# CD138+ spectra and clinical risk

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Code to investigate associations between CD138+ spectra and clinical risk cytogenetics and disease stage

### 0. Setup

```
# Install and load required R packages
library(ggplot2)
library(MASS)
library(survivalAnalysis)
library(dplyr)
library(data.table)
```

Define data directory

```
data_dir = "/path/to/data" # exclude ending "/"
```

Load transcriptome spectra (PC1-PC)

### 1. Cytogenetics

Large somatic chromosomal DNA aberrations detected by cytogenetics are used to define prognostic risk groups in myeloma. Clinical risk categories defined by mSMART7 include: high risk, del(17p) and t(14;16); intermediate risk, amp(1q) and t(4;14); and standard risk, t(11;14).

1.1. del(17p): D\_TRI\_CF\_ABNORMALITYPR11 & D\_TRI\_CF\_17PABNORMALCE (abundance)

```
## [1] "Baseline samples with data: 436 (total), 52 (+), 384 (-)"
DAT$D_TRI_CF_ABNORMALITYPR11 = as.factor(DAT$D_TRI_CF_ABNORMALITYPR11)
# Logistic regression
MOD = glm(data = DAT, formula = D_TRI_CF_ABNORMALITYPR11 ~ .,family = "binomial")
summary(MOD)
##
## Call:
## glm(formula = D_TRI_CF_ABNORMALITYPR11 ~ ., family = "binomial",
##
       data = DAT)
##
## Deviance Residuals:
       Min
                 1Q
                      Median
                                    3Q
                                            Max
## -1.5460
                    -0.3682 -0.2166
                                         3.2274
           -0.5434
## Coefficients:
               Estimate Std. Error z value Pr(>|z|)
## (Intercept) -2.33931
                           0.19885 -11.765
                                              <2e-16 ***
## PC1_SD
               -0.14872
                           0.16816 -0.884
                                              0.3765
## PC2_SD
                                      0.378
               0.05974
                           0.15799
                                              0.7053
## PC3_SD
               0.37944
                           0.18161
                                      2.089
                                              0.0367 *
## PC4 SD
               -0.06789
                           0.16062
                                    -0.423
                                              0.6726
## PC5_SD
               -0.31276
                           0.16150
                                    -1.937
                                              0.0528 .
## PC6_SD
               -0.36238
                           0.16624
                                    -2.180
                                             0.0293 *
## PC7_SD
               0.31209
                           0.17164
                                     1.818
                                              0.0690 .
## PC8_SD
                0.09814
                           0.16563
                                      0.593
                                              0.5535
## PC9_SD
                0.02164
                           0.16421
                                      0.132
                                              0.8951
## PC10_SD
                0.08407
                           0.16914
                                      0.497
                                              0.6192
## PC11_SD
                0.28001
                           0.16681
                                      1.679
                                              0.0932
                                    -0.499
## PC12_SD
               -0.08424
                           0.16878
                                              0.6177
## PC13_SD
               -0.22907
                           0.16579
                                    -1.382
                                              0.1671
               -0.08211
## PC14_SD
                           0.16564
                                    -0.496
                                              0.6201
## PC15_SD
                0.17542
                           0.16406
                                     1.069
                                              0.2849
## PC16 SD
               -0.08960
                           0.15796
                                    -0.567
                                              0.5706
## PC17 SD
               -0.04821
                           0.16229
                                    -0.297
                                              0.7664
## PC18_SD
               0.29209
                           0.17394
                                      1.679
                                              0.0931
## PC19_SD
                0.17987
                           0.15566
                                      1.156
                                              0.2479
## PC20_SD
               -0.04769
                           0.17202 -0.277
                                              0.7816
## PC21_SD
               -0.07767
                           0.15997
                                    -0.486
                                              0.6273
## PC22_SD
               -0.18464
                           0.15887
                                    -1.162
                                              0.2452
## PC23_SD
               -0.04358
                           0.15095
                                    -0.289
                                              0.7728
## PC24_SD
                0.10254
                                     0.654
                           0.15678
                                              0.5131
## PC25_SD
               -0.06833
                           0.15963
                                    -0.428
                                              0.6686
## PC26_SD
                                    -0.264
               -0.04335
                           0.16423
                                              0.7918
## PC27_SD
               -0.11872
                           0.16213
                                    -0.732
                                              0.4640
## PC28_SD
                           0.16361
                                    -0.855
               -0.13992
                                              0.3924
## PC29 SD
               -0.28691
                           0.15851
                                    -1.810
                                              0.0703 .
               -0.17156
## PC30_SD
                           0.15881
                                    -1.080
                                              0.2800
## PC31 SD
                0.07626
                           0.16050
                                     0.475
                                              0.6347
## PC32_SD
               -0.15844
                           0.15977 - 0.992
                                              0.3213
## PC33_SD
                                      0.878
               0.13535
                           0.15422
                                              0.3801
```

