Package ‘multgee’

October 13, 2022

Type Package
Title GEE Solver for Correlated Nominal or Ordinal Multinomial Responses
Version 1.8.0
Depends R (>= 2.15.0), gnm
Imports stats, utils, VGAM, Rcpp
Suggests knitr, rmarkdown, rticles
Description GEE solver for correlated nominal or ordinal multinomial responses using a local odds ratios parameterization.
License GPL-2 | GPL-3
LazyData true
VignetteBuilder knitr
URL https://github.com/AnestisTouloumis/multgee
BugReports https://github.com/AnestisTouloumis/multgee/issues
RoxygenNote 7.1.1
Encoding UTF-8
LinkingTo Rcpp, RcppArmadillo
NeedsCompilation yes
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Repository CRAN
Date/Publication 2021-05-13 17:40:02 UTC

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A GEE Solver For Correlated Nominal Or Ordinal Multinomial Responses

Description

A generalized estimating equations (GEE) solver for fitting marginal regression models with correlated nominal or ordinal multinomial responses based on a local odds ratios parameterization for the association structure.

Details

The package contains two functions that fit GEE models for correlated multinomial responses; ordLORgee for an ordinal response scale and nomLORgee for a nominal response scale.

The main arguments in both functions are: (i) an optional data frame (data), (ii) a model formula (formula), (iii) a cluster identifier variable (id) and (iv) an optional vector that identifies the order of the observations within each cluster (repeated).

Options for the marginal model in the function ordLORgee include cumulative link models or an adjacent categories logit model. A marginal baseline category logit model is offered in the function nomLORgee. For the form of the linear predictor in these models, see the Details sections in nomLORgee and ordLORgee.

The association structure among the correlated multinomial responses is expressed via marginalized local odds ratios (Touloumis et al., 2013). The estimating procedure for the local odds ratios can be summarized as follows: For each level pair of the repeated variable, the available responses are aggregated across clusters to form a square marginalized contingency table. Treating these tables as independent, an RC-G(1) type model (Becker and Clogg, 1989) is fitted in order to estimate the marginalized local odds ratios. The LORstr argument determines the form of the marginalized local odds ratios structure. Since the general RC-G(1) model is closely related to the family of association models (Goodman, 1985), one can instead fit an association model to each of the marginalized contingency tables by setting LORem="2way".

If the underlying association pattern does not change dramatically across the level pairs of repeated then parsimonious marginalized local odds ratios should sufficiently approximate the true underlying association structure. To assess the underlying association structure, one might use the utility function intrinsic.pars.
Instead of estimating the local odds ratios structure, a user-defined structure can be provided by setting LORstr="fixed". In this case, the utility function matrixLOR is useful in constructing the required LORterm argument.

The function waldts provides a goodness-of-fit test between two nested GEE models based on a Wald test statistic.

**Author(s)**

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**References**


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**arthritis**

Rheumatoid Arthritis Clinical Trial

**Description**

Rheumatoid self-assessment scores for 302 patients, measured on a five-level ordinal response scale at three follow-up times.

**Usage**

`arthritis`

**Format**

A data frame with 906 observations on the following 7 variables:

- **id** Patient identifier variable.
- **y** Self-assessment score of rheumatoid arthritis measured on a five-level ordinal response scale.
- **sex** Coded as (1) for female and (2) for male.
- **age** Recorded at the baseline.
- **trt** Treatment group variable, coded as (1) for the placebo group and (2) for the drug group.
- **baseline** Self-assessment score of rheumatoid arthritis at the baseline.
- **time** Follow-up time recorded in months.
**Source**

**Examples**
```r
data(arthritis)
str(arthritis)
```

### Description

Computes confidence intervals for one or more parameters in a fitted LORgee model.

#### Usage
```r
## S3 method for class 'LORgee'
confint(object, parm, level = 0.95, method = "robust", ...)
```

### Arguments
- `object`: a fitted model LORgee object.
- `parm`: a specification of which parameters are to be given confidence intervals, either a vector of numbers or a vector of names. If missing, all parameters are considered.
- `level`: the confidence level required.
- `method`: character indicating whether the sandwich (robust) covariance matrix (method = "robust") or the model–based (naive) covariance matrix (method = "naive") should be used for calculating the confidence intervals.
- `...`: additional argument(s) for methods.

