

Semiparametrically Efficient Population and Permutation Inference in Distribution-free Stratified K -sample Oneway Layouts¹

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Contents

| | | |
|-----------|---|------------|
| 1 | Model and Parameterisation | 1 |
| 2 | Parameter Estimation | 3 |
| 3 | Link Functions | 16 |
| 4 | Optimisation | 22 |
| 5 | ML Estimation | 25 |
| 6 | ML Inference | 37 |
| 6.1 | Wald Statistics | 37 |
| 6.2 | Likelihood-ratio Statistics | 37 |
| 6.3 | Rao Score Statistics | 38 |
| 7 | Permutation Inference | 39 |
| 8 | Distribution-free Tests in Stratified K-sample Oneway Layouts | 46 |
| 8.1 | <code>free1way</code> | 46 |
| 8.2 | <code>free1way</code> Methods | 53 |
| 9 | Special Test Procedures | 64 |
| 9.1 | Wilcoxon Test | 64 |
| 9.2 | Mantel-Haenszel Test | 67 |
| 9.3 | <code>prop.test</code> | 68 |
| 9.4 | Kruskal-Wallis Test | 68 |
| 9.5 | Savage Test | 69 |
| 9.6 | Log-rank Test | 72 |
| 9.7 | van der Waerden Test | 76 |
| 9.8 | Friedman Test | 77 |
| 9.9 | McNemar Test | 79 |
| 9.10 | Incomplete Block Designs | 80 |
| 9.11 | Contrast Tests | 80 |
| 10 | Model Diagnostics | 82 |
| 10.1 | Transformation Plots | 82 |
| 10.2 | Probability-probability Plots | 85 |
| 11 | Random Number Generation | 92 |
| 12 | Power and Sample Size | 97 |
| 13 | Penalisation | 107 |
| 14 | Acknowledgements | 109 |

Abstract

Starting with R 4.6-0, the **stats** package provides infrastructure for distribution-free model-based inference in possibly stratified K -sample oneway layouts via the novel **free1way** model function. Treatment effects to be estimated using **free1way** include odds- and hazard ratios, Lehmann parameters, and a generalised version of Cohen's d for at least ordered and possibly right-censored outcomes.

In addition to nonparametric maximum-likelihood estimators of treatment effects, the procedure allows Wald, Rao score, and likelihood ratio tests with corresponding confidence intervals to be computed. Asymptotic and approximate Monte-Carlo-based permutation tests and confidence intervals are also available. In proportional odds models, exact permutation inference is implemented based on exact permutation distributions derived via the Streitberg-Röhm algorithm.

Graphical tools for model diagnostics, including model-based confidence bands for receiver operating characteristic (ROC) curves in probability-probability plots in the new **ppplot** function, allow data-driven criticism of model assumptions.

Power assessment and sample-size planning is facilitated either in a simulation-based way relying on random number generation via **rfree1way** or based on approximations of the information matrix in **power.free1way.test**, the latter approach being much faster but slightly less accurate.

The new **free1way** function can be understood as a unification and generalisation of some of the classical “nonparametric” test procedures in **stats**, including **kruskal.test**, **wilcox.test**, **friedman.test**, **mantelhaen.test**, **prop.test**, **mcnemar.test**, as well as **power.prop.test**, allowing the magnitude of treatment effects to be interpreted on various scales, providing the possibility to assessment variability by means of confidence intervals and corresponding tests for these parameters, and offering tools for sample-size planning and model criticism.

This document explains the technical underpinnings of the implementation. The **free1way** package provides this vignette as additional documentation and serves as a home for extensive regression tests.

Introduction

Comparing two or more independent samples with respect to some outcome measure is a common task. Many procedures are available in **stats** and other add-on packages, most of these implementations making rather strict assumptions regarding the outcome distribution, the number of samples, the presence of blocks or strata and typically offer either conditional or unconditional (exact or asymptotic) inference.

This document presents a unified, dense, and yet holistic implementation covering many classical procedures as special cases. Leveraging transformation models, likelihood-based parameter estimation as well as permutation- and likelihood-based inference are formulated and implemented. One can understand this contribution as a unification of many of **stats::*.test** procedures, the models available in **MASS::polr**, **rms::orm**, **rms::lm**, **survival::coxph**, or the **tram** add-on package (among many others), and permutation-based inference in **coin**.

This implementation is, however, free of any strong dependencies and only uses functionality available in R itself and the **stats**, **graphics**, and **Matrix** recommended packages.

Chapter 1

Model and Parameterisation

We consider K treatment groups $G \in \{1, \dots, K\}$, $K \geq 2$ for an at least ordered outcome $Y \in \mathcal{Y}$ observed in stratum $S \in \{1, \dots, B\}$ out of $B \geq 1$ blocks with conditional cumulative distribution function (cdf) $F_Y(y \mid G = k, S = b) = \mathbb{P}(Y \leq y \mid G = k, S = b)$. Detecting and describing differential distributions arising from different treatments is our main objective. We refer to the first treatment $G = 1$ as “control”.

Model With model function $m : [0, 1] \times \mathbb{R} \rightarrow [0, 1]$, we describe the conditional distribution under treatment k as a function of the conditional distribution under control and a scalar parameter δ_k :

$$F_Y(y \mid G = k, S = b) = m(F_Y(y \mid G = 1, S = b), \delta_k).$$

The model is assumed to hold for all blocks $b = 1, \dots, B$, treatments $k = 2, \dots, K$, and outcome values $y \in \mathcal{Y}$ based on parameters $\delta_2, \dots, \delta_K \in \mathbb{R}$. For notational convenience, we define $\delta_1 := 0$.

This model formulation gives rise to several specific models, for example, $m_L(p, \delta) = p^{\exp(-\delta)}$ (Lehmann alternatives), $m_{PH}(p, \delta) = 1 - (1 - p)^{\exp(-\delta)}$ (proportional hazards), $m_{PO}(p, \delta) = \text{expit}(\text{logit}(p) - \delta)$ (proportional odds), or $m_{Cd}(p, \delta) = \Phi(\Phi^{-1}(p) - \delta)$ (generalised Cohen’s d).

Instead of directly working with g , we parameterise the model in terms of some absolute continuous cdf F with log-concave density $f = F'$ and corresponding derivative f' . The location model

$$F_Y(y \mid G = k, S = b) = F(F^{-1}(F_Y(y \mid G = 1, S = b)) - \delta_k), \quad k = 2, \dots, K \quad (1.1)$$

describes different distributions by means of shift parameter on a latent scale defined by F . The negative shift term ensures that positive values of δ_k correspond to the situation of outcomes being stochastically larger in group k compared to control. The shift parameters are invariant with respect to monotone transformations of the response values, that is, transforming the observations of all treatment groups by the same function does not affect the values of δ_k .

The choice $F(z) = \exp(-\exp(-z))$ gives rise to m_L , $F(z) = 1 - \exp(-\exp(z))$ corresponds to m_{PH} , $F = \text{expit}$ leads to m_{PO} , and $F = \Phi$ results in m_{Cd} . The choice of F is made a priori and determines the interpretation of δ_k .

This document describes the implementation of estimators of these shift parameters, as well as of confidence intervals and formal hypothesis tests for contrasts thereof under the permutation and population model. Proportional odds models (m_{PO}) are explained in-depths by [Harrell \(2015\)](#), although the models are presented in terms of survivor, not distribution, functions.

Hypothesis We are interested in inference for $\delta_2, \dots, \delta_K$, in terms of confidence intervals and hypothesis tests of the form

$$\begin{aligned} H_0 : \delta_k - \mu_k &= 0, \text{ “two.sided”, } & k = 2, \dots, K, \\ H_0 : \delta_k - \mu_k &\geq 0, \text{ “less”, } & k = K = 2, \\ H_0 : \delta_k - \mu_k &\leq 0, \text{ “greater”, } & k = K = 2, \end{aligned}$$

with the latter two options only for the two-sample case ($K = 2$).

Likelihood For an ordered categorical outcome Y from sample space $\mathcal{Y} = \{v_1 < v_2 < \dots < v_C\}$, we parameterise the model in terms of intercept (ϑ .) and shift (δ .) parameters

$$F_Y(v_c | G = k, S = b) = F(\vartheta_{c,b} - \delta_k), \quad c = 1, \dots, C,$$

that is we replace the transformed control outcome $F^{-1}(F_Y(v_c | G = 1, S = b)) = \vartheta_{c,b}$ with a corresponding intercept parameter. These $C - 1$ intercept parameters are block-specific and monotone increasing $\vartheta_{0,b} = -\infty < \vartheta_{1,b} < \dots < \vartheta_{C,b} = \infty$ within each block $b = 1, \dots, B$.

We collect all model parameters in a vector

$$\begin{aligned} \boldsymbol{\theta} = (\theta_1 &:= \delta_2, \\ &\dots, \\ \theta_{K-1} &:= \delta_K, \\ \theta_K &:= \vartheta_{1,1}, \\ \theta_{K+1} &:= \vartheta_{2,1} > \vartheta_{1,1}, \\ &\dots, \\ \theta_{K+C-2} &:= \vartheta_{C-1,1} > \vartheta_{C-2,1}, \\ \theta_{K+C-1} &:= \vartheta_{1,2}, \\ &\dots, \\ \theta_{B(C-1)+K-1} &:= \vartheta_{C-1,B} > \vartheta_{C-2,B}). \end{aligned}$$

If there is no observation for level c in block b , the corresponding parameter is not identified and removed from $\boldsymbol{\theta}$. The parameter space is defined by all parameter vectors $\boldsymbol{\theta}$ satisfying the monotonicity of the intercept parameters. Violations always lead to the log-likelihood function being undefined and thus taking the value $-\infty$. Harrell (2024) evaluates unconstrained optimisation in this context and recommends Newton-based algorithms leveraging the analytically available Hessian (see below).

For the i th observation ($y_i = v_c, g_i = k, s_i = b$) from block b under treatment k , the log-likelihood contribution is

$$\log(\mathbb{P}(v_{c-1} < Y \leq v_c | G = k, S = b)) = \log(F(\vartheta_{c,b} - \delta_k) - F(\vartheta_{c-1,b} - \delta_k))$$

with $v_0 = -\infty$.

For an absolutely continuous outcome $Y \in \mathbb{R}$, we define $v_c := y_{(c)}$, the c th distinct ordered observation in the sample. The log-likelihood above is then the empirical or nonparametric log-likelihood.

If observations were independently right-censored, the contribution of the event $Y > \tilde{y}$ to the log-likelihood is

$$\log(\mathbb{P}(Y > \tilde{y} | G = k, S = b)) = \log(1 - F(\vartheta_{c-1,b} - \delta_k))$$

where $v_{c-1} = \max\{v \in \mathcal{Y} | v \leq \tilde{y}\}$, that is, observations right-censored between v_{c-1} and v_c correspond to the parameter $\vartheta_{c-1,b}$.

Maximising this form of the log-likelihood leads to semiparametrically efficient estimators (Chapter 15.5 Van der Vaart, 1998). In this framework, tests against deviations from the hypothesis H_0 above are locally most powerful rank tests, for example against proportional odds ($F = \text{expit}$) or proportional hazards alternatives ($F(z) = 1 - \exp(-\exp(z))$, Van der Vaart, 1998, Example 15.16).

We represent the data in form of a $C \times K \times B$ contingency table, whose element (c, k, b) is the number of observations with configuration $(y = y_c, g = k, s = b)$. In the presence of right-censoring, a fourth dimension is added ($C \times K \times B \times 2$) whose first $C \times K \times B$ table presents right-censoring and the second table contains numbers of events.

Chapter 2

Parameter Estimation

We start implementing the log-likelihood function for parameters $\text{parm} = \boldsymbol{\theta}$ (assuming only a single block) with data from a two-way $C \times K$ contingency table \mathbf{x} .

From $\boldsymbol{\theta}$, we first extract the shift parameters $\delta_k, k = 2, \dots, K$ and then the intercept parameters $\vartheta_{c,1}, c = 1, \dots, C-1$ and evaluate the probabilities $\text{prb} = \mathbb{P}(y_{c-1} < Y \leq y_c \mid G = k, S = 1)$ for all groups:

$\langle \text{parm to prob} \rangle \equiv$

```
bidx <- seq_len(ncol(x) - 1L)
delta <- c(0, mu + parm[bidx])
intercepts <- c(-Inf, parm[- bidx], Inf)
tmb <- intercepts - matrix(delta, nrow = length(intercepts),
                           ncol = ncol(x),
                           byrow = TRUE)

Ftmb <- F(tmb)
if (rightcensored) {
  prb <- 1 - Ftmb[- nrow(Ftmb), , drop = FALSE]
} else {
  prb <- Ftmb[- 1L, , drop = FALSE] -
    Ftmb[- nrow(Ftmb), , drop = FALSE]
}
◇
```

Fragment referenced in [?, ?, ?, ?](#).

If the table \mathbf{x} represents right-censored observations, we compute $\text{prb} = 1 - \mathbb{P}(Y \leq y_c \mid G = k, S = 1)$.

With default null values $\mu_k = 0, k = 2, \dots, K$, we define the negative log-likelihood function as the weighted (by number of observations) sum of the log-probabilities

$\langle \text{negative logLik} \rangle \equiv$

```
.nll <- function(parm, x, mu = 0, rightcensored = FALSE)
{
   $\langle \text{parm to prob} \rangle$ 
  if (any(prb < .Machine$double.eps^10))
    return(Inf)
  return(- sum(x * log(prb)))
}
◇
```

Fragment referenced in [?](#).

The code assumes that all elements of the margins of the table \mathbf{x} are larger than zero; otherwise, the corresponding parameter is not identified. We will handle such situation at a higher level later on.

It is important to note that, with F corresponding to distribution with log-concave density f , the negative log-likelihood is a convex function of the parameters θ , and thus we can solve the corresponding constrained minimisation problem quickly and reliably.

Next, we implement the gradient of the negative log-likelihood, the negative score function for the parameters in θ . The score function for the empirical likelihood, evaluated at parameters is given in many places (for example in [Hothorn et al., 2018](#), Formula (2)). We begin computing the ratio of $f(\vartheta_{c,1} - \delta_k)$ and the corresponding likelihood

⟨ density prob ratio ? ⟩ \equiv

```
ftmb <- f(tmb)
zu <- x * ftmb[- 1, , drop = FALSE] / prb
if (rightcensored) zu[] <- 0 ### derivative of a constant
zl <- x * ftmb[- nrow(ftmb), , drop = FALSE] / prb
◇
```

Fragment referenced in [?, ?](#).

and then compute the negative score function:

⟨ negative score ? ⟩ \equiv

```
.nsc <- function(parm, x, mu = 0, rightcensored = FALSE)
{
  ⟨ parm to prob ? ⟩

  ⟨ density prob ratio ? ⟩

  ret <- numeric(length(parm))
  ret[bidx] <- .colSums(zl, m = nrow(zl), n = ncol(zl))[-1L] -
    .colSums(zu[-nrow(zu),,drop = FALSE],
      m = nrow(zu) - 1L, n = ncol(zu))[-1L]
  ret[- bidx] <- .rowSums(zu[-nrow(zu),,drop = FALSE] -
    zl[-1,,drop = FALSE],
    m = nrow(zu) - 1L, n = ncol(zu))

  return(- ret)
}
◇
```

Fragment referenced in [?](#).

In addition, we define negative score residuals, that is, the derivative of the negative log-likelihood with respect to an intercept term constrained to zero:

⟨ negative score residuals ? ⟩ \equiv

```
.nsr <- function(parm, x, mu = 0, rightcensored = FALSE)
{
  ⟨ parm to prob ? ⟩

  ⟨ density prob ratio ? ⟩

  ret <- .rowSums(zl - zu, m = nrow(zl), n = ncol(zl)) /
    .rowSums(x, m = nrow(x), n = ncol(x))
  ret[!is.finite(ret)] <- 0
  return(- ret)
}
◇
```

Fragment referenced in [?](#).

We also need access to the observed Fisher information of the shift parameters. We proceed by implementing the Hessian for the intercept (ϑ .) and shift (δ .) parameters, as given in Formula (4) of [Hothorn et al. \(2018\)](#) first. This partitioned matrix

$$\mathbf{H}(\vartheta_1, \dots, \vartheta_{C-1}, \delta_2, \dots, \delta_K) = \begin{pmatrix} \mathbf{A} & \mathbf{X} \\ \mathbf{X}^\top & \mathbf{Z} \end{pmatrix}$$

consists of a symmetric tridiagonal $\mathbf{A} \sim (C-1, C-1)$, a diagonal $\mathbf{Z} \sim (K-1, K-1)$, and a full $\mathbf{X} \sim (C-1, K-1)$ matrix. In a second step, we compute the Fisher information matrix for the shift parameters only by means of the Schur complement $\mathbf{Z} - \mathbf{X}^\top \mathbf{A}^{-1} \mathbf{X}$.

In addition to probabilities `prb`, the Hessian necessitates the computation of $f(\vartheta_{c,1} - \delta_k)$ and $f'(\vartheta_{c,1} - \delta_k)$. We start preparing these objects, keeping in mind to remove terms not being present under right-censoring:

$\langle \text{Hessian prep ?} \rangle \equiv$

```
ftmb <- f(tmb)
fptmb <- fp(tmb)

dl <- ftmb[- nrow(ftmb), , drop = FALSE]
du <- ftmb[- 1, , drop = FALSE]
if (rightcensored) du[] <- 0
dpl <- fptmb[- nrow(ftmb), , drop = FALSE]
dpu <- fptmb[- 1, , drop = FALSE]
if (rightcensored) dpu[] <- 0
dlm1 <- dl[, -1L, drop = FALSE]
dum1 <- du[, -1L, drop = FALSE]
dplm1 <- dpl[, -1L, drop = FALSE]
dpum1 <- dpu[, -1L, drop = FALSE]
prbm1 <- prb[, -1L, drop = FALSE]

i1 <- length(intercepts) - 1L
i2 <- 1L
◇
```

Fragment referenced in [?](#).

The off-diagonal elements of \mathbf{A} are now available as

$\langle \text{off-diagonal elements for Hessian of intercepts ?} \rangle \equiv$

```
Aoffdiag <- - .rowSums(x * du * dl / prb^2, m = nrow(x), n = ncol(x))[-i2]
Aoffdiag <- Aoffdiag[-length(Aoffdiag)]
◇
```

Fragment referenced in [?](#).

and the diagonal elements of \mathbf{A} as

$\langle \text{diagonal elements for Hessian of intercepts ?} \rangle \equiv$

```
Adiag <- - .rowSums((x * dpu / prb)[-i1, , drop = FALSE] -
  (x * dpl / prb)[-i2, , drop = FALSE] -
  ((x * du^2 / prb^2)[-i1, , drop = FALSE] +
  (x * dl^2 / prb^2)[-i2, , drop = FALSE] ),
  m = nrow(x) - length(i1), n = ncol(x)
)
◇
```

Fragment referenced in [?](#).

For the computation of \mathbf{X} and \mathbf{Z} , the observations corresponding to the control group ($k = 1$) are irrelevant, we remove these first

$\langle \text{intercept} / \text{shift contributions to Hessian} \rangle \equiv$

```

xm1 <- x[,-1L,drop = FALSE]
X <- ((xm1 * dpum1 / prbm1)[-i1,,drop = FALSE] -
      (xm1 * dplm1 / prbm1)[-i2,,drop = FALSE] -
      ((xm1 * dum1^2 / prbm1^2)[-i1,,drop = FALSE] -
       (xm1 * dum1 * dlm1 / prbm1^2)[-i2,,drop = FALSE] -
       (xm1 * dum1 * dlm1 / prbm1^2)[-i1,,drop = FALSE] +
       (xm1 * dlm1^2 / prbm1^2)[-i2,,drop = FALSE]
      )
      )

Z <- - .colSums(xm1 * (dpum1 / prbm1 -
                      dplm1 / prbm1 -
                      (dum1^2 / prbm1^2 -
                       2 * dum1 * dlm1 / prbm1^2 +
                       dlm1^2 / prbm1^2
                      )
                      ),
      m = nrow(xm1), n = ncol(xm1)
      )
if (length(Z) > 1L) Z <- diag(Z)

```

Fragment referenced in [?](#).

We return the three matrices \mathbf{A} , \mathbf{X} , and \mathbf{Z} necessary for two different purposes: We need the `full` Hessian for all parameters $\boldsymbol{\theta}$ as a dense `matrix` such that `nlminb` can compute updates from this object. In addition, the computation of the Fisher information for $\delta_2, \dots, \delta_K$ as the Schur complement $\mathbf{Z} - \mathbf{X}^\top \mathbf{A}^{-1} \mathbf{X}$. Because the matrix \mathbf{A} is symmetric tridiagonal, we use infrastructure from the `Matrix` package to represent this matrix:

$\langle \text{Hessian ?} \rangle \equiv$

```
.hes <- function(parm, x, mu = 0, rightcensored = FALSE, full = FALSE)
{
  < parm to prob ? >

  < Hessian prep ? >

  < off-diagonal elements for Hessian of intercepts ? >
  < diagonal elements for Hessian of intercepts ? >
  < intercept / shift contributions to Hessian ? >

  if (length(Adiag) > 1L) {
    if (!isFALSE(full)) {
      A <- list(Adiag = Adiag, Aoffdiag = Aoffdiag)
    } else {
      A <- Matrix::bandSparse(length(Adiag),
        k = 0:1, diagonals = list(Adiag, Aoffdiag),
        symmetric = TRUE)
    }
  } else {
    if (!isFALSE(full)) {
      A <- list(Adiag = Adiag, Aoffdiag = NULL)
    } else {
      A <- matrix(Adiag)
    }
  }
  return(list(A = A, X = X, Z = Z))
}
◇
```

Fragment referenced in [?](#).

We start with an example involving $K = 3$ groups for a binary outcome and use a binary logistic regression model to estimate the two log-odds ratios δ_2 and δ_3 along with their estimated covariance

```
> library("freelway.docreg")
> (x <- matrix(c(10, 5, 7, 11, 8, 9), nrow = 2))

      [,1] [,2] [,3]
[1,]   10    7    8
[2,]    5   11    9

> d <- expand.grid(y = relevel(g1(2, 1), "2"), t = g1(3, 1))
> d$x <- c(x)
> m <- glm(y ~ t, data = d, weights = x, family = binomial())
> (cf <- coef(m))

(Intercept)          t2          t3
    0.69315    -1.14513    -0.81093
```

Replicating these results requires specification of the inverse link function $F = \text{expit}$ and the density function f of the standard logistic. We use `optim` with numerically approximated Hessian to be able to check the correctness of the analytical Hessian. Note that `glm` operates with a positive linear predictor, so we need to change the sign of the log-odds ratios:

```
> F <- plogis
> f <- dlogis
> op <- optim(par = c("mt2" = 0, "mt3" = 0, "(Intercept)" = 0),
+           fn = .nll, gr = .nsc,
+           x = x, method = "BFGS", hessian = TRUE)
> cbind(glm = c(cf[-1] * -1, cf[1]), freelway = op$par)
```

```

          glm freeway
t2          1.14513  1.14513
t3          0.81093  0.81093
(Intercept) 0.69315  0.69315

```

```
> logLik(m)
```

```
'log Lik.' -33.33 (df=3)
```

```
> -op$value
```

```
[1] -33.33
```

Parameter estimates and the in-sample log-likelihood are practically identical. We now turn to the inverse Hessian of the shift terms, first defining the derivative of the density of the standard logistic distribution

```

> fp <- function(x)
+ {
+   p <- plogis(x)
+   p * (1 - p)^2 - p^2 * (1 - p)
+ }
> H <- .hes(op$par, x)
> ### analytical covariance of parameters
> solve(H$Z - crossprod(H$X, Matrix::solve(H$A, H$X)))

```

```

      [,1]      [,2]
[1,] 0.53377 0.30000
[2,] 0.30000 0.53611

```

```

> ### numerical covariance
> solve(op$hessian)[1:2,1:2]

```

```

      mt2      mt3
mt2 0.53377 0.30000
mt3 0.30000 0.53611

```

```

> ### from glm
> vcov(m)[-1,-1]

```

```

      t2      t3
t2 0.53377 0.30000
t3 0.30000 0.53611

```

Also here we see practically identical results. We will later implement a low-level function `.freewayML` taking a table and an object describing the inverse link F as arguments; these results are also in line with `glm`:

```

> obj <- .freewayML(as.table(x), link = logit())
> obj$coefficients

```

```

      B      C
1.14513 0.81093

```

```
> -obj$value
```

```
[1] -33.33
```

```

> ### analytical covariance
> obj$vcov

```

```

      B      C
B 0.53377 0.30000
C 0.30000 0.53611

```

In the next step, we extend our results to the stratified case. We iterate over all blocks and evaluate the negative log-likelihood for the same values of the shift parameters but block-specific values of the intercept parameters. Because `x` is a `table`, it may happen (especially in the presence of blocks) that some outcome values (rows) were not observed in any group (row sum is zero). Thus, the distribution function does not jump at this value and therefore no parameter for this value is needed. We remove these cases. In the right-censored case, we have to pay attention to censoring happening at these outcome values. We count how many observations were censored between observations and assign the corresponding weights to the subsequent outcome value. Furthermore, we need to be able to undo the removal of observations later, mainly for computing residuals. We store an attribute `idx` for later use in `.snsr` on page 11.

