.)

### Author(s)

Terry Therneau and Elizabeth Atkinson

### See Also

```
cipoisson, pyears, format
```

summary.survexp

Summary function for a survexp object

# Description

Returns a list containing the values of the survival at specified times.

# Usage

```
## S3 method for class 'survexp'
summary(object, times, scale = 1, ...)
```

summary.survfit 125

## **Arguments**

object the result of a call to the survexp function

times vector of times; the returned matrix will contain 1 row for each time. Missing

values are not allowed.

scale numeric value to rescale the survival time, e.g., if the input data to survfit were

in days, scale = 365.25 would scale the output to years.

... For future methods

### **Details**

A primary use of this function is to retrieve survival at fixed time points, which will be properly interpolated by the function.

## Value

a list with the following components:

surv the estimate of survival at time t.
time the timepoints on the curve.

n.risk In expected survival each subject from the data set is matched to a hypothetical

person from the parent population, matched on the characteristics of the parent population. The number at risk is the number of those hypothetical subject who

are still part of the calculation.

## Author(s)

Terry Therneau

### See Also

survexp

summary.survfit

Summary of a Survival Curve

## **Description**

Returns a list containing the survival curve, confidence limits for the curve, and other information.

## Usage

```
## S3 method for class 'survfit'
summary(object, times, censored=FALSE, scale=1,
    extend=FALSE, rmean=getOption('survfit.rmean'), ...)
```

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## **Arguments**

object the result of a call to the survfit function.

times vector of times; the returned matrix will contain 1 row for each time. The

vector will be sorted into increasing order; missing values are not allowed. If censored=T, the default times vector contains all the unique times in fit, oth-

erwise the default times vector uses only the event (death) times.

censored logical value: should the censoring times be included in the output? This is

ignored if the times argument is present.

scale numeric value to rescale the survival time, e.g., if the input data to survfit were

in days, scale = 365.25 would scale the output to years.

extend logical value: if TRUE, prints information for all specified times, even if there

are no subjects left at the end of the specified times. This is only used if the

times argument is present.

rmean Show restricted mean: see print.survfit for details

... for future methods

#### Value

a list with the following components:

surv the estimate of survival at time t+0.

time the timepoints on the curve.

n.risk the number of subjects at risk at time t-0 (but see the comments on weights in

the survfit help file).

n.event if the times argument is missing, then this column is the number of events that

occurred at time t. Otherwise, it is the cumulative number of events that have

occurred since the last time listed until time t+0.

n.entered This is present only for counting process survival data. If the times argument is

missing, this column is the number of subjects that entered at time t. Otherwise, it is the cumulative number of subjects that have entered since the last time listed

until time t.

n.exit.censored

if the times argument is missing, this column is the number of subjects that left without an event at time t. Otherwise, it is the cumulative number of subjects that have left without an event since the last time listed until time t+0. This is

only present for counting process survival data.

std.err the standard error of the survival value.

conf. int level of confidence for the confidence intervals of survival.

lower lower confidence limits for the curve.
upper upper confidence limits for the curve.

strata indicates stratification of curve estimation. If strata is not NULL, there are mul-

tiple curves in the result and the surv, time, n.risk, etc. vectors will contain multiple curves, pasted end to end. The levels of strata (a factor) are the labels

for the curves.

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call the statement used to create the fit object.

na.action same as for fit, if present.

table table of information that is returned from print.survfit function.

type type of data censoring. Passed through from the fit object.

#### **Details**

This routine has two uses: printing out a survival curve at specified time points (often yearly), or extracting the values at specified time points for further processing. In the first case we normally want extend=FALSE, i.e., don't print out data past the end of the curve. If the times option only contains values beyond the last point in the curve then there is nothing to print and an error message will result. For the second usage we almost always want extend=TRUE, so that the results will have a predictable length.

The survfit object itself will have a row of information at each censoring or event time, it does not save information on each unique entry time. For printout at two time points t1, t2, this function will give the number at risk at the smallest event times that are >= t1 and >= t2, respectively, the survival curve at the largest recorded times <= t1 and <= t2, and the number of events and censorings in the interval t1 < t <= t2.

When the routine is called with counting process data many users are confused by counts that are too large. For example, Surv(c(0,0,5,5),c(2,3,8,10),c(1,0,1,0)) followed by a request for the values at time 4. The Surv(i) object has entries only at times 2, 3, 8, and 10; there are 2 subjects at risk at time 8, so that is what will be printed.

### See Also

```
survfit, print.summary.survfit
```

## **Examples**

```
summary( survfit( Surv(futime, fustat)~1, data=ovarian))
summary( survfit( Surv(futime, fustat)~rx, data=ovarian))
```

Surv

Create a Survival Object

## Description

Create a survival object, usually used as a response variable in a model formula. Argument matching is special for this function, see Details below.

### Usage

```
Surv(time, time2, event,
          type=c('right', 'left', 'interval', 'counting', 'interval2', 'mstate'),
          origin=0)
is.Surv(x)
```

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### **Arguments**

time for right censored data, this is the follow up time. For interval data, the first argument is the starting time for the interval. The status indicator, normally 0=alive, 1=dead. Other choices are TRUE/FALSE event (TRUE = death) or 1/2 (2=death). For interval censored data, the status indicator is 0=right censored, 1=event at time, 2=left censored, 3=interval censored. For multiple endpoint data the event variable will be a factor, whose first level is treated as censoring. Although unusual, the event indicator can be omitted, in which case all subjects are assumed to have an event. time2 ending time of the interval for interval censored or counting process data only. Intervals are assumed to be open on the left and closed on the right, (start, end]. For counting process data, event indicates whether an event occurred at the end of the interval. character string specifying the type of censoring. Possible values are "right", type "left", "counting", "interval", "interval2" or "mstate". origin for counting process data, the hazard function origin. This option was intended to be used in conjunction with a model containing time dependent strata in order to align the subjects properly when they cross over from one strata to another, but it has rarely proven useful. any R object. Х

#### **Details**

When the type argument is missing the code assumes a type based on the following rules:

- If there are two unnamed arguments, they will match time and event in that order. If there are three unnamed arguments they match time, time2 and event.
- If the event variable is a factor then type mstate is assumed. Otherwise type right if there is no time2 argument, and type counting if there is.

As a consequence the type argument will normally be omitted.

When the survival type is "mstate" then the status variable will be treated as a factor. The first level of the factor is taken to represent censoring and remaining ones a transition to the given state. (If the status variable is a factor then mstate is assumed.)

Interval censored data can be represented in two ways. For the first use type = "interval" and the codes shown above. In that usage the value of the time2 argument is ignored unless event=3. The second approach is to think of each observation as a time interval with (-infinity, t) for left censored, (t, infinity) for right censored, (t,t) for exact and (t1, t2) for an interval. This is the approach used for type = interval2. Infinite values can be represented either by actual infinity (Inf) or NA. The second form has proven to be the more useful one.

Presently, the only methods allowing interval censored data are the parametric models computed by survreg and survival curves computed by survfit; for both of these, the distinction between open and closed intervals is unimportant. The distinction is important for counting process data and the Cox model.

The function tries to distinguish between the use of 0/1 and 1/2 coding for censored data via the condition if  $(\max(\text{status})==2)$ . If 1/2 coding is used and all the subjects are censored, it will

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guess wrong. In any questionable case it is safer to use logical coding, e.g., Surv(time, status==3) would indicate that '3' is the code for an event. For multi-state survival the status variable will be a factor, whose first level is assumed to correspond to censoring.

Surv objects can be subscripted either as a vector, e.g. x[1:3] using a single subscript, in which case the drop argument is ignored and the result will be a survival object; or as a matrix by using two subscripts. If the second subscript is missing and drop=F (the default), the result of the subscripting will be a Surv object, e.g., x[1:3,,drop=F], otherwise the result will be a matrix (or vector), in accordance with the default behavior for subscripting matrices.

#### Value

An object of class Surv. There are methods for print, is.na, and subscripting survival objects. Surv objects are implemented as a matrix of 2 or 3 columns that has further attributes. These include the type (left censored, right censored, counting process, etc.) and labels for the states for multi-state objects. Any attributes of the input arguments are also preserved in inputAttributes. This may be useful for other packages that have attached further information to data items such as labels; none of the routines in the survival package make use of these values, however.

In the case of is. Surv, a logical value TRUE if x inherits from class "Surv", otherwise an FALSE.

### Note

The use of 1/2 coding for status is an interesting historical artifact. For data contained on punch cards, IBM 360 Fortran treated blank as a zero, which led to a policy within the Mayo Clinic section of Biostatistics to never use "0" as a data value since one could not distinguish it from a missing value. Policy became habit, as is often the case, and the use of 1/2 coding for alive/dead endured long after the demise of the punch cards that had sired the practice. At the time Surv was written many Mayo data sets still used this convention, e.g., the 1994 lung data set found in the package.

### See Also

```
coxph, survfit, survreg, lung.
```

## **Examples**

```
with(aml, Surv(time, status))
survfit(Surv(time, status) ~ ph.ecog, data=lung)
Surv(heart$start, heart$stop, heart$event)
```

Surv-methods

Methods for Surv objects

## Description

The list of methods that apply to Surv objects

Surv-methods

## Usage

```
## S3 method for class 'Surv'
anyDuplicated(x, ...)
    ## S3 method for class 'Surv'
as.character(x, ...)
    ## S3 method for class 'Surv'
as.data.frame(x, ...)
    ## S3 method for class 'Surv'
as.integer(x, ...)
    ## S3 method for class 'Surv'
as.matrix(x, ...)
    ## S3 method for class 'Surv'
as.numeric(x, ...)
    ## S3 method for class 'Surv'
c(...)
    ## S3 method for class 'Surv'
duplicated(x, ...)
    ## S3 method for class 'Surv'
format(x, ...)
    ## S3 method for class 'Surv'
head(x, ...)
    ## S3 method for class 'Surv'
is.na(x)
    ## S3 method for class 'Surv'
length(x)
    ## S3 method for class 'Surv'
mean(x, ...)
    ## S3 method for class 'Surv'
median(x, ...)
    ## S3 method for class 'Surv'
names(x)
    ## S3 replacement method for class 'Surv'
names(x) \leftarrow value
    ## S3 method for class 'Surv'
quantile(x, probs, na.rm=FALSE, ...)
    ## S3 method for class 'Surv'
plot(x, ...)
    ## S3 method for class 'Surv'
rep(x, ...)
    ## S3 method for class 'Surv'
rep.int(x, ...)
    ## S3 method for class 'Surv'
rep_len(x, ...)
    ## S3 method for class 'Surv'
rev(x)
    ## S3 method for class 'Surv'
t(x)
    ## S3 method for class 'Surv'
```

