Package: FastJM (via r-universe)

August 27, 2024

Type Package

Title Semi-Parametric Joint Modeling of Longitudinal and Survival Data

Version 1.4.2

Date 2024-02-29

Maintainer Shanpeng Li lishanpeng0913@ucla.edu>

Description Maximum likelihood estimation for the semi-parametric joint modeling of competing risks and longitudinal data applying customized linear scan algorithms, proposed by Li and colleagues (2022) <doi:10.1155/2022/1362913>. The time-to-event data is modelled using a (cause-specific) Cox proportional hazards regression model with time-fixed covariates. The longitudinal outcome is modelled using a linear mixed effects model. The association is captured by shared random effects. The model is estimated using an Expectation Maximization algorithm.

License GPL (>= 3)

NeedsCompilation yes

Imports Rcpp (>= 1.0.7), dplyr, nlme, caret, survival, timeROC

LinkingTo Rcpp, RcppEigen

Depends R (>= 3.5.0), statmod, MASS

RoxygenNote 7.2.3

LazyData true

Suggests testthat (>= 3.0.0), spelling

Language en-US

Config/testthat/edition 3

Repository https://shanpengli.r-universe.dev

RemoteUrl https://github.com/shanpengli/fastjm

RemoteRef HEAD

RemoteSha c8b70f44fe3cd18e122e5634fc97607b77d81ecf

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anova

Anova Method for Fitted Joint Models

Description

Performs a likelihood ratio test between two nested joint models.

Usage

```
## S3 method for class 'jmcs'
anova(object, object2, digits = 4, ...)
```

Arguments

```
object an object inheriting from class jmcs, nested in object2.

object2 an object inheriting from class jmcs.

digits the number of significant digits to use when printing. Default is 4.

further arguments passed to or from other methods.
```

Value

A table to summarize the likelihood ratio test.

AUCjmcs 3

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

See Also

jmcs

Examples

AUCjmcs

Time-dependent AUC for joint models

Description

Time-dependent AUC for joint models

Usage

```
AUCjmcs(
   seed = 100,
   object,
   landmark.time = NULL,
   horizon.time = NULL,
   obs.time = NULL,
   method = c("Laplace", "GH"),
   quadpoint = NULL,
   maxiter = NULL,
   n.cv = 3,
   survinitial = TRUE,
   ...
)
```

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Arguments

seed a numeric value of seed to be specified for cross validation.

object of class 'jmcs'.

landmark.time a numeric value of time for which dynamic prediction starts..

horizon.time a numeric vector of future times for which predicted probabilities are to be com-

puted.

obs. time a character string of specifying a longitudinal time variable.

method estimation method for predicted probabilities. If Laplace, then the empirical

empirical estimates of random effects is used. If GH, then the pseudo-adaptive

Gauss-Hermite quadrature is used.

quadpoint the number of pseudo-adaptive Gauss-Hermite quadrature points if method =

"GH".

maxiter the maximum number of iterations of the EM algorithm that the function will

perform. Default is 10000.

n.cv number of folds for cross validation. Default is 3.

survinitial Fit a Cox model to obtain initial values of the parameter estimates. Default is

TRUE.

... Further arguments passed to or from other methods.

Value

a list of matrices with conditional probabilities for subjects.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

See Also

jmcs, survfitjmcs

cdata Simulated competing risks data

Description

The cdata data frame has 1000 rows and 7 columns.

Usage

data(cdata)

fitted 5

Format

```
This data frame contains the following columns:
```

```
ID patient identifier.

surv event time.

failure_type event indicator. 0 denotes censoring, 1 risk 1, and 2 risk 2.

x1 continuous variable.

x2 treatment indicator. 0 denotes the placebo group and 1 the treatment group.

gender gender indicator.

race race indicator.
```

fitted

Fitted values for joint models

Description

Extract fitted values for joint models.

Usage

```
## S3 method for class 'jmcs'
fitted(
  object,
  type = c("Marginal", "Subject"),
  process = c("Longitudinal", "Event"),
  ...
)
```

Arguments

object an object inheriting from class jmcs.

type for which type of fitted values to calculate.

process for which sub-model to calculate the fitted values.

further arguments passed to or from other methods.

Value

a numeric vector of fitted values.

Author(s)

Shanpeng Li < lishanpeng 0913@ucla.edu>

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Examples

fixef

Estimated coefficients estimates for joint models

Description

Extracts the fixed effects for a fitted joint model.

