Summary of analysis for survival model meta-analysis

Soumya Banerjee and Tom Bishop

1 January 2021

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1 Summary

This is a document that outlines analysis using survival models and meta-analyzing hazard ratios in the DataSHIELD platform.

2 Survival analysis in DataSHIELD

All code is available here:

- $\bullet \ \ https://github.com/neelsoumya/dsBaseClient/tree/absolute_newbie_client$
- https://github.com/neelsoumya/dsBase/tree/absolute_newbie
- $\bullet \ \ https://github.com/neelsoumya/datashield_testing_basic/blob/master/development_plan.rmd$
- $\bullet \ https://github.com/neelsoumya/datashield_testing_basic/blob/master/development_plan.pdf$
- $\bullet \ \ https://github.com/neelsoumya/datashield_testing_basic/tree/master/gui/survival_models_gui/survival_gui/$

3 Model parameters

This report and the model has been run according to the following parameters.

| Model | Exposure |
|----------------|----------|
| Survival model | poultry |

4 Meta-analysis model summary

A summary of the meta-analyzed model is shown below.

```
##
## Random-Effects Model (k = 7; tau^2 estimator: REML)
## tau^2 (estimated amount of total heterogeneity): 0.0000 (SE = 0.0000)
## tau (square root of estimated tau^2 value):
                                                   0.0026
## I^2 (total heterogeneity / total variability):
                                                   76.31%
## H^2 (total variability / sampling variability): 4.22
##
## Test for Heterogeneity:
## Q(df = 6) = 20.9055, p-val = 0.0019
##
## Model Results:
##
                                pval
## estimate
                        zval
                                       ci.lb
                se
                                              ci.ub
     1.0030 0.0012 845.6114 <.0001
                                     1.0007 1.0053
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
```