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```
tail(x, ...)
    ## S3 method for class 'Surv'
unique(x, ...)
```

## Arguments

x a Surv object
probs a vector of probabilities
na.rm remove missing values from the calculation
value a character vector of up to the same length as x, or NULL
... other arguments to the method

### **Details**

These functions extend the standard methods to Surv objects. The arguments and results from these are mostly as expected, with the following further details:

- The as.character function uses "5+" for right censored at time 5, "5-" for left censored at time 5, "[2,7]" for an observation that was interval censored between 2 and 7, "(1,6]" for a counting process data denoting an observation which was at risk from time 1 to 6, with an event at time 6, and "(1,6+]" for an observation over the same interval but not ending with and event. For a multi-state survival object the type of event is appended to the event time using ":type".
- The print and format methods make use of as.character.
- The as.numeric and as.integer methods perform these actions on the survival times, but do not affect the censoring indicator.
- The as.matrix and t methods return a matrix
- The length of a Surv object is the number of survival times it contains, not the number of items required to encode it, e.g., x <-Surv(1:4,5:9,c(1,0,1,0)); length(x) has a value of 4. Likewise names(x) will be NULL or a vector of length 4. (For technical reasons, any names are actually stored in the rownames attribute of the object.)
- For a multi-state survival object levels returns the names of the endpoints, otherwise it is NULL.
- The median, quantile and plot methods first construct a survival curve using survfit, then apply the appropriate method to that curve.
- The concatonation method c() is asymmetric, its first argument determines the exection path. For instance c(Surv(1:4), Surv(5:6)) will concatonate the two objects, c(Surv(1:4), 5:6) will give an error, and c(5:6, Surv(1:4)) is equivalent to c(5:6, as.vector(Surv(1:4))).

### See Also

Surv

132 Surv2

Surv2	Create a survival object	

## **Description**

Create a survival object from a timeline style data set. This will almost always be the response variable in a formula.

## Usage

```
Surv2(time, event, repeated=FALSE)
```

## **Arguments**

time a timeline variable, such as age, time from enrollment, date, etc.

event the outcome at that time. This can be a 0/1 variable, TRUE/FALSE, or a factor.

If the latter, the first level of the factor corresponds to 'no event was observed at

this time'.

repeated if the same level of the outcome repeats, without an intervening event of another

type, should this be treated as a new event?

## **Details**

This function is still experimental.

When used in a coxph or survfit model, Surv2 acts as a trigger to internally convert a timeline style data set into counting process style data, which is then acted on by the routine.

The repeated argument controls how repeated instances of the same event code are treated. If TRUE, they are treated as new events, an example where this might be desired is repeated infections in a subject. If FALSE, then repeats are not a new event. An example would be a data set where we wanted to use diabetes, say, as an endpoint, but this is repeated at each medical visit.

#### Value

An object of class Surv2. There are methods for print, is.na and subscripting.

# See Also

Surv2data, coxph, survfit

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Surv2data Convert data from timecourse to (time1,time2) style	Surv2data	Convert data from timecourse to (time1,time2) style	
---	-----------	---	--

## Description

The multi-state survival functions coxph and survfit allow for two forms of input data. This routine converts between them. The function is normally called behind the scenes when Surv2 is as the response.

## Usage

```
Surv2data(formula, data, subset, id)
```

### **Arguments**

formula a model formula data a data frame

subset optional, selects rows of the data to be retained

id a variable that identified multiple rows for the same subject, normally found in

the referenced data set

## Details

For timeline style data, each row is uniquely identified by an (identifier, time) pair. The time could be a date, time from entry to a study, age, etc, (there may often be more than one time variable). The identifier and time cannot be missing. The remaining covariates represent values that were observed at that time point. Often, a given covariate is observed at only a subset of times and is missing at others. At the time of death, in particular, often only the identifier, time, and status indicator are known.

In the resulting data set missing covariates are replaced by their last known value, and the response y will be a Surv(time1, time2, endpoint) object.

#### Value

a list with elements

mf an updated model frame (fewer rows, unchanged columns)

S2.y the constructed response variable
S2.state the current state for each of the rows

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survcheck	Checks of a survival data set	

# Description

Perform a set of consistency checks on survival data

## Usage

```
survcheck(formula, data, subset, na.action, id, istate, istate0="(s0)",
timefix=TRUE,...)
```

## **Arguments**

formula	a model formula with a Surv object as the response
data	data frame in which to find the id, istate and formula variables
subset	expression indicating which subset of the rows of data should be used in the fit. All observations are included by default.
na.action	a missing-data filter function. This is applied to the model.frame after any subset argument has been used. Default is options()\$na.action.
id	an identifier that labels unique subjects
istate	an optional vector giving the current state at the start of each interval
istate0	default label for the initial state of each subject (at their first interval) when istate is missing
timefix	process times through the aeqSurv function to eliminate potential roundoff issues.
•••	other arguments, which are ignored (but won't give an error if someone added weights for instance)

#### **Details**

This routine will examine a multi-state data set for consistency of the data. The basic rules are that if a subject is at risk they have to be somewhere, can not be two states at once, and should make sensible transitions from state to state. It reports the number of instances of the following conditions:

**overlap** two observations for the same subject that overlap in time, e.g. intervals of (0, 100) and (90, 120). If y is simple (time, status) survival observation intervals implicitly start at 0, so in that case any duplicate identifiers will generate an overlap.

**jump** a hole in a subject's timeline, where they are in one state at the end of the prior interval, but a new state in the at the start subsequent interval.

**gap** one or more gaps in a subject's timeline; they are presumably in the same state at their return as when they left.

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**teleport** two adjacent intervals for a subject, with the first interval ending in one state and the subsequent interval starting in another. They have instantaneously changed states with experiencing a transition.

The total number of occurences of each is present in the flags vector. Optional components give the location and identifiers of the flagged observations.

#### Value

## a list with components

the vector of possible states states a matrix giving the count of transitions from one state to another transitions statecount table of the number of visits per state, e.g., 18 subjects had 2 visits to the "infection" state flags a vector giving the counts of each check a copy of the istate vector, if it was supplied; otherwise a constructed istate that istate satisfies all the checks overlap a list with the row number and id of overlaps (not present if there are no overlaps) a list with the row number and id of gaps (not present if there are no gaps) gaps

teleport a list with the row number and id of inconsistent rows (not present if there are

none)

jumps a list with the row number and id of jumps (not present if there are no jumps

#### Note

For data sets with time-dependent covariates, a given subject will often have intermediate rows with a status of 'no event at this time'. (numeric value of 0). For instance a subject who started in state 1 at time 0, transitioned to state 2 at time 10, had a covariate x change from 135 to 156 at time 20, and a final transition to state 3 at time 30. The response would be Surv(c(0,10,20),c(10,20,30),c(2,0,3)): the status variable records *changes* in state, and there was no change at time 20. The istate variable would be (1, 2, 2); it contains the *current* state, and so the value is unchanged when status = censored.

Thus, when there are intermediate observations istate is not simply a lagged version of the status, and may be more challenging to create. One approach is to let survcheck do the work: call it with an istate argument that is correct for the first row of each subject, or no istate argument at all, and then insert the returned value into ones data frame.

|--|

### **Description**

Tests if there is a difference between two or more survival curves using the  $G^{\rho}$  family of tests, or for a single curve against a known alternative.

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### Usage

survdiff(formula, data, subset, na.action, rho=0, timefix=TRUE)

### **Arguments**

formula a formula expression as for other survival models, of the form Surv(time, status)

~ predictors. For a one-sample test, the predictors must consist of a single offset(sp) term, where sp is a vector giving the survival probability of each subject. For a k-sample test, each unique combination of predictors defines a subgroup. A strata term may be used to produce a stratified test. To cause missing values in the predictors to be treated as a separate group, rather than

being omitted, use the strata function with its na.group=T argument.

data an optional data frame in which to interpret the variables occurring in the for-

mula.

subset expression indicating which subset of the rows of data should be used in the fit.

This can be a logical vector (which is replicated to have length equal to the number of observations), a numeric vector indicating which observation numbers are to be included (or excluded if negative), or a character vector of row names to

be included. All observations are included by default.

na.action a missing-data filter function. This is applied to the model.frame after any

subset argument has been used. Default is options()\$na.action.

rho a scalar parameter that controls the type of test.

timefix process times through the aeqSurv function to eliminate potential roundoff is-

sues.

### Value

a list with components:

n the number of subjects in each group.

obs the weighted observed number of events in each group. If there are strata, this

will be a matrix with one column per stratum.

exp the weighted expected number of events in each group. If there are strata, this

will be a matrix with one column per stratum.

chisq the chisquare statistic for a test of equality.

var the variance matrix of the test.

strata optionally, the number of subjects contained in each stratum.

### **METHOD**

This function implements the G-rho family of Harrington and Fleming (1982), with weights on each death of  $S(t)^{\rho}$ , where S(t) is the Kaplan-Meier estimate of survival. With rho = 0 this is the log-rank or Mantel-Haenszel test, and with rho = 1 it is equivalent to the Peto & Peto modification of the Gehan-Wilcoxon test.

If the right hand side of the formula consists only of an offset term, then a one sample test is done. To cause missing values in the predictors to be treated as a separate group, rather than being omitted, use the factor function with its exclude argument.

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### References

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

## **Examples**

survexp

Compute Expected Survival

### Description

Returns either the expected survival of a cohort of subjects, or the individual expected survival for each subject.

## Usage

## **Arguments**

formula

formula object. The response variable is a vector of follow-up times and is optional. The predictors consist of optional grouping variables separated by the + operator (as in survfit), and is often ~1, i.e., expected survival for the entire group.

data

data frame in which to interpret the variables named in the formula, subset and weights arguments.

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weights	case weights. This is most useful when conditional survival for a known popula-
	tion is desired, e.g., the data set would contain all unique age/sex combinations and the weights would be the proportion of each.
subset	expression indicating a subset of the rows of data to be used in the fit.
na.action	function to filter missing data. This is applied to the model frame after subset has been applied. Default is options()\$na.action.
rmap	an optional list that maps data set names to the ratetable names. See the details section below.
times	vector of follow-up times at which the resulting survival curve is evaluated. If absent, the result will be reported for each unique value of the vector of times supplied in the response value of the formula.
method	computational method for the creating the survival curves. The individual option does not create a curve, rather it retrieves the predicted survival individual.s or cumulative hazard individual.h for each subject. The default is to use method='ederer' if the formula has no response, and method='hakulinen' otherwise.
cohort	logical value. This argument has been superseded by the method argument. To maintain backwards compatability, if is present and FALSE, it implies method='individual.s'.
conditional	logical value. This argument has been superseded by the method argument. To maintain backwards compatability, if it is present and TRUE it implies method='conditional'.
ratetable	a table of event rates, such as survexp.mn, or a fitted Cox model. Note the survival:: prefix in the default argument is present to avoid the (rare) case of a user who expects the default table but just happens to have an object named "survexp.us" in their own directory.
scale	numeric value to scale the results. If ratetable is in units/day, scale = 365.25 causes the output to be reported in years.
se.fit	compute the standard error of the predicted survival. This argument is currently ignored. Standard errors are not a defined concept for population rate tables (they are treated as coming from a complete census), and for Cox models the calculation is hard. Despite good intentions standard errors for this latter case have not been coded and validated.
model,x,y	flags to control what is returned. If any of these is true, then the model frame, the model matrix, and/or the vector of response times will be returned as components of the final result, with the same names as the flag arguments.

## **Details**

Individual expected survival is usually used in models or testing, to 'correct' for the age and sex composition of a group of subjects. For instance, assume that birth date, entry date into the study, sex and actual survival time are all known for a group of subjects. The survexp.us population tables contain expected death rates based on calendar year, sex and age. Then

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gives for each subject the total hazard experienced up to their observed death time or last follow-up time (variable fu.time) This probability can be used as a rescaled time value in models:

```
glm(status ~ 1 + offset(log(haz)), family=poisson)
glm(status ~ x + offset(log(haz)), family=poisson)
```

In the first model, a test for intercept=0 is the one sample log-rank test of whether the observed group of subjects has equivalent survival to the baseline population. The second model tests for an effect of variable x after adjustment for age and sex.

The ratetable being used may have different variable names than the user's data set, this is dealt with by the rmap argument. The rate table for the above calculation was survexp.us, a call to summary{survexp.us} reveals that it expects to have variables age = age in days, sex, and year = the date of study entry, we create them in the rmap line. The sex variable was not mapped, therefore the function assumes that it exists in mydata in the correct format. (Note: for factors such as sex, the program will match on any unique abbreviation, ignoring case.)

Cohort survival is used to produce an overall survival curve. This is then added to the Kaplan-Meier plot of the study group for visual comparison between these subjects and the population at large. There are three common methods of computing cohort survival. In the "exact method" of Ederer the cohort is not censored, for this case no response variable is required in the formula. Hakulinen recommends censoring the cohort at the anticipated censoring time of each patient, and Verheul recommends censoring the cohort at the actual observation time of each patient. The last of these is the conditional method. These are obtained by using the respective time values as the follow-up time or response in the formula.

## Value

if cohort=TRUE an object of class survexp, otherwise a vector of per-subject expected survival values. The former contains the number of subjects at risk and the expected survival for the cohort at each requested time. The cohort survival is the hypothetical survival for a cohort of subjects enrolled from the population at large, but matching the data set on the factors found in the rate table.

#### References

Berry, G. (1983). The analysis of mortality by the subject-years method. *Biometrics*, 39:173-84.

Ederer, F., Axtell, L. and Cutler, S. (1961). The relative survival rate: a statistical methodology. *Natl Cancer Inst Monogr*, 6:101-21.

Hakulinen, T. (1982). Cancer survival corrected for heterogeneity in patient withdrawal. *Biometrics*, 38:933-942.

Therneau, T. and Grambsch, P. (2000). Modeling survival data: Extending the Cox model. Springer. Chapter 10.

Verheul, H., Dekker, E., Bossuyt, P., Moulijn, A. and Dunning, A. (1993). Background mortality in clinical survival studies. *Lancet*, 341: 872-875.

### See Also

```
survfit, pyears, survexp.us, ratetable, survexp.fit.
```

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## **Examples**

survexp.fit

Compute Expected Survival

## **Description**

Compute expected survival times.

## Usage

```
survexp.fit(group, x, y, times, death, ratetable)
```

# Arguments

group	if there are multiple survival curves this identifies the group, otherwise it is a constant. Must be an integer.
X	A matrix whose columns match the dimensions of the ratetable, in the correct order.
у	the follow up time for each subject.
times	the vector of times at which a result will be computed.
death	a logical value, if TRUE the conditional survival is computed, if FALSE the cohort survival is computed. See survexp for more details.
ratetable	a rate table, such as survexp.uswhite.

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### **Details**

For conditional survival y must be the time of last follow-up or death for each subject. For cohort survival it must be the potential censoring time for each subject, ignoring death.

For an exact estimate times should be a superset of y, so that each subject at risk is at risk for the entire sub-interval of time. For a large data set, however, this can use an inordinate amount of storage and/or compute time. If the times spacing is more coarse than this, an actuarial approximation is used which should, however, be extremely accurate as long as all of the returned values are > .99.

For a subgroup of size 1 and times > y, the conditional method reduces to exp(-h) where h is the expected cumulative hazard for the subject over his/her observation time. This is used to compute individual expected survival.

### Value

A list containing the number of subjects and the expected survival(s) at each time point. If there are multiple groups, these will be matrices with one column per group.

### Warning

Most users will call the higher level routine survexp. Consequently, this function has very few error checks on its input arguments.

### See Also

survexp, survexp.us.

survexp.object Expected Survival Curve Object
---

## Description

This class of objects is returned by the survexp class of functions to represent a fitted survival curve.

Objects of this class have methods for summary, and inherit the print, plot, points and lines methods from survfit.

### **Arguments**

surv	the estimate of survival at time t+0. This may be a vector or a matrix.
n.risk	the number of subjects who contribute at this time.
time	the time points at which the curve has a step.
std.err	the standard error of the cumulative hazard or -log(survival).
strata	if there are multiple curves, this component gives the number of elements of the time etc. vectors corresponding to the first curve, the second curve, and so on. The names of the elements are labels for the curves.
method	the estimation method used. One of "Ederer", "Hakulinen", or "conditional".

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na.action the returned value from the na.action function, if any. It will be used in the

printout of the curve, e.g., the number of observations deleted due to missing

values.

call an image of the call that produced the object.

### Structure

The following components must be included in a legitimate survfit object.

# **Subscripts**

Survexp objects that contain multiple survival curves can be subscripted. This is most often used to plot a subset of the curves.

### **Details**

In expected survival each subject from the data set is matched to a hypothetical person from the parent population, matched on the characteristics of the parent population. The number at risk printed here is the number of those hypothetical subject who are still part of the calculation. In particular, for the Ederer method all hypotheticals are retained for all time, so n.risk will be a constant.

#### See Also

```
plot.survfit, summary.survexp, print.survfit, survexp.
```

survfit

Create survival curves

## Description

This function creates survival curves from either a formula (e.g. the Kaplan-Meier), a previously fitted Cox model, or a previously fitted accelerated failure time model.

# Usage

```
survfit(formula, ...)
```

# Arguments

formula either a formula or a previously fitted model
... other arguments to the specific method

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### **Details**

A survival curve is based on a tabulation of the number at risk and number of events at each unique death time. When time is a floating point number the definition of "unique" is subject to interpretation. The code uses factor() to define the set. For further details see the documentation for the appropriate method, i.e., ?survfit.formula or ?survfit.coxph.

A survfit object may contain a single curve, a set of curves, or a matrix curves. Predicted curves from a coxph model have one row for each stratum in the Cox model fit and one column for each specified covariate set. Curves from a multi-state model have one row for each stratum and a column for each state, the strata correspond to predictors on the right hand side of the equation. The default printing and plotting order for curves is by column, as with other matrices.

Curves can be subscripted using either a single or double subscript. If the set of curves is a matrix, as in the above, and one of the dimensions is 1 then the code allows a single subscript to be used. (That is, it is not quite as general as using a single subscript for a numeric matrix.)

### Value

An object of class survfit containing one or more survival curves.

### Note

Older releases of the code also allowed the specification for a single curve to omit the right hand of the formula, i.e., survfit(Surv(time, status)), in which case the formula argument is not actually a formula. Handling this case required some non-standard and fairly fragile manipulations, and this case is no longer supported.

#### Author(s)

Terry Therneau

#### See Also

survfit.formula, survfit.coxph, survfit.object, print.survfit, plot.survfit, quantile.survfit,
residuals.survfit, summary.survfit

survfit.coxph

Compute a Survival Curve from a Cox model

## **Description**

Computes the predicted survivor function for a Cox proportional hazards model.

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## Usage

### **Arguments**

formula A coxph object.

newdata a data frame with the same variable names as those that appear in the coxph

formula. It is also valid to use a vector, if the data frame would consist of a

single row.

The curve(s) produced will be representative of a cohort whose covariates correspond to the values in newdata. Default is the mean of the covariates used in

the coxph fit.

se.fit a logical value indicating whether standard errors should be computed. Default

is TRUE.

conf. int the level for a two-sided confidence interval on the survival curve(s). Default is

0.95.

individual depricated argument, replaced by the general id

stype computation of the survival curve, 1=direct, 2= exponenial of the cumulative

hazard.

ctype whether the cumulative hazard computation should have a correction for ties,

1=no, 2=yes.

conf.type One of "none", "plain", "log" (the default), "log-log" or "logit". Only

enough of the string to uniquely identify it is necessary. The first option causes confidence intervals not to be generated. The second causes the standard intervals curve +-k \*se(curve), where k is determined from conf.int. The log option calculates intervals based on the cumulative hazard or log(survival). The log-log option uses the log hazard or log(-log(survival)), and the logit log(survival/(1-

survival)).

censor if FALSE time points at which there are no events (only censoring) are not in-

cluded in the result.

id optional variable name of subject identifiers. If this is present, it will be search

for in the newdata data frame. Each group of rows in newdata with the same subject id represents the covariate path through time of a single subject, and the result will contain one curve per subject. If the coxph fit had strata then that must also be specified in newdata. If newid is not present, then each individual

row of newdata is presumed to represent a distinct subject.

start.time optional starting time, a single numeric value. If present the returned curve

contains survival after start.time conditional on surviving to start.time.

influence option to return the influence values

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na. action the na.action to be used on the newdata argument

type older argument that encompassed stype and ctype, now depricated

... for future methods

#### **Details**

This routine produces Pr(state) curves based on a coxph model fit. For single state models it produces the single curve for S(t) = Pr(remain in initial state at time t), known as the survival curve; for multi-state models a matrix giving probabilities for all states. The stype argument states the type of estimate, and defaults to the exponential of the cumulative hazard, better known as the Breslow estimate. For a multi-state Cox model this involves the exponential of a matrix. The argument stype=1 uses a non-exponential or 'direct' estimate. For a single endpoint coxph model the code evaluates the Kalbfleich-Prentice estimate, and for a multi-state model it uses an analog of the Aalen-Johansen estimator. The latter approach is the default in the mstate package.

The ctype option affects the estimated cumulative hazard, and if stype=2 the estimated P(state) curves as well. If not present it is chosen so as to be concordant with the ties option in the coxph call. (For multistate coxphms objects only ctype=1 is currently implemented.) Likewise the choice between a model based and robust variance estimate for the curve will mirror the choice made in the coxph call, any clustering is also inherited from the parent model.

If the newdata argument is missing, then a curve is produced for a single "pseudo" subject with covariate values equal to the means of the data set. The resulting curve(s) almost never make sense, but The default remains due to an unwarranted attachment to the option shown by some users and by other packages. Two particularly egregious examples are factor variables and interactions. Suppose one were studying interspecies transmission of a virus, and the data set has a factor variable with levels ("pig", "chicken") and about equal numbers of observations for each. The "mean" covariate level will be 0.5 – is this a flying pig? As to interactions assume data with sex coded as 0/1, ages ranging from 50 to 80, and a model with age\*sex. The "mean" value for the age:sex interaction term will be about 30, a value that does not occur in the data. Users are strongly advised to use the newdata argument. For these reasons predictions from a multistate coxph model require the newdata argument.

When the original model contains time-dependent covariates, then the path of that covariate through time needs to be specified in order to obtain a predicted curve. This requires newdata to contain multiple lines for each hypothetical subject which gives the covariate values, time interval, and strata for each line (a subject can change strata), along with an id variable which demarks which rows belong to each subject. The time interval must have the same (start, stop, status) variables as the original model: although the status variable is not used and thus can be set to a dummy value of 0 or 1, it is necessary for the response to be recognized as a Surv object. Last, although predictions with a time-dependent covariate path can be useful, it is very easy to create a prediction that is senseless. Users are encouraged to seek out a text that discusses the issue in detail.

When a model contains strata but no time-dependent covariates the user of this routine has a choice. If newdata argument does not contain strata variables then the returned object will be a matrix of survival curves with one row for each strata in the model and one column for each row in newdata. (This is the historical behavior of the routine.) If newdata does contain strata variables, then the result will contain one curve per row of newdata, based on the indicated stratum of the original model. In the rare case of a model with strata by covariate interactions the strata variable must be included in newdata, the routine does not allow it to be omitted (predictions become too confusing).

(Note that the model Surv(time, status) ~ age\*strata(sex) expands internally to strata(sex) + age:sex; the sex variable is needed for the second term of the model.)

See survfit for more details about the counts (number of events, number at risk, etc.)

## Value

an object of class "survfit". See survfit.object for details. Methods defined for survfit objects are print, plot, lines, and points.

#### **Notes**

If the following pair of lines is used inside of another function then the model=TRUE argument must be added to the coxph call: fit <-coxph(...); survfit(fit). This is a consequence of the non-standard evaluation process used by the model.frame function when a formula is involved.

## References

Fleming, T. H. and Harrington, D. P. (1984). Nonparametric estimation of the survival distribution in censored data. *Comm. in Statistics* **13**, 2469-86.

Kalbfleisch, J. D. and Prentice, R. L. (1980). *The Statistical Analysis of Failure Time Data*. New York: Wiley.

Link, C. L. (1984). Confidence intervals for the survival function using Cox's proportional hazards model with covariates. *Biometrics* **40**, 601-610.

Therneau T and Grambsch P (2000), Modeling Survival Data: Extending the Cox Model, Springer-Verlag.

Tsiatis, A. (1981). A large sample study of the estimate for the integrated hazard function in Cox's regression model for survival data. *Annals of Statistics* **9**, 93-108.

## See Also