Usage

```
fixef(object, process = c("Longitudinal", "Event"), ...)
```

Arguments

object an object inheriting from class jmcs.

process for which sub-model to extract the estimated coefficients.

further arguments passed to or from other methods.

Value

A numeric vector or a list of the estimated parameters for the fitted model.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

Examples

```
# fixed effects for the longitudinal process
fixef(fit, process = "Longitudinal")
# fixed effects for the event process
fixef(fit, process = "Event")
```

jmcs

Joint modeling of longitudinal continuous data and competing risks

Description

Joint modeling of longitudinal continuous data and competing risks

Usage

```
jmcs(
  ydata,
  cdata,
  long.formula,
  random = NULL,
  surv.formula,
  REML = TRUE,
  quadpoint = NULL,
  maxiter = 10000,
  print.para = FALSE,
  survinitial = TRUE,
  tol = 1e-04,
  method = "pseudo-adaptive",
  opt = "nlminb"
)
```

Arguments

	- 1itdim-1 data faana in 1	
ydata	a longitudinal data frame in l	ong tormat.

cdata a survival data frame with competing risks or single failure. Each subject has

one data entry.

long, formula a formula object with the response variable and fixed effects covariates to be

included in the longitudinal sub-model.

random a one-sided formula object describing the random effects part of the longitudi-

nal sub-model. For example, fitting a random intercept model takes the form \sim 1 | ID. Alternatively. Fitting a random intercept and slope model takes the form

 $\sim x1 + \ldots + xn | ID.$

surv. formula a formula object with the survival time, event indicator, and the covariates to be

included in the survival sub-model.

REML a logic object that indicates the use of REML estimator. Default is TRUE.

quadpoint the number of pseudo-adaptive Gauss-Hermite quadrature points, to be chosen

for numerical integration. Default is 6 which produces stable estimates in most

dataframes.

maxiter the maximum number of iterations of the EM algorithm that the function will

perform. Default is 10000.

print para Print detailed information of each iteration. Default is FALSE, i.e., not to print

the iteration details.

survinitial Fit a Cox model to obtain initial values of the parameter estimates. Default is

TRUE.

tol Tolerance parameter. Default is 0.0001.

method Method for proceeding numerical integration in the E-step. Default is pseudo-

adaptive.

opt Optimization method to fit a linear mixed effects model, either nlminb (default)

or optim.

Value

Object of class jmcs with elements

beta the vector of fixed effects for the linear mixed effects model.

gamma1 the vector of fixed effects for type 1 failure for the survival model.

gamma2 the vector of fixed effects for type 2 failure for the survival model. Valid only if

CompetingRisk = TRUE.

nu1 the vector of association parameter(s) for type 1 failure.

nu2 the vector of association parameter(s) for type 2 failure. Valid only if CompetingRisk

= TRUE.

H01 the matrix that collects baseline hazards evaluated at each uncensored event time

for type 1 failure. The first column denotes uncensored event times, the second column the number of events, and the third columns the hazards obtained by

Breslow estimator.

H02 the matrix that collects baseline hazards evaluated at each uncensored event

time for type 2 failure. The data structure is the same as H01. Valid only if

CompetingRisk = TRUE.

Sig the variance-covariance matrix of the random effects.

sigma the variance of the measurement error for the linear mixed effects model.

iter the total number of iterations until convergence.

convergence convergence identifier: 1 corresponds to successful convergence, whereas 0 to a

problem (i.e., when 0, usually more iterations are required).

vcov the variance-covariance matrix of all the fixed effects for both models.

sebeta the standard error of beta. segamma1 the standard error of gamma1.

segamma2 the standard error of gamma2. Valid only if CompetingRisk = TRUE.

senu1 the standard error of nu1.

senu2 the standard error of nu2. Valid only if CompetingRisk = TRUE.

seSig the vector of standard errors of covariance of random effects.

sesigma the standard error of variance of measurement error for the linear mixed effects

model.

loglike the log-likelihood value.

fitted a list with the fitted values:

resid the vector of estimated residuals for the linear mixed effects model.

fitted the vector of fitted values for the linear mixed effects model.

fittedmar the vector of marginal fitted values for the linear mixed effects model. **residmar** the vector of estimated marginal residuals for the linear mixed effects

model

fittedSurv the estimated survival rate evaluated at each uncensored event time.

FUNB the estimated random effects for each subject.