5 Cox model summary

A summary of the fitted Cox model for each study is shown below.

```
## Summary of Cox model .....
## $study1
## Call:
## survival::coxph(formula = formula, data = dataTable, weights = weights,
##
       ties = ties, singular.ok = singular.ok, model = model, x = x,
##
       y = y
##
##
     n= 814, number of events= 263
##
##
                 coef exp(coef)
                                 se(coef)
                                                z Pr(>|z|)
## POULTRY
             0.008259
                       1.008293
                                 0.002697
                                           3.062 0.002200 **
## AGEBASE
             0.036657
                       1.037338
                                 0.009654
                                            3.797 0.000146 ***
## GENDERO
                             NA
                                 0.000000
                                                        NA
                   NA
                                               NA
## PA2
            -0.139645
                       0.869667
                                 0.162268 -0.861 0.389469
## PA3
            -0.132103
                                 0.176617 -0.748 0.454482
                       0.876251
                                 0.309900 -2.021 0.043331 *
## PA4
            -0.626155
                       0.534643
## SMOKING2 0.186458
                      1.204974
                                 0.149540 1.247 0.212443
## SMOKING3 -0.192741 0.824696
                                 0.248343 -0.776 0.437687
```

```
## SMOKING4 0.165784 1.180318 0.263561 0.629 0.529338
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
           exp(coef) exp(-coef) lower .95 upper .95
## POULTRY
              1.0083
                         0.9918
                                  1.0030
                                            1.0136
## AGEBASE
              1.0373
                         0.9640
                                  1.0179
                                            1.0572
## GENDERO
                  NA
                            NA
                                      NA
                                                NA
## PA2
              0.8697
                         1.1499
                                  0.6327
                                            1.1953
## PA3
              0.8763
                                  0.6199
                                            1.2387
                        1.1412
## PA4
              0.5346
                        1.8704
                                  0.2913
                                            0.9814
              1.2050
                         0.8299
                                  0.8989
## SMOKING2
                                            1.6153
## SMOKING3
              0.8247
                         1.2126
                                  0.5069
                                           1.3418
## SMOKING4
              1.1803
                         0.8472
                                  0.7041
                                            1.9785
##
## Concordance= 0.623 (se = 0.02)
## Likelihood ratio test= 33.32 on 8 df,
                                          p=5e-05
## Wald test
                      = 33.99 on 8 df,
                                          p = 4e - 05
## Score (logrank) test = 34.41 on 8 df,
                                          p=3e-05
##
## $study2
## Call:
## survival::coxph(formula = formula, data = dataTable, weights = weights,
##
      ties = ties, singular.ok = singular.ok, model = model, x = x,
##
      y = y
##
##
    n= 3045, number of events= 1246
##
     (4 observations deleted due to missingness)
##
##
                coef exp(coef) se(coef)
                                             z Pr(>|z|)
## POULTRY
            0.006312 1.006331 0.001353 4.664 3.10e-06 ***
## AGEBASE
           0.037132 1.037830 0.003827 9.703 < 2e-16 ***
## GENDERO
          0.284419 1.328989 0.065476 4.344 1.40e-05 ***
           -0.351082  0.703926  0.067933  -5.168  2.37e-07 ***
## PA2
## PA3
           -0.387865 0.678504 0.091371 -4.245 2.19e-05 ***
## PA4
           ## SMOKING2 -0.044044 0.956912 0.073785 -0.597
                                                 0.5506
## SMOKING3 0.140839 1.151239 0.070064 2.010
                                                 0.0444 *
## SMOKING4 -0.783791 0.456672 0.710538 -1.103
                                                 0.2700
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
##
           exp(coef) exp(-coef) lower .95 upper .95
## POULTRY
              1.0063
                         0.9937
                                  1.0037
                                           1.0090
## AGEBASE
              1.0378
                                            1.0456
                         0.9635
                                  1.0301
## GENDERO
              1.3290
                         0.7525
                                  1.1689
                                            1.5110
## PA2
              0.7039
                                  0.6162
                                            0.8042
                         1.4206
## PA3
              0.6785
                         1.4738
                                  0.5673
                                            0.8116
## PA4
              0.6735
                                  0.5555
                         1.4847
                                            0.8167
## SMOKING2
              0.9569
                         1.0450
                                  0.8281
                                            1.1058
## SMOKING3
                         0.8686
                                  1.0035
             1.1512
                                            1.3207