```
print.survfit, plot.survfit, lines.survfit, coxph, Surv, strata.
```

survfit.formula

Compute a Survival Curve for Censored Data

## Description

Computes an estimate of a survival curve for censored data using the Aalen-Johansen estimator. For ordinary (single event) survival this reduces to the Kaplan-Meier estimate.

# Usage

#### **Arguments**

formula a formula object, which must have a Surv object as the response on the left of the ~ operator and, if desired, terms separated by + operators on the right. One of the terms may be a strata object. For a single survival curve the right hand side should be ~ 1. data a data frame in which to interpret the variables named in the formula, subset and weights arguments. The weights must be nonnegative and it is strongly recommended that they weights be strictly positive, since zero weights are ambiguous, compared to use of the subset argument. subset expression saying that only a subset of the rows of the data should be used in the a missing-data filter function, applied to the model frame, after any subset arna.action gument has been used. Default is options()\$na.action. the method to be used estimation of the survival curve: 1 = direct,  $2 = \exp(\text{cumulative})$ stype hazard). the method to be used for estimation of the cumulative hazard: 1 = Nelson-Aalen ctype formula, 2 = Fleming-Harrington correction for tied events. id identifies individual subjects, when a given person can have multiple lines of data. cluster used to group observations for the infinitesmal jackknife variance estimate, defaults to the value of id. robust logical, should the function compute a robust variance. For multi-state survival curves this is true by default. For single state data see details, below. istate for multi-state models, identifies the initial state of each subject or observation timefix process times through the aeqSurv function to eliminate potential roundoff isetype a variable giving the type of event. This has been superseded by multi-state Surv objects and is depricated; see example below. this argument is no longer used error The following additional arguments are passed to internal functions called by . . . survfit.

se.fit logical value, default is TRUE. If FALSE then standard error computations are omitted.

conf.type One of "none", "plain", "log" (the default), "log-log", "logit" or "arcsin". Only enough of the string to uniquely identify it is necessary. The first option causes confidence intervals not to be generated. The second causes the standard intervals curve +-k \*se(curve), where k is determined from conf.int. The log option calculates intervals based on the cumulative hazard or log(survival). The log-log option bases the intervals on the log hazard or log(-log(survival)), the logit option on log(survival/(1-survival)) and arcsin on arcsin(survival).

conf.lower a character string to specify modified lower limits to the curve, the upper limit remains unchanged. Possible values are "usual" (unmodified), "peto", and "modified". The modified lower limit is based on an "effective n" argument. The confidence bands will agree with the usual calculation at each death time, but unlike the usual bands the confidence interval becomes wider at each censored observation. The extra width is obtained by multiplying the usual variance by a factor m/n, where n is the number currently at risk and m is the number at risk at the last death time. (The bands thus agree with the un-modified bands at each death time.) This is especially useful for survival curves with a long flat tail.

The Peto lower limit is based on the same "effective n" argument as the modified limit, but also replaces the usual Greenwood variance term with a simple approximation. It is known to be conservative.

**start.time** numeric value specifying a time to start calculating survival information. The resulting curve is the survival conditional on surviving to start.time.

**conf.int** the level for a two-sided confidence interval on the survival curve(s). Default is 0.95.

**se.fit** a logical value indicating whether standard errors should be computed. Default is TRUE.

**influence** a logical value indicating whether to return the infinitesimal jack-knife (influence) values for each subject. These contain the values of the derivative of each value with respect to the case weights of each subject i:  $\partial p/\partial w_i$ , evaluated at the vector of weights. The resulting object will contain influence.surv and influence.chaz components. Alternatively, options of influence=1 or influence=2 will return values for only the survival or hazard curves, respectively.

p0 this applies only to multi-state curves. An optional vector giving the initial probability across the states. If this is missing, then p0 is estimated using the frequency of the starting states of all observations at risk at start.time, or if that is not specified, at the time of the first event.

type an older argument that combined stype and ctype, now depricated. Legal values were "kaplan-meier" which is equivalent to stype=1, ctype=1, "fleming-harrington" which is equivalent to stype=2,ctype=1, and "fh2" which is equivalent to stype=2,ctype=2.

#### Details

If there is a data argument, then variables in the formula, codeweights, subset, id, cluster and istate arguments will be searched for in that data set.

The routine returns both an estimated probability in state and an estimated cumulative hazard estimate. The cumulative hazard estimate is the Nelson-Aalen (NA) estimate or the Fleming-Harrington (FH) estimate, the latter includes a correction for tied event times. The estimated probability in state can estimated either using the exponential of the cumulative hazard, or as a direct estimate using the Aalen-Johansen approach. For single state data the AJ estimate reduces to the Kaplan-Meier and the probability in state to the survival curve; for competing risks data the AJ reduces to the cumulative incidence (CI) estimator. For backward compatability the type argument can be used instead.

When the data set includes left censored or interval censored data (or both), then the EM approach of Turnbull is used to compute the overall curve. Currently this algorithm is very slow, only a survival curve is produced, and it does not support a robust variance.

Robust variance: If a robust is TRUE, or for multi-state curves, then the standard errors of the results will be based on an infinitesimal jackknife (IJ) estimate, otherwise the standard model based estimate will be used. For single state curves, the default for robust will be TRUE if one of: there is a cluster argument, there are non-integer weights, or there is a id statement and at least one of the id values has multiple events, and FALSE otherwise. The default represents our best guess about when one would most often desire a robust variance. When there are non-integer case weights and (time1, time2) survival data the routine is at an impasse: a robust variance likely is called for, but requires either id or cluster information to be done correctly; it will default to robust=FALSE.

With the IJ estimate, the leverage values themselves can be returned as arrays with dimensions: number of subjects, number of unique times, and for a multi-state model, the number of unique states. Be forwarned that these arrays can be huge. If there is a cluster argument this first dimension will be the number of clusters and the variance will be a grouped IJ estimate; this can be an important tool for reducing the size. A numeric value for the influence argument allows finer control: 0= return neither (same as FALSE), 1= return the influence array for probability in state, 2= return the influence array for the cumulative hazard, 3= both (same as TRUE).

#### Value

an object of class "survfit". See survfit.object for details. Methods defined for survfit objects are print, plot, lines, and points.

## References

Dorey, F. J. and Korn, E. L. (1987). Effective sample sizes for confidence intervals for survival probabilities. *Statistics in Medicine* **6**, 679-87.

Fleming, T. H. and Harrington, D. P. (1984). Nonparametric estimation of the survival distribution in censored data. *Comm. in Statistics* **13**, 2469-86.

Kalbfleisch, J. D. and Prentice, R. L. (1980). *The Statistical Analysis of Failure Time Data*. New York: Wiley.

Kyle, R. A. (1997). Moncolonal gammopathy of undetermined significance and solitary plasmacytoma. Implications for progression to overt multiple myeloma}, *Hematology/Oncology Clinics N. Amer.* **11**, 71-87.

Link, C. L. (1984). Confidence intervals for the survival function using Cox's proportional hazards model with covariates. *Biometrics* **40**, 601-610.

Turnbull, B. W. (1974). Nonparametric estimation of a survivorship function with doubly censored data. *J Am Stat Assoc*, **69**, 169-173.

#### See Also

survfit.coxph for survival curves from Cox models, survfit.object for a description of the components of a survfit object, print.survfit, plot.survfit, lines.survfit, coxph, Surv.

## **Examples**