### Details
The (Wald-type) confidence intervals are calculated using either the sandwich (robust) or the model-based (naive) covariance matrix.

### Value
A matrix (or vector) with columns giving lower and upper confidence limits for each parameter. These will be labelled as $(1 - \text{level})/2$ and $1 - (1 - \text{level})/2$ in % (by default 2.5% and 97.5%).

### Examples
```r
fitmod <- ordLORgee(formula = y ~ factor(time) + factor(trt) + factor(baseline),
                     data = arthritis, id = id, LORstr = "uniform", repeated = time)
confint(fitmod)
```
Description

Housing status for 362 severely mentally ill homeless subjects measured at baseline and at three follow-up times.

Usage

housing

Format

A data frame with 1448 observations on the following 4 variables:

id  Subject identifier variable.
y Housing status response, coded as (1) for street living, (2) for community living and (3) for independent housing.
time Time recorded in months.
sec Section 8 rent certificate indicator.

Source


Examples

data(housing)
str(housing)

intrinsic.pars  Intrinsıc Parameters Estimation

Description

Utility function to assess the underlying association pattern.

Usage

intrinsic.pars(y = y, data = parent.frame(), id = id, repeated = NULL, rscale = "ordinal")
Arguments

- **y**: a vector that identifies the response vector of the desired marginal model.
- **data**: an optional data frame containing the variables provided in y, id and repeated.
- **id**: a vector that identifies the clusters.
- **repeated**: an optional vector that identifies the order of observations within each cluster.
- **rscale**: a character string that indicates the nature of the response scale. Options include "ordinal" or "nominal".

Details

Simulation studies in Touloumis et al. (2013) suggested that if the range of the intrinsic parameter estimates is small then simple local odds ratios structures should adequately approximate the association pattern. Otherwise more complicated structures should be employed.

The intrinsic parameters are estimated under the heterogeneous linear-by-linear association model (Agresti, 2013) for ordinal response categories and under the RC-G(1) model (Becker and Clogg, 1989) with homogeneous score parameters for nominal response categories.

A detailed description of the arguments id and repeated can be found in the Details section of nomLORgee or ordLORgee.

Value

Returns a numerical vector with the estimated intrinsic parameters.

Author(s)

Anestis Touloumis

References


See Also

nomLORgee and ordLORgee.

Examples

```r
data(arthritis)
intrinsic.pars(y, arthritis, id, time, rscale = "ordinal")
## The intrinsic parameters do not vary much. The 'uniform' local odds ratios
## structure might be a good approximation for the association pattern.
set.seed(1)
```
data(housing)
intrinsic.pars(y, housing, id, time, rscale = "nominal")
## The intrinsic parameters vary. The 'RC' local odds ratios structure
## might be a good approximation for the association pattern.

---

**ipfp.control**

*IPFP Control*

**Description**

Control variables for the Iterative Proportion Fitting Procedure function `ipfp`.

**Usage**

`ipfp.control(tol = 1e-06, maxit = 200)`

**Arguments**

- **tol**: positive convergence tolerance. The algorithm converges when the absolute difference between the observed and the given row or column totals is less than or equal to `tol`.
- **maxit**: positive integer that indicates the maximum number of iterations.

**Note**

Currently the function `ipfp` is internal.

**Author(s)**

Anestis Touloumis

**See Also**

`nomLORgee` and `ordLORgee`. 
LORgee.control

Control For The GEE Solver

Description

Control variables for the GEE solver in the nomLORgee and ordLORgee functions.

Usage

## S3 method for class 'control'
LORgee(tolerance = 0.001, maxiter = 15, verbose = FALSE, TRACE = FALSE)

Arguments

tolerance  positive convergence tolerance. The algorithm converges when the maximum of the absolute relative difference in parameter estimates is less than or equal to tolerance.

maxiter  positive integer that indicates the maximum number of iterations in the Fisher-scoring iterative algorithm.

verbose  logical that indicates if output should be printed at each iteration.

