Cox PH Models

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Load packages:

```
if (!require("pacman"))
   install.packages("pacman", repos = "http://cran.us.r-project.org/")
p_load("tidyverse", "survival", "kableExtra")

Import data:
breast <- readRDS(file = "breast_final.rds")

# delete all survival time = 0
breast <- subset(breast, SRV_TIME_MON != 0)</pre>
```

Cox Model: All Covariates

convert SEX to dummy: O=male, 1=female
breast\$SEX <- ifelse(breast\$SEX == 1, 0, 1)</pre>

Using the Breslow method of handling ties, we fit a Cox proportional hazards model to the data including all 13 covariates: race, sex, stage, breast subtype, age dx, age, marital status, benign tumor count, malignant tumor count, primary site, pr status, er status, insurance status.

```
##
    n= 55333, number of events= 3033
##
##
                            coef exp(coef)
                                              se(coef)
                                                            z Pr(>|z|)
## factor(SEX)1
                       -0.234365
                                   0.791073
                                              0.213390 -1.098 0.272077
                                              0.709709 2.577 0.009977 **
## factor(stage)1
                                   6.225421
                        1.828641
## factor(stage)2
                        3.106446 22.341492
                                              0.708153 4.387 1.15e-05 ***
## factor(stage)3
                        4.240722 69.457958 0.708109 5.989 2.11e-09 ***
## factor(stage)4
                        5.860086 350.754397
                                              0.707926 8.278 < 2e-16 ***
## factor(RAC_RECY)2
                                              0.067114 3.462 0.000537 ***
                        0.232315
                                   1.261517
## factor(RAC_RECY)3
                        0.081242
                                   1.084634
                                              0.197643 0.411 0.681031
## factor(RAC_RECY)4
                       -0.258741
                                   0.772023
                                              0.077398 -3.343 0.000829 ***
## factor(BRST_SUB)2
                       -0.969818
                                   0.379152
                                              0.152118 -6.375 1.82e-10 ***
                                              0.064187 2.714 0.006642 **
## factor(BRST_SUB)3
                        0.174221
                                   1.190319
## factor(BRST_SUB)4
                       -0.150397
                                   0.860366
                                              0.142171 -1.058 0.290118
```

```
## AGE DX
                          0.033083
                                     1.033637
                                                 0.014080 2.350 0.018794 *
## Age
                         -0.008105
                                                 0.014056 -0.577 0.564208
                                     0.991928
## factor(MAR STAT)2
                         -0.373226
                                     0.688509
                                                 0.049662 -7.515 5.67e-14 ***
                                                 0.146900 -0.958 0.337870
## factor(MAR_STAT)3
                         -0.140786
                                     0.868675
## factor(MAR_STAT)4
                         -0.154337
                                     0.856983
                                                 0.064872 -2.379 0.017354 *
## factor(MAR STAT)5
                          0.075775
                                     1.078719
                                                 0.063998 1.184 0.236410
## factor(MAR STAT)6
                         -0.121050
                                     0.885989
                                                 0.336358 -0.360 0.718933
## MALIGCOUNT
                          0.253995
                                     1.289166
                                                 0.055057
                                                           4.613 3.96e-06 ***
## BENBORDCOUNT
                          0.083414
                                     1.086991
                                                 0.232919
                                                           0.358 0.720252
## factor(PRIMSITE)1
                         -0.244291
                                     0.783260
                                                 0.255573 -0.956 0.339144
## factor(PRIMSITE)2
                         -0.164098
                                     0.848658
                                                 0.254097 -0.646 0.518402
## factor(PRIMSITE)3
                         -0.043522
                                     0.957412
                                                 0.259180 -0.168 0.866645
## factor(PRIMSITE)4
                         -0.276886
                                                 0.247999 -1.116 0.264215
                                     0.758141
## factor(PRIMSITE)5
                         -0.306483
                                     0.736031
                                                 0.256539 -1.195 0.232210
## factor(PRIMSITE)6
                         -0.218727
                                     0.803541
                                                 0.336821 -0.649 0.516089
## factor(PRIMSITE)7
                         -0.095879
                                     0.908574
                                                 0.248255 -0.386 0.699339
## factor(PRIMSITE)8
                                                           0.110 0.912494
                          0.027267
                                     1.027642
                                                 0.248121
## factor(ERSTATUS)1
                          1.080860
                                                           8.375
                                     2.947214
                                                 0.129059
                                                                  < 2e-16 ***
## factor(PRSTATUS)1
                                                 0.053097 12.314 < 2e-16 ***
                          0.653819
                                     1.922871
## factor(INSREC PUB)1
                         -0.144237
                                     0.865682
                                                 0.128615 -1.121 0.262091
## factor(INSREC_PUB)2
                         -0.513753
                                     0.598246
                                                 0.126632 -4.057 4.97e-05 ***
                                                 0.132869 -2.899 0.003745 **
## factor(INSREC_PUB)3
                         -0.385176
                                     0.680331
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
                        exp(coef) exp(-coef) lower .95 upper .95
## factor(SEX)1
                           0.7911
                                    1.264106
                                                 0.5207
                                                           1.2019