```
#fit a Kaplan-Meier and plot it
fit <- survfit(Surv(time, status) ~ x, data = aml)</pre>
plot(fit, lty = 2:3)
legend(100, .8, c("Maintained", "Nonmaintained"), lty = 2:3)
#fit a Cox proportional hazards model and plot the
#predicted survival for a 60 year old
fit <- coxph(Surv(futime, fustat) ~ age, data = ovarian)</pre>
plot(survfit(fit, newdata=data.frame(age=60)),
     xscale=365.25, xlab = "Years", ylab="Survival")
# Here is the data set from Turnbull
# There are no interval censored subjects, only left-censored (status=3),
  right-censored (status 0) and observed events (status 1)
                              Time
                               2
                                  3 4
# Type of observation
           death
                         12
                               6
                                  2
                                       3
#
           losses
                          3
                               2
                                   0
#
      late entry
                          2
                                   2
tdata <- data.frame(time =c(1,1,1,2,2,2,3,3,3,4,4,4),
                    status=rep(c(1,0,2),4),
                    n
                          =c(12,3,2,6,2,4,2,0,2,3,3,5))
fit <- survfit(Surv(time, time, status, type='interval') ~1,
              data=tdata, weight=n)
# Three curves for patients with monoclonal gammopathy.
# 1. KM of time to PCM, ignoring death (statistically incorrect)
# 2. Competing risk curves (also known as "cumulative incidence")
  3. Multi-state, showing Pr(in each state, at time t)
fitKM <- survfit(Surv(stop, event=='pcm') ~1, data=mgus1,</pre>
                    subset=(start==0))
fitCR <- survfit(Surv(stop, event) ~1,</pre>
                    data=mgus1, subset=(start==0))
fitMS <- survfit(Surv(start, stop, event) ~ 1, id=id, data=mgus1)</pre>
## Not run:
# CR curves show the competing risks
plot(fitCR, xscale=365.25, xmax=7300, mark.time=FALSE,
            col=2:3, xlab="Years post diagnosis of MGUS",
            ylab="P(state)")
lines(fitKM, fun='event', xmax=7300, mark.time=FALSE,
            conf.int=FALSE)
text(3652, .4, "Competing risk: death", col=3)
text(5840, .15,"Competing risk: progression", col=2)
text(5480, .30, "KM:prog")
## End(Not run)
```

survfit.matrix 151

survfit.matrix Create Aalen-Johansen estimates of multi-state survival from a matrix of hazards.
--

# Description

This allows one to create the Aalen-Johansen estimate of P, a matrix with one column per state and one row per time, starting with the individual hazard estimates. Each row of P will sum to 1. Note that this routine has been superseded by the use of multi-state Cox models, and will eventually be removed.

## Usage

## **Arguments**

formula	a matrix of lists, each element of which is either NULL or a survival curve object.
p0	the initial state vector. The names of this vector are used as the names of the states in the output object. If there are multiple curves then p0 can be a matrix with one row per curve.
method	use a product of discrete hazards, or a product of matrix exponentials. See details below.
start.time	optional; start the calculations at a given starting point
	further arguments used by other survfit methods

## Details

On input the matrix should contain a set of predicted curves for each possible transition, and NULL in other positions. Each of the predictions will have been obtained from the relevant Cox model. This approach for multistate curves is easy to use but has some caveats. First, the input curves must be consistent. The routine checks as best it can, but can easy be fooled. For instance, if one were to fit two Cox models, obtain predictions for males and females from one, and for treatment A and B from the other, this routine will create two curves but they are not meaningful. A second issue is that standard errors are not produced.

The names of the resulting states are taken from the names of the vector of initial state probabilities. If they are missing, then the dimnames of the input matrix are used, and lacking that the labels '1', '2', etc. are used.

For the usual Aalen-Johansen estimator the multiplier at each event time is the matrix of hazards H (also written as I+dA). When using predicted survival curves from a Cox model, however, it is possible to get predicted hazards that are greater than 1, which leads to probabilities less than 0. If the method argument is not supplied and the input curves are derived from a Cox model this routine instead uses the approximation expm(H-I) as the multiplier, which always gives valid probabilities. (This is also the standard approach for ordinary survival curves from a Cox model.)

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#### Value

a survfitms object

#### Note

The R syntax for creating a matrix of lists is very fussy.

#### Author(s)

Terry Therneau

#### See Also

survfit

## **Examples**

```
etime <- with(mgus2, ifelse(pstat==0, futime, ptime))
event <- with(mgus2, ifelse(pstat==0, 2*death, 1))
event <- factor(event, 0:2, labels=c("censor", "pcm", "death"))

cfit1 <- coxph(Surv(etime, event=="pcm") ~ age + sex, mgus2)
cfit2 <- coxph(Surv(etime, event=="death") ~ age + sex, mgus2)

# predicted competing risk curves for a 72 year old with mspike of 1.2

# (median values), male and female.

# The survfit call is a bit faster without standard errors.
newdata <- expand.grid(sex=c("F", "M"), age=72, mspike=1.2)

AJmat <- matrix(list(), 3,3)

AJmat[1,2] <- list(survfit(cfit1, newdata, std.err=FALSE))
AJmat[1,3] <- list(survfit(cfit2, newdata, std.err=FALSE))
csurv <- survfit(AJmat, p0 =c(entry=1, PCM=0, death=0))</pre>
```

survfit.object

Survival Curve Object

## **Description**

This class of objects is returned by the survfit class of functions to represent a fitted survival curve. For a multi-state model the object has class c('survfitms', 'survfit').

Objects of this class have methods for the functions print, summary, plot, points and lines. The print.survfit method does more computation than is typical for a print method and is documented on a separate page.

survfit.object 153

#### **Arguments**

n total number of subjects in each curve.

time the time points at which the curve has a step.

n.risk the number of subjects at risk at t.

n. event the number of events that occur at time t.

n.enter for counting process data only, the number of subjects that enter at time t.

n. censor for counting process data only, the number of subjects who exit the risk set, with-

out an event, at time t. (For right censored data, this number can be computed

from the successive values of the number at risk).

surv the estimate of survival at time t+0. This may be a vector or a matrix. The

latter occurs when a set of survival curves is created from a single Cox model,

in which case there is one column for each covariate set.

pstate a multi-state survival will have the pstate component instead of surv. It will

be a matrix containing the estimated probability of each state at each time, one

column per state.

std.err for a survival curve this contains standard error of the cumulative hazard or -

log(survival), for a multi-state curve it contains the standard error of prev. This difference is a reflection of the fact that each is the natural calculation for that

case.

cumhaz hazard optional. Contains the cumulative hazard for each possible transtion.

strata if there are multiple curves, this component gives the number of elements of the

time vector corresponding to the first curve, the second curve, and so on. The

names of the elements are labels for the curves.

upper optional upper confidence limit for the survival curve or pstate

lower options lower confidence limit for the survival curve or pstate

start.time optional, the starting time for the curve if other than 0

p0, sp0 for a multistate object, the distribution of starting states. If the curve has a strata

dimension, this will be a matrix one row per stratum. The sp0 element has the

standard error of p0, if p0 was estimated.

newdata for survival curves from a fitted model, this contains the covariate values for the

curves

n.all the total number of observations that were available For counting process data,

and any time that the start.time argument was used, not all may have been used in creating the curve, in which case this value will be larger than n above. The print and plot routines in the package do no use this value, it is for infor-

mation only.

conf. type the approximation used to compute the confidence limits.

conf. int the level of the confidence limits, e.g. 90 or 95%.

transitions for multi-state data, the total number of transitions of each type.

na.action the returned value from the na.action function, if any. It will be used in the

printout of the curve, e.g., the number of observations deleted due to missing

values.

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call an image of the call that produced the object.

type type of survival censoring.

influence.p, influence.c

optional influence matrices for the pstate (or surv) and for the cumhaz estimates. A list with one element per stratum, each element of the list is an array

indexed by subject, time, state.

version the version of the object. Will be missing, 2, or 3

#### Structure

The following components must be included in a legitimate survfit or survfitms object.

## **Subscripts**

Survfit objects can be subscripted. This is often used to plot a subset of the curves, for instance. From the user's point of view the survfit object appears to be a vector, matrix, or array of curves. The first dimension is always the underlying number of curves or "strata"; for multi-state models the state is always the last dimension. Predicted curves from a Cox model can have a second dimension which is the number of different covariate prediction vectors.

#### Details

The survfit object has evolved over time: when first created there was no thought of multi-state models for instance. This evolution has almost entirely been accomplished by the addition of new elements. One change in survival version 3 is the addition of a survfitconf routine which will compute confidence intervals for a survfit object. This allows the computation of CI intervals to be deferred until later, if desired, rather than making them a permanent part of the object. Later iterations of the base routines may omit the confidence intervals.

The survfit object starts at the first observation time, but survival curves are normally plotted from time 0. A helper routine survfit0 can be used to add this first time point and align the data.

## See Also

```
plot.survfit, summary.survfit, print.survfit, survfit, survfit0
```

survfit0

Convert the format of a survfit object.

## Description

Add the point for a starting time (time 0) to a survfit object's elements. This is useful for plotting.

# Usage

```
survfit0(x, start.time=0)
```

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## **Arguments**

x a survfit object

start.time the desired starting time; see details below.

#### **Details**

Survival curves are traditionally plotted forward from time 0, but since the true starting time is not known as a part of the data, the survfit routine does not include a time 0 value in the resulting object. Someone might look at cumulative mortgage defaults versus calendar year, for instance, with the 'time' value a Date object. The plotted curve probably should not start at 0 = 1970/01/01. Due to this uncertainty, it was decided not to include a "time 0" as part of a survfit object. If the original survfit call included a start.time argument, that value is of course retained.

Whether that (1989) decision was wise or foolish, it is now far too late to change it. (We tried it once as a trial, resulting in over 20 errors in the survival test suite. We extrapolate that it might break 1/2 - 2/3 of the other CRAN packages that depend on survival, if made a default.) If the original survfit call included a start.time argument, that value is of course retained.

One problem with this choice is that some functions must choose a starting point, plots for example. This utility function is used by plot.survfit and summary.survfit to do so, adding a new time point at the front of each curve in a consistent way: the optional argument to the survfit0 function as the first choice (if supplied), then the user's start.time if present, otherwise min(0,x\$time). The resulting object is *not* guarranteed to work with functions that further manipulate a survfit object such as subscripting, aggregation, pseudovalues, etc. (remember the 20 errors). Rather it is intended as a penultimate step, most often when creating a plot.

## Value

a reformulated version of the object with an initial data point at start.time added. The time, surv, pstate, cumhaz, std.err, std.cumhaz and other components will all be aligned, so as to make plots and summaries easier to produce.

survfitcoxph.fit

A direct interface to the 'computational engine' of survfit.coxph

## **Description**

This program is mainly supplied to allow other packages to invoke the survfit.coxph function at a 'data' level rather than a 'user' level. It does no checks on the input data that is provided, which can lead to unexpected errors if that data is wrong.

## Usage