## SMOKING4
              0.4567
                         2.1898
                                  0.1134
                                            1.8383
##
```

```
## Concordance= 0.643 (se = 0.01)
## Likelihood ratio test= 183.3 on 9 df,
                                         p=<2e-16
                      = 182 on 9 df, p=<2e-16
## Score (logrank) test = 183.9 on 9 df, p=<2e-16
##
## $study3
## Call:
## survival::coxph(formula = formula, data = dataTable, weights = weights,
##
      ties = ties, singular.ok = singular.ok, model = model, x = x,
##
      y = y
##
##
    n= 5605, number of events= 2360
##
##
                 coef exp(coef)
                                  se(coef)
                                               z Pr(>|z|)
## POULTRY
            0.0019543 1.0019562 0.0006908 2.829 0.00467 **
           0.0388380 1.0396020 0.0027714 14.014 < 2e-16 ***
## AGEBASE
## GENDERO
          0.4063890 1.5013865 0.0517118 7.859 3.88e-15 ***
           -0.0584568  0.9432190  0.0495506  -1.180  0.23810
## PA2
## PA3
           -0.1269046   0.8808177   0.0614389   -2.066   0.03887 *
## PA4
           ## SMOKING2 -0.0143287 0.9857735 0.0628746 -0.228 0.81973
## SMOKING3 0.1160177 1.1230158 0.0535209 2.168 0.03018 *
## SMOKING4 0.5286285 1.6966037 0.7084468 0.746 0.45556
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
           exp(coef) exp(-coef) lower .95 upper .95
## POULTRY
             1.0020
                        0.9980
                                  1.0006
                                           1.0033
## AGEBASE
              1.0396
                        0.9619
                                  1.0340
                                           1.0453
## GENDERO
              1.5014
                        0.6661
                                  1.3567
                                           1.6615
## PA2
              0.9432
                        1.0602
                                  0.8559
                                           1.0394
## PA3
              0.8808
                       1.1353
                                  0.7809
                                           0.9935
## PA4
              0.8124
                        1.2309
                                  0.7017
                                           0.9405
                                           1.1151
## SMOKING2
            0.9858
                        1.0144
                                  0.8715
## SMOKING3
            1.1230
                        0.8905
                                  1.0112
                                           1.2472
## SMOKING4
            1.6966
                        0.5894
                                  0.4232
                                           6.8016
##
## Concordance= 0.64 (se = 0.007)
## Likelihood ratio test= 363.2 on 9 df,
                                         p=<2e-16
                      = 360.3 on 9 df,
## Wald test
                                         p=<2e-16
## Score (logrank) test = 366 on 9 df, p=<2e-16
##
## $study4
## Call:
## survival::coxph(formula = formula, data = dataTable, weights = weights,
##
      ties = ties, singular.ok = singular.ok, model = model, x = x,
##
      y = y
##
##
    n= 2107, number of events= 728
##
     (113 observations deleted due to missingness)
##
##
                 coef exp(coef)
                                  se(coef)
                                             z Pr(>|z|)
```

```
## POULTRY -0.0009743 0.9990261 0.0018951 -0.514 0.607147
## AGEBASE
          0.0216436 1.0218795 0.0041491 5.216 1.82e-07 ***
## GENDERO
           0.2244742 1.2516645 0.0774700 2.898 0.003761 **
           ## PA2
## PA3
           ## PA4
           -0.3029240 0.7386553 0.1283061 -2.361 0.018228 *
## SMOKING2 0.1859179 1.2043234 0.0841615 2.209 0.027170 *
## SMOKING3 0.0898226 1.0939801 0.1180628 0.761 0.446775
## SMOKING4 0.2552638 1.2908021 0.2845787 0.897 0.369725
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
##
           exp(coef) exp(-coef) lower .95 upper .95
## POULTRY
                                 0.9953
                                           1.0027
             0.9990
                        1.0010
## AGEBASE
             1.0219
                        0.9786
                                 1.0136
                                           1.0302
## GENDERO
             1.2517
                        0.7989
                                 1.0753
                                           1.4569
## PA2
             0.7478
                                 0.6225
                        1.3372
                                           0.8984
## PA3
             0.6758
                        1.4797
                                 0.5409
                                           0.8444
## PA4
             0.7387
                                 0.5744
                        1.3538
                                           0.9499
## SMOKING2
             1.2043
                        0.8303
                                 1.0212
                                           1.4203
## SMOKING3
             1.0940
                        0.9141
                                 0.8680
                                           1.3788
## SMOKING4
             1.2908
                        0.7747
                                 0.7390
                                           2.2547
##
## Concordance= 0.629 (se = 0.012)
## Likelihood ratio test= 98.91 on 9 df,
                                         p=<2e-16
## Wald test
                      = 96.89 on 9 df,
                                         p=<2e-16
## Score (logrank) test = 98.55 on 9 df,
                                         p=<2e-16
##
##
## $study5
## Call:
## survival::coxph(formula = formula, data = dataTable, weights = weights,
##
      ties = ties, singular.ok = singular.ok, model = model, x = x,