  factor(stage)1
                           6.2254
                                    0.160632
                                                 1.5490
                                                          25.0192
   factor(stage)2
                          22.3415
                                    0.044760
                                                 5.5761
                                                          89.5144
## factor(stage)3
                          69.4580
                                                17.3372
                                                         278.2695
                                    0.014397
## factor(stage)4
                         350.7544
                                    0.002851
                                                87.5821 1404.7232
  factor(RAC_RECY)2
                           1.2615
                                    0.792697
                                                 1.1060
                                                            1.4389
## factor(RAC_RECY)3
                                    0.921970
                                                 0.7363
                                                            1.5978
                           1.0846
## factor(RAC_RECY)4
                           0.7720
                                                 0.6634
                                    1.295298
                                                           0.8985
## factor(BRST_SUB)2
                                                 0.2814
                           0.3792
                                    2.637465
                                                           0.5109
## factor(BRST_SUB)3
                           1.1903
                                    0.840111
                                                 1.0496
                                                            1.3499
## factor(BRST SUB)4
                           0.8604
                                    1.162296
                                                 0.6511
                                                            1.1368
## AGE_DX
                           1.0336
                                    0.967458
                                                 1.0055
                                                            1.0626
## Age
                           0.9919
                                    1.008138
                                                 0.9650
                                                            1.0196
## factor(MAR_STAT)2
                           0.6885
                                    1.452413
                                                 0.6247
                                                           0.7589
## factor(MAR STAT)3
                           0.8687
                                    1.151178
                                                 0.6514
                                                            1.1585
## factor(MAR STAT)4
                           0.8570
                                    1.166884
                                                 0.7547
                                                           0.9732
## factor(MAR_STAT)5
                           1.0787
                                    0.927025
                                                 0.9516
                                                           1.2229
## factor(MAR_STAT)6
                           0.8860
                                    1.128682
                                                 0.4583
                                                            1.7129
## MALIGCOUNT
                           1.2892
                                    0.775695
                                                 1.1573
                                                            1.4361
## BENBORDCOUNT
                           1.0870
                                    0.919971
                                                 0.6886
                                                            1.7159
## factor(PRIMSITE)1
                           0.7833
                                    1.276716
                                                 0.4746
                                                            1.2926
## factor(PRIMSITE)2
                           0.8487
                                    1.178330
                                                 0.5158
                                                            1.3964
## factor(PRIMSITE)3
                           0.9574
                                    1.044483
                                                 0.5761
                                                            1.5912
## factor(PRIMSITE)4
                           0.7581
                                    1.319017
                                                 0.4663
                                                            1.2327
## factor(PRIMSITE)5
                           0.7360
                                                 0.4452
                                    1.358638
                                                            1.2169
## factor(PRIMSITE)6
                           0.8035
                                    1.244492
                                                 0.4152
                                                            1.5549
## factor(PRIMSITE)7
                           0.9086
                                    1.100626
                                                 0.5585
                                                            1.4780
## factor(PRIMSITE)8
                           1.0276
                                    0.973101
                                                 0.6319
                                                            1.6713
```

```
## factor(ERSTATUS)1
                        2.9472 0.339304
                                             2.2885
                                                       3.7955
## factor(PRSTATUS)1
                         1.9229 0.520056
                                             1.7328
                                                       2.1338
## factor(INSREC PUB)1
                                             0.6728
                         0.8657 1.155158
                                                       1.1139
## factor(INSREC_PUB)2
                         0.5982
                                1.671553
                                             0.4668
                                                       0.7668
## factor(INSREC_PUB)3
                         0.6803
                                 1.469873
                                             0.5244
                                                       0.8827
##
## Concordance= 0.889 (se = 0.006)
## Rsquare= 0.125 (max possible= 0.679 )
## Likelihood ratio test= 7415 on 33 df,
                                          p=<2e-16
## Wald test
                       = 7475 on 33 df,
                                          p=<2e-16
## Score (logrank) test = 15283 on 33 df,
                                         p=<2e-16
```

