

# Package ‘curephEM’

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**Title** NPMLE for Logistic-Cox Cure-Rate Model

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## Description

Expectation-Maximization (EM) algorithm for point estimation and variance estimation to the nonparametric maximum likelihood estimator (NPMLE) for logistic-Cox cure-rate model with left truncation and right-censoring. See Hou, Chambers and Xu (2017) <[doi:10.1007/s10985-017-9415-2](https://doi.org/10.1007/s10985-017-9415-2)>.

**License** GPL-2

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cureph

*NPMLE for Logistic-CoxPH Cure-Rate Model***Description**

Fit NPMLE for Logistic-CoxPH Cure-Rate Model

**Usage**

```
cureph (formula, formula2 , data, subset, na.action, init, control,
        method = c("EM"), singular.ok = TRUE,
        var = c("Louis"),...)
```

**Arguments**

formula	a formula object for cure logistic model,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 <a href="#">Surv.cure</a> function.
formula2	a formula object for event time CoxPH model The Response can be suppressed.
data	a data.frame in which to interpret the variables named in the formula and formula2, 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 <code>options()\$na.action</code> .
init	a list of length 3, initial values of the iteration. Its components are: a vector of formula coefficients,a vector of formula2 coefficients and a step.fun step function of baseline cumulative hazard function. Default initial value is the estimate from naive separate glm and coxph models.
control	Object of class <a href="#">cureph.control</a> specifying iteration limit and other control options. Default is <code>cureph.control(...)</code> .
method	a character string specifying the method for estimation. The default method is the EM algorithm.
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.
var	a character string specifying the method for variance estimation. The default method is derived from the Louis formula.
...	other parameters passed to <code>cureph.control</code> .

## Details

The function can handle left truncation.

"BFGS-Newton" method optimizes the discretized nonparametric likelihood directly. It has 3 steps. First, it roughly search for a local maximal by BFGS method with default setup. Second, it rescales the parameters proportional to the reciprocal of current gradient in another BFGS optimization. Finally, it searches for a solution to the score equations through a Newton's method with Armijo line search.

The "EM" method uses a "ghost copy" algorithm to accomodate left-truncation.

Both variance estimation methods are asymptotically correct, but they have undercoverage issue in case of a small event number. "Louis" is recommended among the two.

See *Hou et al. (2016)* for detail.

## Author(s)

Jue (Marquis) Hou

## References

Hou, J., Chambers, C. and Xu, R. (2016). "A nonparametric maximum likelihood approach for partially observed cured data with left truncation and right-censoring". To be submitted.

## See Also

[cureph.object](#), [summary.cureph](#), [survpred.cureph](#)

## Examples

```
# A simulated data set
data('sim.cureph.data')
# Or you may generate another one
sim.cureph.data = cureph.simgen()

# Check out the true coefficients and baseline survival
attributes(sim.cureph.data)

# Fit logistic-CoxPH cure-rate model
fit=cureph(Surv.cure(time,time2,event,origin=0,end=20)~Z1+Z2+Z3+Z4,data=sim.cureph.data)

# Check out the summary
summary(fit)

# Baseline cummulative hazard
basehaz.cureph(fit)
```

---

 cureph.control

*Ancillary arguments for controlling cureph fits*


---

**Description**

Assign the ancillary arguments for controlling cureph fits

**Usage**

```
cureph.control(n.data,eps = 1e-09, toler.chol = .Machine$double.eps^0.75,
               iter.max = 1000, toler.inf = eps^(1/3),
               line.search=0.5,init.step = 1/n.data)
```

**Arguments**

n.data	Sample size of the data
eps	Iteration continues until the relative change in the log partial likelihood is less than 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.
line.search	The scale of shrinkage in each Armijo line search step.
init.step	The initial size of step in Newton's method.

**Details**

line.search and init.step only matter when method = "BFGS-Newton".

**Value**

a list containing the values of each of the above constants

**Author(s)**

Jue (Marquis) Hou

**See Also**

[cureph](#)

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cureph.object	<i>Cure-Rate Logistic-CoxPH Regression Object</i>
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### Description

This class of objects is returned by the `cureph` class of functions to represent a fitted cure-rate logistic-CoxPH model. Objects of this class have methods for the functions `summary` and `survpred`.

### Arguments

<code>coefficients</code>	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.
<code>var</code>	the variance matrix of the coefficients. Rows and columns corresponding to any missing coefficients are set to zero.
<code>loglik</code>	the log-likelihood with the final values of the coefficients.
<code>baseline</code>	the step-function for baseline cumulative hazard.
<code>wald.test</code>	the Wald test of whether the final coefficients differ from the initial values.
<code>iter</code>	number of iterations used.
<code>means</code>	vector of column means of the X matrix. Subsequent survival curves are adjusted to this value. Factors are set to be the baseline level.
<code>n</code>	the number of observations used in the fit.
<code>nevent</code>	the number of events (usually deaths) used in the fit.
<code>method</code>	the computation method used.
<code>na.action</code>	the <code>na.action</code> attribute, if any, that was returned by the <code>na.action</code> routine.
<code>...</code>	The object will also contain the following: <code>terms.logistic</code> , <code>terms.cox</code> , <code>assign.logistic</code> , <code>assign.cox</code> , <code>df</code> , <code>var.levels</code> , <code>formula.logistic</code> , <code>formula.cox</code> , <code>contrasts.logistic</code> , <code>contrasts.cox</code> , <code>call</code> , <code>data</code> , <code>Y</code> , <code>origin</code> and <code>end</code> .