```
survfitcoxph.fit(y, x, wt, x2, risk, newrisk, strata, se.fit, survtype,
vartype, varmat, id, y2, strata2, unlist=TRUE)
```

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## **Arguments**

У	the response variable used in the Cox model. (Missing values removed of course.)
X	covariate matrix used in the Cox model
wt	weight vector for the Cox model. If the model was unweighted use a vector of 1s.
x2	matrix describing the hypothetical subjects for which a curve is desired. Must have the same number of columns as x.
risk	the risk score exp(X beta) from the fitted Cox model. If the model had an offset, include it in the argument to exp.
newrisk	risk scores for the hypothetical subjects
strata	strata variable used in the Cox model. This will be a factor.
se.fit	if TRUE the standard errors of the curve(s) are returned
survtype	1=Kalbfleisch-Prentice, 2=Nelson-Aalen, 3=Efron. It is usual to match this to the approximation for ties used in the coxph model: KP for 'exact', N-A for 'breslow' and Efron for 'efron'.
vartype	1=Greenwood, 2=Aalen, 3=Efron
varmat	the variance matrix of the coefficients
id	optional; if present and not NULL this should be a vector of identifiers of length nrow(x2). A mon-null value signifies that x2 contains time dependent covariates, in which case this identifies which rows of x2 go with each subject.
y2	survival times, for time dependent prediction. It gives the time range (time1,time2] for each row of x2. Note: this must be a Surv object and thus contains a status indicator, which is never used in the routine, however.
strata2	vector of strata indicators for x2. This must be a factor.
unlist	if FALSE the result will be a list with one element for each strata. Otherwise the strata are "unpacked" into the form found in a survfit object.

# Value

a list containing nearly all the components of a survfit object. All that is missing is to add the confidence intervals, the type of the original model's response (as in a coxph object), and the class.

## Note

The source code for for both this function and survfit.coxph is written using noweb. For complete documentation see the inst/sourcecode.pdf file.

## Author(s)

Terry Therneau

## See Also

survfit.coxph

survival-deprecated 157

survival-deprecated

Deprecated functions in package survival

## **Description**

These functions are temporarily retained for compatability with older programs, and may transition to defunct status.

# Usage

```
survConcordance(formula, data, weights, subset, na.action) # use concordance
survConcordance.fit(y, x, strata, weight)
                                             # use concordancefit
```

## **Arguments**

formula

a formula object, with the response on the left of a ~ operator, and the terms

on the right. The response must be a survival object as returned by the Surv

function.

data

a data frame weights, subset, na.action

as for coxph

x, y, strata, weight

predictor, response, strata, and weight vectors for the direct call

## See Also

Deprecated

survobrien

O'Brien's Test for Association of a Single Variable with Survival

# Description

Peter O'Brien's test for association of a single variable with survival This test is proposed in Biometrics, June 1978.

## Usage

```
survobrien(formula, data, subset, na.action, transform)
```

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## **Arguments**

formula a valid formula for a cox model.

data a data frame in which to interpret the variables named in the formula, or in the

subset and the weights argument.

subset expression indicating which subset of the rows of data should be used in the fit.

All observations are included by default.

na.action a missing-data filter function. This is applied to the model.frame after any subset

argument has been used. Default is options()\$na.action.

transform the transformation function to be applied at each time point. The default is

O'Brien's suggestion logit(tr) where tr = (rank(x)-1/2)/length(x) is the rank

shifted to the range 0-1 and logit(x) = log(x/(1-x)) is the logit transform.

#### Value

a new data frame. The response variables will be column names returned by the Surv function, i.e., "time" and "status" for simple survival data, or "start", "stop", "status" for counting process data. Each individual event time is identified by the value of the variable .strata.. Other variables retain their original names. If a predictor variable is a factor or is protected with I(), it is retained as is. Other predictor variables have been replaced with time-dependent logit scores.

The new data frame will have many more rows that the original data, approximately the original number of rows \* number of deaths/2.

#### Method

A time-dependent cox model can now be fit to the new data. The univariate statistic, as originally proposed, is equivalent to single variable score tests from the time-dependent model. This equivalence is the rationale for using the time dependent model as a multivariate extension of the original paper.

In O'Brien's method, the x variables are re-ranked at each death time. A simpler method, proposed by Prentice, ranks the data only once at the start. The results are usually similar.

#### Note

A prior version of the routine returned new time variables rather than a strata. Unfortunately, that strategy does not work if the original formula has a strata statement. This new data set will be the same size, but the coxph routine will process it slightly faster.

#### References

O'Brien, Peter, "A Nonparametric Test for Association with Censored Data", *Biometrics* 34: 243-250, 1978.

#### See Also

survdiff

survreg 159

## **Examples**

survreg

Regression for a Parametric Survival Model

# Description

Fit a parametric survival regression model. These are location-scale models for an arbitrary transform of the time variable; the most common cases use a log transformation, leading to accelerated failure time models.

# Usage

## **Arguments**

formula	a formula expression as for other regression models. The response is usually a survival object as returned by the Surv function. See the documentation for Surv, lm and formula for details.
data	a data frame in which to interpret the variables named in the formula, weights or the subset arguments.
weights	optional vector of case weights
subset	subset of the observations to be used in the fit
na.action	a missing-data filter function, applied to the model.frame, after any subset argument has been used. Default is options()\$na.action.
dist	assumed distribution for y variable. If the argument is a character string, then it is assumed to name an element from survreg.distributions. These include "weibull", "exponential", "gaussian", "logistic", "lognormal" and "loglogistic". Otherwise, it is assumed to be a user defined list conforming to the format described in survreg.distributions.
parms	a list of fixed parameters. For the t-distribution for instance this is the degrees of freedom; most of the distributions have no parameters.
init	optional vector of initial values for the parameters.
scale	optional fixed value for the scale. If set to <=0 then the scale is estimated.
control	a list of control values, in the format produced by survreg.control. The default value is survreg.control()
scale	optional fixed value for the scale. If set to <=0 then the scale is estimated.  a list of control values, in the format produced by survreg.control. The de-

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model,x,y	flags to control what is returned. If any of these is true, then the model frame, the model matrix, and/or the vector of response times will be returned as components of the final result, with the same names as the flag arguments.
score	return the score vector. (This is expected to be zero upon successful convergence.)
robust	Use robust sandwich error instead of the asymptotic formula. Defaults to TRUE if there is a cluster argument.
cluster	Optional variable that identifies groups of subjects, used in computing the robust variance. Like model variables, this is searched for in the dataset pointed to by the data argument.
	other arguments which will be passed to survreg.control.

#### **Details**

All the distributions are cast into a location-scale framework, based on chapter 2.2 of Kalbfleisch and Prentice. The resulting parameterization of the distributions is sometimes (e.g. gaussian) identical to the usual form found in statistics textbooks, but other times (e.g. Weibull) it is not. See the book for detailed formulas.

#### Value

an object of class survreg is returned.

#### References

Kalbfleisch, J. D. and Prentice, R. L., The statistical analysis of failure time data, Wiley, 2002.

## See Also

```
survreg.object, survreg.distributions, pspline, frailty, ridge
```

# Examples

survreg.control 161

survreg.control

Package options for survreg and coxph

## **Description**

This functions checks and packages the fitting options for survreg

## Usage

```
survreg.control(maxiter=30, rel.tolerance=1e-09,
toler.chol=1e-10, iter.max, debug=0, outer.max=10)
```

# Arguments

maxiter maximum number of iterations

rel.tolerance relative tolerance to declare convergence

toler.chol Tolerance to declare Cholesky decomposition singular

iter.max same as maxiter

debug print debugging information

outer.max maximum number of outer iterations for choosing penalty parameters

## Value

A list with the same elements as the input

## See Also

survreg

162 survreg.distributions

survreg.distributions Parametric Survival Distributions

#### **Description**

List of distributions for accelerated failure models. These are location-scale families for some transformation of time. The entry describes the cdf F and density f of a canonical member of the family.

## Usage

survreg.distributions

## **Format**

There are two basic formats, the first defines a distribution de novo, the second defines a new distribution in terms of an old one.

name: name of distribution

variance: function(parms) returning the variance (currently unused)

init(x,weights,...): Function returning an initial

estimate of the mean and variance (used for initial values in the iteration)

density(x,parms): Function returning a matrix with columns F, 1 - F, f, f'/f, and f''/f

quantile(p,parms): Quantile function

scale: Optional fixed value for the scale parameter

parms: Vector of default values and names for any additional parameters

deviance(y,scale,parms): Function returning the deviance for a

saturated model; used only for deviance residuals.

and to define one distribution in terms of another

name: name of distribution
dist: name of parent distribution
trans: transformation (eg log)
dtrans: derivative of transformation
itrans: inverse of transformation

scale: Optional fixed value for scale parameter

## **Details**

There are four basic distributions:extreme, gaussian, logistic and t. The last three are parametrised in the same way as the distributions already present in R. The extreme value cdf is

$$F = 1 - e^{-e^t}$$
.

survreg.distributions 163

When the logarithm of survival time has one of the first three distributions we obtain respectively weibull, lognormal, and loglogistic. The location-scale parameterization of a Weibull distribution found in surveg is not the same as the parameterization of rweibull.

The other predefined distributions are defined in terms of these. The exponential and rayleigh distributions are Weibull distributions with fixed scale of 1 and 0.5 respectively, and loggaussian is a synonym for lognormal.

For speed parts of the three most commonly used distributions are hardcoded in C; for this reason the elements of survreg.distributions with names of "Extreme value", "Logistic" and "Gaussian" should not be modified. (The order of these in the list is not important, recognition is by name.) As an alternative to modifying survreg.distributions a new distribution can be specified as a separate list. This is the preferred method of addition and is illustrated below.

#### See Also