CompetingRisk logical value; TRUE if a competing event are accounted for.

quadpoint the number of Gauss Hermite quadrature points used for numerical integration.

ydata the input longitudinal dataset for fitting a joint model. It has been re-ordered in

accordance with descending observation times in cdata.

cdata the input survival dataset for fitting a joint model. It has been re-ordered in

accordance with descending observation times.

PropEventType a frequency table of number of events.

LongitudinalSubmodel

the component of the long.formula.

SurvivalSubmodel

the component of the surv.formula.

random the component of the random.

call the matched call.

Quad.method the quadrature rule used for integration. If pseudo-adaptive quadrature rule is

used, then return pseudo-adaptive. Otherwise return standard.

id the grouping vector for the longitudinal outcome.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

See Also

ranef, fixef, fitted.jmcs,residuals.jmcs, survfitjmcs, plot.jmcs,vcov.jmcs

Examples

```
require(FastJM)
require(survival)
# Load a simulated longitudinal dataset
data(ydata)
# Load a simulated survival dataset with two competing events
data(cdata)
# Fit a joint model
fit <- jmcs(ydata = ydata, cdata = cdata,</pre>
            long.formula = response ~ time + gender + x1 + race,
            surv.formula = Surv(surv, failure_type) ~ x1 + gender + x2 + race,
            random = \sim time | ID)
fit
# Extract the parameter estimates of longitudinal sub-model fixed effects
fixef(fit, process = "Longitudinal")
# Extract the parameter estimates of survival sub-model fixed effects
fixef(fit, process = "Event")
# Obtain the random effects estimates for first 6 subjects
head(ranef(fit))
# Obtain the variance-covariance matrix of all parameter estimates
vcov(fit)
# Obtain the result summaries of the joint model fit
summary(fit, process = "Longitudinal")
summary(fit, process = "Event")
# Prediction of cumulative incidence for competing risks data
# Predict the conditional probabilities for two patients who are alive (censored)
ND <- ydata[ydata$ID %in% c(419, 218), ]
ID <- unique(ND$ID)</pre>
NDc <- cdata[cdata$ID %in% ID, ]
survfit <- survfitjmcs(fit,</pre>
                       ynewdata = ND,
                       cnewdata = NDc,
                       u = seq(3, 4.8, by = 0.2),
                       method = "GH"
                       obs.time = "time")
survfit
PE <- PEjmcs(fit, seed = 100, landmark.time = 3, horizon.time = c(3.6, 4, 4.4),
             obs.time = "time", method = "GH",
             quadpoint = NULL, maxiter = 1000, n.cv = 3,
             survinitial = TRUE)
Brier <- summary(PE, error = "Brier")</pre>
Brier
MAEQ <- MAEQjmcs(fit, seed = 100, landmark.time = 3, horizon.time = c(3.6, 4, 4.4),
                 obs.time = "time", method = "GH",
                 quadpoint = NULL, maxiter = 1000, n.cv = 3,
                 survinitial = TRUE)
APE <- summary(MAEQ, digits = 3)
APE
```

evaluate prediction accuracy of fitted joint model using cross-validated mean AUC

MAEQjmcs 11

```
AUC <- AUCjmcs(fit, seed = 100, landmark.time = 3, horizon.time = c(3.6, 4, 4.4), obs.time = "time", method = "GH", quadpoint = NULL, maxiter = 1000, n.cv = 3) summary(AUC, digits = 3)
```

MAEQjmcs

A metric of prediction accuracy of joint model by comparing the predicted risk with the empirical risks stratified on different predicted risk group.

Description

A metric of prediction accuracy of joint model by comparing the predicted risk with the empirical risks stratified on different predicted risk group.

Usage

```
MAEQjmcs(
   seed = 100,
   object,
   landmark.time = NULL,
   horizon.time = NULL,
   obs.time = NULL,
   method = c("Laplace", "GH"),
   quadpoint = NULL,
   maxiter = 1000,
   n.cv = 3,
   survinitial = TRUE,
   quantile.width = 0.25,
   ...
)
```

Arguments

seed a numeric value of seed to be specified for cross validation.

object of class 'jmcs'.

landmark.time a numeric value of time for which dynamic prediction starts..

horizon.time a numeric vector of future times for which predicted probabilities are to be com-

puted.

obs. time a character string of specifying a longitudinal time variable.

method estimation method for predicted probabilities. If Laplace, then the empirical

empirical estimates of random effects is used. If GH, then the pseudo-adaptive

Gauss-Hermite quadrature is used.