0.2296

1.201

## PC34\_SD

0.21123

0.17583

```
## PC35 SD
               0.11168
                           0.17134
                                     0.652
                                            0.5145
## PC36 SD
               0.09137
                           0.15100
                                    0.605 0.5451
               0.12802
## PC37 SD
                           0.16963
                                    0.755 0.4504
## PC38_SD
                0.23489
                           0.15565
                                     1.509
                                            0.1313
## PC39 SD
               0.33491
                           0.15942
                                     2.101
                                            0.0357 *
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
       Null deviance: 318.68 on 435 degrees of freedom
## Residual deviance: 275.47 on 396 degrees of freedom
## AIC: 355.47
##
## Number of Fisher Scoring iterations: 6
# Count number of significant spectra
nsig = data.table(summary(MOD)$coeff[-1,"Pr(>|z|)"]) %>%
  subset(V1<0.05) %>% nrow()
print(pasteO(nsig," of 39 spectra significant (p<.05)"))</pre>
## [1] "3 of 39 spectra significant (p<.05)"
# Save model
mod.risk = list(d17p=MOD)
# Overall p-value
NLL = glm(data = DAT, formula = D_TRI_CF_ABNORMALITYPR11 ~ 1, family = "binomial")
mod.risk$d17p$p = pchisq(deviance(NLL)-deviance(MOD),
                         df.residual(NLL)-df.residual(MOD),
                         lower.tail=FALSE)
mod.risk$d17p$p
## [1] 0.2961634
Run model with significant spectra only
md2 = glm(data = DAT,
          formula = D_TRI_CF_ABNORMALITYPR11 ~ PC3_SD + PC6_SD + PC39_SD,
          family = "binomial")
summary(md2)
##
## glm(formula = D_TRI_CF_ABNORMALITYPR11 ~ PC3_SD + PC6_SD + PC39_SD,
       family = "binomial", data = DAT)
##
## Deviance Residuals:
##
      Min
                 1Q
                     Median
                                   3Q
                                           Max
## -0.9347 -0.5392 -0.4560 -0.3828
                                        2.5694
##
```

```
## Coefficients:
##
             Estimate Std. Error z value Pr(>|z|)
## (Intercept) -2.0722 0.1575 -13.153
## PC3_SD
               0.2493
                           0.1510 1.651
                                            0.0988 .
## PC6 SD
              -0.2745
                           0.1506 -1.823
                                           0.0683 .
## PC39 SD
                0.3362
                           0.1491 2.254 0.0242 *
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
      Null deviance: 318.68 on 435 degrees of freedom
## Residual deviance: 308.03 on 432 degrees of freedom
## AIC: 316.03
##
## Number of Fisher Scoring iterations: 5
pchisq(deviance(NLL)-deviance(md2),
      df.residual(NLL)-df.residual(md2),lower.tail=FALSE)
## [1] 0.01378549
1.2. t(14;16): D TRI CF ABNORMALITYPR8 & D TRI CF T1416ABNORMAL (abundance)
DAT = spectra_clinical %>%
 dplyr::select("D_TRI_CF_ABNORMALITYPR8",starts_with("PC")) %>%
 dplyr::filter(D_TRI_CF_ABNORMALITYPR8!="Not Done" &
               D_TRI_CF_ABNORMALITYPR8!="")
print(paste("Baseline samples with data:",nrow(DAT),"(total),",
           nrow(DAT[DAT$D_TRI_CF_ABNORMALITYPR8=="Yes",]),"(+),",
           nrow(DAT[DAT$D_TRI_CF_ABNORMALITYPR8=="No",]),"(-)"))
## [1] "Baseline samples with data: 554 (total), 53 (+), 501 (-)"
DAT$D_TRI_CF_ABNORMALITYPR8 = as.factor(DAT$D_TRI_CF_ABNORMALITYPR8)
# Logistic regression
MOD <- glm(data = DAT, formula = D_TRI_CF_ABNORMALITYPR8 ~ ., family = "binomial")</pre>
summary(MOD)
##
## glm(formula = D_TRI_CF_ABNORMALITYPR8 ~ ., family = "binomial",
##
      data = DAT)
##
## Deviance Residuals:
       Min
                  1Q
                        Median
                                      3Q
                                               Max
## -1.90797 -0.35808 -0.18434 -0.08671
                                           3.11720
##
## Coefficients:
##
              Estimate Std. Error z value Pr(>|z|)
```

```
## (Intercept) -3.68550
                           0.34193 -10.779 < 2e-16 ***
## PC1 SD
               -0.30672
                           0.21028 -1.459 0.144664
               -0.55403
## PC2 SD
                           0.20849
                                    -2.657 0.007877 **
## PC3_SD
                0.32627
                           0.20724
                                     1.574 0.115408
## PC4 SD
               -0.24505
                           0.18243
                                    -1.343 0.179186
## PC5 SD
                           0.17898
                                    -2.625 0.008669 **
               -0.46980
## PC6 SD
               0.36046
                           0.20485
                                     1.760 0.078472 .
## PC7 SD
               -0.01306
                           0.19061
                                    -0.069 0.945358
## PC8_SD
                0.04631
                           0.19583
                                     0.236 0.813053
## PC9_SD
               -0.24567
                           0.20440
                                    -1.202 0.229401
## PC10_SD
               0.50221