```
survreg, pweibull, pnorm,plogis, pt, survregDtest
```

## **Examples**

```
# time transformation
survreg(Surv(time, status) ~ ph.ecog + sex, dist='weibull', data=lung)
# change the transformation to work in years
# intercept changes by log(365), everything else stays the same
my.weibull <- survreg.distributions$weibull</pre>
my.weibulltrans < function(y) log(y/365)
my.weibull$itrans <- function(y) 365*exp(y)</pre>
survreg(Surv(time, status) ~ ph.ecog + sex, lung, dist=my.weibull)
# Weibull parametrisation
y<-rweibull(1000, shape=2, scale=5)
survreg(Surv(y)~1, dist="weibull")
# survreg scale parameter maps to 1/shape, linear predictor to log(scale)
# Cauchy fit
mycauchy <- list(name='Cauchy',</pre>
                 init= function(x, weights, ...)
                      c(median(x), mad(x)),
                 density= function(x, parms) {
                      temp <-1/(1 + x^2)
                      cbind(.5 + atan(x)/pi, .5 + atan(-x)/pi,
                             temp/pi, -2 *x*temp, 2*temp*(4*x^2*temp -1))
                 quantile= function(p, parms) tan((p-.5)*pi),
                 deviance= function(...) stop('deviance residuals not defined')
survreg(Surv(log(time), status) ~ ph.ecog + sex, lung, dist=mycauchy)
```

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survreg.object

Parametric Survival Model Object

## **Description**

This class of objects is returned by the survreg function to represent a fitted parametric survival model. Objects of this class have methods for the functions print, summary, predict, and residuals.

#### COMPONENTS

The following components must be included in a legitimate survreg object.

coefficients the coefficients of the linear.predictors, which multiply the columns of the model matrix. It does not include the estimate of error (sigma). The names of the coefficients are the names of the single-degree-of-freedom effects (the columns of the model matrix). If the model is over-determined there will be missing values in the coefficients corresponding to non-estimable coefficients.

**icoef** coefficients of the baseline model, which will contain the intercept and log(scale), or multiple scale factors for a stratified model.

var the variance-covariance matrix for the parameters, including the log(scale) parameter(s).

**loglik** a vector of length 2, containing the log-likelihood for the baseline and full models.

iter the number of iterations required

**linear.predictors** the linear predictor for each subject.

**df** the degrees of freedom for the final model. For a penalized model this will be a vector with one element per term.

scale the scale factor(s), with length equal to the number of strata.

idf degrees of freedom for the initial model.

means a vector of the column means of the coefficient matrix.

**dist** the distribution used in the fit.

weights included for a weighted fit.

The object will also have the following components found in other model results (some are optional): linear predictors, weights, x, y, model, call, terms and formula. See lm.

## See Also

survreg, 1m

survregDtest 165

## **Description**

This routine is called by survreg to verify that a distribution object is valid.

## Usage

```
survregDtest(dlist, verbose = F)
```

## **Arguments**

dlist the list describing a survival distribution

verbose return a simple TRUE/FALSE from the test for validity (the default), or a ver-

bose description of any flaws.

## **Details**

If the survreg function rejects your user-supplied distribution as invalid, this routine will tell you why it did so.

## Value

TRUE if the distribution object passes the tests, and either FALSE or a vector of character strings if not.

#### Author(s)

Terry Therneau

#### See Also

```
survreg.distributions, survreg
```

## **Examples**

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```
)
survregDtest(mycauchy, TRUE)
```

survSplit

Split a survival data set at specified times

## **Description**

Given a survival data set and a set of specified cut times, split each record into multiple subrecords at each cut time. The new data set will be in 'counting process' format, with a start time, stop time, and event status for each record.

## Usage

```
survSplit(formula, data, subset, na.action=na.pass,
            cut, start="tstart", id, zero=0, episode,
                              end="tstop", event="event")
```

#### **Arguments**

formula a model formula data a data frame subset, na.action

rows of the data to be retained

the vector of timepoints to cut at cut

character string with the name of a start time variable (will be created if needed) start id

character string with the name of new id variable to create (optional). This can

be useful if the data set does not already contain an identifier.

If start doesn't already exist, this is the time that the original records start. zero

character string with the name of new episode variable (optional) episode

character string with the name of event time variable end character string with the name of censoring indicator event

## **Details**

Each interval in the original data is cut at the given points; if an original row were (15, 60] with a cut vector of (10,30, 40) the resulting data set would have intervals of (15,30], (30,40] and (40, 60].

Each row in the final data set will lie completely within one of the cut intervals. Which interval for each row of the output is shown by the episode variable, where 1= less than the first cutpoint, 2= between the first and the second, etc. For the example above the values would be 2, 3, and 4.

The routine is called with a formula as the first argument. The right hand side of the formula can be used to delimit variables that should be retained; normally one will use ~ . as a shorthand to retain them all. The routine will try to retain variable names, e.g. Surv(adam, joe, fred)~. will tcut 167

result in a data set with those same variable names for tstart, end, and event options rather than the defaults. Any user specified values for these options will be used if they are present, of course. However, the routine is not sophisticated; it only does this substitution for simple names. A call of Surv(time, stat==2) for instance will not retain "stat" as the name of the event variable.

Rows of data with a missing time or status are copied across unchanged, unless the na.action argument is changed from its default value of na.pass. But in the latter case any row that is missing for any variable will be removed, which is rarely what is desired.

#### Value

New, longer, data frame.

#### See Also

Surv, cut, reshape

## **Examples**

tcut

Factors for person-year calculations

## **Description**

Attaches categories for person-year calculations to a variable without losing the underlying continuous representation

## Usage

```
tcut(x, breaks, labels, scale=1)
## S3 method for class 'tcut'
levels(x)
```

168 tmerge

## Arguments

x numeric/date variable
 breaks breaks between categories, which are right-continuous
 labels labels for categories
 scale Multiply x and breaks by this.

## Value

An object of class tcut

## See Also

```
cut, pyears
```

## **Examples**

```
mdy.date <- function(m,d,y)
    as.Date(paste(ifelse(y<100, y+1900, y), m, d, sep='/'))
temp1 <- mdy.date(6,6,36)
temp2 <- mdy.date(6,6,55)# Now compare the results from person-years
#
temp.age <- tcut(temp2-temp1, floor(c(-1, (18:31 * 365.24))),
labels=c('0-18', paste(18:30, 19:31, sep='-')))
temp.yr <- tcut(temp2, mdy.date(1,1,1954:1965), labels=1954:1964)
temp.time <- 3700  #total days of fu
py1 <- pyears(temp.time ~ temp.age + temp.yr, scale=1) #output in days
py1</pre>
```

tmerge

Time based merge for survival data

# Description

A common task in survival analysis is the creation of start, stop data sets which have multiple intervals for each subject, along with the covariate values that apply over that interval. This function aids in the creation of such data sets.

## Usage

```
tmerge(data1, data2, id,..., tstart, tstop, options)
```

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## **Arguments**

data1	the primary data set, to which new variables and/or observation will be added
data2	second data set in which all the other arguments will be found
id	subject identifier
	operations that add new variables or intervals, see below
tstart	optional variable to define the valid time range for each subject, only used on an initial call $% \left( 1\right) =\left( 1\right) \left( 1\right)$
tstop	optional variable to define the valid time range for each subject, only used on an initial call
options	a list of options. Valid ones are idname, tstartname, tstopname, delay, na.rm, and tdcstart. See the explanation below.

## **Details**

The program is often run in multiple passes, the first of which defines the basic structure, and subsequent ones that add new variables to that structure. For a more complete explanation of how this routine works refer to the vignette on time-dependent variables.

There are 4 types of operational arguments: a time dependent covariate (tdc), cumulative count (cumtdc), event (event) or cumulative event (cumevent). Time dependent covariates change their values before an event, events are outcomes.

• newname = tdc(y, x, init) A new time dependent covariate variable will created. The argument y is assumed to be on the scale of the start and end time, and each instance describes the occurrence of a "condition" at that time. The second argument x is optional. In the case where x is missing the count variable starts at 0 for each subject and becomes 1 at the time of the event. If x is present the value of the time dependent covariate is initialized to value of init, if present, or the tdcstart option otherwise, and is updated to the value of x at each observation. If the option na.rm=TRUE missing values of x are first removed, i.e., the update will not create missing values.

newname =  $\operatorname{cumtdc}(y,x,\operatorname{init})$  Similar to tdc, except that the event count is accumulated over time for each subject. The variable x must be numeric.

newname = event(y,x) Mark an event at time y. In the usual case that x is missing the new 0/1 variable will be similar to the 0/1 status variable of a survival time.

newname = cumevent(y,x) Cumulative events.

The function adds three new variables to the output data set: tstart, tstop, and id. The options argument can be used to change these names. If, in the first call, the id argument is a simple name, that variable name will be used as the default for the idname option. If data1 contains the tstart variable then that is used as the starting point for the created time intervals, otherwise the initial interval for each id will begin at 0 by default. This will lead to an invalid interval and subsequent error if say a death time were <= 0.

The na.rm option affects creation of time-dependent covariates. Should a data row in data2 that has a missing value for the variable be ignored (na.rm=FALSE, default) or should it generate an observation with a value of NA? The default value leads to "last value carried forward" behavior. The delay option causes a time-dependent covariate's new value to be delayed, see the vignette for an example.

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#### Value

a data frame with two extra attributes tname and tcount. The first contains the names of the key variables; it's persistence from call to call allows the user to avoid constantly reentering the options argument. The tcount variable contains counts of the match types. New time values that occur before the first interval for a subject are "early", those after the last interval for a subject are "late", and those that fall into a gap are of type "gap". All these are are considered to be outside the specified time frame for the given subject. An event of this type will be discarded. An observation in data2 whose identifier matches no rows in data1 is of type "missid" and is also discarded. A time-dependent covariate value will be applied to later intervals but will not generate a new time point in the output.

The most common type will usually be "within", corresponding to those new times that fall inside an existing interval and cause it to be split into two. Observations that fall exactly on the edge of an interval but within the (min, max] time for a subject are counted as being on a "leading" edge, "trailing" edge or "boundary". The first corresponds for instance to an occurrence at 17 for someone with an intervals of (0,15] and (17, 35]. A tdc at time 17 will affect this interval but an event at 17 would be ignored. An event occurrence at 15 would count in the (0,15] interval. The last case is where the main data set has touching intervals for a subject, e.g. (17, 28] and (28,35] and a new occurrence lands at the join. Events will go to the earlier interval and counts to the latter one. A last column shows the number of additions where the id and time point were identical. When this occurs, the tdc and event operators will use the final value in the data (last edit wins), but ignoring missing, while cumtdc and cumevent operators add up the values.

These extra attributes are ephemeral and will be discarded if the dataframe is modified. This is intentional, since they will become invalid if for instance a subset were selected.

#### Author(s)

Terry Therneau

#### See Also

neardate

## **Examples**

tobin 171

tobin

Tobin's Tobit data

# Description

Economists fit a parametric censored data model called the 'tobit'. These data are from Tobin's original paper.