PEjmcs

quadpoint	the number of pseudo-adaptive Gauss-Hermite quadrature points if method = "GH".
maxiter	the maximum number of iterations of the EM algorithm that the function will perform. Default is 10000 .
n.cv	number of folds for cross validation. Default is 3.
survinitial	Fit a Cox model to obtain initial values of the parameter estimates. Default is TRUE.
quantile.width	a numeric value of width of quantile to be specified. Default is 0.25.
	Further arguments passed to or from other methods.

Value

a list of matrices with conditional probabilities for subjects.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

See Also

```
jmcs, survfitjmcs
```

PEjmcs A metric of prediction accuracy of joint model by comparing the predicted risk with the counting process.

Description

A metric of prediction accuracy of joint model by comparing the predicted risk with the counting process.

Usage

```
PEjmcs(
  object,
  seed = 100,
  landmark.time = NULL,
  horizon.time = NULL,
  obs.time = NULL,
  method = c("Laplace", "GH"),
  quadpoint = NULL,
  maxiter = NULL,
  n.cv = 3,
  survinitial = TRUE,
  ...
)
```

plot.jmcs 13

Arguments

object of class 'jmcs'.

seed a numeric value of seed to be specified for cross validation.

landmark.time a numeric value of time for which dynamic prediction starts..

horizon.time a numeric vector of future times for which predicted probabilities are to be com-

puted.

obs. time a character string of specifying a longitudinal time variable.

method estimation method for predicted probabilities. If Laplace, then the empirical

empirical estimates of random effects is used. If GH, then the pseudo-adaptive

Gauss-Hermite quadrature is used.

quadpoint the number of pseudo-adaptive Gauss-Hermite quadrature points if method =

"GH".

maxiter the maximum number of iterations of the EM algorithm that the function will

perform. Default is 10000.

n.cv number of folds for cross validation. Default is 3.

survinitial Fit a Cox model to obtain initial values of the parameter estimates. Default is

TRUE.

. . . Further arguments passed to or from other methods.

Value

a list of matrices with conditional probabilities for subjects.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

See Also

```
jmcs, survfitjmcs
```

plot.jmcs

Fitted values for joint models

Description

Plot Diagnostics for Joint Models.

Usage

```
## S3 method for class 'jmcs'
plot(x, add.smooth = getOption("add.smooth"), ...)
```

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Arguments

```
    x x of class 'jmcs'.
    add. smooth logical; if TRUE a smooth line is superimposed in the "Residuals vs Fitted" plot.
    further arguments passed to or from other methods.
```

Value

The first two plots are longitudinal sub-model diagnostics and the last two are marginal survival function and marginal cumulative hazard.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

Examples

print

Print jmcs

Description

Print jmcs

Usage

```
## S3 method for class 'jmcs'
print(x, digits = 4, ...)
```

Arguments

```
x Object of class 'jmcs'.digits the number of significant digits to use when printing.... Further arguments passed to or from other methods.
```

Value

a summary of data, joint model, log likelihood, and parameter estimates.

print.survfitjmcs 15

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

See Also

jmcs

print.survfitjmcs

Print survfitjmcs

Description

Print survfitjmcs

Usage

```
## S3 method for class 'survfitjmcs' print(x, ...)
```

Arguments

x x of class 'survfitjmcs'.

. . . Further arguments passed to or from other methods.

Value

a list of matrices with conditional probabilities for subjects.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

See Also

```
jmcs, survfitjmcs
```

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ranef

Random effects estimates for joint models

Description

Extracts the posterior mean of the random effects for a fitted joint model.

Usage

```
ranef(object, ...)
```

Arguments

object an object inheriting from class jmcs.

... further arguments passed to or from other methods.

Value

a matrix of random effects estimates.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

See Also

jmcs

Examples

residuals 17

residuals

Residuals for joint models

Description

Extract residuals for joint models.

Usage

```
## S3 method for class 'jmcs'
residuals(object, type = c("Marginal", "Subject"), ...)
```

Arguments

object an object inheriting from class jmcs.
type what type of residuals to calculate.

... further arguments passed to or from other methods.

Value

a vector of residuals of the longitudinal sub-model.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

See Also

jmcs

Examples

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summary

Anova Method for Fitted Joint Models

Description

Produce result summaries of a joint model fit.