(determine steps in blocks?) \equiv

```
xlist <- xrclist <- vector(mode = "list", length = B)

for (b in seq_len(B)) {
  xb <- matrix(x[,b, drop = TRUE], ncol = K)
  xw <- rowSums(abs(xb)) > 0
  if (sum(xw) > 1L) {
    ### do not remove last parameter if there are corresponding
    ### right-censored observations
    wm <- which(xw)[sum(xw)]
    if (!is.null(xrc) && any(xrc[wm:dx[1],b,drop = TRUE] > 0))
      xw[length(xw)] <- TRUE
    xlist[[b]] <- xb[xw,,drop = FALSE]
    Cidx <- rep.int(1L, times = C)
    Cidx[xw] <- Cidx[xw] + seq_len(sum(xw))
    attr(xlist[[b]], "idx") <- Cidx
    if (!is.null(xrc)) {
      ### count right-censored observations between distinct event
      ### times
      cs <- apply(xrc[,b,drop = TRUE] * (!xw), 2, function(x)
        diff(c(0, cumsum(x)[xw])))
      xrclist[[b]] <- matrix(xrc[xw,,b,drop = TRUE], ncol = K) + cs
      idx <- seq_len(C)[xw]
      idx <- rep(seq_len(sum(xw)), times = c(idx[1], diff(idx)))
      Cidx <- rep.int(1L, times = C)
      Cidx[seq_along(idx)] <- Cidx[seq_along(idx)] + idx
      attr(xrclist[[b]], "idx") <- Cidx
    }
  }
}

### remove empty blocks
strata <- !vapply(xlist, is.null, NA)
xlist <- xlist[strata]
xrclist <- xrclist[strata]
◇
```

Fragment referenced in ?.

Before we begin, we convert the table $C \times K \times B(\times 2)$ table `x` into a list of non-empty $C' \times K$ tables (yet still allowing zero row sums):

$\langle \text{table2list body ?} \rangle \equiv$

```

dx <- dim(x)
if (length(dx) == 1L)
  stop("incorrect dimensions")
if (length(dx) == 2L)
  x <- as.table(array(x, dim = c(dx, 1)))
dx <- dim(x)
if (length(dx) < 3L)
  stop("incorrect dimensions")
C <- dim(x)[1L]
K <- dim(x)[2L]
B <- dim(x)[3L]
if (C < 2L)
  stop("at least two response categories required")
if (K < 2L)
  stop("at least two groups required")
xrc <- NULL
if (length(dx) == 4L) {
  if (dx[4] == 2L) {
    xrc <- array(x[,,"FALSE", drop = TRUE], dim = dx[1:3])
    x <- array(x[,,"TRUE", drop = TRUE], dim = dx[1:3])
  } else {
    stop(gettextf("%s currently only allows independent right-censoring",
                  "freelway"),
         domain = NA)
  }
}

```

$\langle \text{determine steps in blocks ?} \rangle$

◇

Fragment referenced in [?](#).

We first extract the shift parameters δ . and then, separately for each stratum, the corresponding contrasts of the intercept parameters:

$\langle \text{stratum prep ?} \rangle \equiv$

```

C <- vapply(x, NROW, 0L) ### might differ by stratum
K <- unique(do.call("c", lapply(x, ncol))) ### the same
B <- length(x)
sidx <- factor(rep(seq_len(B), times = pmax(0, C - 1L)),
              levels = seq_len(B))
bidx <- seq_len(K - 1L)
delta <- parm[bidx]
intercepts <- split(parm[-bidx], sidx)

```

◇

Fragment referenced in [?](#), [?](#), [?](#), [?](#), [?](#).

before we loop over the non-empty strata and return the sum of the corresponding log-likelihoods:

$\langle \text{stratified negative logLik ?} \rangle \equiv$

```
.snll <- function(parm, x, mu = 0, rightcensored = FALSE)
{
   $\langle \text{stratum prep ?} \rangle$ 

  ret <- 0
  for (b in seq_len(B))
    ret <- ret + .nll(c(delta, intercepts[[b]]), x[[b]], mu = mu,
                     rightcensored = rightcensored)
  return(ret)
}
◇
```

Fragment referenced in [?](#).

In a similar way, we evaluate the gradients for each block and sum-up the contributions by the shift parameters whereas the gradients for the intercept parameters are only concatenated:

$\langle \text{stratified negative score ?} \rangle \equiv$

```
.snsc <- function(parm, x, mu = 0, rightcensored = FALSE)
{
   $\langle \text{stratum prep ?} \rangle$ 

  ret <- numeric(length(bidx))
  for (b in seq_len(B)) {
    nsc <- .nsc(c(delta, intercepts[[b]]), x[[b]], mu = mu,
               rightcensored = rightcensored)
    ret[bidx] <- ret[bidx] + nsc[bidx]
    ret <- c(ret, nsc[-bidx])
  }
  return(ret)
}
◇
```

Fragment referenced in [?](#).

The score residuum is zero for an observation with weight zero, that is, a row of zeros in the table:

$\langle \text{stratified negative score residual ?} \rangle \equiv$

```
.snsr <- function(parm, x, mu = 0, rightcensored = FALSE)
{
   $\langle \text{stratum prep ?} \rangle$ 

  ret <- c()
  for (b in seq_len(B)) {
    idx <- attr(x[[b]], "idx")
    ### idx == 1L means zero residual, see definition of idx
    sr <- c(0, .nsr(c(delta, intercepts[[b]]), x[[b]], mu = mu,
                   rightcensored = rightcensored))
    ret <- c(ret, sr[idx])
  }
  return(ret)
}
◇
```

Fragment referenced in [?](#).

```

> (x <- as.table(array(c(10, 5, 7, 11, 8, 9,
+                       9, 4, 8, 15, 5, 4), dim = c(2, 3, 2))))
, , A
      A  B  C
A 10   7   8
B  5  11   9

, , B
      A  B  C
A  9   8   5
B  4  15   4

> d <- expand.grid(y = relevel(gl(2, 1), "2"), t = gl(3, 1), s = gl(2, 1))
> d$x <- c(x)
> m <- glm(y ~ 0 + s + t, data = d, weights = x, family = binomial())
> logLik(m)

'log Lik.' -62.523 (df=4)

> (cf <- coef(m))

      s1      s2      t2      t3
0.72433 0.77385 -1.30224 -0.74147

> x1 <- .table2list(x)$xlist
> op <- optim(par = c("mt2" = 0, "mt3" = 0, "(Intercept 1)" = 0, "(Intercept 2)" = 0),
+          fn = .snll, gr = .snsc,
+          x = x1,
+          method = "BFGS",
+          hessian = TRUE)
> cbind(glm = c(cf[-(1:2)] * -1, cf[1:2]), free1way = op$par)

      glm free1way
t2 1.30224  1.30225
t3 0.74147  0.74149
s1 0.72433  0.72435
s2 0.77385  0.77386

> logLik(m)

'log Lik.' -62.523 (df=4)

> -op$value

[1] -62.523

```

For the analytical Hessian, we sum-up over the stratum-specific Hessians of the shift parameters. For right-censored observations, we need to compute the contributions by the events and obtain the joint Hessian for shift- and intercept parameters first. We differentiate between computing the null Hessian for θ as a dense matrix:

$\langle \text{full Hessian ?} \rangle \equiv$

```

for (b in seq_len(B)) {
  H <- .hes(c(delta, intercepts[[b]]), x[[b]], mu = mu, full = full)
  if (!is.null(xrc)) {
    Hrc <- .hes(c(delta, intercepts[[b]]), xrc[[b]], mu = mu,
      rightcensored = TRUE, full = full)
    H$X <- H$X + Hrc$X
    H$A$Adiag <- H$A$Adiag + Hrc$A$Adiag
    H$A$Aoffdiag <- H$A$Aoffdiag + Hrc$A$Aoffdiag
    H$Z <- H$Z + Hrc$Z
  }
  if (b == 1L) {
    Adiag <- H$A$Adiag
    Aoffdiag <- H$A$Aoffdiag
    X <- H$X
    Z <- H$Z
  } else {
    Adiag <- c(Adiag, H$A$Adiag)
    Aoffdiag <- c(Aoffdiag, 0, H$A$Aoffdiag)
    X <- rbind(X, H$X)
    Z <- Z + H$Z
  }
}

if (length(Adiag) > 1L) {
  A <- Matrix::bandSparse(length(Adiag),
    k = 0:1, diagonals = list(Adiag, Aoffdiag),
    symmetric = TRUE)
} else {
  A <- matrix(Adiag)
}

ret <- cbind(Z, t(X))
ret <- rbind(ret, cbind(X, A))
if (retMatrix) return(ret)
return(as.matrix(ret))
◇

```

Fragment referenced in [?](#).

and the computation of the Hessian for the shift parameters using **Matrix** technology:

$\langle \text{stratified Hessian ?} \rangle \equiv$

```
.shes <- function(parm, x, mu = 0, xrc = NULL, full = FALSE,
                  retMatrix = FALSE)
{
   $\langle \text{stratum prep ?} \rangle$ 

  if (!isFALSE(ret <- full)) {
     $\langle \text{full Hessian ?} \rangle$ 
  }
  ret <- matrix(0, nrow = length(bidx), ncol = length(bidx))
  for (b in seq_len(B)) {
    H <- .hes(c(delta, intercepts[[b]]), x[[b]], mu = mu)
    if (!is.null(xrc)) {
      Hrc <- .hes(c(delta, intercepts[[b]]), xrc[[b]], mu = mu,
                  rightcensored = TRUE)
      H$X <- H$X + Hrc$X
      H$A <- H$A + Hrc$A
      H$Z <- H$Z + Hrc$Z
    }
    sAH <- tryCatch(Matrix::solve(H$A, H$X), error = function(e) NULL)
    if (is.null(sAH))
      stop(gettextf("error computing the Hessian in %s",
                    "freelway"),
           domain = NA)
    ret <- ret + (H$Z - crossprod(H$X, sAH))
  }
  as.matrix(ret)
}
 $\diamond$ 
```

Fragment referenced in [?](#).

```
> ### analytical covariance of parameters
> solve(.shes(op$par, xl))

      [,1]      [,2]
[1,] 0.27095 0.16154
[2,] 0.16154 0.32006

> ### numerical covariance
> solve(op$hessian)[1:2,1:2]

      mt2      mt3
mt2 0.27095 0.16154
mt3 0.16154 0.32006

> ### from glm
> vcov(m)[- (1:2), - (1:2)]

      t2      t3
t2 0.27095 0.16153
t3 0.16153 0.32006

> obj <- .freelwayML(as.table(x), link = logit())
> obj$coefficients

      B      C
1.30224 0.74147

> -obj$value
```

```
[1] -62.523

> ### analytical covariance
> obj$vcov

      B      C
B 0.27095 0.16153
C 0.16153 0.32006
```

Chapter 3

Link Functions

Similar to `family` objects, we provide some infrastructure for `link` functions F^{-1} and derived quantities (`linkinv` F , `dlinkinv` f , and `ddlinkinv` f'). If not provided, we also set-up the ratio f'/f in the constructor.

Although there is some overlap with `family` objects for binomial outcomes, it doesn't seem beneficial to extend this richer class.

```
"linkfun.R" ?≡
```

```
# File src/library/stats/R/linkfun.R
# generated by running nuweb on
#   system.file("free1way.w", package = "free1way.docreg")
# Part of the R package, https://www.R-project.org
#
# Copyright (C) 2026 The R Core Team
#
# This program is free software; you can redistribute it and/or modify
# it under the terms of the GNU General Public License as published by
# the Free Software Foundation; either version 2 of the License, or
# (at your option) any later version.
#
# This program is distributed in the hope that it will be useful,
# but WITHOUT ANY WARRANTY; without even the implied warranty of
# MERCHANTABILITY or FITNESS FOR A PARTICULAR PURPOSE. See the
# GNU General Public License for more details.
#
# A copy of the GNU General Public License is available at
# https://www.R-project.org/Licenses/

< linkfun ? >
< logit ? >
< probit ? >
< cloglog ? >
< loglog ? >
◇
```

$\langle \text{linkfun ?} \rangle \equiv$

```

### distribution function
.p <- function(link, q, ...)
  link$linkinv(q = q, ...)

### quantile function
.q <- function(link, p, ...)
  link$link(p = p, ...)

### density function
.d <- function(link, x, ...)
  link$dlinkinv(x = x, ...)

### derivative of density function
.dd <- function(link, x, ...)
  link$ddlinkinv(x = x, ...)

### 2nd derivative of density function
.ddd <- function(link, x, ...)
  link$dddlinkinv(x = x, ...)

### ratio of derivative of density to
### density function
.dd2d <- function(link, x, ...)
  link$dd2dlinkinv(x = x, ...)

### constructor
linkfun <- function(name,      ### nickname
                    alias,    ### char
                    model,    ### char, semiparametric model name
                    parm,     ### char, parameter name
                    link,     ### quantile function
                    linkinv,  ### distribution function
                    dlinkinv, ### density function
                    ddlinkinv, ### derivative of density function
                    ...)
{
  ret <- list(name = name,
             alias = alias,
             model = model,
             parm = parm,
             link = link,
             linkinv = linkinv,
             dlinkinv = dlinkinv,
             ddlinkinv = ddlinkinv)
  if (is.null(ret$dd2d))
    ret$dd2d <- function(x)
      ret$dddlinkinv(x) / ret$dlinkinv(x)
  ret <- c(ret, list(...))
  class(ret) <- "linkfun"
  ret
}

```

Fragment referenced in [?](#).

We start with the logit link, that is $F(z) = (1 + \exp(-z))^{-1}$, giving rise to Wilcoxon or Kruskal-Wallis type score residuals:

$\langle \text{logit ?} \rangle \equiv$

```
logit <- function()
  linkfun(name = "Logit",
    alias = c("Wilcoxon", "Kruskal-Wallis"),
    model = "proportional odds",
    parm = "log-odds ratio",
    link = qlogis,
    linkinv = plogis,
    dlinkinv = dlogis,
    ddlinkinv = function(x) {
      p <- plogis(x)
      p * (1 - p)^2 - p^2 * (1 - p)
    },
    dddlinkinv = function(x) {
      ex <- exp(x)
      ifelse(is.finite(x), (ex - 4 * ex^2 + ex^3) / (1 + ex)^4, 0.0)
    },
    dd2d = function(x) {
      ex <- exp(x)
      (1 - ex) / (1 + ex)
    },
    parm2PI = function(x) {
      OR <- exp(x)
      ret <- OR * (OR - 1 - x) / (OR - 1)^2
      ret[abs(x) < .Machine$double.eps] <- 0.5
      return(ret)
    },
    PI2parm = function(p) {
      f <- function(x, PI)
        x + (exp(-x) * (PI +
          exp(2 * x) * (PI - 1) +
          exp(x) * (1 - 2 * PI)))
      ret <- vapply(p, function(p)
        uniroot(f, PI = p, interval = 50 * c(-1, 1))$root, 0)
      return(ret)
    },
    parm2OVL = function(x) 2 * plogis(-abs(x / 2))
  )
  )
```

◇

Fragment referenced in ?.

The **parm2PI** function converts log-odds ratios to probabilistic indices (or AUCs) and the inverse operation is implemented by **PI2parm**. The overlap coefficient can be obtained from a log-odds ratio via **parm2OVL**.

The log-log link, with $F(z) = \exp(-\exp(-z))$, is used to construct tests against Lehmann alternatives:

$\langle \log\log ? \rangle \equiv$

```
loglog <- function()
  linkfun(name = "Log-log",
    alias = "Lehmann",
    model = "Lehmann",
    parm = "log-reverse time hazard ratio",
    link = function(p, log.p = FALSE) {
      if (!log.p) p <- log(p)
      -log(-p)
    },
  linkinv = function(q, lower.tail = TRUE, log.p = FALSE) {
    ### p = exp(-exp(-q))
    if (log.p) {
      if (lower.tail)
        return(-exp(-q))
      return(log1p(-exp(-exp(-q))))
    }
    if (lower.tail)
      return(exp(-exp(-q)))
    -expm1(-exp(-q))
  },
  dlinkinv = function(x)
    ifelse(is.finite(x), exp(- x - exp(-x)), 0.0),
  ddlinkinv = function(x) {
    ex <- exp(-x)
    ifelse(is.finite(x), exp(-ex - x) * (ex - 1.0), 0.0)
  },
  dddlinkinv = function(x) {
    ex <- exp(-x)
    ifelse(is.finite(x), exp(-x - ex) * (ex - 1)^2 -
      exp(-ex - 2 * x),
      0.0)
  },
  dd2d = function(x)
    expm1(-x),
  parm2PI = plogis,
  PI2parm = qlogis,
  parm2OVL = function(x) {
    x <- abs(x)
    rt <- exp(-x / (exp(x) - 1))
    ret <- rt^exp(x) + 1 - rt
    ret[abs(x) < .Machine$double.eps] <- 1
    x[] <- ret
    return(x)
  }
)
```

◇

Fragment referenced in [?](#).

The complementary log-log link, with $F(z) = 1 - \exp(-\exp(z))$, provides log-rank or Savage score residuals against proportional hazards alternatives:

$\langle \text{cloglog ?} \rangle \equiv$

```
cloglog <- function()
  linkfun(name = "Complementary Log-log",
    alias = "Savage",
    model = "proportional hazards",
    parm = "log-hazard ratio",
    link = function(p, log.p = FALSE) {
      if (log.p) p <- exp(p)
      log(-log1p(- p))
    },
    linkinv = function(q, lower.tail = TRUE, log.p = FALSE) {
      ### p = 1 - exp(-exp(q))
      ret <- exp(-exp(q))
      if (log.p) {
        if (lower.tail)
          return(log1p(-ret))
        return(-exp(q))
      }
      if (lower.tail)
        return(-expm1(-exp(q)))
      return(ret)
    },
    dlinkinv = function(x)
      ifelse(is.finite(x), exp(x - exp(x)), 0.0),
    ddlinkinv = function(x) {
      ex <- exp(x)
      ifelse(is.finite(x), (ex - ex^2) / exp(ex), 0.0)
    },
    dddlinkinv = function(x) {
      ex <- exp(x)
      ifelse(is.finite(x), (ex - 3*ex^2 + ex^3) / exp(ex), 0.0)
    },
    dd2d = function(x)
      -expm1(x),
    parm2PI = plogis,
    PI2parm = qlogis,
    parm2OVL = function(x) {
      x <- abs(x)
      ret <- exp(x / (exp(-x) - 1)) - exp(-x / (exp(x) - 1)) + 1
      ret[abs(x) < .Machine$double.eps] <- 1
      x[] <- ret
      return(x)
    }
  )
  )
```

◇

Fragment referenced in ?.

The probit link, with $F(z) = \Phi$, leads to normal scores tests, where the shift effect can be interpreted as a generalised version of Cohen's d , that is, differences on a latent normal scale with variance one:

$\langle \text{probit ?} \rangle \equiv$

```
probit <- function()
  linkfun(name = "Probit",
    alias = "van der Waerden normal scores",
    model = "latent normal shift",
    parm = "generalised Cohen's d",
    link = qnorm,
    linkinv = pnorm,
    dlinkinv = dnorm,
    ddlinkinv = function(x)
      ifelse(is.finite(x), -dnorm(x = x) * x, 0.0),
    dddlinkinv = function(x)
      ifelse(is.finite(x), dnorm(x = x) * (x^2 - 1), 0.0),
    dd2d = function(x) -x,
    parm2PI = function(x) pnorm(x, sd = sqrt(2)),
    PI2parm = function(p) qnorm(p, sd = sqrt(2)),
    parm2OVL = function(x) 2 * pnorm(-abs(x / 2))
  )
  ◇
```

Fragment referenced in [?](#).

Chapter 4

Optimisation

[Harrell \(2024\)](#) reports on experiments with a number of optimisers for the specific optimisation problem arising here and recommends a Newton-Raphson algorithm leveraging the sparse matrix structure of the observed Fisher information matrix. The following code was adopted from his **rms** package, function `rms:::lrm.fit`.

⟨ Newton update ? ⟩ \equiv

```
gradthe <- gradient(theta)      # Compute the gradient vector
hessthe <- hessian(theta)       # Compute the Hessian matrix

delta <- Matrix::solve(hessthe, gradthe, tol = control$tol.solve)

if (control$trace)
  cat(iter, ': ', theta, "\n", sep = "")

step_size <- 1L                 # Initialize step size for step-halving
◇
```

Fragment referenced in [?](#).

$\langle \text{Newton step halving ?} \rangle \equiv$

```

new_theta <- theta - step_size * delta # Update parameter vector
objnew_the <- objective(new_theta)

if (control$trace)
  cat("Old, new, old - new objective:",
      objthe, objnew_the, objthe - objnew_the, "\n")

# Objective function failed to be reduced or is infinite
if (!is.finite(objnew_the) || (objnew_the > objthe + 1e-6)) {
  step_size <- step_size / 2      # Reduce the step size

  if (control$trace)
    cat("Step size reduced to", step_size, "\n")

  if (step_size <= control$minstepsize) {
    msg <- paste("Step size ", step_size,
                 " has reduced below minstepsize")
    return(list(par = theta, objective = objthe, convergence = 1,
                message = msg))
  }
} else {
  theta <- new_theta # accept the new parameter vector
  oldobj <- objthe
  objthe <- objnew_the
  break
}

```

Fragment referenced in [?](#).

$\langle \text{Newton convergence ?} \rangle \equiv$

```

# Convergence check - must meet 3 criteria
if ((objthe <= oldobj + 1e-6 && (oldobj - objthe < control$objtol)) &&
    (max(abs(gradthe)) < control$gradtol) &&
    (max(abs(delta)) < control$paramtol))

  return(list(par      = theta,
              objective = objthe,
              convergence = 0,
              message   = "Normal convergence"))

```

Fragment referenced in [?](#).

$\langle \text{NewtonRaphson} ? \rangle \equiv$

```
### adopted from rms:::lrn.fit
.NewtonRaphson <- function(start, objective, gradient, hessian,
                           control = list(iter.max = 150, trace = trace,
                                           objtol = 5e-4, gradtol = 1e-5,
                                           paramtol = 1e-5, minstepsize = 1e-2,
                                           tolsolve = .Machine$double.eps),
                           trace = FALSE)
{
  theta <- start # Initialize the parameter vector
  oldobj <- Inf
  objthe <- objective(theta)
  if (!is.finite(objthe)) {
    msg <- "Infeasible starting values"
    return(list(par = theta, objective = objthe, convergence = 1,
               message = msg))
  }

  if(!suppressPackageStartupMessages(requireNamespace("Matrix")))
    stop(gettextf("%s needs package 'Matrix' correctly installed",
                  ".NewtonRaphson"),
         domain = NA)

  for (iter in seq_len(control$iter.max)) {

     $\langle \text{Newton update} ? \rangle$ 

    # Step-halving loop
    while (TRUE) {
       $\langle \text{Newton step halving} ? \rangle$ 
    }

     $\langle \text{Newton convergence} ? \rangle$ 
  }

  msg <- paste("Reached", control$iter.max, "iterations without convergence")
  return(list(par = theta, objective = objthe, convergence = 1, message = msg))
}
◇
```

Fragment referenced in ?.