ANOVA Table: All Covariates

We constructed an Analysis of Variance table to summarize estimates of the risk coefficients and the results of the one degree of freedom tests for each covariate in the model:

	Coefficient	Exp. Coeff.	Std. Error	Z-Score	P-Value
factor(SEX)1	-0.2343650	0.7910730	0.2133903	-1.0982930	0.2720765
factor(stage)1	1.8286411	6.2254212	0.7097085	2.5766087	0.0099775
factor(stage)2	3.1064456	22.3414916	0.7081526	4.3866896	0.0000115
factor(stage)3	4.2407216	69.4579580	0.7081091	5.9887970	0.0000000
factor(stage)4	5.8600863	350.7543972	0.7079259	8.2778240	0.0000000
$factor(RAC_RECY)2$	0.2323147	1.2615166	0.0671137	3.4615099	0.0005372
$factor(RAC_RECY)3$	0.0812425	1.0846338	0.1976429	0.4110568	0.6810309
$factor(RAC_RECY)4$	-0.2587411	0.7720229	0.0773981	-3.3429909	0.0008288
$factor(BRST_SUB)2$	-0.9698181	0.3791520	0.1521181	-6.3754270	0.0000000
$factor(BRST_SUB)3$	0.1742212	1.1903188	0.0641868	2.7142821	0.0066420
$factor(BRST_SUB)4$	-0.1503972	0.8603662	0.1421707	-1.0578634	0.2901177
AGE_DX	0.0330834	1.0336367	0.0140805	2.3495887	0.0187942
Age	-0.0081050	0.9919278	0.0140565	-0.5766028	0.5642078
factor(MAR_STAT)2	-0.3732264	0.6885093	0.0496616	-7.5153933	0.0000000
factor(MAR_STAT)3	-0.1407861	0.8686751	0.1468997	-0.9583822	0.3378701
$factor(MAR_STAT)4$	-0.1543374	0.8569829	0.0648718	-2.3791128	0.0173544
factor(MAR_STAT)5	0.0757746	1.0787194	0.0639984	1.1840084	0.2364097
factor(MAR_STAT)6	-0.1210502	0.8859895	0.3363584	-0.3598847	0.7189334
MALIGCOUNT	0.2539954	1.2891659	0.0550571	4.6133054	0.0000040
BENBORDCOUNT	0.0834136	1.0869913	0.2329193	0.3581223	0.7202518
factor(PRIMSITE)1	-0.2442908	0.7832598	0.2555725	-0.9558572	0.3391444
factor(PRIMSITE)2	-0.1640985	0.8486585	0.2540967	-0.6458110	0.5184017
factor(PRIMSITE)3	-0.0435219	0.9574116	0.2591802	-0.1679214	0.8666451
factor(PRIMSITE)4	-0.2768864	0.7581406	0.2479985	-1.1164840	0.2642150
factor(PRIMSITE)5	-0.3064831	0.7360310	0.2565386	-1.1946861	0.2322097
factor(PRIMSITE)6	-0.2187273	0.8035408	0.3368214	-0.6493866	0.5160885
factor(PRIMSITE)7	-0.0958792	0.9085738	0.2482549	-0.3862127	0.6993392
factor(PRIMSITE)8	0.0272669	1.0276420	0.2481207	0.1098937	0.9124937
factor(ERSTATUS)1	1.0808603	2.9472139	0.1290589	8.3749408	0.0000000
factor(PRSTATUS)1	0.6538192	1.9228707	0.0530969	12.3137000	0.0000000
$factor(INSREC_PUB)1$	-0.1442372	0.8656823	0.1286153	-1.1214622	0.2620912
$factor(INSREC_PUB)2$	-0.5137528	0.5982462	0.1266319	-4.0570570	0.0000497
factor(INSREC_PUB)3	-0.3851760	0.6803309	0.1328691	-2.8989135	0.0037446

Cox Model: Top 9 Significant Variables

Using variable selection methods (LASSO, SCAD, MCP), we decided the top 9 significant variables were:

- Stage
- ERSTATUS
- PRSTATUS
- MALIGCOUNT
- RAC_RECY
- PRIMSITE
- BRST_SUB
- MAR_STAT
- INSREC_PUB

We fit a Cox model with these covariates plus sex:

```
fit2 <- coxph(Surv(SRV_TIME_MON, delta) ~ factor(SEX) + factor(stage) + factor(RAC_RECY) +
             factor(BRST_SUB) + factor(MAR_STAT) + MALIGCOUNT + factor(PRIMSITE) +
             factor(ERSTATUS) + factor(PRSTATUS) + factor(INSREC_PUB) ,
             data = breast, ties = "breslow" )
summary(fit2)
## Call:
## coxph(formula = Surv(SRV_TIME_MON, delta) ~ factor(SEX) + factor(stage) +
       factor(RAC_RECY) + factor(BRST_SUB) + factor(MAR_STAT) +
##
       MALIGCOUNT + factor(PRIMSITE) + factor(ERSTATUS) + factor(PRSTATUS) +
##
       factor(INSREC_PUB), data = breast, ties = "breslow")
##
##
    n=55333, number of events= 3033
##
##
                                               se(coef)
                             coef
                                   exp(coef)
                                                             z Pr(>|z|)
## factor(SEX)1
                        -0.349810
                                    0.704822
                                               0.213794 -1.636 0.10180
## factor(stage)1
                         1.856788
                                    6.403136
                                               0.709702 2.616 0.00889 **
## factor(stage)2
                         3.076275
                                   21.677495
                                               0.708156 4.344 1.40e-05 ***
## factor(stage)3
                                               0.708103 5.909 3.45e-09 ***
                         4.184014 65.628743
## factor(stage)4
                         5.831913 341.010252
                                               0.707915 8.238 < 2e-16 ***
## factor(RAC RECY)2
                         0.191037
                                    1.210504
                                               0.066904 2.855
                                                                0.00430 **
## factor(RAC RECY)3
                         0.067350
                                    1.069670
                                               0.197577 0.341 0.73319
## factor(RAC RECY)4
                        -0.312395
                                    0.731692
                                               0.077236 -4.045 5.24e-05 ***
## factor(BRST_SUB)2
                        -0.965328
                                    0.380858
                                               0.152375 -6.335 2.37e-10 ***
## factor(BRST_SUB)3
                         0.251263
                                    1.285648
                                               0.063903 3.932 8.43e-05 ***
                                               0.142382 -0.728 0.46644
## factor(BRST SUB)4
                        -0.103694
                                    0.901502
## factor(MAR STAT)2
                        -0.315727
                                    0.729258
                                               0.049500 -6.378 1.79e-10 ***
## factor(MAR_STAT)3
                        -0.099548
                                    0.905246
                                               0.146908 -0.678 0.49801
## factor(MAR_STAT)4
                        -0.008995
                                    0.991045
                                               0.064307 -0.140
                                                                0.88876
## factor(MAR_STAT)5
                         0.515853
                                    1.675066
                                               0.057982 8.897
                                                                < 2e-16 ***
## factor(MAR_STAT)6
                        -0.126694
                                    0.881003
                                               0.336125 -0.377
                                                                0.70623
## MALIGCOUNT
                         0.288007
                                    1.333767
                                               0.054876 5.248 1.54e-07 ***
## factor(PRIMSITE)1
                                               0.255827 -0.961
                        -0.245851
                                    0.782039
                                                                0.33655
## factor(PRIMSITE)2
                        -0.244650
                                    0.782979
                                               0.254533 -0.961
                                                                0.33647
## factor(PRIMSITE)3
                        -0.120621
                                    0.886370
                                               0.259590 -0.465
                                                                0.64217
## factor(PRIMSITE)4
                        -0.363928
                                               0.248432 -1.465
                                    0.694941
                                                                0.14295
## factor(PRIMSITE)5
                        -0.370951
                                    0.690078
                                               0.256980 -1.443
                                                                0.14888
## factor(PRIMSITE)6
                        -0.175356
                                               0.336763 -0.521
                                                                0.60257
                                    0.839158
## factor(PRIMSITE)7
                        -0.185623
                                    0.830586
                                               0.248680 -0.746
                                                                0.45541
## factor(PRIMSITE)8
                        -0.044644
                                    0.956338
                                               0.248489 -0.180
                                                                0.85742
## factor(ERSTATUS)1
                         1.033302
                                    2.810329
                                               0.128667 8.031 9.68e-16 ***
## factor(PRSTATUS)1
                         0.678659
                                    1.971233
                                               0.053083 12.785
                                                               < 2e-16 ***
## factor(INSREC PUB)1 -0.107959
                                    0.897665
                                               0.128455 -0.840
                                                                0.40066
## factor(INSREC PUB)2
                        -0.357259
                                    0.699591
                                               0.126129 -2.832
                                                                0.00462 **
## factor(INSREC_PUB)3 -0.148713
                                    0.861816
                                               0.131929 -1.127 0.25965
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
##
                       exp(coef) exp(-coef) lower .95 upper .95