                           0.20863
                                     2.407 0.016076 *
## PC11_SD
               -0.08352
                           0.17871
                                    -0.467 0.640253
## PC12_SD
                0.60753
                           0.19491
                                     3.117 0.001827 **
## PC13_SD
               -0.41082
                           0.17773
                                    -2.311 0.020806 *
## PC14_SD
               -0.31970
                           0.18560
                                    -1.723 0.084969 .
## PC15_SD
                0.02333
                           0.19011
                                     0.123 0.902330
## PC16_SD
                0.80028
                           0.22187
                                     3.607 0.000310 ***
## PC17 SD
                0.04611
                           0.19884
                                     0.232 0.816632
## PC18_SD
                                    -1.272 0.203369
               -0.22539
                           0.17719
## PC19 SD
               -0.89008
                           0.22412
                                    -3.971 7.14e-05 ***
## PC20_SD
                0.25206
                           0.20866
                                     1.208 0.227043
## PC21 SD
               -0.22007
                           0.17950
                                    -1.226 0.220210
## PC22_SD
                0.02942
                           0.20551
                                     0.143 0.886173
## PC23 SD
               -0.10984
                           0.19196
                                    -0.572 0.567199
## PC24 SD
               -0.16723
                           0.18541
                                    -0.902 0.367077
## PC25 SD
               -0.72354
                           0.20843
                                    -3.471 0.000518 ***
## PC26_SD
                0.30567
                           0.18513
                                     1.651 0.098721
## PC27_SD
               -0.12525
                           0.19136
                                    -0.655 0.512773
## PC28_SD
                0.56144
                           0.17975
                                     3.123 0.001787 **
                                     0.422 0.672958
## PC29_SD
                           0.19796
                0.08356
## PC30_SD
               -0.17086
                           0.17983
                                    -0.950 0.342045
## PC31_SD
               -0.27408
                           0.18853
                                    -1.454 0.146007
## PC32_SD
                0.17124
                           0.17858
                                     0.959 0.337619
## PC33_SD
                0.19644
                           0.18058
                                     1.088 0.276670
## PC34 SD
               -0.12919
                           0.19111
                                    -0.676 0.499049
## PC35_SD
               -0.37854
                           0.19025
                                    -1.990 0.046623 *
## PC36 SD
               -0.07250
                           0.16971
                                    -0.427 0.669217
## PC37_SD
               0.40193
                           0.19317
                                     2.081 0.037465 *
## PC38 SD
               -0.01291
                           0.17264 -0.075 0.940380
## PC39_SD
               -0.40072
                           0.19526 -2.052 0.040147 *
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 349.53 on 553 degrees of freedom
## Residual deviance: 227.27 on 514 degrees of freedom
## AIC: 307.27
## Number of Fisher Scoring iterations: 7
# Count sig spectra in model
nsig = data.table(summary(MOD)$coeff[-1,"Pr(>|z|)"]) %>%
 subset(V1<0.05) %>% nrow()
```

```
print(pasteO(nsig," of 39 spectra significant (p<.05)"))</pre>
## [1] "12 of 39 spectra significant (p<.05)"
# Save model
mod.risk = c(mod.risk,list(t1416=MOD))
# Overall p-value
NLL = glm(data = DAT, formula = D_TRI_CF_ABNORMALITYPR8 ~ 1, family = "binomial")
mod.risk$t1416$p = pchisq(deviance(NLL)-deviance(MOD),
                       df.residual(NLL)-df.residual(MOD),
                       lower.tail=FALSE)
mod.risk$t1416$p
## [1] 1.608411e-10
1.3. amp(1q): D_TRI_CF_ABNORMALITYPR13 & D_TRI_CF_1PAMPLIFICATI2 (abundance)
DAT = spectra_clinical %>%
  dplyr::select("D_TRI_CF_ABNORMALITYPR13",starts_with("PC")) %>%
  dplyr::filter(D TRI CF ABNORMALITYPR13!="Not Done" &
               D_TRI_CF_ABNORMALITYPR13!="")
print(paste("Baseline samples with data:",nrow(DAT),"(total),",
           nrow(DAT[DAT$D_TRI_CF_ABNORMALITYPR13=="Yes",]),"(+),",
           nrow(DAT[DAT$D_TRI_CF_ABNORMALITYPR13=="No",]),"(-)"))
## [1] "Baseline samples with data: 525 (total), 202 (+), 323 (-)"
DAT$D TRI CF ABNORMALITYPR13 = as.factor(DAT$D TRI CF ABNORMALITYPR13)
# Logistic regression
MOD <- glm(data = DAT,formula = D_TRI_CF_ABNORMALITYPR13 ~ .,family = "binomial")
summary(MOD)
##
## Call:
## glm(formula = D_TRI_CF_ABNORMALITYPR13 ~ ., family = "binomial",
      data = DAT)
##
##
## Deviance Residuals:
      Min
##
               1Q