TRACE  logical that indicates if the parameter estimates and the convergence criterion at each iteration should be saved.

Author(s)

Anestis Touloumis

See Also

nomLORgee and ordLORgee.

Examples

data(arthritis)
fitmod <- ordLORgee(y ~ factor(trt) + factor(baseline) + factor(time),
  data = arthritis, id = id, repeated = time)

## A one-step GEE estimator
fitmod1 <- update(fitmod, control = LORgee.control(maxiter = 1))
coef(fitmod)
coef(fitmod1)
Description

Utility function to create a square probability matrix that satisfies the specified local odds ratios structure.

Usage

matrixLOR(x)

Arguments

x

a square matrix with positive entries that describes the desired local odds ratios matrix.

Details

It is designed to ease the construction of the argument LORterm in the nomLORgee and ordLORgee functions.

Value

Returns a square probability matrix that satisfies the local odds ratios structure defined by x.

Warning

Caution is needed for local odds ratios close to zero.

Author(s)

Anestis Touloumis

See Also

nomLORgee and ordLORgee.

Examples

## Illustrating the construction of a "fixed" local odds ratios structure
## using the arthritis dataset. Here, we assume a uniform local odds ratios
## structure equal to 2 for each time pair.

## Create the uniform local odds ratios structure.
lorterm <- matrixLOR(matrix(2, 4, 4))

## Create the LORterm argument.
lorterm <- c(lorterm)
lorterm <- matrix(c(lorterm), 3, 25, TRUE)

## Fit the marginal model.
data(arthritis)
fitmod <- ordLORgee(y ~ factor(trt) + factor(time) + factor(baseline),
  data = arthritis, id = id, repeated = time, LORstr = "fixed",
  LORterm = lorterm)
fitmod

---

**nomLORgee**

*Marginal Models For Correlated Nominal Multinomial Responses*

**Description**

Solving the generalized estimating equations for correlated nominal multinomial responses assuming a baseline category logit model for the marginal probabilities.

**Usage**

nomLORgee(formula = formula(data), data = parent.frame(), id = id,
  repeated = NULL, bstart = NULL, LORstr = "time.exch", LORem = "3way",
  LORterm = NULL, add = 0, homogeneous = TRUE,
  control = LORgee.control(), ipfp.ctrl = ipfp.control(), IM = "solve")

**Arguments**

- **formula**: a formula expression as for other regression models for multinomial responses. An intercept term must be included.
- **data**: an optional data frame containing the variables provided in formula, id and repeated.
- **id**: a vector that identifies the clusters.
- **repeated**: an optional vector that identifies the order of observations within each cluster.
- **bstart**: a vector that includes an initial estimate for the marginal regression parameter vector.
- **LORstr**: a character string that indicates the marginalized local odds ratios structure. Options include "independence", "time.exch", "RC" or "fixed".
- **LORem**: a character string that indicates if the marginalized local odds ratios structure is estimated simultaneously ("3way") or independently at each level pair of repeated ("2way").
- **LORterm**: a matrix that satisfies the user-defined local odds ratios structure. It is ignored unless LORstr="fixed".
- **add**: a positive constant to be added at each cell of the full marginalized contingency table in the presence of zero observed counts.
nomLORgee

homogeneous a logical that indicates homogeneous score parameters when LORstr="time.exch" or "RC".
control a vector that specifies the control variables for the GEE solver.
ipfp.ctrl a vector that specifies the control variables for the function ipfp.
IM a character string that indicates the method used for inverting a matrix. Options include "solve", "qr.solve" or "cholesky".