## factor(SEX)1
                          0.7048
                                               0.4636
                                   1.418799
                                                         1.0717
## factor(stage)1
                          6.4031
                                   0.156173
                                               1.5933
                                                        25.7331
## factor(stage)2
                         21.6775
                                   0.046131
                                               5.4104
                                                        86.8545
## factor(stage)3
                         65.6287
                                   0.015237
                                              16.3816 262.9256
## factor(stage)4
                        341.0103
                                   0.002932
                                              85.1508 1365.6711
```

```
## factor(RAC_RECY)2
                            1.2105
                                     0.826102
                                                  1.0617
                                                             1.3801
## factor(RAC_RECY)3
                            1.0697
                                                  0.7262
                                                             1.5755
                                     0.934868
## factor(RAC_RECY)4
                            0.7317
                                     1.366695
                                                  0.6289
                                                             0.8513
## factor(BRST_SUB)2
                            0.3809
                                     2.625650
                                                  0.2825
                                                             0.5134
## factor(BRST_SUB)3
                            1.2856
                                     0.777818
                                                  1.1343
                                                             1.4572
  factor(BRST SUB)4
                                                  0.6820
                                                             1.1917
                            0.9015
                                     1.109260
## factor(MAR STAT)2
                            0.7293
                                     1.371256
                                                  0.6618
                                                             0.8036
## factor(MAR_STAT)3
                            0.9052
                                     1.104672
                                                  0.6788
                                                             1.2073
  factor(MAR_STAT)4
                            0.9910
                                     1.009035
                                                  0.8737
                                                             1.1242
   factor(MAR_STAT)5
                            1.6751
                                     0.596991
                                                  1.4951
                                                             1.8767
## factor(MAR_STAT)6
                            0.8810
                                     1.135070
                                                  0.4559
                                                             1.7025
## MALIGCOUNT
                            1.3338
                                     0.749756
                                                  1.1978
                                                             1.4852
## factor(PRIMSITE)1
                            0.7820
                                                  0.4737
                                                             1.2912
                                     1.278709
                                                             1.2895
## factor(PRIMSITE)2
                            0.7830
                                     1.277174
                                                  0.4754
                                                  0.5329
## factor(PRIMSITE)3
                            0.8864
                                     1.128197
                                                             1.4743
## factor(PRIMSITE)4
                            0.6949
                                     1.438971
                                                  0.4271
                                                             1.1309
## factor(PRIMSITE)5
                            0.6901
                                                  0.4170
                                     1.449112
                                                             1.1419
## factor(PRIMSITE)6
                            0.8392
                                                  0.4337
                                                             1.6237
                                     1.191671
## factor(PRIMSITE)7
                            0.8306
                                                  0.5102
                                                             1.3523
                                     1.203969
## factor(PRIMSITE)8
                            0.9563
                                     1.045655
                                                  0.5876
                                                             1.5564
## factor(ERSTATUS)1
                            2.8103
                                     0.355830
                                                  2.1839
                                                             3.6164
## factor(PRSTATUS)1
                            1.9712
                                     0.507297
                                                  1.7765
                                                             2.1874
## factor(INSREC_PUB)1
                            0.8977
                                                  0.6979
                                                             1.1547
                                     1.114002
## factor(INSREC_PUB)2
                            0.6996
                                     1.429407
                                                  0.5464
                                                             0.8958
## factor(INSREC_PUB)3
                            0.8618
                                     1.160340
                                                  0.6655
                                                             1.1161
## Concordance= 0.882
                        (se = 0.006)
   Rsquare= 0.121
                     (max possible= 0.679 )
## Likelihood ratio test= 7165
                                  on 30 df,
                                               p=<2e-16
                                  on 30 df,
## Wald test
                         = 7211
                                               p=<2e-16
## Score (logrank) test = 15003 on 30 df,
                                                p = < 2e - 16
```