##
      y = y
##
##
    n= 2069, number of events= 745
##
     (144 observations deleted due to missingness)
##
##
                 coef exp(coef)
                                 se(coef)
                                               z Pr(>|z|)
            5.189e-03 1.005e+00 2.379e-03 2.181 0.029210 *
## POULTRY
           4.871e-02 1.050e+00 4.801e-03 10.146 < 2e-16 ***
## AGEBASE
## GENDERO
           4.775e-01 1.612e+00 1.035e-01 4.614 3.95e-06 ***
           -3.852e-01 6.803e-01 1.162e-01 -3.315 0.000918 ***
## PA2
## PA3
           -3.489e-01 7.055e-01 1.228e-01 -2.841 0.004503 **
           -5.570e-01 5.729e-01 1.174e-01 -4.745 2.09e-06 ***
## SMOKING2 7.371e-02 1.076e+00 8.550e-02 0.862 0.388637
## SMOKING3 2.365e-02 1.024e+00 9.636e-02 0.245 0.806124
## SMOKING4 -1.315e+01 1.939e-06 7.212e+02 -0.018 0.985449
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
##
           exp(coef) exp(-coef) lower .95 upper .95
## POULTRY 1.005e+00 9.948e-01
                                 1.0005
                                           1.0099
## AGEBASE 1.050e+00 9.525e-01
                                 1.0401
                                           1.0598
```

```
## GENDERO 1.612e+00 6.203e-01
                                1.3161
                                         1.9745
## PA2
          6.803e-01 1.470e+00
                                0.5417
                                         0.8543
          7.055e-01 1.418e+00
                                         0.8975
## PA3
                                0.5545
## PA4
          5.729e-01 1.745e+00
                                0.4552
                                         0.7212
## SMOKING2 1.076e+00 9.289e-01
                                0.9104
                                         1.2729
## SMOKING3 1.024e+00 9.766e-01
                                0.8477
                                         1.2368
## SMOKING4 1.939e-06 5.157e+05
                                0.0000
                                            Inf
##
## Concordance= 0.659 (se = 0.012)
## Likelihood ratio test= 167.9 on 9 df,
                                       p=<2e-16
## Wald test = 154.9 on 9 df,
                                      p=<2e-16
## Score (logrank) test = 158.1 on 9 df,
                                       p=<2e-16
##
## $study7
## Call:
## survival::coxph(formula = formula, data = dataTable, weights = weights,
      ties = ties, singular.ok = singular.ok, model = model, x = x,
##
      y = y
##
    n= 3420, number of events= 1497
##
##
##
               coef exp(coef) se(coef)
                                          z Pr(>|z|)
           0.001944 1.001945 0.001821 1.068 0.285735
## POULTRY
          0.053138 1.054575 0.003537 15.025 < 2e-16 ***
## AGEBASE
          0.343834 1.410345 0.056900 6.043 1.51e-09 ***
## GENDERO
## PA2
          ## PA3
## PA4
          ## SMOKING2 0.242791 1.274802 0.061597 3.942 8.09e-05 ***
                             0.072198 3.658 0.000254 ***
## SMOKING3 0.264102 1.302261
## SMOKING4 -0.495348 0.609359 1.001980 -0.494 0.621045
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
          exp(coef) exp(-coef) lower .95 upper .95
##
## POULTRY
            1.0019
                       0.9981
                                0.9984
                                         1.0055
## AGEBASE
             1.0546
                       0.9482
                                1.0473
                                         1.0619
## GENDERO
             1.4103
                       0.7090
                                1.2615
                                         1.5767
## PA2
             0.8221
                       1.2164
                                0.7196
                                         0.9391
## PA3
             0.7473
                       1.3381
                                0.6434
                                         0.8680
## PA4
             0.7383
                       1.3545
                                0.6259
                                         0.8708
## SMOKING2
             1.2748
                       0.7844
                                1.1298
                                         1.4384
## SMOKING3
             1.3023
                       0.7679
                                1.1304
                                         1.5002
             0.6094
## SMOKING4
                       1.6411
                                0.0855
                                         4.3427
##
## Concordance= 0.696 (se = 0.009)
## Likelihood ratio test= 413.6 on 9 df,
                                      p=<2e-16
## Wald test
                     = 384.5 on 9 df,
                                       p=<2e-16
## Score (logrank) test = 397.2 on 9 df,
                                       p=<2e-16
##
##
## $study8
## Call:
```

```
## survival::coxph(formula = formula, data = dataTable, weights = weights,
##
      ties = ties, singular.ok = singular.ok, model = model, x = x,
##
      y = y
##
##
    n= 5167, number of events= 2363
##
     (19 observations deleted due to missingness)
##
##
                 coef
                      exp(coef)
                                  se(coef)
                                               z Pr(>|z|)
## POULTRY
            0.0002034
                      1.0002034 0.0013437 0.151 0.879697
## AGEBASE
            0.0176385
                      1.0177950 0.0022050 7.999 1.25e-15 ***
## GENDERO
            0.2582253 1.2946305 0.0420403 6.142 8.13e-10 ***
## PA2
                      -0.2179843
## PA3
           -0.1970206
                      0.8211737
                                0.0567991 -3.469 0.000523 ***
## PA4
           -0.2922857
                      0.7465552  0.0686133  -4.260  2.05e-05 ***