## Usage

```
tobin
data(tobin, package="survival")
```

## **Format**

A data frame with 20 observations on the following 3 variables.

```
durable Durable goods purchaseage Age in yearsquant Liquidity ratio (x 1000)
```

## Source

J Tobin (1958), Estimation of relationships for limited dependent variables. *Econometrica* **26**, 24–36.

## **Examples**

transplant

Liver transplant waiting list

# Description

Subjects on a liver transplant waiting list from 1990-1999, and their disposition: received a transplant, died while waiting, withdrew from the list, or censored.

#### Usage

```
transplant
data(transplant, package="survival")
```

172 transplant

#### **Format**

A data frame with 815 (transplant) observations on the following 6 variables.

```
age age at addition to the waiting list
sex m or f
abo blood type: A, B, AB or 0
year year in which they entered the waiting list
futime time from entry to final disposition
event final disposition: censored, death, ltx or withdraw
```

#### **Details**

This represents the transplant experience in a particular region, over a time period in which liver transplant became much more widely recognized as a viable treatment modality. The number of liver transplants rises over the period, but the number of subjects added to the liver transplant waiting list grew much faster. Important questions addressed by the data are the change in waiting time, who waits, and whether there was an consequent increase in deaths while on the list.

Blood type is an important consideration. Donor livers from subjects with blood type O can be used by patients with A, B, AB or 0 blood types, whereas an AB liver can only be used by an AB recipient. Thus type O subjects on the waiting list are at a disadvantage, since the pool of competitors is larger for type O donor livers.

This data is of historical interest and provides a useful example of competing risks, but it has little relevance to current practice. Liver allocation policies have evolved and now depend directly on each individual patient's risk and need, assessments of which are regularly updated while a patient is on the waiting list. The overall organ shortage remains acute, however.

The transplant data set was a version used early in the analysis, transplant2 has several additions and corrections, and was the final data set and matches the paper.

## References

Kim WR, Therneau TM, Benson JT, Kremers WK, Rosen CB, Gores GJ, Dickson ER. Deaths on the liver transplant waiting list: An analysis of competing risks. Hepatology 2006 Feb; 43(2):345-51.

#### **Examples**

udca 173

udca

Data from a trial of usrodeoxycholic acid

## **Description**

Data from a trial of ursodeoxycholic acid (UDCA) in patients with primary biliary cirrohosis (PBC).

## Usage

```
udca
udca2
data(udca, package="survival")
```

#### **Format**

A data frame with 170 observations on the following 15 variables.

```
id subject identifier

trt treatment of 0=placebo, 1=UDCA

entry.dt date of entry into the study

last.dt date of last on-study visit

stage stage of disease

bili bilirubin value at entry

riskscore the Mayo PBC risk score at entry

death.dt date of death

tx.dt date of liver transplant

hprogress.dt date of histologic progression

varices.dt appearance of esphogeal varices

ascites.dt appearance of ascites

enceph.dt appearance of encephalopathy

double.dt doubling of initial bilirubin

worsen.dt worsening of symptoms by two stages
```

## **Details**

This data set is used in the Therneau and Grambsh. The udcal data set contains the baseline variables along with the time until the first endpoint (any of death, transplant, ..., worsening). The udcal data set treats all of the endpoints as parallel events and has a stratum for each.

174 untangle.specials

#### References

T. M. Therneau and P. M. Grambsch, Modeling survival data: extending the Cox model. Springer, 2000.

K. D. Lindor, E. R. Dickson, W. P Baldus, R.A. Jorgensen, J. Ludwig, P. A. Murtaugh, J. M. Harrison, R. H. Weisner, M. L. Anderson, S. M. Lange, G. LeSage, S. S. Rossi and A. F. Hofman. Ursodeoxycholic acid in the treatment of primary biliary cirrhosis. Gastroenterology, 106:1284-1290, 1994.

## **Examples**

untangle.specials

Help Process the 'specials' Argument of the 'terms' Function.

## Description

Given a terms structure and a desired special name, this returns an index appropriate for subscripting the terms structure and another appropriate for the data frame.

## Usage

```
untangle.specials(tt, special, order=1)
```

## **Arguments**

tt a terms object.

special the name of a special function, presumably used in the terms object.

order the order of the desired terms. If set to 2, interactions with the special function

will be included.

## Value

a list with two components:

vars a vector of variable names, as would be found in the data frame, of the specials.

terms a numeric vector, suitable for subscripting the terms structure, that indexes the

terms in the expanded model formula which involve the special.

uspop2 175

## **Examples**

```
formula<-Surv(tt,ss)~x+z*strata(id)
tms<-terms(formula, specials="strata")
## the specials attribute
attr(tms,"specials")
## main effects
untangle.specials(tms,"strata")
## and interactions
untangle.specials(tms,"strata",order=1:2)</pre>
```

uspop2

Projected US Population

## **Description**

US population by age and sex, for 2000 through 2020

#### **Format**

The data is a matrix with dimensions age, sex, and calendar year. Age goes from 0 through 100, where the value for age 100 is the total for all ages of 100 or greater.

## **Details**

This data is often used as a "standardized" population for epidemiology studies.

# Source

NP2008\_D1: Projected Population by Single Year of Age, Sex, Race, and Hispanic Origin for the United States: July 1, 2000 to July 1, 2050, www.census.gov/population/projections.

## See Also

uspop

## **Examples**

176 veteran

Variance-covariance matrix	nce matrix	Variance-co	cov.coxph
----------------------------	------------	-------------	-----------

## **Description**

Extract and return the variance-covariance matrix.

## Usage

```
## S3 method for class 'coxph'
vcov(object, complete=TRUE, ...)
## S3 method for class 'survreg'
vcov(object, complete=TRUE, ...)
```

## Arguments

object a fitted model object

complete logical indicating if the full variance-covariance matrix should be returned. This

has an effect only for an over-determined fit where some of the coefficients are undefined, and coef(object) contains corresponding NA values. If complete=TRUE the returned matrix will have row/column for each coefficient, if FALSE it will contain rows/columns corresponding to the non-missing coefficients. The coef()

function has a simpilar complete argument.

. . . additional arguments for method functions

## **Details**

For the coxph and survreg functions the returned matrix is a particular generalized inverse: the row and column corresponding to any NA coefficients will be zero. This is a side effect of the generalized cholesky decomposion used in the unerlying computation.

## Value

a matrix

veteran Veterans' Administration Lung Cancer study

# **Description**

Randomised trial of two treatment regimens for lung cancer. This is a standard survival analysis data set.

xtfrm.Surv 177

## Usage

```
veteran
data(cancer, package="survival")
```

#### **Format**

trt: 1=standard 2=test

celltype: 1=squamous, 2=smallcell, 3=adeno, 4=large

time: survival time status: censoring status

karno: Karnofsky performance score (100=good) diagtime: months from diagnosis to randomisation

age: in years

prior: prior therapy 0=no, 10=yes

#### **Source**

D Kalbfleisch and RL Prentice (1980), *The Statistical Analysis of Failure Time Data*. Wiley, New York.

xtfrm.Surv

Sorting order for Surv objects

# Description

Sort survival objects into a partial order, which is the same one used internally for many of the calculations.

## Usage

```
## S3 method for class 'Surv'
xtfrm(x)
```

#### **Arguments**

Х

a Surv object

## **Details**

This creates a partial ordering of survival objects. The result is sorted in time order, for tied pairs of times right censored events come after observed events (censor after death), and left censored events are sorted before observed events. For counting process data (tstart,tstop,status) the ordering is by stop time, status, and start time, again with censoring last. Interval censored data is sorted using the midpoint of each interval.

The xtfrm routine is used internally by order and sort, so these results carry over to those routines.

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## Value

a vector of integers which will have the same sort order as x.

## Author(s)

Terry Therneau

## See Also

```
sort, order
```

## **Examples**

```
test <- c(Surv(c(10, 9,9, 8,8,8,7,5,5,4), rep(1:0, 5)), Surv(6.2, NA)) test sort(test)
```

yates

Population prediction

## **Description**

Compute population marginal means (PMM) from a model fit, for a chosen population and statistic.

## Usage

```
yates(fit, term, population = c("data", "factorial", "sas"),
levels, test = c("global", "trend", "pairwise"), predict = "linear",
options, nsim = 200, method = c("direct", "sgtt"))
```

# Arguments

fit	a model fit. Examples using lm, glm, and coxph objects are given in the vignette.
term	the term from the model whic is to be evaluated. This can be written as a character string or as a formula.
population	the population to be used for the adjusting variables. User can supply their own data frame or select one of the built in choices. The argument also allows "empirical" and "yates" as aliases for data and factorial, respectively, and ignores case.
levels	optional, what values for term should be used.
test	the test for comparing the population predictions.
predict	what to predict. For a glm model this might be the 'link' or 'response'. For a coxph model it can be linear, risk, or survival. User written functions are allowed.
options	optional arguments for the prediction method.

yates\_setup 179

nsim number of simulations used to compute a variance for the predictions. This is

not needed for the linear predictor.

method the computational approach for testing equality of the population predictions.

Either the direct approach or the algorithm used by the SAS glim procedure for

"type 3" tests.

## **Details**

The many options and details of this function are best described in a vignette on population prediction.

#### Value

an object of class yates with components of

estimate a data frame with one row for each level of the term, and columns containing the

level, the mean population predicted value (mppv) and its standard deviation.

tests a matrix giving the test statistics

mvar the full variance-covariance matrix of the mppv values

summary optional: any further summary if the values provided by the prediction method.

#### Author(s)

Terry Therneau

## **Examples**

```
fit1 <- lm(skips ~ Solder*Opening + Mask, data = solder)
yates(fit1, ~Opening, population = "factorial")

fit2 <- coxph(Surv(time, status) ~ factor(ph.ecog)*sex + age, lung)
yates(fit2, ~ ph.ecog, predict="risk") # hazard ratio</pre>
```

yates\_setup

Method for adding new models to the yates function.

## **Description**

This is a method which is called by the yates function, in order to setup the code to handle a particular model type. Methods for glm, coxph, and default are part of the survival package.

## Usage

```
yates_setup(fit, ...)
```

180 yates\_setup

## **Arguments**

fit a fitted model object

... optional arguments for some methods

## **Details**

If the predicted value should be the linear predictor, the function should return NULL. The yates routine has particularly efficient code for this case. Otherwise it should return a prediction function or a list of two elements containing the prediction function and a summary function. The prediction function will be passed the linear predictor as a single argument and should return a vector of predicted values.

## Note

See the vignette on population prediction for more details.

## Author(s)

Terry Therneau

## See Also

yates

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