Test PH Assumption for Sex

2

4

0

0

To test the proportional hazards assumption for Sex (a fixed-time covariate), we can create a time-dependent covariate $Z_2(t)$, defined as $Z_2(t) = Z_1 \times g(t)$, where g(t) is a known function of the time t. In most applications, we take $g(t) = \ln(t)$. A test of $H_0: \beta_2 = 0$ is a test of the proportional hazards assumption.

```
# convert dataset into a counting process-like dataset
cut.points <- unique(breast$SRV_TIME_MON[breast$delta == 1])</pre>
breast2 <- survSplit(data = breast, cut = cut.points, end = "SRV_TIME_MON", start = "t0",
                       event = "delta")
head(breast2)
##
     stage RAC_RECY SEX BRST_SUB AGE_DX Age MAR_STAT MALIGCOUNT BENBORDCOUNT
                                                         2
## 1
          1
                                  3
                                         45
                                              49
                                                                                    0
                    1
                        1
                                                                     1
                                                         2
## 2
          1
                        1
                                  3
                                         45
                                              49
                                                                     1
                                                                                    0
                    1
                                                         2
## 3
          1
                    1
                        1
                                  3
                                         45
                                              49
                                                                     1
                                                                                    0
                                  3
                                                         2
                                                                                    0
##
          1
                    1
                        1
                                         45
                                              49
                                                                     1
                                                         2
##
   5
          1
                        1
                                  3
                                         45
                                              49
                                                                     1
                                                                                    0
                    1
                                  3
                                                         2
##
   6
          1
                    1
                        1
                                         45
                                              49
                                                                     1
                                                                                    0
     PRIMSITE ERSTATUS PRSTATUS INSREC_PUB tO SRV_TIME_MON delta
##
## 1
             4
                       0
                                 0
                                              2
                                                 0
                                                                      0
```

2

0

2 1

```
## 3
                                           2 2
                                                           3
## 4
            4
                               0
                                           2 3
                                                           4
                                                                  0
                      0
## 5
            4
                      0
                               0
                                           2
                                             4
                                                           5
                                                                  0
## 6
            4
                      0
                               0
                                           2 5
                                                           6
                                                                  Ω
# create time-dependent covariate
breast2$tdc_sex <- breast2$SEX * log(breast2$SRV_TIME_MON)</pre>
coxph(Surv(t0, SRV_TIME_MON, delta) ~ SEX + tdc_sex, data = breast2, ties = "breslow")
## coxph(formula = Surv(t0, SRV_TIME_MON, delta) ~ SEX + tdc_sex,
##
       data = breast2, ties = "breslow")
##
             coef exp(coef) se(coef)
##
                                          Z
## SEX
            2.777
                      16.070
                                1.165 2.38 0.0171
                       0.377
                                0.342 -2.85 0.0044
## tdc_sex -0.976
## Likelihood ratio test=12.61 on 2 df, p=0.002
## n= 1742258, number of events= 3033
```

Using $g(t) = \ln(t)$, the Wald p-value for the test of H_0 : $\beta_2 = 0$ is 0.0044, which is significant at $\alpha = 0.05$. Thus, there is evidence that SEX covariate has nonproportional hazards. Therefore, we should stratify on Sex.