We can now test the optimiser on a least-squares problem

```
> N <- 10000
> P <- 30
> X <- matrix(rnorm(N * P), ncol = P)
> y <- X %*% runif(P) + rnorm(nrow(X))
> f <- function(par) sum((y - X %*% par)^2)
> g <- function(par) colSums(- 2 * c(y - X %*% par) * X)
> h <- function(par) 2 * crossprod(X)
> start <- runif(P)
> cf <- .NewtonRaphson(start = start, objective = f, gradient = g, hessian = h)
> cf2 <- coef(m <- lm(y ~ 0 + X))
> all.equal(sum((y - fitted(m))^2), cf$objective)

[1] TRUE

> all.equal(unname(cf$par), unname(cf2))

[1] TRUE
```

Chapter 5

ML Estimation

We use two internal `stats` functions, define here

"utils.R" ?≡

```
### gives warnings but no diffs
C_dpermdist2 <- stats:::C_dpermdist2
assert_NULL_or_prob <- stats:::assert_NULL_or_prob
◇
```

The file `free1way.R` goes into `src/library/stats/R`, so add copyright statement here as well (such that we can simply copy the file in case of updates).

```
"free1way.R" ?≡
```

```
# File src/library/stats/R/free1way.R
# generated by running nuweb on
#   system.file("free1way.w", package = "free1way.docreg")
# Part of the R package, https://www.R-project.org
#
# Copyright (C) 2026 The R Core Team
#
# This program is free software; you can redistribute it and/or modify
# it under the terms of the GNU General Public License as published by
# the Free Software Foundation; either version 2 of the License, or
# (at your option) any later version.
#
# This program is distributed in the hope that it will be useful,
# but WITHOUT ANY WARRANTY; without even the implied warranty of
# MERCHANTABILITY or FITNESS FOR A PARTICULAR PURPOSE. See the
# GNU General Public License for more details.
#
# A copy of the GNU General Public License is available at
# https://www.R-project.org/Licenses/

<NewtonRaphson ?>
<ML estimation ?>
<free1way generic and table method (main workhorse) ?>
<free1way methods ?>
<free1way print ?>
<free1way summary ?>
<free1way confint ?>
<free1way formula ?>
<free1way numeric ?>
<free1way factor ?>
<plot free1way ?>
<ppplot ?>
<rfree1way ?>
<power ?>
◇
```

We now put together a low-level function for parameter estimation and evaluation of scores, Hessians, and residuals. We also set-up a profile likelihood function for later re-use.

Assuming all shift effects been zero, we compute starting values for the intercept parameters from the empirical cumulative distribution function after merging all treatment groups:

$\langle \text{setup and starting values ?} \rangle \equiv$

```



```

Fragment referenced in [?](#).

The profile negative log-likelihood can be evaluated for some of the parameters in θ (denoted as **fix**), the remaining parameters are updated. Note that **start** can either just contain a subset of the shift parameter or must contain the full and feasible (meeting monotonicity constraints for the intercept parameters) parameter vector θ .

We call **nlminb** and will increase the **eval.max** and **iter.max** control parameters if we encounter optimisation issues and restart at the current solution:

$\langle do\ optim\ ? \rangle \equiv$

```

maxit <- control[[1L]]$iter.max
while(maxit < 10001) {
  ret <- do.call(names(control)[[1L]], opargs)
  maxit <- 5 * maxit
  if (ret$convergence > 0) {
    opargs$control$eval.max <- maxit
    opargs$control$iter.max <- maxit
    opargs$start <- ret$par
  } else {
    break()
  }
}

if (isTRUE(MPL_Jeffreys)) {
   $\langle Jeffreys\ penalisation\ ? \rangle$ 
} else {
  if (ret$convergence > 0) {
    if (is.na(MPL_Jeffreys)) { ### only after failure
      warning(gettextf("Jeffreys penalisation was applied in %s because initial optimisation failed
                        "free1way"),
              "\n ", ret$message, domain = NA)
      MPL_Jeffreys <- TRUE
       $\langle Jeffreys\ penalisation\ ? \rangle$ 
    }
  }
}

if (ret$convergence > 0)
  warning(gettextf("unsuccessful optimisation in %s", "free1way"),
          ": ", ret$message, domain = NA)

ret$MPL_Jeffreys <- MPL_Jeffreys
ret$value <- ret$objective
ret$objective <- NULL

```

Fragment referenced in [?](#), [?](#).

We first set-up the target function (the negative log-likelihood, also dealing with right-censoring) and the corresponding gradient. We then add the profile negative log-likelihood, which in turn calls the two functions defined first. We start with the log-likelihood, its gradient and Hessian

$\langle \logLik, gradient, Hessian ? \rangle \equiv$

```
fn <- function(par)
{
  ret <- .snll(par, x = xlist, mu = mu)
  if (!is.null(xrc))
    ret <- ret + .snll(par, x = xrclist, mu = mu,
                      rightcensored = TRUE)
  return(ret)
}
gr <- function(par)
{
  ret <- .snsc(par, x = xlist, mu = mu)
  if (!is.null(xrc))
    ret <- ret + .snsc(par, x = xrclist, mu = mu,
                      rightcensored = TRUE)
  return(ret)
}

### allocate memory for hessian
Hess <- Matrix::Matrix(0, nrow = length(start), ncol = length(start))

he <- function(par)
{
  if (!is.null(xrc)) {
    ret <- .shes(par, x = xlist, mu = mu, xrc = xrclist, full = Hess,
                retMatrix = names(control)[1L] == ".NewtonRaphson")
  } else {
    ret <- .shes(par, x = xlist, mu = mu, full = Hess,
                retMatrix = names(control)[1L] == ".NewtonRaphson")
  }
  return(ret)
}
◇
```

Fragment referenced in [?](#).

and define the profile log-likelihood based on these functions

$\langle profile ? \rangle \equiv$

$\langle logLik, gradient, Hessian ? \rangle$

```
.profile <- function(start, fix = seq_len(K - 1))
{
  if (!all(fix %in% seq_len(K - 1)))
    stop(gettextf("invalid argument '%s'", "fix"), domain = NA)
  delta <- start[fix]
  opargs <- list(start = start[-fix],
    objective = function(par) {
      p <- numeric(length(par) + length(fix))
      p[fix] <- delta
      p[-fix] <- par
      fn(p)
    },
    gradient = function(par) {
      p <- numeric(length(par) + length(fix))
      p[fix] <- delta
      p[-fix] <- par
      gr(p)[-fix]
    },
    hessian = function(par) {
      p <- numeric(length(par) + length(fix))
      p[fix] <- delta
      p[-fix] <- par
      he(p)[-fix, -fix, drop = FALSE]
    })
  opargs$control <- control[[1L]]
  MPL_Jeffreys <- FALSE ### turn off Jeffreys penalisation in .profile

   $\langle do optim ? \rangle$ 

  p <- numeric(length(start))
  p[fix] <- delta
  p[-fix] <- ret$par
  ret$par <- p
  ret
}
◇
```

Fragment referenced in ?.

Chapter 13 introduces a bias correction (Firth, 1993), essentially by adding a penalty (Jeffreys prior) term to the log-likelihood. The `MPL_Jeffreys` argument triggers this bias correction via penalisation with Jeffreys prior to be applied when `TRUE`, not to be applied when `FALSE`, and to be applied in case the unpenalised ML estimation resulted in a convergence issue (`NA`). This part is still experimental and needs more testing. It also seems unclear if and how the Fisher information needs additional correction and it is certainly unclear if one can proceed with permutation testing after correcting the bias.

$\langle \text{Jeffreys penalisation ?} \rangle \equiv$

```
.pll_Jeffreys <- function(cf, start)
{
  fix <- seq_along(cf)
  start[fix] <- cf
  ### compute profile likelihood w/o warnings
  ret <- suppressWarnings(.profile(start, fix = fix))
  Hfull <- he(ret$par)
  Hfix <- as.matrix(solve(solve(Hfull)[fix, fix]))
  return(ret$value -
    .5 * determinant(Hfix, logarithm = TRUE)$modulus)
}
if (K == 2) {
  MLcf <- ret$par[seq_len(K - 1)]
  Fret <- optim(MLcf, fn = .pll_Jeffreys, start = ret$par,
    method = "Brent", lower = MLcf - 5,
    upper = MLcf + 5)
} else {
  ### Nelder-Mead
  Fret <- optim(ret$par[seq_len(K - 1)], fn = .pll_Jeffreys,
    start = ret$par)
}
if (Fret$convergence == 0) {
  start <- ret$par
  start[seq_len(K - 1)] <- Fret$par
  ret <- .profile(start, fix = seq_len(K - 1))
  ret$value <- ret$value
}
}
```

Fragment referenced in [?](#).

The heart of the function is a call to `nlminb`, trying to obtain parameter estimates of θ by minimising the negative log-likelihood. We allow some (or all) parameters to be fixed at some constants, and provide a profile version of the likelihood:

$\langle \text{optim ?} \rangle \equiv$

```
if (!length(fix)) {
  opargs <- list(start = start,
    objective = fn,
    gradient = gr,
    hessian = he)
  opargs$control <- control[[1L]]
   $\langle \text{do optim ?} \rangle$ 
} else if (length(fix) == length(start)) {
  ret <- list(par = start,
    value = fn(start))
} else {
  ret <- .profile(start, fix = fix)
}
}
```

Fragment referenced in [?](#).

After parameter estimation, we evaluate negative scores, the Hessian, and negative residuals as requested:

< post processing ? > \equiv

```

if (is.null(fix) || (length(fix) == length(start)))
  parm <- seq_len(K - 1)
else
  parm <- fix
if (any(parm >= K)) return(ret)

ret$coefficients <- ret$par[parm]
dn2 <- dimnames(xt)[2L]
names(ret$coefficients) <- cnames <- paste0(names(dn2), dn2[[1L]][1L + parm])

par <- ret$par
intercepts <- function(parm, x)
{
  < stratum prep ? >

  return(intercepts)
}
ret$intercepts <- intercepts(par, x = xlist)

if (score) {
  ret$negscore <- .snsc(par, x = xlist, mu = mu)[parm]
  if (!is.null(xrc))
    ret$negscore <- ret$negscore + .snsc(par, x = xrc, mu = mu,
                                          rightcensored = TRUE)[parm]
}
if (hessian) {
  if (!is.null(xrc)) {
    ret$hessian <- .shes(par, x = xlist, mu = mu, xrc = xrc)
  } else {
    ret$hessian <- .shes(par, x = xlist, mu = mu)
  }
  ret$vcov <- solve(ret$hessian)
  if (length(parm) != nrow(ret$hessian))
    ret$hessian <- solve(ret$vcov <- ret$vcov[parm, parm, drop = FALSE])
  rownames(ret$vcov) <- colnames(ret$vcov) <- rownames(ret$hessian) <-
    colnames(ret$hessian) <- cnames
}
if (residuals) {
  ret$negresiduals <- .snsr(par, x = xlist, mu = mu)
  if (!is.null(xrc)) {
    rcr <- .snsr(par, x = xrc, mu = mu, rightcensored = TRUE)
    ret$negresiduals <- c(rbind(matrix(ret$negresiduals, nrow = C),
                                     matrix(rcr, nrow = C)))
  }
}
ret$profile <- function(start, fix)
  .freelwayML(xt, link = link, mu = mu, start = start, fix = fix, tol = tol,
             ...)
ret$table <- xt

ret$strata <- strata
ret$mu <- mu
if (length(ret$mu) == 1) {
  names(ret$mu) <- link$parm
} else {
  names(ret$mu) <- c(paste(link$parm, cnames[1L], sep = ":"), cnames[-1L])
}
◇

```

Fragment referenced in [?](#).

Finally, we put everything into one function which returns an object of class `free1wayML` for later use. The control parameters for `.NewtonRaphson` and `stats::nlminb` are the ones suggested by [Harrell \(2024\)](#). By default, the internal Newton-Raphson implementation is used, we can switch to `stats::nlminb` by specifying `doptim = "nlminb"`. The latter option cannot handle Fisher information matrices in form of a `Matrix` object and thus computing the updates takes more time whenever a larger number of intercept parameters is present in the problem.

< ML estimation ? > \equiv

```
.freelwayML <- function(x, link, mu = 0, start = NULL, fix = NULL,
  residuals = TRUE, score = TRUE, hessian = TRUE,
  MPL_Jeffreys = FALSE,
  ### use nlminb for small sample sizes
  dooptim = c(".NewtonRaphson", "nlminb")[1 + (sum(x) < 20)],
  control = list(
    "nlminb" = list(trace = trace, iter.max = 200,
      eval.max = 200, rel.tol = 1e-10,
      abs.tol = 1e-20, xf.tol = 1e-16),
    ".NewtonRaphson" = list(iter.max = 200, trace = trace,
      objtol = 5e-4,
      gradtol = 1e-5 * sum(x) / 1000,
      paramtol = 1e-5, minstepsize = 1e-2,
      tolsolve = .Machine$double.eps)
  )[dooptim],
  trace = FALSE,
  tol = sqrt(.Machine$double.eps), ...)
{
  ### convert to three-way table
  xt <- x
  if (!is.table(x))
    stop(gettextf("invalid argument '%s'", "x"), domain = NA) # 'y' in freelway ...
  dx <- dim(x)
  dn <- dimnames(x)
  if (length(dx) == 2L) {
    x <- as.table(array(c(x), dim = dx <- c(dx, 1L)))
    dimnames(x) <- dn <- c(dn, list(A = "A"))
  }

  ### short-cuts for link functions
  F <- function(q) .p(link, q = q)
  Q <- function(p) .q(link, p = p)
  f <- function(q) .d(link, x = q)
  fp <- function(q) .dd(link, x = q)

  if(!suppressPackageStartupMessages(requireNamespace("Matrix")))
    stop(gettextf("%s needs package 'Matrix' correctly installed",
      ".freelwayML"),
      domain = NA)

  < setup and starting values ? >
  < negative logLik ? >
  < negative score ? >
  < negative score residuals ? >
  < Hessian ? >
  < stratified negative logLik ? >
  < stratified negative score ? >
  < stratified Hessian ? >
  < stratified negative score residual ? >
  < profile ? >
  < optim ? >
  < post processing ? >

  class(ret) <- "freelwayML"
  ret
}
◇
```

Fragment referenced in [?](#).

As an example, consider a stratified (two stata) 3×3 problem where outcome category B is missing from the second stratum:

```
> N <- 10
> a <- matrix(c(5, 6, 4,
+             3, 5, 7,
+             3, 4, 5,
+             3, 5, 6,
+             0, 0, 0,
+             4, 6, 5), ncol = 3, byrow = TRUE)
> x <- as.table(array(c(a[1:3,], a[-(1:3),]), dim = c(3, 3, 2)))
> x

, , A
  A B C
A 5 6 4
B 3 5 7
C 3 4 5

, , B
  A B C
A 3 5 6
B 0 0 0
C 4 6 5

> ret <- .freelwayML(x, logit())
> ret[c("value", "par")]

$value
[1] 65.932

$par
[1] 0.052627 0.204750 -0.489670 1.020466 0.028488

> cf <- ret$par
> cf[1:2] <- cf[1:2] + .5
> ### new2old parameterisation
> c(cf[1:2], cf[3], log(cf[4] - cf[3]), cf[5])

[1] 0.552627 0.704750 -0.489670 0.412200 0.028488

> ### profile for cf[1:2]
> .freelwayML(x, logit(), start = cf, fix = 1:2)[c("value", "par")]

$value
[1] 66.381

$par
[1] 0.55263 0.70475 -0.11885 1.40796 0.40841

> ### profile for cf[2]
> .freelwayML(x, logit(), start = cf, fix = 2)[c("value", "par")]

$value
[1] 66.299

$par
[1] 0.36095 0.70475 -0.18771 1.34153 0.33443
```

```

> ### evaluate log-likelihood at cf
> .freelwayML(x, logit(), start = cf,
+           fix = seq_along(ret$par))[c("value", "par")]

$value
[1] 67.771

$par
[1] 0.552627 0.704750 -0.489670 1.020466 0.028488

```


Chapter 6

ML Inference

Based on an object of class `free1wayML`, we can setup different test statistics and obtain the limiting null distribution based on classical ML theory under the population model:

$\langle statistics ? \rangle \equiv$

```

if (test == "Wald") {
   $\langle Wald statistic ? \rangle$ 
} else if (test == "LRT") {
   $\langle LRT ? \rangle$ 
} else if (test == "Rao") {
   $\langle Rao ? \rangle$ 
} else if (test == "Permutation") {
   $\langle Permutation statistics ? \rangle$ 
}

```

Fragment referenced in [?, ?](#).

6.1 Wald Statistics

We only need access to the parameter estimates $\hat{\delta}_2, \dots, \hat{\delta}_K$ and the corresponding Hessian:

$\langle Wald statistic ? \rangle \equiv$

```

if (alternative == "two.sided") {
  STATISTIC <- c("Wald chi-squared" =
    c(crossprod(cf, x$hessian %*% cf)))
  DF <- c("df" = length(parm))
  PVAL <- pchisq(STATISTIC, df = DF, lower.tail = FALSE)
} else {
  STATISTIC <- c("Wald Z" = unname(c(cf * sqrt(c(x$hessian)))))
  PVAL <- pnorm(STATISTIC, lower.tail = alternative == "less")
}

```

Fragment referenced in [?](#).

6.2 Likelihood-ratio Statistics

In addition to the log-likelihood evaluated at the ML estimates, we need to evaluate the profile log-likelihood at some value corresponding the null hypothesis to be tested:

$\langle LRT ? \rangle \equiv$

```
par <- x$par
par[parm] <- value
unll <- x$value ### neg logLik
rnll <- x$profile(par, parm)$value ### neg logLik
STATISTIC <- c("logLR chi-squared" = - 2 * (unll - rnll))
DF <- c("df" = length(parm))
PVAL <- pchisq(STATISTIC, df = DF, lower.tail = FALSE)
◇
```

Fragment referenced in [?](#).

6.3 Rao Score Statistics

For the Rao score test, the inverse of the Hessian as well as the score function of the shift parameters evaluated for some null values need to be computed:

$\langle Rao ? \rangle \equiv$

```
par <- x$par
par[parm] <- value
ret <- x$profile(par, parm)
if (alternative == "two.sided") {
  STATISTIC <- c("Rao chi-squared" = c(crossprod(ret$negscore,
                                                  ret$vcov %*%
                                                  ret$negscore)))

  DF <- c("df" = length(parm))
  PVAL <- pchisq(STATISTIC, df = DF, lower.tail = FALSE)
} else {
  STATISTIC <- c("Rao Z" = unname(- ret$negscore *
                                   sqrt(c(ret$vcov))))
  PVAL <- pnorm(STATISTIC, lower.tail = alternative == "less")
}
◇
```

Fragment referenced in [?](#).

Chapter 7

Permutation Inference

Under the permutation model, that is, in randomised experiments where the random treatment allocation is the only relevant source of randomness, we compute a permutation variant of the Rao score test, based on the conditional asymptotic distribution or based on a Monte-Carlo estimate of the reference distribution:

$\langle \text{Permutation statistics ?} \rangle \equiv$

```
par <- x$par
par[parm] <- value
ret <- x$profile(par, parm)
sc <- - ret$negscore
if (length(cf) == 1L)
  sc <- sc / sqrt(c(ret$hessian))
if (!is.null(x$exact)) {
  STATISTIC = c("W" = sc)
} else {
  Esc <- sc - x$perm$Expectation

  if (alternative == "two.sided" && length(cf) > 1L) {
    STATISTIC <- c("Perm chi-squared" =
      sum(Esc * solve(x$perm$Covariance, Esc)))
  } else {
    STATISTIC <- c("Perm Z" = Esc / sqrt(c(x$perm$Covariance)))
  }
}
◇
```

Fragment referenced in [?](#).

In addition, we compute permutation p -values

$\langle \text{Permutation } p\text{-values?} \rangle \equiv$

```

if (!is.null(x$exact)) {
  PVAL <- switch(alternative,
    "two.sided" = 2 * min(c(x$exact$ple(sc),
                           x$exact$pgr(sc))),
    "less" = x$exact$ple(sc),
    "greater" = x$exact$pgr(sc))
} else {
  .pm <- function(x) sum(x) / length(x)
  ps <- x$perm$permStat

  .GE <- function(x, y)
    (y - x) <= sqrt(.Machine$double.eps)

  .LE <- function(x, y)
    (x - y) <= sqrt(.Machine$double.eps)

  if (alternative == "two.sided" && length(cf) > 1L) {
    if (!is.null(ps)) {
      PVAL <- .pm(.GE(ps, STATISTIC))
    } else {
      DF <- c("df" = x$perm$DF)
      PVAL <- pchisq(STATISTIC, df = DF, lower.tail = FALSE)
    }
  } else {
    if (!is.null(ps)) {
      PVALle <- .pm(.LE(ps, STATISTIC))
      PVALge <- .pm(.GE(ps, STATISTIC))
      if (alternative == "two.sided")
        PVAL <- 2 * min(c(PVALle, PVALge))
      else if (alternative == "less")
        PVAL <- PVALle
      else
        PVAL <- PVALge
    } else {
      if (alternative == "two.sided")
        PVAL <- pchisq(STATISTIC^2, df = 1, lower.tail = FALSE)
      else
        PVAL <- pnorm(STATISTIC,
                      lower.tail = alternative == "less")
    }
  }
}

```

Fragment referenced in [?](#).

The mean and variance of the linear permutation statistic under the null was given by [Strasser and Weber \(1999\)](#):

$\langle \text{Strasser Weber ?} \rangle \equiv$

```
.SW <- function(res, xt)
{
  if (length(dim(xt)) == 3L) {
    res <- matrix(res, nrow = dim(xt)[1L], ncol = dim(xt)[3])
    STAT <- Exp <- Cov <- 0
    for (b in seq_len(dim(xt)[3L])) {
      sw <- .SW(res[,b, drop = TRUE], xt[,b, drop = TRUE])
      STAT <- STAT + sw$Statistic
      Exp <- Exp + sw$Expectation
      Cov <- Cov + sw$Covariance
    }
    return(list(Statistic = STAT, Expectation = as.vector(Exp),
               Covariance = Cov))
  }

  Y <- matrix(res, ncol = 1, nrow = length(xt))
  weights <- c(xt)
  x <- gl(ncol(xt), nrow(xt))
  X <- model.matrix(~ x, data = data.frame(x = x))[, -1L, drop = FALSE]

  w. <- sum(weights)
  wX <- weights * X
  wY <- weights * Y
  ExpX <- colSums(wX)
  ExpY <- colSums(wY) / w.
  CovX <- crossprod(X, wX)
  Yc <- t(t(Y) - ExpY)
  CovY <- crossprod(Yc, weights * Yc) / w.
  Exp <- kronecker(ExpY, ExpX)
  Cov <- w. / (w. - 1) * kronecker(CovY, CovX) -
    1 / (w. - 1) * kronecker(CovY, tcrossprod(ExpX))
  STAT <- crossprod(X, wY)
  list(Statistic = STAT, Expectation = as.vector(Exp),
       Covariance = Cov)
}
◇
```

Fragment referenced in [?](#).

For small samples, we used the `r2dtable` function to sample from tables with fixed marginal distributions:

$\langle \text{resampling?} \rangle \equiv$

```
.resample <- function(res, xt, B = 10000)
{
  if (length(dim(xt)) == 2L)
    xt <- as.table(array(xt, dim = c(dim(xt), 1)))

  res <- matrix(res, nrow = dim(xt)[1L], ncol = dim(xt)[3L])
  stat <- 0
  ret <- .SW(res, xt)
  if (dim(xt)[2L] == 2L) {
    ret$testStat <- c((ret$Statistic - ret$Expectation) /
                      sqrt(c(ret$Covariance)))
  } else {
    ES <- ret$Statistic - ret$Expectation
    ret$testStat <- sum(ES * solve(ret$Covariance, ES))
  }
  ret$DF <- dim(xt)[2L] - 1L

  if (B) {
    for (j in 1:dim(xt)[3L]) {
      rt <- r2dtable(B, r = rowSums(xt[,j]), c = colSums(xt[,j]))
      stat <- stat + vapply(rt,
                           function(x) .colSums(x[, -1L, drop = FALSE] * res[,j],
                                                    m = nrow(x), n = ncol(x) - 1L),
                           FUN.VALUE = rep(0, dim(xt)[[2L]] - 1L))
    }
    if (dim(xt)[2L] == 2L) {
      ret$permStat <- (stat - ret$Expectation) /
                      sqrt(c(ret$Covariance))
    } else {
      ES <- matrix(stat, ncol = B) - ret$Expectation
      ret$permStat <- .colSums(ES * solve(ret$Covariance, ES),
                              m = dim(xt)[[2L]] - 1L, n = B)
    }
  }
  ret
}
◇
```

Fragment referenced in [?](#).