## SMOKING2 0.1347059
                      1.1442002 0.0496762 2.712 0.006694 **
## SMOKING3 0.0895323
                      1.0936626
                                 0.0513898 1.742 0.081470 .
## SMOKING4 -0.0459120 0.9551260 0.2908950 -0.158 0.874591
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
##
           exp(coef) exp(-coef) lower .95 upper .95
## POULTRY
              1.0002
                        0.9998
                                  0.9976
## AGEBASE
              1.0178
                                           1.0222
                        0.9825
                                  1.0134
## GENDERO
              1.2946
                                           1.4058
                        0.7724
                                  1.1922
## PA2
              0.8041
                        1.2436
                                  0.7268
                                           0.8897
## PA3
              0.8212
                        1.2178
                                  0.7347
                                           0.9179
## PA4
              0.7466
                        1.3395
                                  0.6526
                                           0.8540
## SMOKING2
              1.1442
                        0.8740
                                  1.0380
                                           1.2612
              1.0937
                        0.9144
                                  0.9889
## SMOKING3
                                           1.2096
## SMOKING4
              0.9551
                        1.0470
                                  0.5401
                                           1.6892
##
## Concordance= 0.598 (se = 0.008)
                                         p=<2e-16
## Likelihood ratio test= 165.5 on 9 df,
## Wald test
                      = 163.6 on 9 df,
                                         p=<2e-16
## Score (logrank) test = 164.6 on 9 df,
                                         p = < 2e - 16
##
##
## $study9
## Call:
## survival::coxph(formula = formula, data = dataTable, weights = weights,
      ties = ties, singular.ok = singular.ok, model = model, x = x,
##
      y = y
##
    n= 5167, number of events= 2363
##
##
     (19 observations deleted due to missingness)
##
##
                 coef
                      exp(coef)
                                  se(coef)
                                               z Pr(>|z|)
## POULTRY
            0.0002034
                      1.0002034
                                0.0013437 0.151 0.879697
## AGEBASE
            0.0176385
                      1.0177950
                                 0.0022050 7.999 1.25e-15 ***
## GENDERO
            0.2582253
                      1.2946305
                                 0.0420403 6.142 8.13e-10 ***
                                0.0515820 -4.226 2.38e-05 ***
## PA2
                      0.8041381
           -0.2179843
## PA3
           -0.1970206
                      ## PA4
           -0.2922857
                      ## SMOKING2 0.1347059 1.1442002 0.0496762 2.712 0.006694 **
```

```
## SMOKING3 0.0895323
                        1.0936626 0.0513898 1.742 0.081470 .
## SMOKING4 -0.0459120
                         0.9551260
                                    0.2908950 -0.158 0.874591
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
##
            exp(coef) exp(-coef) lower .95 upper .95
##
                           0.9998
                                     0.9976
## POULTRY
               1.0002
                                                1.0028
## AGEBASE
               1.0178
                           0.9825
                                     1.0134
                                                1.0222
## GENDERO
               1.2946
                           0.7724
                                     1.1922
                                                1.4058
## PA2
               0.8041
                           1.2436
                                     0.7268
                                                0.8897
## PA3
               0.8212
                           1.2178
                                     0.7347
                                                0.9179
               0.7466
                           1.3395
                                     0.6526
## PA4
                                                0.8540
## SMOKING2
               1.1442
                           0.8740
                                     1.0380
                                                1.2612
## SMOKING3
               1.0937
                           0.9144
                                     0.9889
                                                1.2096
## SMOKING4
               0.9551
                           1.0470
                                     0.5401
                                                1.6892
##
## Concordance= 0.598
                       (se = 0.008)
## Likelihood ratio test= 165.5
                                              p = < 2e - 16
                                  on 9 df,
## Wald test
                         = 163.6
                                              p=<2e-16
## Score (logrank) test = 164.6
                                  on 9 df,
                                              p=<2e-16
```

6 Forest plot of meta-analyzed hazard ratios

We now outline the hazard ratios from the survival models which are meta-analyzed. We use the *metafor* package for meta-analysis. We show a forest plot below.

7 References

- https://github.com/datashield
- http://www.metafor-project.org
- https://github.com/neelsoumya/datashield_testing_basic/tree/master/gui/survival_models_gui

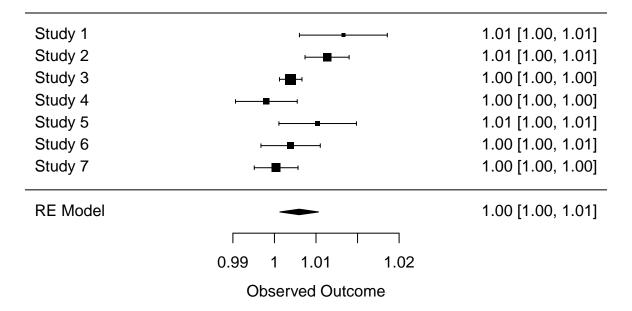


Figure 1: Forest plot of meta-analyzed hazard ratios.