2014627

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(event rate) Peto(Yusuf et. al., 1985) DerSimonian-Laird(DerSimonian and Laird, 1986)Peto Mental-Haenzel(odds ratio)DSL(DerSimonian-Laird)Peto(within-study variation)DSL(among-study variation)

Peto

Peto Mentel-Haenzel $i(i=1,\cdots,K)Nn_iN_i-n_i2d_i(\text{event})O_i(d_i-O_i)$ $E_i=(n_i/N_i)d_iN_i,n_i,d_iO_i(\text{ hypergeometric distribution})O_i-E_i0V_i=E_i[(N_i-n_i)/N_i][(N_i-d_i)(N_i-1)]$

$$\frac{\left[\sum_{i=1}^{K} (O_i - E_i)\right]^2}{\sum_{i=1}^{K} V_i}$$

 $1\chi^2 K$ (pooled)

$$\widehat{OR} = exp\left[\frac{\sum_{i=1}^{K} (O_i - E_i)}{\sum_{i=1}^{K} V_i}\right]$$

 $\ln(\widehat{OR})~SE(\ln\widehat{OR}) = (\sum V_i)^{-1/2} \ln(\widehat{OR}) \chi^2 \text{-}OR100 (1-\alpha)\%$

$$exp\left[\frac{\sum (O_i - E_i)}{\sum V_i} \pm \frac{Z_{\alpha/2}}{\sqrt{\sum V_i}}\right]$$

$$Q = \sum \left[\frac{(O_i - E_i)^2}{V_i} \right] - \frac{\left[\sum (O_i - E_i)\right]^2}{\sum V_i}$$

$$K - 1\chi^2\chi^2_\alpha(K-1)$$

DerSimonian-Laird

$$i(i=1,\cdots,K)d_{t_i}d_{c_i}n_{t_i}n_{c_i}$$

$$\hat{\theta}_i = \hat{p_{t_i}} - \hat{p_{c_i}} = d_{t_i}/n_{t_i} - d_{c_i}/n_{c_i}$$

$$S_i = \frac{\hat{p_{t_i}}(1 - \hat{p_{t_i}})}{n_{t_i}} + \frac{\hat{p_{c_i}}(1 - \hat{p_{c_i}})}{n_{c_i}}$$

$$Q = \sum w_i (\hat{\theta}_i - \bar{\theta}_w)^2$$

$$w_i = S_i^{-1} \bar{\theta}_w = \sum w_i \hat{\theta}_i / \sum w_i n_{c_i} n_{t_i} QK - 1\chi^2$$

$$\hat{\tau}^2 = \max \left[0, \frac{Q - (K - 1)}{\sum w_i - (\sum w_i^2 / \sum w_i)} \right]$$

$$w_i^* = (S_i + \hat{\tau}^2)^{-1}$$

$$\hat{\mu} = \frac{\sum w_i^* \hat{\theta}_i}{\sum w_i^*}$$

$$SE(\hat{\mu}) = (\sum w_i^*)^{-1/2}$$

Rmeta package

R metaFleiss(1993)Aspirin meta

```
library(meta)
## Loading required package: grid
## Loading 'meta' package (version 3.6-0).
data("Fleiss93")
Fleiss93
##
     study year event.e n.e event.c n.c
## 1 MRC-1 1974
                49 615
                                 67 624
      CDP 1976
                   44 758
                                 64 771
## 3 MRC-2 1979
                   102 832
                                126 850
     GASP 1979
                  32 317
                                 38 309
## 5 PARIS 1980
                   85 810
                                52 406
                  246 2267
## 6 AMIS 1980
                                219 2257
                1570 8587
## 7 ISIS-2 1988
                               1720 8600
```

metabin

```
m1 <- metabin(event.e, n.e, event.c, n.c, data=Fleiss93, studlab=paste(study,year), sm="OR")
m1</pre>
```

```
##
                  OR
                               95%-CI %W(fixed) %W(random)
## MRC-1 1974 0.7197 [0.4890; 1.0593]
                                           3.18
                                                      8.21
              0.6808 [0.4574; 1.0132]
                                                      7.85
## CDP 1976
                                           3.10
## MRC-2 1979 0.8029 [0.6065; 1.0629]
                                          5.68
                                                     13.23
              0.8007 [0.4863; 1.3186]
## GASP 1979
                                          1.80
                                                     5.36
## PARIS 1980 0.7981 [0.5526; 1.1529]
                                          3.22
                                                     8.89
## AMIS 1980
              1.1327 [0.9347; 1.3728]
                                                     20.70
                                          10.15
## ISIS-2 1988 0.8950 [0.8294; 0.9657]
                                          72.88
                                                     35.77
## Number of studies combined: k=7
##
##
                                        95%-CI
                           OR
                                                    z p.value
                       0.8969 [0.8405; 0.9570] -3.288
## Fixed effect model
                                                        0.001
## Random effects model 0.8763 [0.7743; 0.9917] -2.092
```

```
##
## Quantifying heterogeneity:
## tau^2 = 0.0096; H = 1.29 [1; 1.99]; I^2 = 39.7% [0%; 74.6%]
##
## Test of heterogeneity:
       Q d.f. p.value
##
##
   9.95
            6
               0.1269
##
## Details on meta-analytical method:
## - Mantel-Haenszel method
## - DerSimonian-Laird estimator for tau^2
Fixed effect modelPetoRandom effects modelDSL95%ORTest of HomogeneityTest of heterogeneity5%
1980
summary(m1, byvar=Fleiss93$year<1980, bylab="year<1980")</pre>
## Number of studies combined: k=7
##
##
                           OR
                                       95%-CI
                                                   z p.value
## Fixed effect model
                        0.897 [0.841; 0.957] -3.288
                                                       0.001
## Random effects model 0.876 [0.774; 0.992] -2.092
                                                       0.0365
##
## Quantifying heterogeneity:
## tau^2 = 0.0096; H = 1.29 [1; 1.99]; I^2 = 39.7% [0%; 74.6%]
##
## Test of heterogeneity:
       Q d.f. p.value
##
           6
               0.1269
##
  9.95
##
## Details on meta-analytical method:
## - Mantel-Haenszel method
## - DerSimonian-Laird estimator for tau^2
```

(Forest Plot)OR95%

```
forest(m1, comb.fixed = FALSE, leftcols = "studlab", rightcol = FALSE)
```

Study Odds Ratio MRC-1 1974 CDP 1976 MRC-2 1979 GASP 1979 PARIS 1980 AMIS 1980 ISIS-2 1988 Random effects model Heterogeneity: I-squared=39.7%, tau-squared=0.0096, p=0.1269