Details
The data must be provided in case level or equivalently in ‘long’ format. See details about the ‘long’ format in the function reshape.
A term of the form offset(expression) is allowed in the right hand side of formula.
The default set for the response categories is \{1, \ldots, J\}, where \(J > 2\) is the maximum observed response category. If otherwise, the function recodes the observed response categories onto this set.
The \(J\)-th response category is treated as baseline.
The default set for the id labels is \{1, \ldots, N\}, where \(N\) is the sample size. If otherwise, the function recodes the given labels onto this set.
The argument repeated can be ignored only when data is written in such a way that the \(t\)-th observation in each cluster is recorded at the \(t\)-th measurement occasion. If this is not the case, then the user must provide repeated. The suggested set for the levels of repeated is \{1, \ldots, T\}, where \(T\) is the number of observed levels. If otherwise, the function recodes the given levels onto this set.
The variables id and repeated do not need to be pre-sorted. Instead the function reshapes data in an ascending order of id and repeated.
The fitted marginal baseline category logit model is

\[ \log \Pr(Y_{it} = j | x_{it}) = \beta_{0j} + \beta_j' x_{it} \]

where \(Y_{it}\) is the \(t\)-th multinomial response for cluster \(i\), \(x_{it}\) is the associated covariates vector, \(\beta_{0j}\) is the \(j\)-th response category specific intercept and \(\beta_j\) is the \(j\)-th response category specific parameter vector.

The formula is easier to read from either the Vignette or the Reference Manual (both available here).
The LORterm argument must be an \(L \times J^2\) matrix, where \(L\) is the number of level pairs of repeated. These are ordered as \((1, 2), (1, 3), \ldots, (1, T), (2, 3), \ldots, (T - 1, T)\) and the rows of LORterm are supposed to preserve this order. Each row is assumed to contain the vectorized form of a probability table that satisfies the desired local odds ratios structure.

Value
Returns an object of the class "LORgee". This has components:
call the matched call.
title title for the GEE model.
version the current version of the GEE solver.
link the marginal link function.
local.odds.ratios
the marginalized local odds ratios structure variables.

terms
the terms structure describing the marginal model.

contrasts
the contrasts used for the factors.

nobs
the number of observations.

convergence
the values of the convergence variables.

coefficients
the estimated regression parameter vector of the marginal model.

linear.pred
the estimated linear predictor of the marginal regression model. The $j$-th column corresponds to the $j$-th response category.

fitted.values
the estimated fitted values of the marginal regression model. The $j$-th column corresponds to the $j$-th response category.

residuals
the residuals of the marginal regression model based on the binary responses. The $j$-th column corresponds to the $j$-th response category.

y
the multinomial response variables.

id
the id variable.

max.id
the number of clusters.

clusz
the number of observations within each cluster.

robust.variance
the estimated sandwich (robust) covariance matrix.

naive.variance
the estimated model-based (naive) covariance matrix.

xnames
the regression coefficients’ symbolic names.

categories
the number of observed response categories.

occasions
the levels of the repeated variable.

LORgee.control
the control values for the GEE solver.

ipfp.control
the control values for the function ipfp.

inverse.method
the method used for inverting matrices.

adding.constant
the value used for add.

pvalue
the p-value based on a Wald test that no covariates are statistically significant.

Generic coef, summary, print, fitted and residuals methods are available. The pvalue of the Null model corresponds to the hypothesis $H_0 : \beta_1 = \ldots = \beta_{J-1} = 0$ based on the Wald test statistic.

Author(s)
Anestis Touloumis

References
See Also

For an ordinal response scale use the function `ordLORgee`.

Examples

```r
## See the interpretation in Touloumis (2011).
data(housing)
fitmod <- nomLORgee(y ~ factor(time) * sec, data = housing, id = id,
                   repeated = time)
summary(fitmod)
```

Description

Solving the generalized estimating equations for correlated ordinal multinomial responses assuming a cumulative link model or an adjacent categories logit model for the marginal probabilities.