                   Median
                                 3Q
                                        Max
## -2.4620 -0.6314 -0.3313 0.6043
                                     3.0382
##
## Coefficients:
##
              Estimate Std. Error z value Pr(>|z|)
## (Intercept) -0.690992  0.126291 -5.471 4.46e-08 ***
## PC1_SD -0.007713 0.122192 -0.063 0.949672
## PC2 SD
             ## PC3 SD
```

```
## PC4 SD
                0.109691
                           0.125437
                                      0.874 0.381861
## PC5 SD
               -0.802967
                           0.135274 -5.936 2.92e-09 ***
## PC6 SD
                0.482744
                           0.127773
                                      3.778 0.000158 ***
## PC7_SD
                           0.119897
                0.117559
                                      0.980 0.326841
## PC8_SD
               -0.230686
                           0.122601
                                     -1.882 0.059890
## PC9 SD
                           0.123028
                                     -1.986 0.047004 *
               -0.244367
## PC10 SD
               -0.408167
                           0.125689 -3.247 0.001164 **
## PC11_SD
               -0.284010
                           0.123191
                                     -2.305 0.021141 *
## PC12 SD
                0.148954
                           0.121227
                                      1.229 0.219178
## PC13_SD
                0.144725
                           0.116883
                                      1.238 0.215641
## PC14_SD
               -0.446631
                           0.124381
                                     -3.591 0.000330 ***
## PC15_SD
                0.178899
                           0.118319
                                      1.512 0.130534
## PC16_SD
                0.127909
                           0.120903
                                      1.058 0.290083
## PC17_SD
                                     -0.642 0.520947
               -0.076199
                           0.118711
## PC18_SD
                                     -1.570 0.116334
               -0.193285
                           0.123084
## PC19_SD
               -0.202939
                           0.125611
                                     -1.616 0.106177
## PC20_SD
                           0.125938
                                      0.383 0.701557
                0.048262
## PC21 SD
               -0.287935
                           0.117308
                                     -2.455 0.014107 *
## PC22_SD
                           0.124526
                                     -1.312 0.189610
               -0.163345
## PC23 SD
               -0.308160
                           0.119422
                                     -2.580 0.009868 **
## PC24_SD
                0.393140
                           0.128223
                                      3.066 0.002169 **
## PC25_SD
                0.288189
                           0.127105
                                      2.267 0.023370 *
## PC26 SD
                           0.123295
                                     -0.853 0.393528
               -0.105200
## PC27 SD
               -0.453369
                           0.124358 -3.646 0.000267 ***
## PC28 SD
                0.015377
                           0.116773
                                      0.132 0.895233
## PC29 SD
               -0.065995
                           0.128013
                                     -0.516 0.606179
## PC30_SD
               -0.065103
                           0.113445
                                     -0.574 0.566053
## PC31_SD
               -0.010491
                           0.115031
                                     -0.091 0.927330
## PC32_SD
               -0.183180
                           0.128897
                                     -1.421 0.155280
               -0.082332
## PC33_SD
                                     -0.661 0.508701
                           0.124583
## PC34_SD
                0.221452
                           0.124709
                                      1.776 0.075773
## PC35_SD
                0.291681
                           0.122200
                                      2.387 0.016991 *
## PC36_SD
                0.065219
                           0.120091
                                      0.543 0.587076
## PC37_SD
               -0.252369
                           0.127726
                                     -1.976 0.048170
## PC38 SD
                0.140748
                           0.117811
                                      1.195 0.232208
## PC39 SD
                0.038282
                           0.125684
                                      0.305 0.760681
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
       Null deviance: 699.66 on 524 degrees of freedom
## Residual deviance: 443.73 on 485 degrees of freedom
## AIC: 523.73
## Number of Fisher Scoring iterations: 5
# Count sig spectra in model
nsig = data.table(summary(MOD)$coeff[-1,"Pr(>|z|)"]) %>%
  subset(V1<0.05) %>% nrow()
print(pasteO(nsig," of 39 spectra significant (p<.05)"))</pre>
```