Stratify on Sex

Fix Cox model with the top 9 covariates, stratified on SEX.

```
fit3 <- coxph(Surv(SRV_TIME_MON, delta) ~ strata(SEX) + factor(stage) + factor(RAC_RECY) +
             factor(BRST_SUB) + factor(MAR_STAT) + MALIGCOUNT + factor(PRIMSITE) +
             factor(ERSTATUS) + factor(PRSTATUS) + factor(INSREC_PUB) ,
             data = breast, ties = "breslow" )
summary(fit3)
## Call:
  coxph(formula = Surv(SRV_TIME_MON, delta) ~ strata(SEX) + factor(stage) +
       factor(RAC_RECY) + factor(BRST_SUB) + factor(MAR_STAT) +
##
       MALIGCOUNT + factor(PRIMSITE) + factor(ERSTATUS) + factor(PRSTATUS) +
##
##
       factor(INSREC_PUB), data = breast, ties = "breslow")
##
##
     n= 55333, number of events= 3033
##
##
                                   exp(coef)
                                               se(coef)
                                                              z Pr(>|z|)
                             coef
## factor(stage)1
                         1.857621
                                    6.408470
                                               0.709702 2.617 0.00886 **
## factor(stage)2
                         3.077618
                                   21.706631
                                               0.708156 4.346 1.39e-05 ***
## factor(stage)3
                         4.184598
                                   65.667066
                                               0.708103 5.910 3.43e-09 ***
## factor(stage)4
                         5.832492 341.208011
                                               0.707915 8.239 < 2e-16 ***
## factor(RAC_RECY)2
                         0.189521
                                    1.208670
                                               0.066937
                                                         2.831
                                                                 0.00464 **
## factor(RAC_RECY)3
                         0.067450
                                    1.069777
                                               0.197574 0.341
                                                                 0.73281
## factor(RAC_RECY)4
                                               0.077238 -4.039 5.38e-05 ***
                        -0.311933
                                    0.732030
## factor(BRST_SUB)2
                        -0.965198
                                    0.380908
                                               0.152381 -6.334 2.39e-10 ***
## factor(BRST_SUB)3
                         0.251216
                                    1.285588
                                               0.063901 3.931 8.45e-05 ***
## factor(BRST_SUB)4
                                    0.900793
                                               0.142392 -0.734 0.46310
                        -0.104480
## factor(MAR_STAT)2
                        -0.314578
                                    0.730097
                                               0.049510 -6.354 2.10e-10 ***
## factor(MAR_STAT)3
                        -0.099928
                                    0.904903
                                               0.146909 -0.680 0.49638
```

```
## factor(MAR_STAT)4
                         -0.008725
                                     0.991312
                                                 0.064331 -0.136
                                                                   0.89211
## factor(MAR_STAT)5
                                                 0.057990 8.904
                          0.516334
                                     1.675872
                                                                   < 2e-16 ***
## factor(MAR STAT)6
                         -0.125466
                                     0.882086
                                                 0.336123 -0.373
                                                                   0.70895
## MALIGCOUNT
                          0.288792
                                     1.334814
                                                 0.054880
                                                           5.262 1.42e-07 ***
## factor(PRIMSITE)1
                         -0.221547
                                     0.801278
                                                 0.256714 - 0.863
                                                                   0.38813
## factor(PRIMSITE)2
                         -0.218838
                                     0.803452
                                                 0.255639 - 0.856
                                                                   0.39197
## factor(PRIMSITE)3
                         -0.093926
                                     0.910350
                                                 0.260677 - 0.360
                                                                   0.71861
## factor(PRIMSITE)4
                         -0.337854
                                     0.713299
                                                 0.249553 - 1.354
                                                                   0.17579
## factor(PRIMSITE)5
                         -0.345290
                                     0.708015
                                                 0.258044 -1.338
                                                                   0.18086
## factor(PRIMSITE)6
                         -0.149278
                                     0.861330
                                                 0.337596 - 0.442
                                                                   0.65836
## factor(PRIMSITE)7
                         -0.158763
                                     0.853198
                                                 0.249813 -0.636
                                                                   0.52508
## factor(PRIMSITE)8
                         -0.019407
                                     0.980780
                                                 0.249559 - 0.078
                                                                   0.93801
## factor(ERSTATUS)1
                          1.033526
                                     2.810961
                                                 0.128666 8.033 9.54e-16 ***
## factor(PRSTATUS)1
                          0.679725
                                     1.973336
                                                 0.053092 12.803
                                                                   < 2e-16 ***
## factor(INSREC_PUB)1
                         -0.107544
                                     0.898037
                                                 0.128449 -0.837
                                                                   0.40245
## factor(INSREC_PUB)2
                         -0.357413
                                     0.699484
                                                 0.126123 -2.834
                                                                   0.00460 **
  factor(INSREC_PUB)3
                         -0.149441
                                     0.861189
                                                 0.131925 -1.133
                                                                   0.25731
##
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
##
                        exp(coef) exp(-coef) lower .95 upper .95
## factor(stage)1
                           6.4085
                                    0.156043
                                                 1.5946
                                                          25.7545
                                    0.046069
## factor(stage)2
                          21.7066