For the special case of the unstratified Wilcoxon two-sample test, we can also provide exact p -values computed via the Streitberg-Röhmle shift algorithm, mainly because the scores can be mapped to integers:

$\langle \text{exact proportional odds ?} \rangle \equiv$

```
.exact <- function(z, grp, w = rep.int(1, length(z)))
{
  z <- rep(z, times = w)
  grp <- rep(grp, times = w)
  x <- rank(z)
  f <- 2 - all(x == floor(x))
  x <- as.integer(x * f)
  x <- x - min(x) + 1L
  sx <- sort(x)

  m <- as.integer(sum(grp > 0))
  stopifnot(m > 1)
  stopifnot(m < length(x))

  d <- .Call(stats::C_dpermdist2, sx, m)
  s <- seq.int(from = 1L, to = sum(rev(sx)[seq_len(m)]), by = 1L)

  STATISTIC <- sum(x[grp > 0])
  F <- cumsum(d)
  S <- rev(cumsum(rev(d)))
  cf <- lm.fit(x = cbind(1, x), y = as.double(z))$coefficients

  z2x <- function(z) round((z - m * cf[1]) / cf[2])

  c(ple = function(z) sum(d[s <= z2x(z)]), # s and STATISTIC are integers
    pgr = function(z) sum(d[s >= z2x(z)]),
    qle = function(q) c(m, max(s[F < q + 1e-08])) %*% cf,
    qgr = function(q) c(m, min(s[S < q + 1e-08])) %*% cf)
}
```

◇

Fragment referenced in ?.

As an example, consider the Wilcoxon rank sum test, where the scores under the null are a linear function of the ranks of the data. We compute the asymptotic and approximated reference distribution and corresponding p -values for a test statistics in quadratic form:

```
> set.seed(29)
> w <- gl(2, 15)
> (s <- .SW(r <- rank(u <- runif(length(w))), model.matrix(~ 0 + w)))

$Statistic
  [,1]
x2 287

$Expectation
[1] 232.5

$Covariance
  [,1]
[1,] 581.25

> ps <- .resample(r, model.matrix(~ 0 + w), B = 100000)
> ps$testStat^2

[1] 5.1101

> mean(abs(ps$permStat) > abs(ps$testStat) - .Machine$double.eps)

[1] 0.02435
```

```
> pchisq(ps$testStat~ifelse(ps$DF == 1, 2, 1), df = ps$DF, lower.tail = FALSE)

[1] 0.023787
```

```
> ### exactly the same
> kruskal.test(u ~ w)
```

Kruskal-Wallis rank sum test

```
data: u by w
Kruskal-Wallis chi-squared = 5.11, df = 1, p-value = 0.024
```

```
> library("coin")
> ### almost the same
> kruskal_test(u ~ w, distribution = approximate(100000))
```

Approximative Kruskal-Wallis Test

```
data: u by w (1, 2)
chi-squared = 5.11, p-value = 0.024
```

and the exact versions are

```
> wilcox_test(u ~ w, distribution = "exact")
```

Exact Wilcoxon-Mann-Whitney Test

```
data: u by w (1, 2)
Z = -2.26, p-value = 0.023
alternative hypothesis: true mu is not equal to 0
```

```
> free1way(u ~ w, exact = TRUE)
```

2-sample Wilcoxon test against proportional odds alternatives

```
data: u by w (1, 2)
W = 2.38, p-value = 0.023
alternative hypothesis: true log-odds ratio is not equal to 0
```

```
> wilcox_test(u ~ w, distribution = "exact", alternative = "less")
```

Exact Wilcoxon-Mann-Whitney Test

```
data: u by w (1, 2)
Z = -2.26, p-value = 0.012
alternative hypothesis: true mu is less than 0
```

```
> print(free1way(u ~ w, exact = TRUE), alternative = "greater")
```

2-sample Wilcoxon test against proportional odds alternatives

```
data: u by w (1, 2)
W = 2.38, p-value = 0.012
alternative hypothesis: true log-odds ratio is greater than 0
```

```
> wilcox_test(u ~ w, distribution = "exact", alternative = "greater")
```

Exact Wilcoxon-Mann-Whitney Test

```
data: u by w (1, 2)
Z = -2.26, p-value = 0.99
alternative hypothesis: true mu is greater than 0
```



```
> print(freelway(u ~ w, exact = TRUE), alternative = "less")

      2-sample Wilcoxon test against proportional odds alternatives

data:  u by w (1, 2)
W = 2.38, p-value = 0.99
alternative hypothesis: true log-odds ratio is less than 0

<TH>Ordered alternatives: Use contrast based tests in multcomp</TH>
```

Chapter 8

Distribution-free Tests in Stratified K -sample Oneway Layouts

8.1 `free1way`

We provide a new test procedure in a generic `free1way`, featuring a method for tables (the main workhorse) and additional user interfaces.

$\langle \text{link2fun ?} \rangle \equiv$

```
if (!inherits(link, "linkfun")) {  
  link <- match.arg(link)  
  link <- do.call(link, list())  
}  
◇
```

Fragment referenced in [?](#), [?](#), [?](#), [?](#).

We use the positive residuals for defining a permutation test with treatment effect coding using the first group as control, that is, the test statistic is defined through the sum of the positive residuals in all but the control group. Unfortunately, most `stats::*.test` procedures use the second group as control, so factors need to be releveled to obtain identical results (this is relevant for the one-sided case).

< freeway generic and table method (main workhorse) ? > \equiv

```

freeway <- function(y, ...)
  UseMethod("freeway")

freeway.table <- function(y, link = c("logit", "probit", "cloglog", "loglog"),
  mu = 0, B = 0, exact = FALSE, ...)
{
  cl <- match.call()

  d <- dim(y)
  dn <- dimnames(y)
  DNAME <- NULL
  if (!is.null(dn)) {
    DNAME <- paste(names(dn)[1], "by", names(dn)[2],
      paste0("(", paste0(dn[2], collapse = ", "), ")"))
    if (length(dn) == 3L)
      DNAME <- paste(DNAME, "\n\t stratified by", names(dn)[3])
  }

  < link2fun ? >

  if (!(length(mu) == 1L || length(mu) == d[2L] - 1L)) {
    warning(gettextf("incompatible length of argument 'mu' in %s",
      "freeway"),
      domain = NA)
    mu <- rep(mu, length.out = d[2L] - 1L)
  }

  ret <- .freewayML(y, link = link, mu = mu, ...)
  ret$link <- link
  ret$data.name <- DNAME
  ret$call <- cl

  < freeway permutation tests ? >

  if (ret$MPL_Jeffreys)
    ret$method <- paste(ret$method,
      "with Jeffreys prior penalisation", sep = ", ")

  class(ret) <- "freeway"
  return(ret)
}

```

Fragment referenced in [?](#).

where preparations for permutations tests are performed before returning the object

<free1way permutation tests ?> \equiv

```
alias <- link$alias
if (length(link$alias) == 2L) alias <- alias[1L + (d[2] > 2L)]
stratified <- FALSE
if (length(d) == 3L) stratified <- d[3L] > 1
ret$method <- paste(ifelse(stratified, "Stratified", ""),
                    paste0(d[2L], "-sample"), alias,
                    "test against", link$model, "alternatives")

cf <- ret$par
### compute the permutation distribution always
### for H0: delta = 0, not delta = mu
### otherwise, permutation confidence intervals
### are not aligned with permutation p-values
cf[idx <- seq_len(d[2L] - 1L)] <- -mu
pr <- ret$profile(cf, idx)
res <- -pr$negresiduals
if (d[2L] == 2L)
  res <- res / sqrt(c(pr$hessian))

<Strasser Weber ?>
<resampling ?>

if (length(dim(y)) == 3L) y <- y[,ret$strata, drop = FALSE]
if (length(dim(y)) == 4L) {
  y <- y[,ret$strata,, drop = FALSE]
  dy <- dim(y)
  dy[1] <- dy[1] * 2
  y <- apply(y, 3, function(x) rbind(x[, "TRUE"], x[, "FALSE"]), simplify = FALSE)
  y <- array(unlist(y), dim = dy[1:3])
}

### exact two-sample Wilcoxon w/o stratification
if (exact) {
  if (!stratified && link$model == "proportional odds" && d[2L] == 2L) {
    <exact proportional odds ?>
    ret$exact <- .exact(c(res, res), grp = unclass(gl(2, d[1L])) - 1L,
                      w = c(y))

    B <- 0
  } else {
    warning(gettextf("cannot compute exact permutation distribution in %s",
                    "free1way"),
           domain = NA)
  }
}
ret$perm <- .resample(res, y, B = B)
◇
```

Fragment referenced in [?](#).

The `formula` method allows formulae

```
> y ~ groups / blocks
```

for model specification. We start handling the formula

$\langle \text{formula business ?} \rangle \equiv$

```

if(missing(formula) || (length(formula) != 3L))
  stop("'formula' missing or incorrect")

if (stratum <- (length(formula[[3L]]) > 1)) {
  if ((length(formula[[3L]]) != 3L) ||
      (formula[[3L]][[1L]] != as.name("|")) ||
      (length(formula[[3L]][[2L]]) != 1L) ||
      (length(formula[[3L]][[3L]]) != 1L))
    stop(gettextf("incorrect specification for '%s'", "formula"),
         domain = NA)
  formula[[3L]][[1L]] <- as.name("+")
}

formula <- terms(formula)
if (length(attr(formula, "term.labels")) > 1L + stratum)
  stop("'formula' missing or incorrect")
group <- attr(formula, "term.labels")[1L]

m <- match.call(expand.dots = FALSE)
m$formula <- formula
if (is.matrix(eval(m$data, parent.frame()))))
  m$data <- as.data.frame(data)
## need stats:: for non-standard evaluation
m[[1L]] <- quote(stats::model.frame)
m$... <- NULL
mf <- eval(m, parent.frame())

```

Fragment referenced in [?](#).

$\langle \text{freelway formula ?} \rangle \equiv$

```

freelway.formula <- function(formula, data, weights, subset, na.action = na.pass,
                             event = NULL, ...)
{
  cl <- match.call()

   $\langle \text{formula business ?} \rangle$ 

  response <- attr(attr(mf, "terms"), "response")
  DNAME <- paste(vn <- c(names(mf)[response], group),
                 collapse = " by ") # works in all cases
  w <- as.vector(model.weights(mf))
  y <- mf[[response]]
  if (inherits(y, "Surv")) {
    if (!is.null(event))
      stop(gettextf("cannot have both a 'Surv()' response and an 'event' argument in %s",
                    "freelway"),
           domain = NA)
    if (attr(y, "type") != "right")
      stop(gettextf("%s currently only allows independent right-censoring",
                    "freelway"),
           domain = NA)
    event <- (y[,2] > 0)
    y <- y[,1]
  }
  g <- factor(mf[[group]])
  mf[[group]] <- g
  lev <- levels(g)
  DNAME <- paste(DNAME, paste0("(", paste0(lev, collapse = ", "), ")"))
  if (nlevels(g) < 2L)
    stop(gettextf("incorrect argument 'groups' in %s: at least two groups needed",
                  "freelway"),
         domain = NA)
  if (stratum) {
    st <- factor(mf[[3L]])
    mf[[3L]] <- st
    ### nlevels(st) == 1L is explicitly allowed
    vn <- c(vn, names(mf)[3L])
    RVAL <- freelway(y = y, groups = g, blocks = st, event = event,
                    weights = w, varnames = vn, ...)
    DNAME <- paste(DNAME, paste("\n\t stratified by", names(mf)[3L]))
  } else {
    ## Call the corresponding method
    RVAL <- freelway(y = y, groups = g, event = event, weights = w,
                    varnames = vn, ...)
  }
  RVAL$data <- mf
  RVAL$data.name <- DNAME
  RVAL$call <- cl
  RVAL
}

```

Fragment referenced in [?](#).

The method for numeric outcomes provides a discretisation at the unique observed outcome values, or (for very large sample sizes), for binned outcomes. The `event` argument is a logical where TRUE is interpreted as an event and FALSE as right-censored observation

< variable names and checks ? > \equiv

```
cl <- match.call()
if (is.null(varnames))
  varnames <- c(deparse1(substitute(y)),
                deparse1(substitute(groups)),
                deparse1(substitute(blocks)))

DNAME <- paste(varnames[1], "by", varnames[2])
groups <- factor(groups)
if (nlevels(groups) < 2L)
  stop(gettextf("incorrect argument 'groups' in %s: at least two groups needed",
                "free1way"),
       domain = NA)
DNAME <- paste(DNAME, paste0("(", paste0(levels(groups), collapse = ", "),
                               ")"))

if (!is.null(blocks)) {
  if (length(unique(blocks)) < 2L) {
    blocks <- NULL
  } else {
    blocks <- factor(blocks)
    DNAME <- paste(DNAME, "\n\t stratified by", varnames[3])
  }
}
varnames <- varnames[varnames != "NULL"]
◇
```

Fragment referenced in [?](#), [?](#).

Note that the return value of `unique` might differ between platforms. Because users can decide about the unique values in the vector `y` (by using `round` or `trunc`, for example), before calling this function, we refrain from handling this issue internally. However, we offer an `nbins` argument for binning response observations at sample quantiles in the absence of right-censoring. Note that we ignore the blocks when calling `cut`. This is inefficient for many blocks with non-overlapping support of the outcome distribution, as large sparse tables are resulting. We remove the corresponding elements from the first dimension of such a table later on (in `.free1wayML`). The reason for this inconvenience is that all the data going into `.free1wayML` can be stored as a `table` (and not as a list of things).

<free1way numeric ?> \equiv

```
free1way.numeric <- function(y, groups, blocks = NULL, event = NULL,
                             weights = NULL, nbins = 0, varnames = NULL, ...)
{
  <variable names and checks ?>

  if (!is.null(event)) {
    if (!is.logical(event))
      stop(gettextf("%s currently only allows independent right-censoring",
                    "free1way"),
           domain = NA)
    uy <- sort(unique(y[event]))
  } else {
    uy <- sort(unique(y))
  }
  if (nbins && nbins < length(uy) && is.null(event)) {
    nbins <- ceiling(nbins)
    breaks <- c(-Inf, quantile(y, probs = seq_len(nbins) / (nbins + 1L)),
               Inf)
  } else {
    breaks <- c(-Inf, uy, Inf)
  }
  r <- ordered(cut(y, breaks = breaks, ordered_result = TRUE,
                  labels = FALSE)) ### avoids costly formatC call
  RVAL <- free1way(y = r, groups = groups, blocks = blocks,
                 event = event, weights = weights,
                 varnames = varnames, ...)
  RVAL$data.name <- DNAME
  RVAL$call <- c1
  RVAL
}
```

◇

Fragment referenced in [?](#).

The **factor** method also allows right-censoring but otherwise is just a call to **xtabs**:

$\langle \text{free1way factor ?} \rangle \equiv$

```
free1way.factor <- function(y, groups, blocks = NULL, event = NULL,
                           weights = NULL, varnames = NULL, ...)
{
   $\langle \text{variable names and checks ?} \rangle$ 

  if (nlevels(y) > 2L && !is.ordered(y))
    stop(gettextf("%s is not defined for unordered responses",
                  "free1way"),
         domain = NA)
  d <- data.frame(w = 1, y = y, groups = groups)
  if (!is.null(weights)) d$w <- weights
  if (is.null(blocks)) blocks <- gl(1, nrow(d))
  d$blocks <- blocks
  if (!is.null(event)) {
    if (!is.logical(event))
      stop(gettextf("%s currently only allows independent right-censoring",
                    "free1way"),
           domain = NA)
    d$event <- factor(event, levels = c(FALSE, TRUE),
                      labels = c("FALSE", "TRUE"))
  }
  tab <- xtabs(w ~ ., data = d)
  dn <- dimnames(tab)
  names(dn)[seq_along(varnames)] <- varnames
  dimnames(tab) <- dn
  RVAL <- free1way(tab, ...)
  RVAL$data.name <- DNAME
  RVAL$call <- cl
  RVAL
}
◇
```

Fragment referenced in [?](#).

8.2 free1way Methods

We start with `coef`, `vcov`, and `model.frame/model.matrix` methods such that multiple comparison procedures from **multcomp** will work out of the box. The `coef` method allows to obtain effects at alternative scales: probabilistic indices ($\text{AUC} = \text{PI}$) or the overlap coefficient:

$\langle \text{free1way methods?} \rangle \equiv$

```

coef.free1way <- function(object, what = c("shift", "PI", "AUC", "OVL"), ...)
{
  what <- match.arg(what)
  cf <- object$coefficients
  return(switch(what, "shift" = cf,
                "PI" = object$link$parm2PI(cf),
                "AUC" = object$link$parm2PI(cf),      ### same as PI
                "OVL" = object$link$parm2OVL(cf)))
}
vcov.free1way <- function(object, ...)
  object$vcov
logLik.free1way <- function(object, ...)
  -object$value
model.frame.free1way <- function(formula, ...) {
  if (!is.null(formula[["data"]])) return(formula[["data"]])
  ret <- as.data.frame(formula$table)
  ret <- ret[rep(seq_len(nrow(ret)), ret$Freq),,drop = FALSE]
  ret
}
### the next two might go into multcomp
terms.free1way <- function(x, ...)
{
  mf <- model.frame(x)
  terms(as.formula(paste(names(mf)[1:2], collapse = "~")),
        data = mf)
}
model.matrix.free1way <- function (object, ...)
{
  mf <- model.frame(object)
  tm <- terms(object)
  mm <- model.matrix(delete.response(tm), data = mf)
  at <- attributes(mm)
  mm <- mm[, -1]
  at$dim[2] <- at$dim[2] - 1
  at$dimnames[[2]] <- at$dimnames[[2]][-1]
  at$assign <- at$assign[-1]
  attributes(mm) <- at
  mm
}
◇

```

Fragment referenced in [?](#).

We use the `print` method to report different test statistics and corresponding p -values via the `test` and `alternative` arguments. The reason for doing so is that the parameter estimation only needs to be performed once in cases users are interested in different tests or (see below) confidence intervals. By default, an asymptotic permutation test is performed, mainly because the p -values coincide with some special cases (`wilcox`, `kruskal`, `friedman.test`):

< free1way print ? > \equiv

```
.print.free1way <- function(x, test = c("Permutation", "Wald", "LRT", "Rao"),
                             alternative = c("two.sided", "less", "greater"),
                             tol = sqrt(.Machine$double.eps),
                             mu = 0, ### allow permutation testing non-null hypotheses
                             ### in alignment with confint(free1way(, B > 0))
                             ...)
{
  test <- match.arg(test)
  alternative <- match.arg(alternative)

  ### global
  cf <- coef(x)
  if ((length(cf) > 1L || test == "LRT") && alternative != "two.sided")
    stop(gettextf("cannot compute one-sided p-values in %s",
                  "free1way"),
         domain = NA)

  DF <- NULL
  parm <- seq_along(cf)
  value <- mu

  < statistics ? >

  if (test == "Permutation") {
    < Permutation p-values ? >
  }

  RVAL <- list(statistic = STATISTIC, parameter = DF, p.value = PVAL,
               null.value = x$mu, alternative = alternative, method = x$method,
               data.name = x$data.name)
  class(RVAL) <- "htest"
  return(RVAL)
}

print.free1way <- function(x, ...)
{
  print(ret <- .print.free1way(x, ...))
  return(invisible(x))
}
◇
```

Fragment referenced in [?](#).

The `summary` method performs population Wald inference unless the `test` argument is specified:

$\langle \text{free1way summary ?} \rangle \equiv$

```
summary.free1way <- function(object, test,
                             alternative = c("two.sided", "less", "greater"),
                             tol = .Machine$double.eps, ...)
{
  if (!missing(test))
    return(.print.free1way(object, test = test,
                           alternative = alternative, tol = tol, ...))

  alternative <- match.arg(alternative)

  ESTIMATE <- coef(object)
  SE <- sqrt(diag(vcov(object)))
  STATISTIC <- unname(ESTIMATE / SE)
  if (alternative == "less") {
    PVAL <- pnorm(STATISTIC)
  } else if (alternative == "greater") {
    PVAL <- pnorm(STATISTIC, lower.tail = FALSE)
  } else {
    PVAL <- 2 * pnorm(-abs(STATISTIC))
  }
  cformat <- cbind(ESTIMATE, SE, STATISTIC, PVAL)
  colnames(cformat) <- c(object$link$parm, "Std. Error", "z value",
                         switch(alternative, "two.sided" = "P(>|z|)",
                                "less" = "P(<z)",
                                "greater" = "P(>z)"))
  ret <- list(call = object$call, coefficients = cformat)
  class(ret) <- "summary.free1way"
  return(ret)
}

.print.summary.free1way <- function(x, ...)
{
  cat("\nCall:\n", paste(deparse(x$call), sep = "\n", collapse = "\n"),
      "\n\n", sep = "")
  cat("Coefficients:\n")
  printCofmat(x$coefficients)
}
◇
```

Fragment referenced in [?](#).

Confidence intervals are computed by inversion of the corresponding test statistics. Because LRT and Rao confidence intervals are invariant with respect to transformations, proper LRT or Rao confidence intervals for probabilistic indices or overlap coefficients can also be computed.

We begin computing the critical values for permutation tests, making sure the confidence intervals will be in line with one- and two-sided p -values:

$\langle \text{permutation confint?} \rangle \equiv$

```

if (length(cf) > 1L)
  stop(gettextf("permutation confidence intervals only available for 2-sample comparisons in %s",
    "confint.freelway"),
    domain = NA)
if (!is.null(object$exact)) {
  qu <- c(object$exact$qle(1 - conf.level),
    object$exact$qgr(1 - conf.level))
} else {
  if (is.null(object$perm$permStat)) {
    qu <- qnorm(conf.level) * c(-1, 1)
  } else {
    .pq <- function(s, alpha)
    {
      su <- sort(unique(s))
      ### F = P(T <= t), S = P(T >= t)
      Fs <- cumsum(st <- table(match(s, su)))
      Ss <- length(s) - Fs + st
      c(max(su[Fs <= alpha * length(s)]),
        min(su[Ss <= alpha * length(s)]))
    }
    ### cf PVAL computation!!!
    rs <- object$perm$permStat
    qu <- .pq(round(rs, 10), alpha = 1 - conf.level)
    att.level <- mean(rs > qu[1] & rs < qu[2])
    attr(CINT, "Attained level") <- att.level
  }
}

```

Fragment referenced in [?](#).

The `confint` method starts with Wald intervals, which are either returned or used as starting values for the inversion. We start with the lower bound. Sometimes (for example in case of complete separation), the information is very low so the Wald intervals are extremely wide and the profile log-likelihood cannot be computed. So we try to make the starting interval wider in a step-wise manner. However, this may still fail and we thus exit gently, returning NA and issuing a warning.

$\langle \text{confint lower?} \rangle \equiv$

```

CINT[p,1] <- max(CINT[p, 1], cf[p] - 1)
sdlwr <- sign(sfun(cf[p], parm = p, quantile = qu[2]))
slwr <- try(sfun(CINT[p,1], parm = p, quantile = qu[2]))
k <- 1
if (inherits(slwr, "try-error")) {
  CINT[p,1] <- NA
} else {
  while ((is.na(slwr) ||
    sign(slwr) == sdlwr) && k < 30) {
    CINT[p,1] <- CINT[p,1] - 1
    slwr <- try(sfun(CINT[p,1], parm = p, quantile = qu[2]))
    if (inherits(slwr, "try-error")) {
      CINT[p,1] <- NA
      break()
    }
    k <- k + 1
  }
}
if (k == 30) {
  CINT[p,1] <- NA
} else {
  lwr <- try(uniroot(sfun, interval = c(CINT[p,1], cf[p]),
    parm = p, quantile = qu[2])$root)
  if (inherits(lwr, "try-error")) {
    CINT[p,1] <- NA
  } else {
    CINT[p,1] <- lwr
  }
}
if (is.na(CINT[p,1]))
  warning(gettextf("failed to compute confidence interval in %s",
    "confint.freelway"),
    domain = NA)

```

◇

Fragment referenced in [?](#).

The upper bound works in the very same way.

$\langle \text{confint upper ?} \rangle \equiv$

```

CINT[p,2] <- min(CINT[p, 2], cf[p] + 1)
sdupr <- sign(sfun(cf[p], parm = p, quantile = qu[1]))
supr <- try(sfun(CINT[p,2], parm = p, quantile = qu[1]))
k <- 1
if (inherits(supr, "try-error")) {
  CINT[p,2] <- NA
} else {
  while ((is.na(supr) ||
    sign(supr) == sdupr) && k < 30) {
    CINT[p,2] <- CINT[p,2] + 1
    supr <- try(sfun(CINT[p,2], parm = p, quantile = qu[1]))
    if (inherits(supr, "try-error")) {
      CINT[p,2] <- NA
      break()
    }
    k <- k + 1
  }
}
if (k == 30) {
  CINT[p,2] <- NA
} else {
  upr <- try(uniroot(sfun, interval = c(cf[p], CINT[p, 2]),
    parm = p, quantile = qu[1])$root)
  if (inherits(upr, "try-error")) {
    CINT[p, 2] <- NA
  } else {
    CINT[p, 2] <- upr
  }
}
if (is.na(CINT[p,2]))
  warning(gettextf("failed to compute confidence interval in %s",
    "confint.freelway"),
    domain = NA)

```

◇

Fragment referenced in [?](#).