## [1] "15 of 39 spectra significant (p<.05)"

```
# Save model
mod.risk = c(mod.risk,list(a1q=MOD))
# Overall p-value
NLL = glm(data = DAT, formula = D TRI CF ABNORMALITYPR13 ~ 1, family = "binomial")
mod.risk$a1q$p = pchisq(deviance(NLL)-deviance(MOD),
                     df.residual(NLL)-df.residual(MOD),
                     lower.tail=FALSE)
mod.risk$a1q$p
## [1] 1.071633e-33
1.4. t(4;14): D_TRI_CF_ABNORMALITYPR3 & D_TRI_CF_T414ABNORMALC (abundance)
DAT = spectra_clinical %>%
 dplyr::select("D_TRI_CF_ABNORMALITYPR3",starts_with("PC")) %>%
 dplyr::filter(D_TRI_CF_ABNORMALITYPR3!="Not Done" &
              D_TRI_CF_ABNORMALITYPR3!="")
print(paste("Baseline samples with data:",nrow(DAT),"(total),",
          nrow(DAT[DAT$D_TRI_CF_ABNORMALITYPR3=="Yes",]),"(+),",
          nrow(DAT[DAT$D_TRI_CF_ABNORMALITYPR3=="No",]),"(-)"))
## [1] "Baseline samples with data: 594 (total), 108 (+), 486 (-)"
DAT$D_TRI_CF_ABNORMALITYPR3 = as.factor(DAT$D_TRI_CF_ABNORMALITYPR3)
# Logistic regression
MOD <- glm(data = DAT, formula = D_TRI_CF_ABNORMALITYPR3 ~ ., family = "binomial")
summary(MOD)
##
## Call:
## glm(formula = D TRI CF ABNORMALITYPR3 ~ ., family = "binomial",
      data = DAT)
##
## Deviance Residuals:
      Min 1Q Median
                            3Q
                                     Max
## -2.4577 -0.4685 -0.3038 -0.1423
                                  2.9473
##
## Coefficients:
             Estimate Std. Error z value Pr(>|z|)
## (Intercept) -2.286504   0.184652 -12.383   < 2e-16 ***
## PC1_SD
             ## PC2_SD
            ## PC3 SD
            0.246057 0.144001 1.709 0.087504 .
## PC4_SD
            0.148729 0.137047 1.085 0.277816
## PC5 SD
## PC6_SD
            0.235319  0.138274  1.702  0.088787 .
## PC7 SD
            ## PC8 SD
```

```
## PC9 SD
              -0.789087
                          0.160637 -4.912
                                             9e-07 ***
## PC10 SD
              -0.590438
                          0.158077 -3.735 0.000188 ***
## PC11 SD
              0.066273
                          0.142892
                                   0.464 0.642794
## PC12_SD
                          0.142266
                                   0.057 0.954846
               0.008056
## PC13 SD
              -0.410435
                          0.142481 -2.881 0.003969 **
## PC14 SD
              0.040884
                          0.146781
                                   0.279 0.780602
## PC15 SD
               0.286835
                          0.141114
                                   2.033 0.042088 *
## PC16 SD
              -0.067236
                          0.145373 -0.463 0.643717
## PC17_SD
              -0.110613
                          0.149501 -0.740 0.459371
## PC18_SD
              0.111623
                          0.156538
                                   0.713 0.475802
## PC19_SD
              -0.261750
                          0.152306 -1.719 0.085690
## PC20_SD
               0.292234
                          0.142093
                                   2.057 0.039721 *
## PC21_SD
              -0.220324
                          0.153225 -1.438 0.150459
## PC22_SD
              -0.025535
                          0.152651 -0.167 0.867150
## PC23_SD
                          0.144174