### See Also

[cureph](#), [summary.cureph](#), [survpred.cureph](#).

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<code>sim.cureph.data</code>	<i>Simulated Data for cureph</i>
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### Description

A simulated data for `cureph` example.

### Usage

```
data("sim.cureph.data")

cureph.simgen()
```

**Format**

A data frame with 200 observations on the following 7 variables.

time entry time.

time2 exit time.

event binary event indicator.

Z1 numerical vector.

Z2 factor with levels '0' and '1'.

Z3 factor with levels 'A', 'B' and 'C'.

Z4 factor with levels '0', '1' and '2', colinear with Z2 and Z3.

attr(,"true.coef") the true coefficients.

attr(,"true.surv0") the true baseline survival function.

---

summary.cureph

*Summarizing Logistic-CoxPH Cure-Rate Model Fit*

---

**Description**

These functions are all methods for class cureph or summary.cureph objects.

**Usage**

```
## S3 method for class 'cureph'
summary(object,combine = T,...)

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

**Arguments**

object	an object of class "cureph", usually, a result of a call to <a href="#">cureph</a> .
x	an object of class "summary.glm", usually, a result of a call to <code>summary.glm</code> .
combine	logical; if TRUE, a Wald test table of each variable appearing in both models is returned.
digits	the number of significant digits to use when printing.
signif.stars	logical. If TRUE, "significance stars" are printed for each coefficient.
...	Other parameters to match the generic S3 method.

**See Also**

[cureph](#)

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Surv.cure	<i>Survival Object with Cure Portion</i>
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**Description**

Create a Survival Object with Cure Portion

**Usage**

```
Surv.cure(time, time2, event, type=c("right","counting"), origin = 0, end = Inf)
```

```
## S3 method for class 'Surv.cure'  
print(x, digit = getOption("digits"),...)
```

**Arguments**

time	for right censored data, this is the follow up time. For left truncated data, the first argument is the truncation time.
event	The status indicator, normally 0=alive, 1=dead. Other choices are TRUE/FALSE (TRUE = death) or 1/2 (2=death). Although unusual, the event indicator can be omitted, in which case all subjects are assumed to have an event.
time2	ending time of the counting process data only. Intervals are assumed to be open on the left and closed on the right, (start, end]. Event indicates whether an event occurred at the end of the interval.
type	character string specifying the type of censoring. Possible values are "right" and "counting".
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.
end	the upper bound for event time. Any subject survives beyond this time is considered as an observed cure.
x	a 'Surv.cure' object.
digit	number of decimal digits in the output.
...	Other parameters to match the generic S3 method.

**See Also**

[Surv](#)

**Examples**

```
# A simulated data set
data("sim.cureph.data")
attach(sim.cureph.data)

# Construct and print a Surv.cure object
Surv.cure(time,time2,event,origin=0,end=20)

# Detach the simulated data
detach(sim.cureph.data)
```

---

survpred.cureph

*Prediction for a Logistic-CoxPH Cure-Rate Model*


---

**Description**

Compute Predictions and Survival Curves from a Logistic-CoxPH Cure-Rate Model

**Usage**

```
## S3 method for class 'cureph'
survpred(object, newdata , time, center = F)
basehaz.cureph(object)

## S3 method for class 'survpred.cureph'
plot(x, pooled = T, censor = x$censored,...)
```

**Arguments**

object	a cureph object.
newdata	a data frame with the same variable names as those that appear in the cureph 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 data frame used in the original fit for prediction and the mean of the covariates for survival curve. See <a href="#">cureph.object</a> .
time	a numeric vector; all the time points to be evaluated in the survival table in addition to the observed event times.
center	logical; if TRUE, the baseline survival is evaluated at the mean of the covariates. Otherwise, it is evaluated at zero.
x	a survpred.cureph object.
pooled	logical; if TRUE, the marginal survival curve(s) for the cure-rate model will be plotted. Otherwise, the conditional survival curve(s) in the Cox part will be plotted.



censor            a numeric vector of the observed censoring times. If not NULL, a cross will mark each censored point on the survival curve.  
 ...                Other parameters to match the generic S3 method.

### Details

The means of factor variables are not computed. Instead, the baseline level is used.  
 The NPMLE baseline survival is estimated in a Breslow-Nelson-Aalen fashion.

### Value

logistic.linear.predict            the linear predictions for cure indicator logistic model.  
 logistic.linear.prob                the estimated probability for NOT being cured.  
 cox.linear.predict                  the linear predictions for event time CoxPH model.  
 cox.cumhaz                          the baseline cumulative hazard for event time CoxPH model.  
 surv.cox                             the marginal survival table for the cure-rate model.  
 surv.cureph                         the conditional survival table for event time CoxPH model.  
 ...                                  The object will also contain the following: censored, origin and end.

### Author(s)

Jue (Marquis) Hou

### See Also

[cureph](#)

### Examples

```

library(curephEM)

# A simulated data set
data("sim.cureph.data")

# Fit logistic-CoxPH cure-rate model
fit=cureph(Surv.cure(time,time2,event,origin=0,end=20)~Z1+Z2+Z3+Z4,data=sim.cureph.data)

# The survival-prediction object
mysurv = survpred(fit,center=TRUE)

oldpar = par()
par(mfrow=c(1,2))
plot(mysurv)
plot(mysurv,FALSE)
par(oldpar)

```

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