$\langle \text{free1way confint ?} \rangle \equiv$

```

confint.free1way <- function(object, parm,
  level = .95, test = c("Permutation", "Wald", "LRT", "Rao"),
  what = c("shift", "PI", "AUC", "OVL"), ...)
{
  test <- match.arg(test)
  conf.level <- 1 - (1 - level) / 2

  cf <- coef(object)
  if (missing(parm))
    parm <- seq_along(cf)

  CINT <- confint.default(object, level = level)
  if (test != "Wald") {
    wlevel <- level
    wlevel <- 1 - (1 - level) / 2
    CINT[] <- confint.default(object, level = wlevel)

    sfun <- function(value, parm, quantile)
    {
      x <- object
      alternative <- "two.sided"
      tol <- .Machine$double.eps
       $\langle \text{statistics ?} \rangle$ 
      ### we also could invert p-values, but the
      ### p-value function might be discrete for permutation
      ### tests, in contrast to the test statistic
      return(STATISTIC - quantile)
    }

    if (test == "Permutation") {
       $\langle \text{permutation confint ?} \rangle$ 
    } else {
      qu <- rep.int(qchisq(level, df = 1), 2) ### always two.sided
    }

    for (p in parm) {
       $\langle \text{confint lower ?} \rangle$ 
       $\langle \text{confint upper ?} \rangle$ 
    }
  }

  what <- match.arg(what)
  CINT <- switch(what, "shift" = CINT,
    "PI" = object$link$parm2PI(CINT),
    "AUC" = object$link$parm2PI(CINT), ### same as PI
    "OVL" = object$link$parm2OVL(CINT))

  return(CINT)
}

```

Fragment referenced in [?](#).

As an example, we compute log-odds ratios for the table introduced above and report some tests and confidence intervals:

```

> x
, , A
A B C

```



```

A 5 6 4
B 3 5 7
C 3 4 5

, , B

  A B C
A 3 5 6
B 0 0 0
C 4 6 5

> ### asymptotic permutation test
> (ft <- free1way(x))

      Stratified 3-sample Kruskal-Wallis test against proportional odds
      alternatives

data:   by   (c("A", "B", "C"))
       stratified by
Perm chi-squared = 0.145, df = 2, p-value = 0.93
alternative hypothesis: true log-odds ratio is not equal to 0

> coef(ft)

      B      C
0.052627 0.204750

> vcov(ft)

      B      C
B 0.34916 0.20834
C 0.20834 0.33850

> ### Wald per parameter
> summary(ft)

Call:
free1way.table(y = x)

Coefficients:
  log-odds ratio Std. Error z value P(>|z|)
B      0.0526      0.5909  0.0891   0.93
C      0.2047      0.5818  0.3519   0.72

> library("multcomp")
> summary(glht(ft), test = univariate())

      Simultaneous Tests for General Linear Hypotheses

Fit: free1way.table(y = x)

Linear Hypotheses:
      Estimate Std. Error z value Pr(>|z|)
B == 0   0.0526      0.5909   0.09   0.93
C == 0   0.2047      0.5818   0.35   0.72
(Univariate p values reported)

> ### global Wald
> summary(ft, test = "Wald")

```

```

Stratified 3-sample Kruskal-Wallis test against proportional odds
alternatives

data:  by (c("A", "B", "C"))
       stratified by
Wald chi-squared = 0.148, df = 2, p-value = 0.93
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(glht(ft), test = Chisqtest())

General Linear Hypotheses

Linear Hypotheses:
      Estimate
B == 0  0.0526
C == 0  0.2047

Global Test:
      Chisq DF Pr(>Chisq)
1 0.148  2      0.929

> ### Rao score, Permutation score, LRT
> summary(ft, test = "Rao")

Stratified 3-sample Kruskal-Wallis test against proportional odds
alternatives

data:  by (c("A", "B", "C"))
       stratified by
Rao chi-squared = 0.148, df = 2, p-value = 0.93
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft, test = "Permutation")

Stratified 3-sample Kruskal-Wallis test against proportional odds
alternatives

data:  by (c("A", "B", "C"))
       stratified by
Perm chi-squared = 0.145, df = 2, p-value = 0.93
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft, test = "LRT")

Stratified 3-sample Kruskal-Wallis test against proportional odds
alternatives

data:  by (c("A", "B", "C"))
       stratified by
logLR chi-squared = 0.148, df = 2, p-value = 0.93
alternative hypothesis: true log-odds ratio is not equal to 0

> ### Wald confidence intervals, unadjusted
> confint(glht(ft), calpha = univariate_calpha())

Simultaneous Confidence Intervals

Fit: free1way.table(y = x)

Quantile = 1.96
95% confidence level

```

```

Linear Hypotheses:
      Estimate lwr      upr
B == 0  0.0526  -1.1055  1.2108
C == 0  0.2047  -0.9356  1.3451

> confint(ft, test = "Wald")

      2.5 % 97.5 %
B -1.10552 1.2108
C -0.93558 1.3451

> ### Rao and LRT intervals
> confint(ft, test = "Rao")

      2.5 % 97.5 %
B -1.09075 1.1954
C -0.92072 1.3301

> confint(ft, test = "LRT")

      2.5 % 97.5 %
B -1.1113 1.2205
C -0.9387 1.3573

```

Chapter 9

Special Test Procedures

We now demonstrate that `free1way` produces the exact same results as some of the classical test procedures implemented in the `stats` package, and how the new implementation extends the existing functionality.

9.1 Wilcoxon Test

The first example is a Wilcoxon test for a single log-odds ratio comparing to treatment groups. The Wilcoxon test is the score test in a 2-sample proportional odds model

```
> N <- 25
> w <- gl(2, N)
> y <- rlogis(length(w), location = c(0, 1)[w])
> #### link = logit is default
> ft <- free1way(y ~ w)
> ### Wald
> summary(ft)

Call:
free1way.formula(formula = y ~ w)

Coefficients:
      log-odds ratio Std. Error z value P(>|z|)
w2           1.290      0.524    2.462    0.01

> ### Permutation test
> wilcox.test(y ~ w, alternative = "greater", correct = FALSE, exact = FALSE)$p.value

[1] 0.99215

> pvalue(wilcox_test(y ~ w, alternative = "greater"))

[1] 0.99215

> summary(ft, test = "Permutation", alternative = "less")$p.value

Perm Z
0.99215

> wilcox.test(y ~ w, alternative = "less", correct = FALSE, exact = FALSE)$p.value

[1] 0.0078535

> pvalue(wilcox_test(y ~ w, alternative = "less"))

[1] 0.0078535
```

```

> summary(ft, test = "Permutation", alternative = "greater")$p.value

    Perm Z
0.0078535

> wilcox.test(y ~ w, correct = FALSE, exact = FALSE)$p.value

[1] 0.015707

> kruskal.test(y ~ w)$p.value

[1] 0.015707

> pvalue(wilcox_test(y ~ w))

[1] 0.015707

> summary(ft, test = "Permutation")$p.value

    Perm Z
0.015707

> ### Wald tests
> summary(ft, test = "Wald", alternative = "less")

    2-sample Wilcoxon test against proportional odds alternatives

data:  y by w (1, 2)
Wald Z = 2.46, p-value = 0.99
alternative hypothesis: true log-odds ratio is less than 0

> summary(ft, test = "Wald", alternative = "greater")

    2-sample Wilcoxon test against proportional odds alternatives

data:  y by w (1, 2)
Wald Z = 2.46, p-value = 0.0069
alternative hypothesis: true log-odds ratio is greater than 0

> summary(ft, test = "Wald")

    2-sample Wilcoxon test against proportional odds alternatives

data:  y by w (1, 2)
Wald chi-squared = 6.06, df = 1, p-value = 0.014
alternative hypothesis: true log-odds ratio is not equal to 0

> ### Rao score tests
> summary(ft, test = "Rao", alternative = "less")

    2-sample Wilcoxon test against proportional odds alternatives

data:  y by w (1, 2)
Rao Z = 2.5, p-value = 0.99
alternative hypothesis: true log-odds ratio is less than 0

> summary(ft, test = "Rao", alternative = "greater")

    2-sample Wilcoxon test against proportional odds alternatives

data:  y by w (1, 2)
Rao Z = 2.5, p-value = 0.0063
alternative hypothesis: true log-odds ratio is greater than 0

```

```

> summary(ft, test = "Rao")

      2-sample Wilcoxon test against proportional odds alternatives

data:  y by w (1, 2)
Rao chi-squared = 6.24, df = 1, p-value = 0.013
alternative hypothesis: true log-odds ratio is not equal to 0

> ### LRT (only two-sided)
> summary(ft, test = "LRT")

      2-sample Wilcoxon test against proportional odds alternatives

data:  y by w (1, 2)
logLR chi-squared = 6.33, df = 1, p-value = 0.012
alternative hypothesis: true log-odds ratio is not equal to 0

> ### confidence intervals for log-odds ratios
> confint(ft, test = "Permutation")

      2.5 % 97.5 %
w2 0.23829 2.3423

> confint(ft, test = "LRT")

      2.5 % 97.5 %
w2 0.28151 2.3446

> confint(ft, test = "Wald")

      2.5 % 97.5 %
w2 0.26308 2.3168

> confint(ft, test = "Rao")

      2.5 % 97.5 %
w2 0.27214 2.3084

> ### confidence interval for "Wilcoxon Parameter" = PI = AUC
> confint(ft, test = "Rao", what = "AUC")

      2.5 % 97.5 %
w2 0.54524 0.82742

> ### comparison with rms::orm
> library("rms")
> rev(coef(or <- orm(y ~ w)))[1]

w=2
1.29

> coef(ft)

w2
1.29

> logLik(or)

'log Lik.' -192.44 (df=50)

> logLik(ft)

[1] -192.44

```

```

> vcov(or)[2,2]

[1] 0.2745

> vcov(ft)

      w2
w2 0.2745

> ci <- confint(or)
> ci[nrow(ci),]

      2.5 % 97.5 %
0.26308 2.31683

> confint(ft, test = "Wald")

      2.5 % 97.5 %
w2 0.26308 2.3168

```

9.2 Mantel-Haenszel Test

The Cochran-Mantel-Haenszel test for conditional independence in 2×2 tables also relies on a proportional odds model.

```

> example(mantelhaen.test, echo = FALSE)
> mantelhaen.test(UCBAdmissions, correct = FALSE)

      Mantel-Haenszel chi-squared test without continuity correction

data:  UCBAdmissions
Mantel-Haenszel X-squared = 1.52, df = 1, p-value = 0.22
alternative hypothesis: true common odds ratio is not equal to 1
95 percent confidence interval:
 0.77191 1.06033
sample estimates:
common odds ratio
      0.9047

> ft <- free1way(UCBAdmissions)
> summary(ft, test = "Wald")

      Stratified 2-sample Wilcoxon test against proportional odds
      alternatives

data:  Admit by Gender (c("Male", "Female"))
      stratified by Dept
Wald chi-squared = 1.53, df = 1, p-value = 0.22
alternative hypothesis: true log-odds ratio is not equal to 0

> exp(coef(ft))

GenderFemale
      0.90495

> exp(confint(ft, test = "Wald"))

      2.5 % 97.5 %
GenderFemale 0.77234 1.0603

> exp(sapply(dimnames(UCBAdmissions)[[3L]], function(dept)
+      confint(free1way(UCBAdmissions[,dept]), test = "Permutation")))

```

```

      A      B      C      D      E      F
[1,] 0.20967 0.34796 0.8547 0.68644 0.82584 0.4583
[2,] 0.58184 1.85295 1.5021 1.23646 1.80738 1.4956

> sapply(dimnames(UCBAdmissions)[[3L]], function(dept)
+       fisher.test(UCBAdmissions[,dept], conf.int = TRUE)$conf.int)

      A      B      C      D      E      F
[1,] 0.19704 0.2945 0.84522 0.67896 0.80648 0.43329
[2,] 0.59204 2.0040 1.51629 1.25047 1.83852 1.57563

```

9.3 prop.test

For a single 2×2 table, all tests are nonparametric (as the model is saturated) and therefore also inference procedures result in the same p -values, for example.

```

> prop.test(UCBAdmissions[,1], correct = FALSE)

2-sample test for equality of proportions without continuity correction

data:  UCBAdmissions[, , 1]
X-squared = 17.2, df = 1, p-value = 3.3e-05
alternative hypothesis: two.sided
95 percent confidence interval:
 -0.128682 -0.053034
sample estimates:
 prop 1  prop 2
0.85191 0.94277

> summary(free1way(UCBAdmissions[,1]), test = "Rao")

2-sample Wilcoxon test against proportional odds alternatives

data:  Admit by Gender (c("Male", "Female"))
Rao chi-squared = 17.2, df = 1, p-value = 3.3e-05
alternative hypothesis: true log-odds ratio is not equal to 0

```

9.4 Kruskal-Wallis Test

The Kruskal-Wallis test is the score test in a K -sample proportional odds model

```

> example(kruskal.test, echo = FALSE)
> kruskal.test(x ~ g)

Kruskal-Wallis rank sum test

data:  x by g
Kruskal-Wallis chi-squared = 0.771, df = 2, p-value = 0.68

> free1way(x ~ g)

3-sample Kruskal-Wallis test against proportional odds alternatives

data:  x by g (Normal subjects, Subjects with obstructive airway disease, Subjects with asbestosis)
Perm chi-squared = 0.771, df = 2, p-value = 0.68
alternative hypothesis: true log-odds ratio is not equal to 0

```


9.5 Savage Test

The Savage test assumes proportional odds and, consequently, is the score test in a proportional odds model. We start without censoring (Savage test) and add strata

```
> library("survival")
> N <- 10
> nd <- expand.grid(g = gl(3, N), s = gl(3, N))
> nd$tm <- rexp(nrow(nd))
> nd$ev <- TRUE
> cm <- coxph(Surv(tm, ev) ~ g + strata(s), data = nd)
> (ft <- freeWay(tm ~ g | s, data = nd, link = "cloglog"))

      Stratified 3-sample Savage test against proportional hazards
      alternatives

data:  tm by g (1, 2, 3)
      stratified by s
Perm chi-squared = 4.68, df = 2, p-value = 0.096
alternative hypothesis: true log-hazard ratio is not equal to 0

> coef(cm)

      g2      g3
-0.15981 -0.16021

> coef(ft)

      g2      g3
0.16038 0.16058

> vcov(cm)

      g2      g3
g2 0.0067826 0.0034163
g3 0.0034163 0.0067785

> vcov(ft)

      g2      g3
g2 0.0067888 0.0034204
g3 0.0034204 0.0067851

> ### Rao score tests
> summary(cm)$sctest

      test      df  pvalue
5.03230 2.00000 0.08077

> summary(ft, test = "Rao")

      Stratified 3-sample Savage test against proportional hazards
      alternatives

data:  tm by g (1, 2, 3)
      stratified by s
Rao chi-squared = 5.06, df = 2, p-value = 0.08
alternative hypothesis: true log-hazard ratio is not equal to 0

> ### likelihood ratio tests
> summary(cm)$logtest
```

```

      test      df    pvalue
4.933154 2.000000 0.084875

> summary(ft, test = "LRT")

      Stratified 3-sample Savage test against proportional hazards
      alternatives

data:  tm by g (1, 2, 3)
      stratified by s
logLR chi-squared = 4.96, df = 2, p-value = 0.084
alternative hypothesis: true log-hazard ratio is not equal to 0

> ### Wald tests
> summary(cm)$waldtest

      test      df    pvalue
5.020000 2.000000 0.081193

> summary(ft, test = "Wald")

      Stratified 3-sample Savage test against proportional hazards
      alternatives

data:  tm by g (1, 2, 3)
      stratified by s
Wald chi-squared = 5.05, df = 2, p-value = 0.08
alternative hypothesis: true log-hazard ratio is not equal to 0

> ### asymptotic permutation tests
> survdiff(Surv(tm, ev) ~ g + strata(s), data = nd, rho = 0)[c("chisq", "pvalue")]

$chisq
[1] 5.0323

$pvalue
[1] 0.08077

> summary(ft, test = "Permutation")

      Stratified 3-sample Savage test against proportional hazards
      alternatives

data:  tm by g (1, 2, 3)
      stratified by s
Perm chi-squared = 4.68, df = 2, p-value = 0.096
alternative hypothesis: true log-hazard ratio is not equal to 0

> library("coin")
> independence_test(Surv(tm, ev) ~ g | s, data = nd, ytrafo = function(...)
+                   trafo(..., numeric_trafo = logrank_trafo, block = nd$s),
+                   teststat = "quad")

      Asymptotic General Independence Test

data:  Surv(tm, ev) by g (1, 2, 3)
      stratified by s
chi-squared = 4.75, df = 2, p-value = 0.093

      Wilcoxon against proportional odds

> survdiff(Surv(tm, ev) ~ g + strata(s), data = nd, rho = 1)[c("chisq", "pvalue")]

```

```

$chisq
[1] 5.3318

$pvalue
[1] 0.069535

> (ft <- free1way(tm ~ g | s, data = nd, link = "logit"))
      Stratified 3-sample Kruskal-Wallis test against proportional odds
      alternatives

data:  tm by g (1, 2, 3)
      stratified by s
Perm chi-squared = 5.2, df = 2, p-value = 0.074
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft)

Call:
free1way.formula(formula = tm ~ g | s, data = nd, link = "logit")

Coefficients:
      log-odds ratio Std. Error z value P(>|z|)
g2           0.301      0.142    2.119    0.03
g3           0.261      0.142    1.830    0.07

> summary(ft, test = "Rao")
      Stratified 3-sample Kruskal-Wallis test against proportional odds
      alternatives

data:  tm by g (1, 2, 3)
      stratified by s
Rao chi-squared = 5.27, df = 2, p-value = 0.072
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft, test = "LRT")
      Stratified 3-sample Kruskal-Wallis test against proportional odds
      alternatives

data:  tm by g (1, 2, 3)
      stratified by s
logLR chi-squared = 5.27, df = 2, p-value = 0.072
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft, test = "Wald")
      Stratified 3-sample Kruskal-Wallis test against proportional odds
      alternatives

data:  tm by g (1, 2, 3)
      stratified by s
Wald chi-squared = 5.26, df = 2, p-value = 0.072
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft, test = "Permutation")
      Stratified 3-sample Kruskal-Wallis test against proportional odds
      alternatives

data:  tm by g (1, 2, 3)
      stratified by s
Perm chi-squared = 5.2, df = 2, p-value = 0.074
alternative hypothesis: true log-odds ratio is not equal to 0

```

9.6 Log-rank Test

And now with censoring. We cannot expect this to be identical with what **survival** reports, as this package is based on the partial likelihood and we operate on the full likelihood.

```
> library("survival")
> data("GBSG2", package = "TH.data")
> cm <- coxph(Surv(time, cens) ~ horTh + strata(tgrade), data = GBSG2)
> ft <- with(GBSG2, flex1way(Surv(time, cens) ~ horTh | tgrade,
+                             link = "cloglog"))
> coef(cm)

horThyes
-0.33972

> coef(ft)

horThyes
 0.34056

> vcov(cm)

           horThyes
horThyes 0.015754

> vcov(ft)

           horThyes
horThyes 0.015751

> ### Rao score tests
> summary(cm)$sctest

      test      df    pvalue
7.3935325 1.0000000 0.0065459

> summary(ft, test = "Rao")

      2-sample Savage test against proportional hazards alternatives

data:  Surv(time, cens) by horTh (no, yes)
      stratified by tgrade
Rao chi-squared = 7.43, df = 1, p-value = 0.0064
alternative hypothesis: true log-hazard ratio is not equal to 0

> ### likelihood ratio tests
> summary(cm)$logtest

      test      df    pvalue
7.6027550 1.0000000 0.0058279

> summary(ft, test = "LRT")

      2-sample Savage test against proportional hazards alternatives

data:  Surv(time, cens) by horTh (no, yes)
      stratified by tgrade
logLR chi-squared = 7.64, df = 1, p-value = 0.0057
alternative hypothesis: true log-hazard ratio is not equal to 0

> ### Wald tests
> summary(cm)$waldtest
```

```

      test      df    pvalue
7.3300000 1.0000000 0.0067971

```

```
> summary(ft, test = "Wald")
```

2-sample Savage test against proportional hazards alternatives

```

data:  Surv(time, cens) by horTh (no, yes)
      stratified by tgrade
Wald chi-squared = 7.36, df = 1, p-value = 0.0067
alternative hypothesis: true log-hazard ratio is not equal to 0

```

```
> ### asymptotic permutation tests
```

```
> survdiff(Surv(time, cens) ~ horTh + strata(tgrade), data = GBSG2, rho = 0)[c("chisq", "pvalue")]
```

```

$chisq
[1] 7.3958

```

```

$pvalue
[1] 0.0065376

```

```
> summary(ft, test = "Permutation")
```

2-sample Savage test against proportional hazards alternatives

```

data:  Surv(time, cens) by horTh (no, yes)
      stratified by tgrade
Perm Z = 2.76, p-value = 0.0058
alternative hypothesis: true log-hazard ratio is not equal to 0

```

```

> independence_test(Surv(time, cens) ~ horTh | tgrade, data = GBSG2, ytrafo = function(...)
+                   trafo(..., numeric_trafo = logrank_trafo, block = GBSG2$tgrade),
+                   teststat = "quad")

```

Asymptotic General Independence Test

```

data:  Surv(time, cens) by horTh (no, yes)
      stratified by tgrade
chi-squared = 7.63, df = 1, p-value = 0.0057

```

Wilcoxon against proportional odds

```
> survdiff(Surv(time, cens) ~ horTh + strata(tgrade), data = GBSG2, rho = 1)[c("chisq", "pvalue")]
```

```

$chisq
[1] 7.5938

```

```

$pvalue
[1] 0.0058569

```

```

> (ft <- with(GBSG2, freeway(Surv(time, cens) ~ horTh | tgrade,
+                             link = "logit")))

```

2-sample Wilcoxon test against proportional odds alternatives

```

data:  Surv(time, cens) by horTh (no, yes)
      stratified by tgrade
Perm Z = 2.78, p-value = 0.0054
alternative hypothesis: true log-odds ratio is not equal to 0

```

```
> summary(ft, test = "Rao")
```

2-sample Wilcoxon test against proportional odds alternatives

```
data:  Surv(time, cens) by horTh (no, yes)
       stratified by tgrade
Rao chi-squared = 7.8, df = 1, p-value = 0.0052
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft, test = "LRT")
```

2-sample Wilcoxon test against proportional odds alternatives

```
data:  Surv(time, cens) by horTh (no, yes)
       stratified by tgrade
logLR chi-squared = 7.89, df = 1, p-value = 0.005
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft, test = "Wald")
```

2-sample Wilcoxon test against proportional odds alternatives

```
data:  Surv(time, cens) by horTh (no, yes)
       stratified by tgrade
Wald chi-squared = 7.75, df = 1, p-value = 0.0054
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft, test = "Permutation")
```

2-sample Wilcoxon test against proportional odds alternatives

```
data:  Surv(time, cens) by horTh (no, yes)
       stratified by tgrade
Perm Z = 2.78, p-value = 0.0054
alternative hypothesis: true log-odds ratio is not equal to 0
```

And now with more and smaller blocks

```
> library("survival")
> GBSG2$str <- cut(GBSG2$tsize, breaks = c(0, 1:9 * 10, Inf))
> cm <- coxph(Surv(time, cens) ~ horTh + strata(str), data = GBSG2)
> ft <- with(GBSG2, flexway(Surv(time, cens) ~ horTh | str,
+                           link = "cloglog"))
> coef(cm)

horThyes
-0.33921

> coef(ft)

horThyes
 0.33879

> vcov(cm)

horThyes
horThyes 0.016147

> vcov(ft)

horThyes
horThyes 0.016204

> ### Rao score tests
> summary(cm)$sctest
```

```

      test      df    pvalue
7.1917181 1.0000000 0.0073241