                                   1.402 0.160935
              0.202122
## PC24_SD
              -0.010786
                          0.143987 -0.075 0.940289
## PC25_SD
              -0.085848
                          0.151531 -0.567 0.571027
## PC26 SD
              0.014047
                          0.141659 0.099 0.921013
## PC27_SD
               0.321734
                          0.146496 2.196 0.028078 *
## PC28 SD
               0.056878
                          0.145160 0.392 0.695181
## PC29_SD
               0.126763
                          ## PC30 SD
              -0.237363
                          0.144678 -1.641 0.100875
## PC31_SD
                                   1.690 0.090979 .
               0.244815
                          0.144839
## PC32 SD
               0.224236
                          0.151767
                                    1.478 0.139541
## PC33 SD
              0.224455
                          0.143789 1.561 0.118524
## PC34 SD
              0.040400
                          0.146833
                                   0.275 0.783205
## PC35_SD
              -0.228223
                          0.146554 -1.557 0.119408
## PC36_SD
              -0.195065
                          0.143619 -1.358 0.174398
## PC37_SD
               0.210835
                          0.146764
                                   1.437 0.150844
## PC38_SD
              -0.383261
                          0.139350 -2.750 0.005953 **
## PC39_SD
              -0.289167
                          0.154489 -1.872 0.061239 .
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
      Null deviance: 563.28 on 593 degrees of freedom
## Residual deviance: 355.97 on 554 degrees of freedom
## AIC: 435.97
##
## Number of Fisher Scoring iterations: 6
# Count sig spectra in model
nsig = data.table(summary(MOD)$coeff[-1,"Pr(>|z|)"]) %>%
 subset(V1<0.05) %>% nrow()
print(paste0(nsig," of 39 spectra significant (p<.05)"))</pre>
## [1] "10 of 39 spectra significant (p<.05)"
# Save model
mod.risk = c(mod.risk,list(t414=MOD))
# Overall p-value
```

```
NLL = glm(data = DAT,formula = D_TRI_CF_ABNORMALITYPR3 ~ 1,family = "binomial")
mod.risk$t414$p = pchisq(deviance(NLL)-deviance(MOD),
                      df.residual(NLL)-df.residual(MOD),
                      lower.tail=FALSE)
mod.risk$t414$p
## [1] 8.207055e-25
1.5. t(11;14): D_TRI_CF_ABNORMALITYPR6 & D_TRI_CF_T1114ABNORMAL (abundance)
DAT = spectra_clinical %>%
 dplyr::select("D_TRI_CF_ABNORMALITYPR6",starts_with("PC")) %>%
 dplyr::filter(D_TRI_CF_ABNORMALITYPR6!="Not Done" &
              D_TRI_CF_ABNORMALITYPR6!="")
print(paste("Baseline samples with data:",nrow(DAT),"(total),",
          nrow(DAT[DAT$D_TRI_CF_ABNORMALITYPR6=="Yes",]),"(+),",
           nrow(DAT[DAT$D TRI CF ABNORMALITYPR6=="No",]),"(-)"))
## [1] "Baseline samples with data: 594 (total), 153 (+), 441 (-)"
DAT$D_TRI_CF_ABNORMALITYPR6 = as.factor(DAT$D_TRI_CF_ABNORMALITYPR6)
# Logistic regression
MOD <- glm(data = DAT, formula = D_TRI_CF_ABNORMALITYPR6 ~ ., family = "binomial")
summary(MOD)
##
## glm(formula = D_TRI_CF_ABNORMALITYPR6 ~ ., family = "binomial",
##
      data = DAT)
##
## Deviance Residuals:
      Min
               1Q
                   Median
                               3Q
                                       Max