Usage

```r
ordLORgee(formula = formula(data), data = parent.frame(), id = id,
           repeated = NULL, link = "logit", bstart = NULL,
           LORstr = "category.exch", LORem = "3way", LORterm = NULL, add = 0,
           homogeneous = TRUE, restricted = FALSE, control = LORgee.control(),
           ipfp.ctrl = ipfp.control(), IM = "solve")
```

Arguments

- **formula**: a formula expression as for other regression models for multinomial responses. An intercept term must be included.
- **data**: an optional data frame containing the variables provided in `formula`, `id` and `repeated`.
- **id**: a vector that identifies the clusters.
- **repeated**: an optional vector that identifies the order of observations within each cluster.
- **link**: a character string that specifies the link function. Options include "logit", "probit", "cauchit", "cloglog" or "acl".
- **bstart**: a vector that includes an initial estimate for the marginal regression parameter vector.
- **LORstr**: a character string that indicates the marginalized local odds ratios structure. Options include "independence", "uniform", "category.exch", "time.exch", "RC" or "fixed".
- **LORem**: a character string that indicates if the marginalized local odds ratios structure is estimated simultaneously ("3way") or independently at each level pair of `repeated` ("2way").
LORterm

a matrix that satisfies the user-defined local odds ratios structure. It is ignored unless LORstr="fixed".

add

a positive constant to be added at each cell of the full marginalized contingency table in the presence of zero observed counts.

homogeneous

a logical that indicates homogeneous score parameters when LORstr="time.exch" or "RC".

restricted

a logical that indicates monotone score parameters when LORstr="time.exch" or "RC".

control

a vector that specifies the control variables for the GEE solver.

ipfp.ctrl

a vector that specifies the control variables for the function ipfp.

IM

a character string that indicates the method used for inverting a matrix. Options include "solve", "qr.solve" or "cholesky".

Details

The data must be provided in case level or equivalently in ‘long’ format. See details about the ‘long’ format in the function reshape.

A term of the form offset(expression) is allowed in the right hand side of formula.

The default set for the response categories is \(\{1, \ldots, J\}\), where \(J > 2\) is the maximum observed response category. If otherwise, the function recodes the observed response categories onto this set.

The \(J\)-th response category is omitted.

The default set for the id labels is \(\{1, \ldots, N\}\), where \(N\) is the sample size. If otherwise, the function recodes the given labels onto this set.

The argument repeated can be ignored only when data is written in such a way that the \(t\)-th observation in each cluster is recorded at the \(t\)-th measurement occasion. If this is not the case, then the user must provide repeated. The suggested set for the levels of repeated is \(\{1, \ldots, T\}\), where \(T\) is the number of observed levels. If otherwise, the function recodes the given levels onto this set.

The variables id and repeated do not need to be pre-sorted. Instead the function reshapes data in an ascending order of id and repeated.

The fitted marginal cumulative link model is

\[
Pr(Y_{it} \leq j|x_{it}) = F(\beta_{j0} + \beta' x_{it})
\]

where \(Y_{it}\) is the \(t\)-th multinomial response for cluster \(i\), \(x_{it}\) is the associated covariates vector, \(F\) is the cumulative distribution function determined by link, \(\beta_{j0}\) is the \(j\)-th response category specific intercept and \(\beta\) is the marginal regression parameter vector excluding intercepts.

The marginal adjacent categories logit model

\[
\log \frac{Pr(Y_{it} = j|x_{it})}{Pr(Y_{it} = j+1|x_{it})} = \beta_{j0} + \beta' x_{it}
\]

is fitted if and only if link="acl". In contrast to a marginal cumulative link model, here the intercepts do not need to be monotone increasing.

The formulae are easier to read from either the Vignette or the Reference Manual (both available here).
The LORterm argument must be an $L \times J^2$ matrix, where $L$ is the number of level pairs of repeated. These are ordered as $(1, 2), (1, 3), \ldots, (1, T), (2, 3), \ldots, (T - 1, T)$ and the rows of LORterm are supposed to preserve this order. Each row is assumed to contain the vectorized form of a probability table that satisfies the desired local odds ratios structure.