```

```
> summary(ft, test = "Rao")
```

2-sample Savage test against proportional hazards alternatives

```

data:  Surv(time, cens) by horTh (no, yes)
      stratified by str
Rao chi-squared = 7.15, df = 1, p-value = 0.0075
alternative hypothesis: true log-hazard ratio is not equal to 0

```

```

> ### likelihood ratio tests
> summary(cm)$logtest

```

```

      test      df    pvalue
7.3938622 1.0000000 0.0065447

```

```
> summary(ft, test = "LRT")
```

2-sample Savage test against proportional hazards alternatives

```

data:  Surv(time, cens) by horTh (no, yes)
      stratified by str
logLR chi-squared = 7.35, df = 1, p-value = 0.0067
alternative hypothesis: true log-hazard ratio is not equal to 0

```

```

> ### Wald tests
> summary(cm)$waldtest

```

```

      test      df    pvalue
7.1300000 1.0000000 0.0075971

```

```
> summary(ft, test = "Wald")
```

2-sample Savage test against proportional hazards alternatives

```

data:  Surv(time, cens) by horTh (no, yes)
      stratified by str
Wald chi-squared = 7.08, df = 1, p-value = 0.0078
alternative hypothesis: true log-hazard ratio is not equal to 0

```

```

> ### asymptotic permutation tests
> survdiff(Surv(time, cens) ~ horTh + strata(str), data = GBSG2, rho = 0)[c("chisq", "pvalue")]

```

```

$chisq
[1] 7.1868

```

```

$pvalue
[1] 0.0073441

```

```
> summary(ft, test = "Permutation")
```

2-sample Savage test against proportional hazards alternatives

```

data:  Surv(time, cens) by horTh (no, yes)
      stratified by str
Perm Z = 2.67, p-value = 0.0076
alternative hypothesis: true log-hazard ratio is not equal to 0

```

Wilcoxon against proportional odds

```

> survdiff(Surv(time, cens) ~ horTh + strata(str), data = GBSG2, rho = 1)[c("chisq", "pvalue")]

$chisq
[1] 8.2045

$pvalue
[1] 0.0041787

> (ft <- with(GBSG2, free1way(Surv(time, cens) ~ horTh | str,
+                             link = "logit")))

      2-sample Wilcoxon test against proportional odds alternatives

data:  Surv(time, cens) by horTh (no, yes)
      stratified by str
Perm Z = 2.88, p-value = 0.0039
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft, test = "Rao")

      2-sample Wilcoxon test against proportional odds alternatives

data:  Surv(time, cens) by horTh (no, yes)
      stratified by str
Rao chi-squared = 8.47, df = 1, p-value = 0.0036
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft, test = "LRT")

      2-sample Wilcoxon test against proportional odds alternatives

data:  Surv(time, cens) by horTh (no, yes)
      stratified by str
logLR chi-squared = 8.57, df = 1, p-value = 0.0034
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft, test = "Wald")

      2-sample Wilcoxon test against proportional odds alternatives

data:  Surv(time, cens) by horTh (no, yes)
      stratified by str
Wald chi-squared = 8.42, df = 1, p-value = 0.0037
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft, test = "Permutation")

      2-sample Wilcoxon test against proportional odds alternatives

data:  Surv(time, cens) by horTh (no, yes)
      stratified by str
Perm Z = 2.88, p-value = 0.0039
alternative hypothesis: true log-odds ratio is not equal to 0

```

9.7 van der Waerden Test

Normal scores test against a generalised Cohen's d :

```

> nd$y <- rnorm(nrow(nd))
> free1way(y ~ g | s, data = nd, link = "probit")

```


Stratified 3-sample van der Waerden normal scores test against latent normal shift alternatives

```
data: y by g (1, 2, 3)
      stratified by s
Perm chi-squared = 1.01, df = 2, p-value = 0.6
alternative hypothesis: true generalised Cohen's d is not equal to 0

> independence_test(y ~ g | s, data = nd, ytrafo = function(...)
+                   trafo(..., numeric_trafo = normal_trafo, block = nd$s),
+                   teststat = "quad")
```

Asymptotic General Independence Test

```
data: y by g (1, 2, 3)
      stratified by s
chi-squared = 1.06, df = 2, p-value = 0.59
```

9.8 Friedman Test

Each observation is a block in a K -sample proportional odds model

```
> friedman.test(RoundingTimes)
```

Friedman rank sum test

```
data: RoundingTimes
Friedman chi-squared = 11.1, df = 2, p-value = 0.0038
```

```
> (ft <- free1way(time ~ me | id, data = d))
```

Stratified 3-sample Kruskal-Wallis test against proportional odds alternatives

```
data: time by me (Round Out, Narrow Angle, Wide Angle)
      stratified by id
Perm chi-squared = 11.1, df = 2, p-value = 0.0038
alternative hypothesis: true log-odds ratio is not equal to 0

> summary(ft)
```

Call:

```
free1way.formula(formula = time ~ me | id, data = d)
```

Coefficients:

| | log-odds ratio | Std. Error | z value | P(> z) |
|----------------|----------------|------------|---------|---------|
| meNarrow Angle | -1.265 | 0.683 | -1.854 | 0.06 |
| meWide Angle | -3.289 | 0.794 | -4.144 | 0.00 |

```
> library("multcomp")
> glht(ft, linfct = mcp(me = "Tukey"))
```

General Linear Hypotheses

Multiple Comparisons of Means: Tukey Contrasts

Linear Hypotheses:

| | Estimate |
|--------------------------------|----------|
| Narrow Angle - Round Out == 0 | -1.27 |
| Wide Angle - Round Out == 0 | -3.29 |
| Wide Angle - Narrow Angle == 0 | -2.02 |

```

> example(friedman.test, echo = FALSE)
> ### Myles Hollander & Wolfe (2014, Example 7.1, page 294)
> boxplot(RoundingTimes, xlab = "Method", ylab = "Rounding-First-Base Time",
+         las = 1)
> matplot(t(RoundingTimes), add = TRUE, type = "l",
+         lty = 1, lwd = 2, col = rgb(.1, .1, .1, .1))
> me <- colnames(RoundingTimes)
> d <- expand.grid(me = factor(me, labels = me, levels = me),
+                 id = factor(seq_len(nrow(RoundingTimes))))
> d$time <- c(t(RoundingTimes))

```

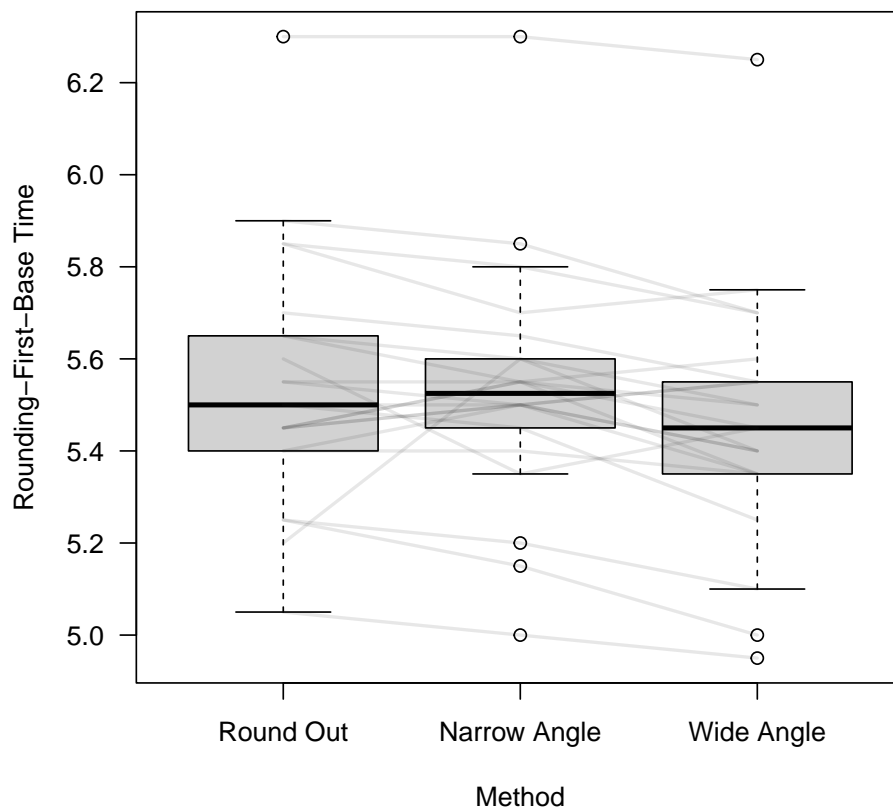


Figure 9.1: Rounding-first-base time data.

```

> logLik(ft)
[1] -55.857
> logLik(free1way(time ~ me | id, data = d, link = "probit"))
[1] -56.579
> logLik(free1way(time ~ me | id, data = d, link = "cloglog"))
[1] -53.993
> logLik(free1way(time ~ me | id, data = d, link = "loglog"))
[1] -55.624

```

Maybe proportional-hazards model better?

9.9 McNemar Test

```

> example(mcnemar.test, echo = FALSE)
> # set-up data frame with survey outcomes for voters
> s <- gl(2, 1, labels = dimnames(Performance)[[1L]])
> survey <- gl(2, 1, labels = c("1st", "2nd"))
> nvoters <- c(Performance)
> x <- expand.grid(survey = survey, voter = factor(seq_len(sum(nvoters))))
> x$performance <- c(rep(s[c(1, 1)], nvoters[1]), rep(s[c(2, 1)], nvoters[2]),
+                    rep(s[c(1, 2)], nvoters[3]), rep(s[c(2, 2)], nvoters[4]))
> # note that only those voters changing their minds are relevant
> mcn <- free1way(xtabs(~ performance + survey + voter, data = x))
> # same result as mcnemar.test w/o continuity correction
> print(mcn)

```

Stratified 2-sample Wilcoxon test against proportional odds
alternatives

```

data: performance by survey (c("1st", "2nd"))
      stratified by voter
Perm Z = 4.17, p-value = 3.1e-05
alternative hypothesis: true log-odds ratio is not equal to 0

```

```

> # X^2 statistic
> summary(mcn, test = "Permutation")$statistic^2

```

```

Perm Z
17.356

```

```

> mcnemar.test(Performance, correct = FALSE)

```

McNemar's Chi-squared test

```

data: Performance
McNemar's chi-squared = 17.4, df = 1, p-value = 3.1e-05

```

```

> # Wald inference
> summary(mcn)

```

Call:

```
free1way.table(y = xtabs(~performance + survey + voter, data = x))
```

Coefficients:

| | log-odds | ratio | Std. Error | z value | P(> z) |
|-----------|----------|-------|------------|---------|---------|
| survey2nd | 1.113 | | 0.191 | 5.816 | 0 |

```
> confint(mcn, test = "Wald")

                2.5 % 97.5 %
survey2nd 0.73767 1.4875

> ### because the model is saturated, the link function doesn't affect the
> ### p-value (but the coefficients are of course different)
> free1way(xtabs(~ performance + survey + voter, data = x), link = "probit")

Stratified 2-sample van der Waerden normal scores test against latent
normal shift alternatives

data: performance by survey (c("1st", "2nd"))
      stratified by voter
Perm Z = 4.17, p-value = 3.1e-05
alternative hypothesis: true generalised Cohen's d is not equal to 0
```

9.10 Incomplete Block Designs

`friedman.test` expects all blocks to be complete and `kruskal.test` has no idea about blocks. When blocks are incomplete, `free1way` can be employed. Replacing the normality assumption `inherit` in `aov` (Chapter 8.3.1. in [Meier, 2022](#)) with a semiparametric proportional odds model, we get

```
> data("taste", package = "daewr")
> ### highly discrete
> table(taste$score)

4 5 6 7 8
4 6 6 6 2

> summary(free1way(score ~ recipe | panelist, data = taste))

Call:
free1way.formula(formula = score ~ recipe | panelist, data = taste)
```

Coefficients:

| | log-odds | ratio | Std. Error | z value | P(> z) |
|---------|----------|-------|------------|---------|---------|
| recipeB | 2.483 | | 2.118 | 1.172 | 0.24 |
| recipeC | 2.930 | | 1.933 | 1.515 | 0.13 |
| recipeD | -1.696 | | 1.793 | -0.946 | 0.34 |

9.11 Contrast Tests

`free1way` output can be used to define multiple contrast tests and corresponding confidence intervals via the **multcomp** package. For example, Tukey-style simultaneous all-pair comparisons can be implemented via

```
> tk <- free1way(Ozone ~ Month, data = airquality)
> library("multcomp")
> confint(glht(tk, linfct = mcp(Month = "Tukey")))
```

Simultaneous Confidence Intervals

Multiple Comparisons of Means: Tukey Contrasts

```
Fit: free1way.formula(formula = Ozone ~ Month, data = airquality)
```

```
Quantile = 2.718
95% family-wise confidence level
```

Linear Hypotheses:

| | Estimate | lwr | upr |
|------------|----------|---------|---------|
| 6 - 5 == 0 | 0.8124 | -0.9382 | 2.5629 |
| 7 - 5 == 0 | 2.5282 | 1.0837 | 3.9726 |
| 8 - 5 == 0 | 2.3826 | 0.9322 | 3.8330 |
| 9 - 5 == 0 | 0.7513 | -0.5308 | 2.0334 |
| 7 - 6 == 0 | 1.7158 | -0.0692 | 3.5008 |
| 8 - 6 == 0 | 1.5702 | -0.2219 | 3.3624 |
| 9 - 6 == 0 | -0.0610 | -1.7487 | 1.6266 |
| 8 - 7 == 0 | -0.1456 | -1.4745 | 1.1834 |
| 9 - 7 == 0 | -1.7768 | -3.1206 | -0.4331 |
| 9 - 8 == 0 | -1.6313 | -2.9843 | -0.2783 |

Chapter 10

Model Diagnostics

10.1 Transformation Plots

The model formulation (1.1) suggests a simple graphical check of the main model assumption, that is, the existence of a constant shift on a latent scale defined by F . For one block and two samples, the model reads

$$\begin{aligned} F_Y(y \mid G = 1) &= F(F^{-1}(F_Y(y \mid G = 1))) = F(h(y)) \\ F_Y(y \mid G = 2) &= F(F^{-1}(F_Y(y \mid G = 1)) - \delta_2) = F(h(y) - \delta_2). \end{aligned}$$

We can now contrast the conditional distributions obtained from this model with the marginally estimated distribution functions of Y , that is, nonparametric estimates for the two distribution functions obtained within each treatment group separately. These latter estimates are typically simply the ECDF or Kaplan-Meier estimators in the presence of right-censoring. It is easier to see deviations from model (1.1) when the plot is presented on the scale of the link function F^{-1} . If the control distribution is close to $\hat{h}(y)$ and the distribution of those treated close to $\hat{h}(y) - \hat{\delta}_2$, model (1.1) provides a good approximation. If the two curves cross or if their horizontal distance varies considerably across the sample space, we should be concerned.

The model is assumed to hold in each block with overall treatment effects δ_k , but the intercept function h may differ between blocks. Therefore, we produce such a plot for each block separately. We also do not pay attention to the original observations of the outcome but plot the model on the scale of the ranked outcomes (the model is invariant with respect to monotone and therefore rank transformations).

All the necessary information can be extracted from an **object** of class **free1way**. We extract the sub-table containing the data for **block**, paying attention to possible right-censoring (in the four dimension)

< extract plot data ? > \equiv

```
object <- x
x <- object$table
if (RC <- (length(dim(x)) == 4L)) {
  x <- x[, , block, , drop = FALSE]
  x <- x[marginSums(x, margin = 1) > 0, , , drop = FALSE]
} else {
  x <- x[, , block, drop = FALSE]
  x <- x[marginSums(x, margin = 1) > 0, , , drop = FALSE]
}
K <- dim(x)[2L]
ret0 <- matrix(NA, nrow = dim(x)[1L], ncol = K)
ln <- object$link
◇
```

Fragment referenced in ?.

We then refit the intercept parameters (that is, $\hat{h}(y)$) for this block re-using the treatment effects already contained in **object**. We will plot them later on as a means to directly compare the model to the data

< refit block intercepts ? > \equiv

```
### refit for this block only
m1 <- .freelwayML(x, link = ln, start = coef(object),
                 fix = seq_along(coef(object)),
                 residuals = FALSE, hessian = FALSE)
intercepts <- m1$intercepts[[1L]]
j1 <- which(attr(get("xlist", environment(m1$profile)))[[1L]], "idx") > 1)
j1 <- j1[-length(j1)]
cf <- c(0, coef(object))
◇
```

Fragment referenced in [?](#).

Last, we compute the marginal distributions, that is, the distribution function of the outcome separately for each group. We could have used `ecdf` or `survfit` here, but since we have everything available in `.freelwayML`, we simply remove the observations corresponding to other groups and refit the intercept parameters.

< marginal fit ? > \equiv

```
for (k in seq_len(K)) {
  y <- x
  if (RC) {
    y[, -k, 1, ] <- 0
  } else {
    y[, -k, 1] <- 0
  }
  start <- numeric(K - 1)
  m0 <- .freelwayML(y, link = ln, start = start,
                  fix = seq_len(K - 1), residuals = FALSE,
                  hessian = FALSE)
  j <- which(attr(get("xlist", environment(m0$profile)))[[1L]], "idx") > 1)
  ret0[j[-length(j)], k] <- m0$intercepts[[1L]]
}
◇
```

Fragment referenced in [?](#).

We can now setup a `plot` method for `freelway` objects, we begin with an empty plot with appropriate axes annotations (we allow plotting on the scale of the CDF as well):

< setup canvas ? > \equiv

```
if (cdf) {
  ylim <- c(0, 1)
  FUN <- function(x) ln$linkinv(x)
} else {
  ylim <- range(c(ret0, intercepts), na.rm = TRUE)
  FUN <- function(x) x
}

idx <- seq_len(nrow(x))
main <- list(...)$main
if (is.null(main) && dim(object$table)[3L] > 1L)
  main <- paste(names(dimnames(x))[3L], dimnames(x)[[3L]][1L], sep = "=")
plot(idx, rep(0, length(idx)), type = "n", ylim = ylim,
     xlab = paste("Rank(", names(dimnames(x))[1L], ")", sep = ""),
     ylab = ifelse(cdf, "Probability", paste(ln$name, "Link")),
     main = main, ...)
◇
```

Fragment referenced in [?](#).

The marginally estimated functions are plotted first

⟨ marginal plot ? ⟩ ≡

```
out <- sapply(seq_len(K), function(k)
  lines(which(!is.na(ret0[,k])), FUN(ret0[!is.na(ret0[,k]),k]),
    type = "s", col = col[k], lty = lty[1]))
```

◇

Fragment referenced in [?](#).

followed by a plot of the model-based functions (which can be switched off)

⟨ model plot ? ⟩ ≡

```
if (model)
  out <- sapply(seq_len(K), function(k)
    lines(j1, FUN(intercepts - cf[k]), type = "s", col = col[k],
      lty = lty[2]))
```

◇

Fragment referenced in [?](#).

and finally we add a legend

⟨ add legend ? ⟩ ≡

```
if (legend) {
  legend("topleft", lty = lty[1], col = col,
    legend = paste(names(dimnames(x))[2L], dimnames(x)[[2L]]),
    title = "Nonparametric", bty = "n")
  if (model)
    legend("bottomright", lty = lty[2], col = col,
      legend = paste(names(dimnames(x))[2L], dimnames(x)[[2L]]),
      title = "Semiparametric", bty = "n")
}
```

◇

Fragment referenced in [?](#).

We put everything together in a plot method

⟨ plot free1way ? ⟩ ≡

```
plot.free1way <- function(x, ..., block = 1L, cdf = FALSE, model = TRUE,
  col = seq_len(length(coef(object)) + 1L),
  lty = 1:2, legend = TRUE)
{
```

```
  ⟨ extract plot data ? ⟩
  ⟨ refit block intercepts ? ⟩
  ⟨ marginal fit ? ⟩
```

```
  ⟨ setup canvas ? ⟩
  ⟨ marginal plot ? ⟩
  ⟨ model plot ? ⟩
  ⟨ add legend ? ⟩
```

}

◇

Fragment referenced in [?](#).


```
> tk <- free1way(Ozone ~ Month, data = airquality)
> plot(tk, las = 1)
```

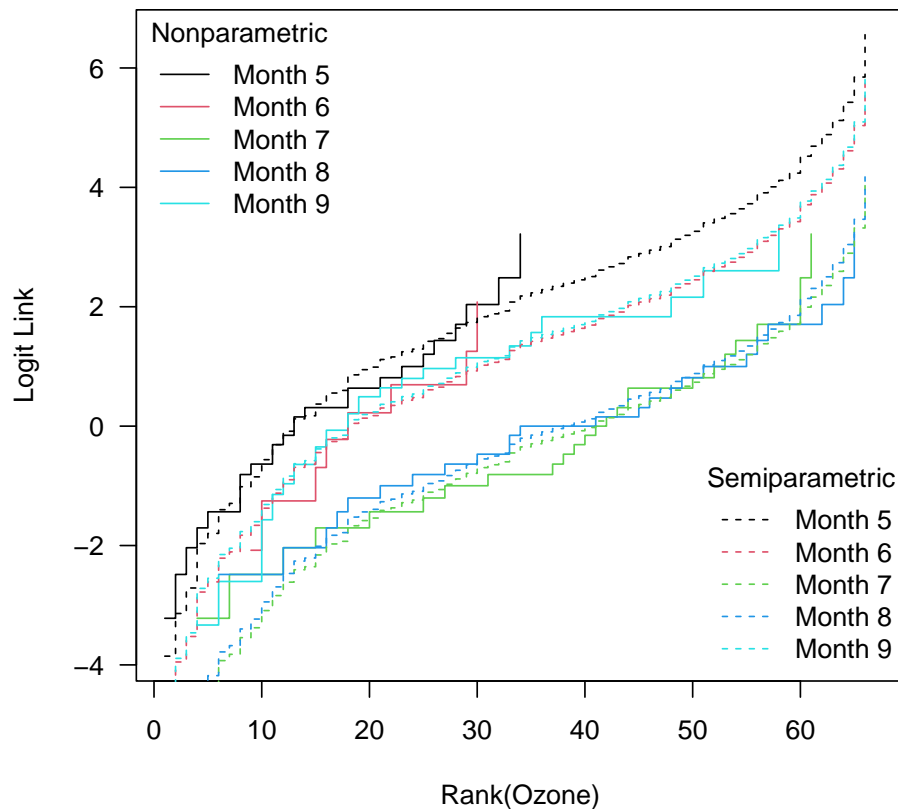


Figure 10.1: Model diagnostics for proportional odds model comparing ozone concentrations for different months.

By default, the plot shows the marginal (“nonparametric”) and model-based (“semiparametric”) estimates in the sample plot. For the ozone concentrations in different months in Figure 10.1, we see that the distributions differ, but the effects can be well understood as shifts on a log-odds scale. Note that the solid nonparametric curves agree quite well with the dashed model-based ones.

We can check if the plot is correct by comparing the result (Figure 10.2) to the one obtained with `ecdfplot` after rank transformation on the scale of the distribution functions (Figure 10.3)

10.2 Probability-probability Plots

The classical shift model $F_Y(y | T = 2) = F_Y(y - \mu | T = 1)$ can be criticised using confidence bands for QQ-plots in `qqplot`, because the parameter μ shows up as a vertical shift of the diagonal if the model is appropriate.