                                                 5.4176
                                                          86.9713
## factor(stage)3
                          65.6671
                                    0.015228
                                                16.3911
                                                         263.0789
## factor(stage)4
                         341.2080
                                    0.002931
                                                85.2002 1366.4628
## factor(RAC_RECY)2
                           1.2087
                                    0.827355
                                                 1.0601
                                                           1.3781
## factor(RAC_RECY)3
                           1.0698
                                    0.934774
                                                 0.7263
                                                           1.5757
                                                           0.8517
## factor(RAC_RECY)4
                           0.7320
                                    1.366064
                                                 0.6292
## factor(BRST_SUB)2
                           0.3809
                                    2.625306
                                                 0.2826
                                                           0.5135
## factor(BRST_SUB)3
                           1.2856
                                    0.777854
                                                 1.1343
                                                           1.4571
## factor(BRST_SUB)4
                           0.9008
                                    1.110133
                                                 0.6814
                                                           1.1908
## factor(MAR_STAT)2
                           0.7301
                                                 0.6626
                                                           0.8045
                                    1.369681
## factor(MAR_STAT)3
                           0.9049
                                                 0.6785
                                                           1.2068
                                    1.105091
## factor(MAR_STAT)4
                           0.9913
                                    1.008764
                                                 0.8739
                                                           1.1245
## factor(MAR_STAT)5
                           1.6759
                                    0.596704
                                                 1.4958
                                                           1.8776
## factor(MAR_STAT)6
                           0.8821
                                    1.133676
                                                 0.4565
                                                           1.7046
## MALIGCOUNT
                           1.3348
                                    0.749168
                                                 1.1987
                                                           1.4864
## factor(PRIMSITE)1
                           0.8013
                                    1.248006
                                                 0.4845
                                                           1.3253
## factor(PRIMSITE)2
                           0.8035
                                    1.244630
                                                 0.4868
                                                           1.3261
## factor(PRIMSITE)3
                           0.9104
                                    1.098478
                                                 0.5462
                                                           1.5174
## factor(PRIMSITE)4
                           0.7133
                                    1.401936
                                                 0.4374
                                                           1.1633
## factor(PRIMSITE)5
                           0.7080
                                    1.412399
                                                 0.4270
                                                           1.1741
## factor(PRIMSITE)6
                           0.8613
                                    1.160996
                                                 0.4444
                                                           1.6693
## factor(PRIMSITE)7
                           0.8532
                                    1.172060
                                                 0.5229
                                                           1.3922
## factor(PRIMSITE)8
                           0.9808
                                    1.019597
                                                 0.6014
                                                           1.5995
## factor(ERSTATUS)1
                           2.8110
                                    0.355750
                                                 2.1844
                                                           3.6172
## factor(PRSTATUS)1
                           1.9733
                                    0.506756
                                                 1.7783
                                                           2.1897
## factor(INSREC_PUB)1
                           0.8980
                                    1.113539
                                                 0.6982
                                                           1.1551
## factor(INSREC_PUB)2
                           0.6995
                                    1.429626
                                                 0.5463
                                                           0.8956
## factor(INSREC_PUB)3
                           0.8612
                                                 0.6650
                                    1.161185
                                                           1.1153
##
## Concordance= 0.883
                       (se = 0.006)
## Rsquare= 0.121
                     (max possible= 0.677 )
## Likelihood ratio test= 7163 on 29 df,
                                              p=<2e-16
```

```
## Wald test
                         = 7208 on 29 df,
                                              p = < 2e - 16
## Score (logrank) test = 14991 on 29 df,
                                               p=<2e-16
Use LRT to test whether the covariate effects are different between the 2 strata
breastSEX0 <- breast[breast$SEX == 0, ]</pre>
breastSEX1 <- breast[breast$SEX == 1, ]</pre>
fit0 <- coxph(Surv(SRV_TIME_MON, delta) ~ factor(stage) + factor(RAC_RECY) +
              factor(BRST_SUB) + factor(MAR_STAT) + MALIGCOUNT + factor(PRIMSITE) +
              factor(ERSTATUS) + factor(PRSTATUS) + factor(INSREC_PUB),
              data = breastSEXO, ties = "breslow")
## Warning in fitter(X, Y, strats, offset, init, control, weights = weights, :
## Ran out of iterations and did not converge
fit1 <- coxph(Surv(SRV TIME MON, delta) ~ factor(stage) + factor(RAC RECY) +
              factor(BRST_SUB) + factor(MAR_STAT) + MALIGCOUNT + factor(PRIMSITE) +
              factor(ERSTATUS) + factor(PRSTATUS) + factor(INSREC_PUB),
              data = breastSEX1, ties = "breslow")
X2 \leftarrow -2*(fit2\$loglik[2] - (fit0\$loglik[2] + fit1\$loglik[2])); X2
## [1] 174.0883
1 - pchisq(X2, 9) #9 degrees of freedom for each covariate
```

The p-value is <<0.0001, so the assumption of using a stratified model is not met; the covariate effects are not the same between the two strata. So a stratified model is not appropriate.

[1] 0