**Value**

Returns an object of the class "LORgee". This has components:

- **call**: the matched call.
- **title**: title for the GEE model.
- **version**: the current version of the GEE solver.
- **link**: the marginal link function.
- **local.odds.ratios**: the marginalized local odds ratios structure variables.
- **terms**: the terms structure describing the model.
- **contrasts**: the contrasts used for the factors.
- **nobs**: the number of observations.
- **convergence**: the values of the convergence variables.
- **coefficients**: the estimated regression parameter vector of the marginal model.
- **linear.pred**: the estimated linear predictor of the marginal regression model. The $j$-th column corresponds to the $j$-th response category.
- **fitted.values**: the estimated fitted values of the marginal regression model. The $j$-th column corresponds to the $j$-th response category.
- **residuals**: the residuals of the marginal regression model. The $j$-th column corresponds to the $j$-th response category.
- **y**: the multinomial response variables.
- **id**: the id variable.
- **max.id**: the number of clusters.
- **clusz**: the number of observations within each cluster.
- **robust.variance**: the estimated sandwich (robust) covariance matrix.
- **naive.variance**: the estimated model-based (naive) covariance matrix.
- **xnames**: the regression coefficients’ symbolic names.
- **categories**: the number of observed response categories.
- **occasions**: the levels of the repeated variable.
- **LORgee.control**: the control values for the GEE solver.
- **ipfp.control**: the control values for the function ipfp.
- **inverse.method**: the method used for inverting matrices.
- **adding.constant**: the value used for add.
- **pvalue**: the p-value based on a Wald test that no covariates are statistically significant.

Generic coef, summary, print, fitted and residuals methods are available. The pvalue of the Null model corresponds to the hypothesis $H_0 : \beta = 0$ based on the Wald test statistic.
Author(s)

Anestis Touloumis

References


See Also

For a nominal response scale use the function *nomLORgee*.

Examples

```r
data(arthritis)
intrinsic.pars(y, arthritis, id, time)
fitmod <- ordLORgee(formula = y ~ factor(time) + factor(trt) + factor(baseline),
data = arthritis, id = id, repeated = time, LORstr = "uniform")
summary(fitmod)
```

### vcov.LORgee

*Calculate Variance-Covariance Matrix for a Fitted LORgee Object.*

Description

Returns the variance-covariance matrix of the main parameters of a fitted model LORgee object.

Usage

```r
## S3 method for class 'LORgee'
vcov(object, method = "robust", ...)
```

Arguments

- **object**: a fitted model LORgee object.
- **method**: character indicating whether the sandwich (robust) covariance matrix (method = "robust") or the model–based (naive) covariance matrix (method = "naive") should be returned.
- **...**: additional argument(s) for methods.

Details

Default is to obtain the estimated sandwich (robust) covariance matrix and method = "naive" obtains the estimated model-based (naive) covariance matrix.
Value

A matrix of the estimated covariances between the parameter estimates in the linear predictor of the GEE model. This should have row and column names corresponding to the parameter names given by the \texttt{coef} method.

Examples

\begin{verbatim}
fitmod <- ordLORgee(formula = y ~ factor(time) + factor(trt) + factor(baseline),
  data = arthritis, id = id, repeated = time, LORstr = "uniform")
vcov(fitmod, method = "robust")
vcov(fitmod, method = "naive")
\end{verbatim}

---

\textbf{waldts} \hfill \textit{Wald Test of Nested GEE Models}

\section*{Description}

Comparing two nested GEE models by carrying out a Wald test.

\section*{Usage}

\begin{verbatim}
waldts(object0, object1)
\end{verbatim}

\section*{Arguments}

- \texttt{object0} \hfill A GEE model of the class "LORgee".
- \texttt{object1} \hfill A GEE model of the class "LORgee".

\section*{Details}

The two GEE models implied by \texttt{object0} and \texttt{object1} must be nested.

\section*{Author(s)}

Anestis Touloumis

\section*{Examples}

\begin{verbatim}
data(housing)
set.seed(1)
fitmod1 <- nomLORgee(y ~ factor(time) * sec, data = housing, id = id,
  repeated = time)
set.seed(1)
fitmod0 <- update(fitmod1, formula = y ~ factor(time) + sec)
waldts(fitmod0, fitmod1)
\end{verbatim}
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