Likewise, model (1.1) can be graphically assessed using the P-P-plot. We concentrate on the two-sample case. The shift parameter δ_2 gives rise to the model-based P-P graph $(p, F(F^{-1}(p) - \delta_2))$ and a confidence *band* can be obtained from a confidence *interval* for δ_2 . The P-P-plot is, up to rescalings, identical to the ROC curve.

```
> plot(tk, cdf = TRUE, model = FALSE, las = 1)
```

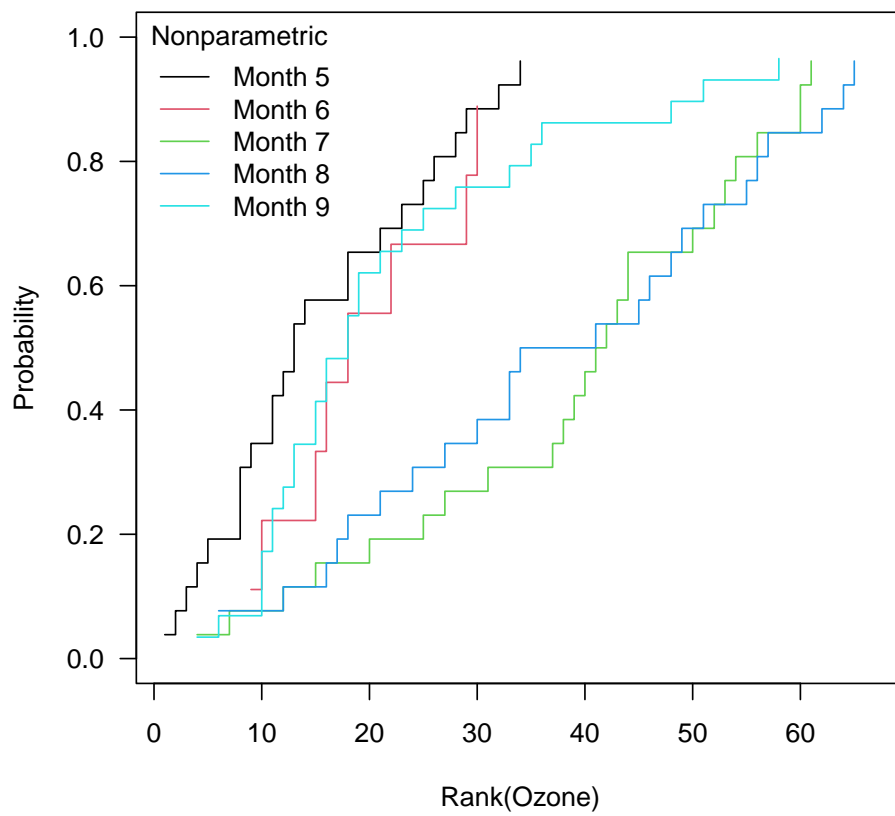


Figure 10.2: Nonparametric distributions of ozone concentrations for different months.

```

> aq <- subset(airquality, !is.na(Ozone))
> aq$r <- match(aq$Ozone, sort(unique(aq$Ozone)))
> library("latticeExtra")
> plot(ecdfplot(~ r, data = aq, groups = Month, col = 1:5))

```

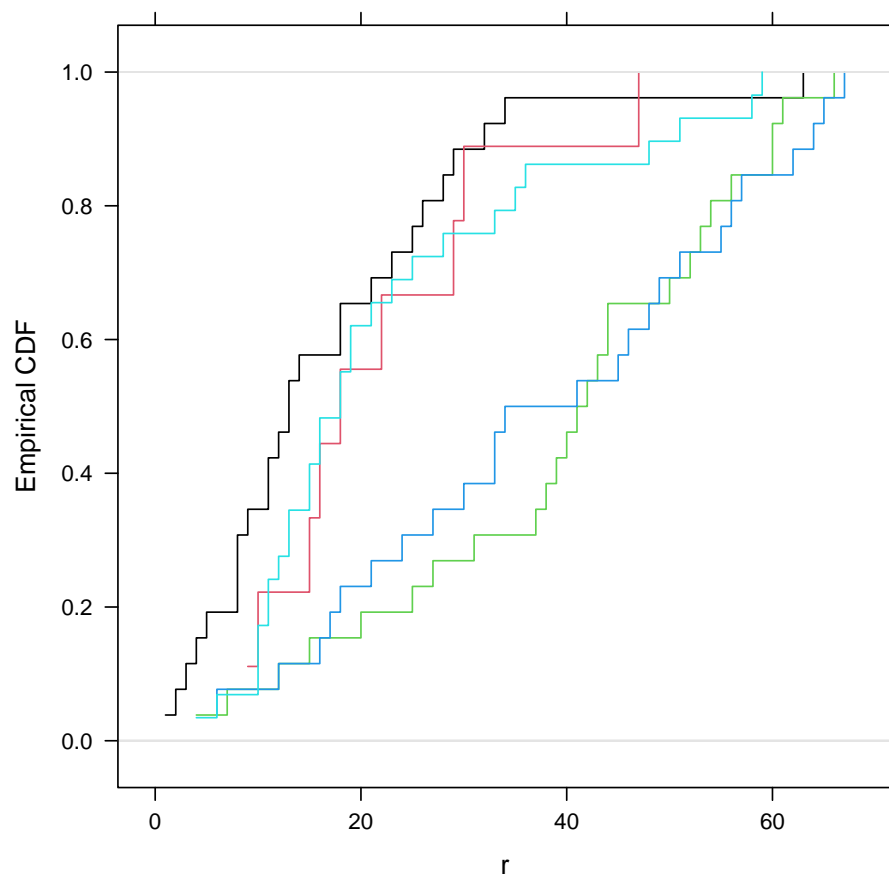


Figure 10.3: Nonparametric distributions of ozone concentrations for different months.

$\langle \text{ROC bands ?} \rangle \equiv$

```

if (!is.null(conf.level)) {
  prb <- seq_len(1000) / 1001
  res <- c(x, y)
  grp <- gl(2, 1, labels = c(xlab, ylab))
  grp <- grp[rep(1:2, c(length(x), length(y)))]
  args <- conf.args
  args$y <- res
  args$groups <- grp
  args$border <- args$col <- args$type <- NULL
  f1w <- do.call("free1way", args)

  ci <- confint(f1w, level = conf.level, type = args$type)
  lwr <- .p(f1w$link, .q(f1w$link, prb) - ci[1,1])
  upr <- .p(f1w$link, .q(f1w$link, prb) - ci[1,2])
  x <- c(prb, rev(prb))
  y <- c(lwr, rev(upr))
  xn <- c(x[1L], rep(x[-1L], each = 2))
  yn <- c(rep(y[-length(y)], each = 2), y[length(y)])
  polygon(x = xn, y = yn, col = conf.args$col, border = conf.args$border)
  lines(prb, .p(f1w$link, .q(f1w$link, prb) - coef(f1w)))
}

```

Fragment referenced in [?](#).

We introduce a new function `ppplot`, closely following the implementation of `qqplot`, allowing to plot the empirical ([Wilk and Gnanadesikan, 1968](#)) and corresponding model-based ([Sewak and Hothorn, 2023](#)) probability-probability plot, the latter for a certain choice of link function:

$\langle ppplot ? \rangle \equiv$

```
ppplot <- function(x, y, plot.it = TRUE,
                  xlab = paste("Cumulative probabilities for",
                               deparse1(substitute(x))),
                  ylab = paste("Cumulative probabilities for",
                               deparse1(substitute(y))),
                  main = "P-P plot",
                  ..., conf.level = NULL,
                  conf.args = list(link = "logit", type = "Wald",
                                   col = NA, border = NULL))
{
  force(xlab)
  force(ylab)
  if (xlab == ylab) {
    xlab <- paste0("x = ", xlab)
    ylab <- paste0("y = ", ylab)
  }

  ex <- ecdf(x)
  sy <- sort(unique(c(x, y)))
  py <- ecdf(y)(sy)
  px <- ex(sy)
  ret <- stepfun(px, c(0, py))
  if (!plot.it)
    return(ret)

  plot(ret, xlim = c(0, 1), ylim = c(0, 1),
       xlab = xlab, ylab = ylab, main = main,
       verticals = FALSE, ...)

   $\langle ROC\ bands ? \rangle$ 

  plot(ret, add = TRUE, verticals = FALSE, ...)

  return(invisible(ret))
}
◇
```

Fragment referenced in [?](#).

A correct logistic model with log-odds ratio three is shown in [Figure 10.4](#) and an incorrect proportional hazards model for the same data in [Figure 10.5](#).

```

> y <- rlogis(50)
> x <- rlogis(50, location = 3)
> ppplot(y, x, conf.level = .95, las = 1)

```

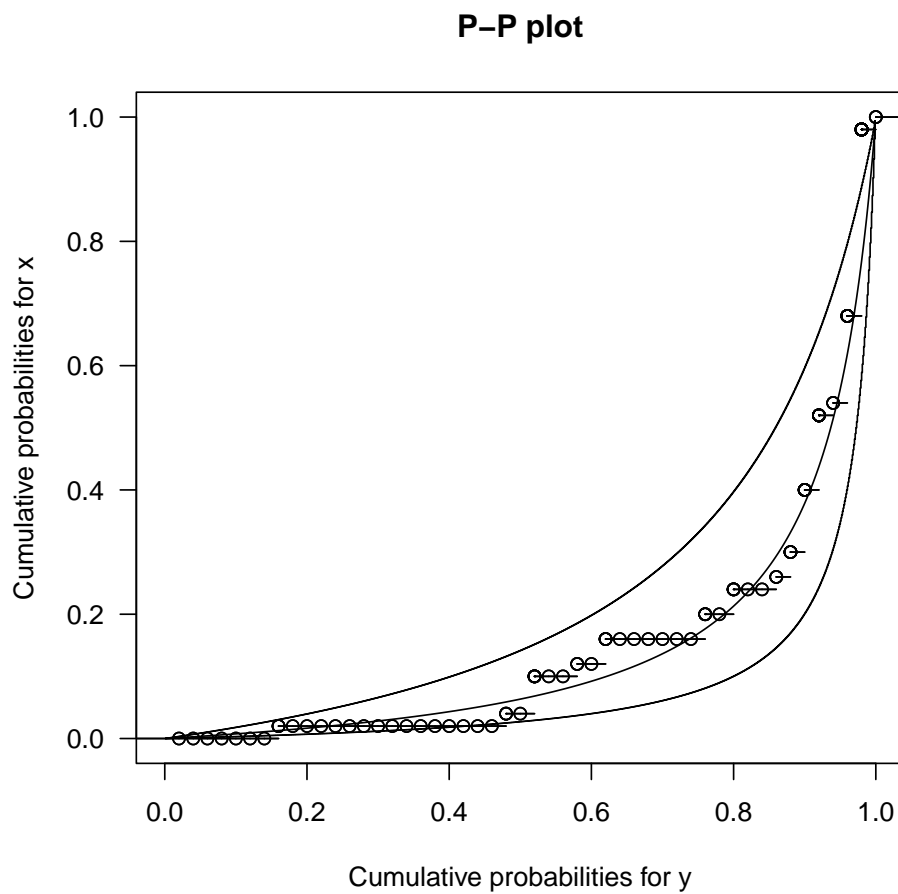


Figure 10.4: Data sampled from a proportional-odds model with probability-probability (P-P) curve and 95% confidence band obtained from a proportional-odds model.

```
> ppplot(y, x, conf.args = list(link = "cloglog", type = "Wald",
+                               col = NA, border = NULL),
+         conf.level = .95, las = 1)
```

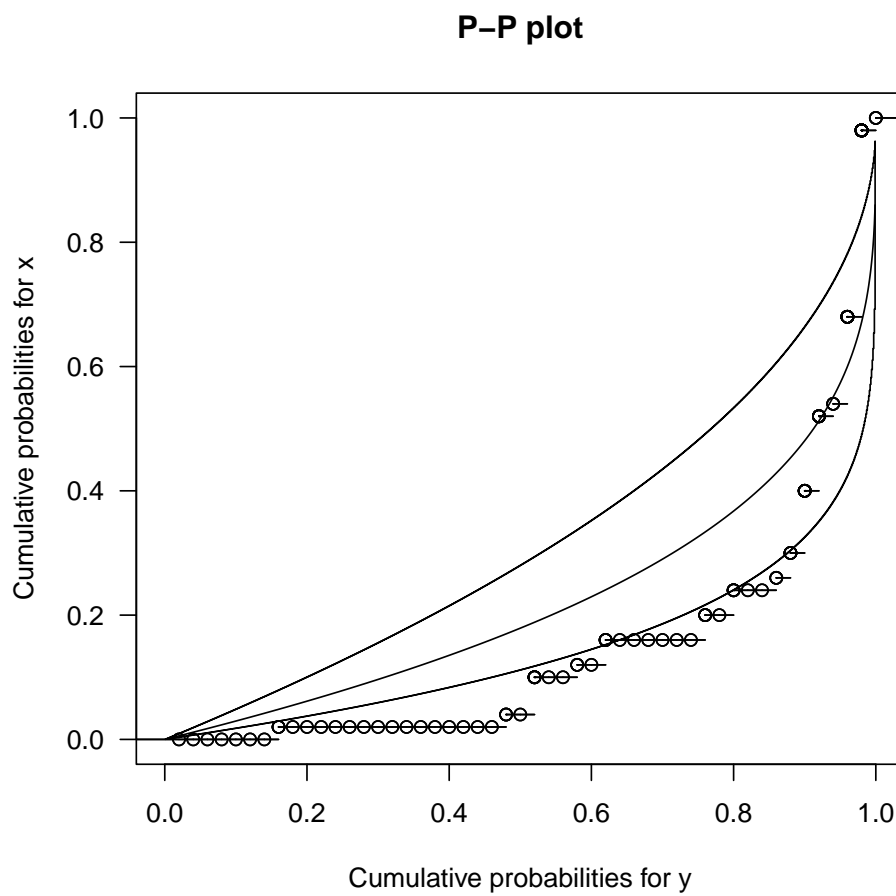


Figure 10.5: Data sampled from a proportional-odds model with probability-probability (P-P) curve and 95% confidence band obtained from a proportional-hazards model.

Chapter 11

Random Number Generation

With 1.1 we know that for an absolutely continuous random variable Y

$$U = F_Y(Y \mid G = k, S = b) = F(F^{-1}(F_Y(Y \mid G = 1, S = b)) - \delta_k), \quad k = 2, \dots, K$$

follows a standard uniform distribution on the unit interval. This means that we can sample from the distribution of Y using

$$F_Y^{-1}(F(F^{-1}(U) + \delta_k) \mid G = 1, S = b).$$

It is therefore enough to draw samples from $F(F^{-1}(U) + \delta_k)$, that is, assuming a uniform distribution for F_Y in each control group. Because of the invariance with respect to monotone transformations, transforming all observations by the same quantile function changes the outcome distributions but not the shift effects. Discrete outcomes can be generated by post-hoc categorisation.

$\langle \text{design args ?} \rangle \equiv$

```
K <- length(delta) + 1L
if (is.null(names(delta)))
  names(delta) <- LETTERS[seq_len(K)[-1]]
if (length(alloc_ratio) == 1L)
  alloc_ratio <- rep_len(alloc_ratio, K - 1)
if (length(alloc_ratio) != K - 1L)
  stop(gettextf("invalid argument '%s'", "alloc_ratio"), domain = NA)
if (length(strata_ratio) == 1L)
  strata_ratio <- rep_len(strata_ratio, B - 1)
if (length(strata_ratio) != B - 1L)
  stop(gettextf("invalid argument '%s'", "strata_ratio"), domain = NA)
### sample size per group (columns) and stratum (rows)
N <- n * matrix(c(1, alloc_ratio), nrow = B, ncol = K, byrow = TRUE) *
  matrix(c(1, strata_ratio), nrow = B, ncol = K)
```

◇

Fragment referenced in ?, ?.

$\langle rfree1way ? \rangle \equiv$

```
.rfree1way <- function(n, delta = 0, link = c("logit", "probit",
                                             "cloglog", "loglog"))
{
  logU <- log(ret <- runif(n))

   $\langle link2fun ? \rangle$ 

  trt <- (abs(delta) > 0)
  ret[trt] <- .p(link, .q(link, logU[trt], log.p = TRUE) + delta[trt])

  return(ret)
}

rfree1way <- function(n, prob = NULL, alloc_ratio = 1,
                     blocks = ifelse(is.null(prob), 1, NCOL(prob)),
                     strata_ratio = 1, delta = 0, offset = 0,
                     link = c("logit", "probit", "cloglog", "loglog"))
{
  B <- blocks

   $\langle design\ args ? \rangle$ 

  rownames(N) <- paste0("block", seq_len(B))
  ctrl <- "Control"
  colnames(N) <- c(ctrl, names(delta))

  if (length(offset) != K)
    offset <- rep_len(offset, K)

  trt <- gl(K, 1, labels = colnames(N))
  blk <- gl(B, 1, labels = rownames(N))
  ret <- expand.grid(blocks = blk, groups = trt)
  if (B == 1L) ret$blocks <- NULL
  ret <- ret[rep(seq_len(nrow(ret)), times = N), , drop = FALSE]
  ret$y <- .rfree1way(nrow(ret),
                    delta = offset[ret$groups] + c(0, delta)[ret$groups],
                    link = link)
  if (is.null(prob)) return(ret)

  ### return discrete distribution
  if (!is.matrix(prob))
    prob <- matrix(prob, nrow = NROW(prob), ncol = B)
  if (ncol(prob) != B)
    stop(gettextf("incorrect number of columns for 'prob' in %s",
                  "rfree1way"),
         domain = NA)
  prob <- prop.table(prob, margin = 2L)
  ret <- do.call("rbind", lapply(1:ncol(prob), function(b) {
    if (B > 1)
      ret <- subset(ret, blocks == levels(blocks)[b])
    ret$y <- cut(ret$y, breaks = c(-Inf, cumsum(prob[,b])),
                labels = paste0("Y", 1:nrow(prob)), ordered_result = TRUE)

    ret
  })))
  return(ret)
}
◇
```

Fragment referenced in [?](#).

```

> (logOR <- c(log(1.5), log(2)))

[1] 0.40547 0.69315

> nd <- rfree1way(150, delta = logOR)
> coef(ft <- free1way(y ~ groups, data = nd))

groupsB groupsC
0.48984 0.74855

> sqrt(diag(vcov(ft)))

groupsB groupsC
0.20077 0.20210

> logLik(ft)

[1] -2742

> nd$y <- qchisq(nd$y, df = 3)
> coef(ft <- free1way(y ~ groups, data = nd))

groupsB groupsC
0.48984 0.74855

> sqrt(diag(vcov(ft)))

groupsB groupsC
0.20077 0.20210

> logLik(ft)

[1] -2742

> N <- 25
> pvals <- replicate(Nsim,
+ {
+   nd <- rfree1way(n = N, blocks = 2, delta = c(.25, .5), alloc_ratio = 2)
+   summary(free1way(y ~ groups | blocks, data = nd), test = "Permutation")$p.value
+ })
> power.free1way.test(n = N, blocks = 2, delta = c(.25, .5), alloc_ratio = 2)

```

Stratified 3-sample Kruskal-Wallis test against proportional odds alternatives

```

      n = 25
Total sample size = 50 (Control) + 100 (B) + 100 (C) = 250
      power = 0.31044
      sig.level = 0.05
      log-odds ratio = 0.25, 0.50

```

NOTE: 'n' is sample size in control group of first stratum

```

> mean(pvals < .05)

[1] 0.36

```

This function can also be used to simulate survival times, for example, from a proportional hazards model with a censoring probability of .25 in the control arm and of .5 in the treated arm, under random censoring (that is, event and censoring times are independent given treatment).

```

> N <- 1000
> nd <- rfreelway(N, delta = 1, link = "cloglog")
> nd$C <- rfreelway(n = N, delta = 1, offset = -c(qlogis(.25), qlogis(.5)),
+               link = "cloglog")$y
> nd$y <- Surv(pmin(nd$y, nd$C), nd$y < nd$C)
> ### check censoring probability
> 1 - tapply(nd$y[,2], nd$groups, mean)

Control      B
0.258      0.524

> summary(freelway(y ~ groups, data = nd, link = "cloglog"))

Call:
freelway.formula(formula = y ~ groups, data = nd, link = "cloglog")

Coefficients:
      log-hazard ratio Std. Error z value P(>|z|)
groupsB           1.034      0.061  16.957      0

> summary(coxph(y ~ groups, data = nd))

Call:
coxph(formula = y ~ groups, data = nd)

      n= 2000, number of events= 1218

      coef exp(coef) se(coef)      z Pr(>|z|)
groupsB -1.033      0.356      0.061 -16.9  <2e-16 ***
---
Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1

      exp(coef) exp(-coef) lower .95 upper .95
groupsB      0.356      2.81      0.316      0.401

Concordance= 0.627 (se = 0.007 )
Likelihood ratio test= 299 on 1 df,  p=<2e-16
Wald test              = 287 on 1 df,  p=<2e-16
Score (logrank) test = 310 on 1 df,  p=<2e-16

```

Next we start implementing a function for simulating $C \times K$ tables. We need to specify the number of observations in each treatment group (`c`), the discrete distribution of the control (`r`), a model (`link`), and a treatment effect (`delta`, in line with `power.XYZ.test`). In essence, we draw samples from the multinomial distribution after computing the relevant discrete density.

$\langle r2dsim ? \rangle \equiv$

```
.r2dsim <- function(n, r, c, delta = 0,
                    link = c("logit", "probit", "cloglog", "loglog"))
{
  if (length(n <- as.integer(n)) == 0L || (n < 0) || is.na(n))
    stop(gettextf("invalid argument '%s'", "n"), domain = NA)
  colsums <- c
  if (length(colsums[] <- as.integer(c)) <= 1L ||
      any(colsums < 0) || anyNA(colsums))
    stop(gettextf("invalid argument '%s'", "c"), domain = NA)

  prob <- r
  if (length(prob[] <- as.double(r / sum(r))) <= 1L ||
      any(prob < 0) || anyNA(prob))
    stop(gettextf("invalid argument '%s'", "r"), domain = NA)

  if (is.null(names(prob)))
    names(prob) <- paste0("i", seq_along(prob))

  K <- length(colsums)
  if (is.null(names(colsums)))
    names(colsums) <- LETTERS[seq_len(K)]
  delta <- rep_len(delta, K - 1L)

   $\langle link2fun ? \rangle$ 

  p0 <- cumsum(prob) / sum(prob)
  h0 <- .q(link, p0[-length(p0)]) ### last element of p0 is one

  h1 <- h0 - matrix(delta, nrow = length(prob) - 1L, ncol = K - 1,
                    byrow = TRUE)
  p1 <- rbind(.p(link, h1), 1)
  p <- cbind(p0, p1)
  ret <- vector(mode = "list", length = n)

  for (i in seq_len(n)) {
    tab <- sapply(seq_len(K), function(k) {
      ### deal with zero prob categories,
      ### b/c cut() below will complain if p[,k] has duplicates
      ndp <- which(!duplicated(p[,k]))
      pu <- sort(p[ndp,k])
      ret <- integer(length(prob))
      ret[ndp] <- unclass(table(cut(runif(colsums[k]), breaks = c(-Inf, pu))))
      ret
    })
    ret[[i]] <- as.table(array(unlist(tab), dim = c(length(prob), K),
                              dimnames = list(names(prob),
                                                names(colsums))))
  }
  return(ret)
}
◇
```

Fragment referenced in [?](#).

Chapter 12

Power and Sample Size

The term “distribution-free” refers to the invariance of the reference distribution with respect to the distribution of an absolutely continuous outcome under control. Unfortunately, this is no longer true for non-continuous outcomes (due to ties) and under the alternative. That means that sample size assessments always take place under certain assumptions regarding the outcome distribution.

With the infrastructure from Chapter 11, we are now ready to put together a function for power evaluation and sample size assessment. The core idea is to draw samples from the relevant data (under a specific model in the alternative) and to estimate the Fisher information of the treatment effect parameters for this configuration. The power of the global Wald test can then be approximated by a non-central χ^2 distribution. This is much faster than approximating the power directly. Nevertheless, this is a random experiment, so we first make computations reproducible:

random seed ? \equiv

```
if (!exists(".Random.seed", envir = .GlobalEnv, inherits = FALSE))
  runif(1)
if (is.null(seed))
  seed <- RNGstate <- get(".Random.seed", envir = .GlobalEnv)
else {
  R.seed <- get(".Random.seed", envir = .GlobalEnv)
  set.seed(seed)
  RNGstate <- structure(seed, kind = as.list(RNGkind()))
  on.exit(assign(".Random.seed", R.seed, envir = .GlobalEnv))
}
◇
```

Fragment referenced in ?.

$\langle \text{power setup ?} \rangle \equiv$

$\langle \text{link2fun ?} \rangle$

```
### matrix means control distributions in different strata
if (!is.matrix(prob))
  prob <- matrix(prob, nrow = NROW(prob), ncol = blocks)
prob <- prop.table(prob, margin = 2L)
C <- nrow(prob)
B <- ncol(prob)
if (is.null(colnames(prob)))
  colnames(prob) <- paste0("stratum", seq_len(B))
p0 <- apply(prob, 2, function(x) cumsum(x) / sum(x))
h0 <- .q(link, p0[-nrow(p0)], drop = FALSE)
```

$\langle \text{design args ?} \rangle$

```
rownames(N) <- colnames(prob)
ctrl <- "Control"
dn <- dimnames(prob)
if (!is.null(names(dn)[1L]))
  ctrl <- names(dn)[1L]
colnames(N) <- c(ctrl, names(delta))
◇
```

Fragment referenced in [?](#).