Usage

```
## S3 method for class 'jmcs'
summary(object, process = c("Longitudinal", "Event"), digits = 4, ...)
```

Arguments

object an object inheriting from class jmcs.

process for which model (i.e., longitudinal model or survival model) to extract the esti-

mated coefficients.

digits the number of significant digits to use when printing. Default is 4.

. . . further arguments passed to or from other methods.

Value

A table to summarize the model results.

See Also

jmcs

summary.AUCjmcs

Print AUCjmcs

Description

Print AUCjmcs

Usage

```
## S3 method for class 'AUCjmcs'
summary(object, digits = 4, ...)
```

Arguments

object of class 'AUCjmcs'.

digits number of digits of decimal to be printed.

Further arguments passed to or from other methods.

summary.MAEQjmcs 19

Value

a list of matrices with conditional probabilities for subjects.

Author(s)

```
Shanpeng Li lishanpeng0913@ucla.edu>
```

See Also

```
jmcs, survfitjmcs
```

summary.MAEQjmcs

Print MAEQjmcs

Description

Print MAEQimcs

Usage

```
## S3 method for class 'MAEQjmcs'
summary(object, digits = 3, ...)
```

Arguments

object of class 'MAEQjmcs'.

digits number of decimal points to be rounded.

... Further arguments passed to or from other methods.

Value

a list of matrices with conditional probabilities for subjects.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

See Also

```
jmcs, survfitjmcs
```

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summary.PEjmcs

Print PEjmcs

Description

Print PEjmcs

Usage

```
## S3 method for class 'PEjmcs'
summary(object, error = c("MAE", "Brier"), ...)
```

Arguments

object of class 'PEjmcs'.

error a character string that specifies the loss function.

. . . Further arguments passed to or from other methods.

Value

a list of matrices with conditional probabilities for subjects.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

See Also

```
jmcs, survfitjmcs
```

survfitjmcs

Prediction in Joint Models

Description

This function computes the conditional probability of surviving later times than the last observed time for which a longitudinal measurement was available.

survfitjmcs 21

Usage

```
survfitjmcs(
  object,
  seed = 100,
  ynewdata = NULL,
  cnewdata = NULL,
  u = NULL,
  Last.time = NULL,
  obs.time = NULL,
  method = c("Laplace", "GH"),
  quadpoint = NULL,
  ...
)
```

Arguments

object an object inheriting from class jmcs.

seed a random seed number to proceed Monte Carlo simulation. Default is 100.

ynewdata a data frame that contains the longitudinal and covariate information for the

subjects for which prediction of survival probabilities is required.

cnewdata a data frame that contains the survival and covariate information for the subjects

for which prediction of survival probabilities is required.

u a numeric vector of times for which prediction survival probabilities are to be

computed.

Last.time a numeric vector or character string. This specifies the known time at which

each of the subjects in cnewdata was known to be alive. If NULL, then this is automatically taken as the survival time of each subject. If a numeric vector, then it is assumed to be greater than or equals to the last available longitudinal time point for each subject. If a character string, then it should be a variable in

cnewdata.

obs.time a character string of specifying a longitudinal time variable in ynewdata.

method a character string specifying the type of probability approximation; if Laplace,

then a first order estimator is computed. If GH, then the standard Gauss-Hermite

quadrature is used instead.

quadpoint number of quadrature points used for estimating conditional probabilities when

method = "GH". Default is NULL. If method = "GH", then use the same amount

of quadrature points obtained from object.

... further arguments passed to or from other methods.

Value

a list of matrices with conditional probabilities for subjects.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

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See Also

jmcs

vcov

Variance-covariance matrix of the estimated parameters for joint models

Description

Extract variance-covariance matrix for joint models.

Usage

```
## S3 method for class 'jmcs'
vcov(object, ...)
```

Arguments

object an object inheriting from class jmcs.

... further arguments passed to or from other methods.

Value

a matrix of variance covariance of all parameter estimates.

Author(s)

Shanpeng Li lishanpeng0913@ucla.edu>

See Also

jmcs

ydata

Simulated longitudinal data

Description

The ydata data frame has 3067 rows and 6 columns.

Usage

```
data(ydata)
```

ydata 23

Format

This data frame contains the following columns:

ID patient identifier.

response response variable.

time visit time.

x1 treatment indicator. 0 denotes the placebo group and 1 the treatment group.

gender gender indicator.

race race indicator.

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