For estimating the Fisher information, we draw samples from the discrete outcome distribution and evaluate the observed Fisher information for the, here and now known true parameters. The average of these Fisher information matrices is then used as an estimate for the expected Fisher information. For small sample sizes less than 100, we draw larger samples (at least 1000) and adjust the obtained Fisher information accordingly to reduce sampling error.

$\langle \text{estimate Fisher information ?} \rangle \equiv$

```
he <- 0
deltamu <- delta - mu
Nboost <- ifelse(n < 100, ceiling(1000 / n), 1)
for (i in seq_len(nsim)) {
  parm <- deltamu
  x <- as.table(array(0, dim = c(C, K, B)))
  for (b in seq_len(B)) {
    x[,b] <- .r2dsim(1L, r = prob[, b], c = Nboost * N[b,],
                    delta = delta, link = link)[[1L]]
    rs <- which(.rowSums(x[,b], m = dim(x)[1L], n = dim(x)[2L]) > 0)
    theta <- h0[pmin(nrow(h0), rs), b]
    parm <- c(parm, theta[-length(theta)])
  }
  ### evaluate observed hessian for true parameters parm and x data
  he <- he + .free1wayML(x, link = link, mu = mu, start = parm,
                        fix = seq_along(parm))$hessian / Nboost
}
### estimate expected Fisher information
he <- he / nsim
◇
```

Fragment referenced in [?](#).

The power function now depends on sample size (n ; the number of control observations in the first stratum), a discrete control distribution (prob , this can be a $C \times B$ matrix for stratum-specific control

distributions), a vector of allocation ratios (`alloc_ratio = 2` means control:treatment = 1:2) and the sample size ratios between strata.

The treatment effects are contained in $K - 1$ vector `delta`:

< power call ? > \equiv

```
power.freelway.test(n = n, prob = prob,
                   alloc_ratio = alloc_ratio,
                   blocks = blocks,
                   strata_ratio = strata_ratio,
                   delta = delta, mu = mu,
                   sig.level = sig.level, link = link,
                   alternative = alternative,
                   nsim = nsim, seed = seed,
                   tol = tol)$power - power
```

◇

Fragment referenced in [?](#).

< power args check ? > \equiv

```
if (sum(vapply(list(n, delta, power, sig.level), is.null,
                NA)) != 1)
  stop("exactly one of 'n', 'delta', 'power', and 'sig.level' must be NULL")
assert_NULL_or_prob(sig.level)
assert_NULL_or_prob(power)
```

◇

Fragment referenced in [?](#).

< power htest output ? > \equiv

```
ss <- paste(colSums(N), paste0("(", colnames(N), ")"), collapse = " + ")
ret <- list(n = n,
           "Total sample size" = paste(ss, "=", sum(N)),
           power = power,
           sig.level = sig.level)
if (mu != 0) ret$mu <- mu
if (K == 2L) ret[["Standard error"]] <- se
ret[[link$parm]] <- delta
ret$note <- "'n' is sample size in control group"
if (B > 1) ret$note <- paste(ret$note, "of first stratum")
alias <- link$alias
if (length(link$alias) == 2L) alias <- alias[1L + (K > 2L)]
ret$method <- paste(ifelse(B > 1L, "Stratified", ""),
                   paste0(K, "-sample"), alias,
                   "test against", link$model, "alternatives")
class(ret) <- "power.htest"
```

◇

Fragment referenced in [?](#).

We can invert the power function for finding nominal levels, sample or effect sizes necessary to achieve a certain power. The option is available because the power approximation is relatively fast.

$\langle \text{power inversion ?} \rangle \equiv$

```

if (is.null(n))
  n <- ceiling(uniroot(function(n) {
     $\langle \text{power call ?} \rangle$ 
  }, interval = c(5, 1e+03), tol = tol, extendInt = "upX")$root)
else if (is.null(delta)) {
  ### 2-sample only
  if (length(alloc_ratio) > 1L)
    stop(gettextf("effect size can only be computed for two-sample problems in %s",
      "power.free1way.test"),
      domain = NA)
  delta <- uniroot(function(delta) {
     $\langle \text{power call ?} \rangle$ 
  }, interval = c(0, 10), tol = tol, extendInt = "upX")$root
}
else if (is.null(sig.level))
  sig.level <- uniroot(function(sig.level) {
     $\langle \text{power call ?} \rangle$ 
  }, interval = c(1e-10, 1 - 1e-10), tol = tol, extendInt = "yes")$root

```

◇

Fragment referenced in [?](#).

$\langle \text{power ?} \rangle \equiv$

```
power.freelway.test <- function(n = NULL,
                                prob = if (is.null(n)) NULL else
                                      rep.int(1 / n, n),
                                alloc_ratio = 1,
                                blocks = if (is.null(prob)) 1 else NCOL(prob),
                                strata_ratio = 1,
                                delta = NULL, mu = 0,
                                sig.level = .05, power = NULL,
                                link = c("logit", "probit", "cloglog", "loglog"),
                                alternative = c("two.sided", "less", "greater"),
                                nsim = 100, seed = NULL,
                                tol = .Machine$double.eps^0.25)
{
   $\langle \text{power args check ?} \rangle$ 

   $\langle \text{random seed ?} \rangle$ 

   $\langle \text{r2dsim ?} \rangle$ 

   $\langle \text{power inversion ?} \rangle$ 

  ### n is available now
  if (is.null(prob)) prob <- rep(1 / n, n)

   $\langle \text{power setup ?} \rangle$ 
   $\langle \text{estimate Fisher information ?} \rangle$ 

  alternative <- match.arg(alternative)
  if (K == 2L) {
    se <- 1 / sqrt(c(he))
    power <- switch(alternative,
      "two.sided" = pnorm(qnorm(sig.level / 2) + deltam / se) +
                    pnorm(qnorm(sig.level / 2) - deltam / se),
      "less" = pnorm(qnorm(sig.level) - deltam / se),
      "greater" = pnorm(qnorm(sig.level) + deltam / se)
    )
  } else {
    if (alternative != "two.sided")
      stop(gettextf("%s only allows two-sided alternatives in the presence of more than two groups",
                    "power.freelway.test"),
           domain = NA)
    ncp <- sum((chol(he) %*% deltam)^2)
    qsig <- qchisq(sig.level, df = K - 1L, lower.tail = FALSE)
    power <- pchisq(qsig, df = K - 1L, ncp = ncp, lower.tail = FALSE)
  }

   $\langle \text{power htest output ?} \rangle$ 

  ret
}
```

Fragment referenced in [?](#).

We start with the power of a binomial experiment with $N = 2 \times 25$ observations. In the control group, the odds of winning is 1. Under treatment, we increase this odds by 50%. We compare the results with `power.prop.test`:

```
> delta <- log(1.5)
> power.prop.test(n = 25, p1 = .5, p2 = plogis(qlogis(.5) - delta))
```

Two-sample comparison of proportions power calculation

```
n = 25
p1 = 0.5
p2 = 0.4
sig.level = 0.05
power = 0.10462
alternative = two.sided
```

NOTE: n is number in *each* group

```
> power.free1way.test(n = 25, prob = c(.5, .5), delta = delta)
```

2-sample Wilcoxon test against proportional odds alternatives

```
n = 25
Total sample size = 25 (Control) + 25 (B) = 50
power = 0.10935
sig.level = 0.05
Standard error = 0.57155
log-odds ratio = 0.40547
```

NOTE: 'n' is sample size in control group

Under stratification (twice as many observations in the second stratum) and with an ordered outcome at four levels, we might want to compare four groups, with 25%, 50%, and 75% increase compared to the odds of the control:

```
> prb <- matrix(c(.25, .25, .25, .25,
+               .10, .20, .30, .40), ncol = 2)
> colnames(prb) <- c("s1", "s2")
> power.free1way.test(n = 20, prob = prb,
+                   strata_ratio = 2,
+                   alloc_ratio = c(1.5, 2, 2),
+                   delta = log(c("low" = 1.25, "med" = 1.5, "high" = 1.75)))
```

Stratified 4-sample Kruskal-Wallis test against proportional odds alternatives

```
n = 20
Total sample size = 60 (Control) + 90 (low) + 120 (med) + 120 (high) = 390
power = 0.37911
sig.level = 0.05
log-odds ratio = 0.22314, 0.40547, 0.55962
```

NOTE: 'n' is sample size in control group of first stratum

We now estimate the power of a Wilcoxon test with, first by simulation from a logistic distribution, and then by our power function:

```
> delta <- log(3)
> N <- 15
> w <- gl(2, N)
> pw <- numeric(Nsim)
> for (i in seq_along(pw)) {
+   y <- rlogis(length(w), location = c(0, delta)[w])
+   pw[i] <- wilcox.test(y ~ w)$p.value
+ }
> mean(pw < .05)

[1] 0.37
```

```
> power.freelway.test(n = N, delta = delta)

2-sample Wilcoxon test against proportional odds alternatives

      n = 15
Total sample size = 15 (Control) + 15 (B) = 30
      power = 0.38378
      sig.level = 0.05
Standard error = 0.66021
log-odds ratio = 1.0986
```

NOTE: 'n' is sample size in control group

```
> ### approximate formula in Hmisc::popower
> library("Hmisc")
> popower(p = rep(1 / N, N), odds.ratio = exp(delta), n = 2 * N)
```

```
Power: 0.389
Efficiency of design compared with continuous response: 0.997
Approximate standard error of log odds ratio: 0.655
```

The power of the Kruskal-Wallis test only needs one additional treatment effect

```
> delta <- c("B" = log(2), "C" = log(3))
> N <- 15
> w <- gl(3, N)
> pw <- numeric(Nsim)
> for (i in seq_along(pw)) {
+   y <- rlogis(length(w), location = c(0, delta)[w])
+   pw[i] <- kruskal.test(y ~ w)$p.value
+ }
> mean(pw < .05)

[1] 0.31

> power.freelway.test(n = N, delta = delta)
```

```
3-sample Kruskal-Wallis test against proportional odds alternatives

      n = 15
Total sample size = 15 (Control) + 15 (B) + 15 (C) = 45
      power = 0.31247
      sig.level = 0.05
log-odds ratio = 0.69315, 1.09861
```

NOTE: 'n' is sample size in control group

We next use the `rfree1way` function to sample from 4×3 tables with odds ratios 2 and 3 and compare the resulting power with result obtained from the approximated Fisher information. By default, the continuous control distribution is uniform on the unit interval, thus cut with breaks defined by the target control discrete probability distribution generates the outcome. The plot shows the distribution of the parameter estimates and the corresponding population values as red dots (Figure 12.1).

In the last example, we sample from 4×3 tables with odds ratios 2 and 3 for three strata with different control distributions, see Figure 12.2, and again compare the simulation results to the power function.

```
> power.freelway.test(n = N, prob = prb, delta = delta, seed = 3)

Stratified 3-sample Kruskal-Wallis test against proportional odds alternatives

      n = 15
Total sample size = 45 (Ctrl) + 45 (B) + 45 (C) = 135
```

```

> prb <- rep.int(1, 4) / 4
> pw <- numeric(Nsim)
> cf <- matrix(0, nrow = Nsim, ncol = length(delta))
> colnames(cf) <- names(delta)
> for (i in seq_along(pw)) {
+   nd <- rfreelway(n = N, prob = prb, delta = delta)
+   ft <- freelway(y ~ groups, data = nd)
+   cf[i,] <- coef(ft)
+   pw[i] <- summary(ft, test = "Permutation")$p.value
+ }
> mean(pw < .05)

```

```
[1] 0.26
```

```

> boxplot(cf, las = 1, ylab = expression(hat(delta)))
> points(c(1:2), delta, pch = 19, col = "red")
> power.freelway.test(n = N, prob = prb, delta = delta)

```

3-sample Kruskal-Wallis test against proportional odds alternatives

```

      n = 15
Total sample size = 15 (Control) + 15 (B) + 15 (C) = 45
      power = 0.29158
      sig.level = 0.05
log-odds ratio = 0.69315, 1.09861

```

NOTE: 'n' is sample size in control group

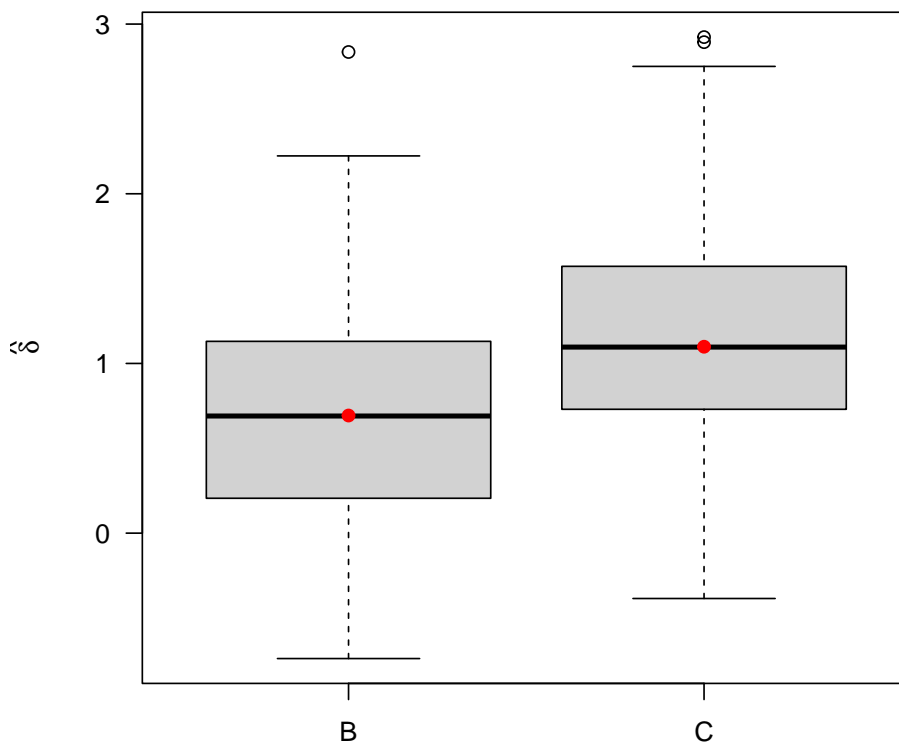


Figure 12.1: Power simulation for proportional-odds model and corresponding power approximation.

```

> prb <- cbind(S1 = rep(1, 4),
+             S2 = c(1, 2, 1, 2),
+             S3 = 1:4)
> dimnames(prb) <- list(Ctrl = paste0("i", seq_len(nrow(prb))),
+                       Strata = colnames(prb))
> pw <- numeric(Nsim)
> cf <- matrix(0, nrow = Nsim, ncol = length(delta))
> colnames(cf) <- names(delta)
> for (i in seq_along(pw)) {
+   nd <- rfree1way(n = N, prob = prb, delta = delta)
+   ft <- free1way(y ~ groups | blocks, data = nd)
+   cf[i,] <- coef(ft)
+   pw[i] <- summary(ft, test = "Permutation")$p.value
+ }
> mean(pw < .05)

[1] 0.79

> boxplot(cf, las = 1, ylab = expression(hat(delta)))
> points(c(1:2), delta, pch = 19, col = "red")

```

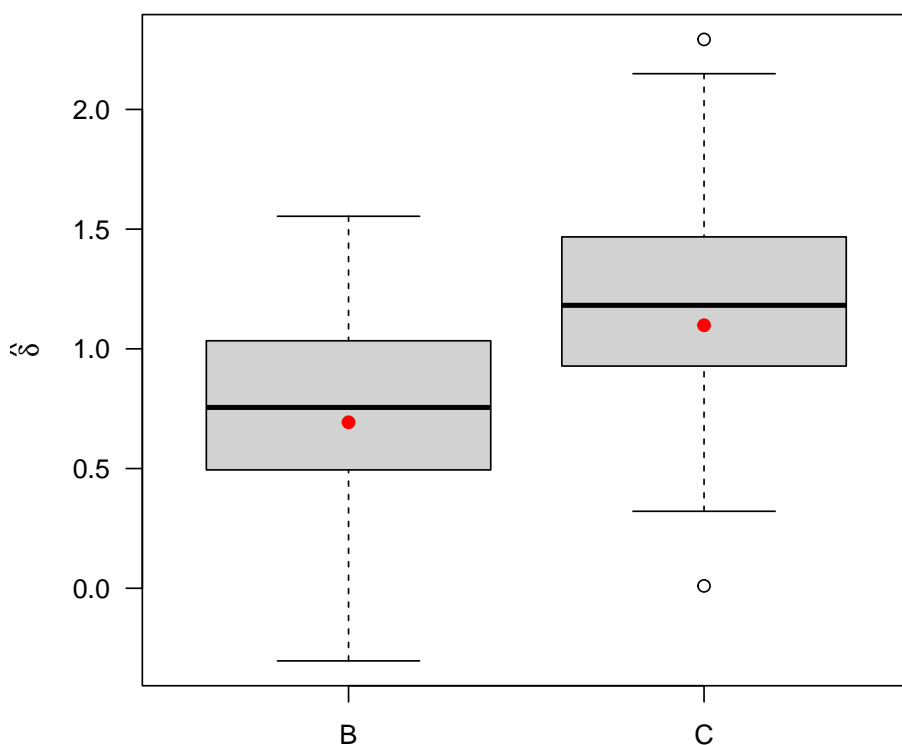


Figure 12.2: Power simulation for stratified proportional-odds model and corresponding power approximation.

```

        power = 0.69905
        sig.level = 0.05
        log-odds ratio = 0.69315, 1.09861

NOTE: 'n' is sample size in control group of first stratum

> power.freelway.test(power = .8, prob = prb, delta = delta, seed = 3)

Stratified 3-sample Kruskal-Wallis test against proportional odds alternatives

        n = 19
Total sample size = 57 (Ctrl) + 57 (B) + 57 (C) = 171
        power = 0.80451
        sig.level = 0.05
        log-odds ratio = 0.69315, 1.09861

NOTE: 'n' is sample size in control group of first stratum

> power.freelway.test(n = 19, prob = prb, delta = delta, seed = 3)

Stratified 3-sample Kruskal-Wallis test against proportional odds alternatives

        n = 19
Total sample size = 57 (Ctrl) + 57 (B) + 57 (C) = 171
        power = 0.80451
        sig.level = 0.05
        log-odds ratio = 0.69315, 1.09861

NOTE: 'n' is sample size in control group of first stratum

```

Chapter 13

Penalisation

Sometimes, especially under complete separation, the maximum likelihood estimator does not exist. We could think of offering the option to add a penalty term to the log-likelihood, for example half of the log-determinant of the Hessian (Jeffreys prior) as suggested by [Firth \(1993\)](#) and studied in [Kosmidis and Firth \(2020\)](#). Here is an example

```
> N <- 20
> w <- gl(2, N)
> y <- rnorm(length(w), mean = c(-2, 3)[w])
> x <- freelway(y ~ w, link = "probit")
> coef(x)

      w2
6.6457

> logLik(x)

[1] -119.85

> pll <- function(cf) {
+
+   start <- x$par
+   start[1] <- cf
+   x$profile(start, fix = 1)
+ }
> ### https://doi.org/10.1111/j.0006-341X.2001.00114.x
> ### https://doi.org/10.1111/j.1467-9876.2012.01057.x
> ### https://doi.org/10.1186/s12874-017-0313-9
> ### https://files.osf.io/v1/resources/fet4d_v3/providers/osfstorage/682fb176db88f967facacb5a?format=pdf
> ### https://doi.org/10.1002/sim.6537
> ### https://doi.org/10.1007/s11222-023-10217-3
> ### https://arxiv.org/abs/2510.06465
> fun <- function(cf) {
+   ret <- pll(cf)
+   ret$value - .5 * determinant(ret$hessian, logarithm = TRUE)$modulus
+ }
> ci <- confint(x, level = .99, test = "Wald")
> grd <- seq(from = ci[1], to = ci[2], length.out = 50)
> optim(coef(x), fn = fun, method = "Brent",
+       lower = min(grd), upper = max(grd))[c("par", "value")]

$par
[1] 4.7194

$value
[1] 120.46
```

```
attr("logarithm")  
[1] TRUE
```

The `MPL_Jeffreys` argument can be used to request this type of penalisation from `free1way` (this argument should be added to `free1way.table` and documented)

```
> free1way(y ~ w, link = "probit", MPL_Jeffreys = TRUE)
```

```
2-sample van der Waerden normal scores test against latent normal shift  
alternatives, with Jeffreys prior penalisation
```

```
data: y by w (1, 2)
```

```
Perm Z = 5, p-value = 5.7e-07
```

```
alternative hypothesis: true generalised Cohen's d is not equal to 0
```


Chapter 14

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Index

Files

"free1way.R" Defined by ?.

"linkfun.R" Defined by ?.

"utils.R" Defined by ?.

Fragments

<add legend ?> Referenced in ?.

<cloglog ?> Referenced in ?.

<confint lower ?> Referenced in ?.

<confint upper ?> Referenced in ?.

<density prob ratio ?> Referenced in ?, ?.

<design args ?> Referenced in ?, ?.

<determine steps in blocks ?> Referenced in ?.

<diagonal elements for Hessian of intercepts ?> Referenced in ?.

<do optim ?> Referenced in ?, ?.

<estimate Fisher information ?> Referenced in ?.

<exact proportional odds ?> Referenced in ?.

<extract plot data ?> Referenced in ?.

<formula business ?> Referenced in ?.

<free1way confint ?> Referenced in ?.

<free1way factor ?> Referenced in ?.

<free1way formula ?> Referenced in ?.

<free1way generic and table method (main workhorse) ?> Referenced in ?.

<free1way methods ?> Referenced in ?.

<free1way numeric ?> Referenced in ?.

<free1way permutation tests ?> Referenced in ?.

<free1way print ?> Referenced in ?.

<free1way summary ?> Referenced in ?.

<full Hessian ?> Referenced in ?.

<Hessian ?> Referenced in ?.

<Hessian prep ?> Referenced in ?.

<intercept / shift contributions to Hessian ?> Referenced in ?.

<Jeffreys penalisation ?> Referenced in ?.

<link2fun ?> Referenced in ?, ?, ?, ?.

<linkfun ?> Referenced in ?.

<logit ?> Referenced in ?.

<logLik, gradient, Hessian ?> Referenced in ?.

<loglog ?> Referenced in ?.

<LRT ?> Referenced in ?.

<marginal fit ?> Referenced in ?.

<marginal plot ?> Referenced in ?.

<ML estimation ?> Referenced in ?.

<model plot ?> Referenced in ?.

<negative logLik ?> Referenced in ?.

<negative score ?> Referenced in ?.

<negative score residuals ?> Referenced in ?.

<Newton ?> Not referenced.

<Newton convergence ?> Referenced in ?.

<Newton step halving ?> Referenced in ?.
 <Newton update ?> Referenced in ?.
 <NewtonRaphson ?> Referenced in ?.
 <off-diagonal elements for Hessian of intercepts ?> Referenced in ?.
 <optim ?> Referenced in ?.
 <parm to prob ?> Referenced in ?, ?, ?, ?.
 <permutation confint ?> Referenced in ?.
 <Permutation p-values ?> Referenced in ?.
 <Permutation statistics ?> Referenced in ?.
 <plot free1way ?> Referenced in ?.
 <post processing ?> Referenced in ?.
 <power ?> Referenced in ?.
 <power args check ?> Referenced in ?.
 <power call ?> Referenced in ?.
 <power htest output ?> Referenced in ?.
 <power inversion ?> Referenced in ?.
 <power setup ?> Referenced in ?.
 <ppplot ?> Referenced in ?.
 <probit ?> Referenced in ?.
 <profile ?> Referenced in ?.
 <r2dsim ?> Referenced in ?.
 <random seed ?> Referenced in ?.
 <Rao ?> Referenced in ?.
 <refit block intercepts ?> Referenced in ?.
 <resampling ?> Referenced in ?.
 <rfree1way ?> Referenced in ?.
 <ROC bands ?> Referenced in ?.
 <setup and starting values ?> Referenced in ?.
 <setup canvas ?> Referenced in ?.
 <statistics ?> Referenced in ?, ?.
 <Strasser Weber ?> Referenced in ?.
 <stratified Hessian ?> Referenced in ?.
 <stratified negative logLik ?> Referenced in ?.
 <stratified negative score ?> Referenced in ?.
 <stratified negative score residual ?> Referenced in ?.
 <stratum prep ?> Referenced in ?, ?, ?, ?, ?.
 <table2list body ?> Referenced in ?.
 <variable names and checks ?> Referenced in ?, ?.
 <Wald statistic ?> Referenced in ?.

Identifiers

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