## -2.6025 -0.5794 -0.3609 0.4043
                                    2.9983
##
## Coefficients:
              Estimate Std. Error z value Pr(>|z|)
## (Intercept) -1.629511 0.144002 -11.316 < 2e-16 ***
## PC1 SD
                                6.118 9.5e-10 ***
             0.757514 0.123825
                        0.121617 -2.752 0.00592 **
## PC2_SD
             -0.334711
## PC3 SD
             ## PC4_SD
             0.173529 0.129039 1.345 0.17869
## PC5 SD
             -0.124781
                        0.126141 -0.989 0.32256
## PC6_SD
             -0.335438
                      0.136356 -2.460 0.01389 *
## PC7_SD
             ## PC8_SD
             0.003280 0.125356 0.026 0.97912
## PC9 SD
             0.219560 0.139354 1.576 0.11513
             ## PC10 SD
## PC11 SD
             0.243246 0.134069 1.814 0.06963 .
```

```
## PC12 SD
              -0.075078
                         0.122723 -0.612 0.54069
## PC13 SD
              -0.131966 0.127691 -1.033 0.30138
## PC14 SD
              -0.152805
                         0.128630 -1.188 0.23486
## PC15_SD
                         0.130661 -0.073 0.94193
              -0.009517
## PC16 SD
              0.211483 0.130026
                                  1.626 0.10385
## PC17 SD
              -0.050028 0.129911 -0.385 0.70017
## PC18 SD
              0.360956
                         0.125101
                                  2.885 0.00391 **
## PC19 SD
              -0.104462
                         0.125673 -0.831 0.40585
## PC20 SD
              -0.108394
                         0.129358 -0.838 0.40206
## PC21_SD
             -0.267156
                         0.128115 -2.085 0.03704 *
## PC22_SD
              0.067931
                         0.122739
                                  0.553 0.57995
## PC23_SD
              -0.149114
                         0.127474 -1.170 0.24210
                                  0.999 0.31766
## PC24_SD
              0.126206
                         0.126297
                         0.121039 -1.025 0.30529
## PC25_SD
              -0.124085
## PC26_SD
                                  0.240 0.81004
              0.030517
                         0.126957
## PC27_SD
              -0.142821
                         0.121754 -1.173 0.24079
## PC28_SD
              0.276628
                         0.127636 2.167 0.03021 *
## PC29 SD
              0.142448
                         0.128801
                                  1.106 0.26875
## PC30_SD
              0.074267
                         0.120390
                                  0.617 0.53731
## PC31 SD
              -0.149381
                         0.121115 -1.233 0.21743
## PC32_SD
              ## PC33 SD
              -0.058201
                         0.125441 -0.464 0.64267
## PC34_SD
                         0.124263 -0.641 0.52144
              -0.079668
## PC35 SD
                                   0.941 0.34674
              0.116446
                         0.123757
## PC36 SD
              0.277852 0.125477
                                   2.214 0.02680 *
## PC37 SD
              -0.139475
                         0.135170 -1.032 0.30214
## PC38_SD
               0.119632
                                  0.960 0.33730
                         0.124680
## PC39_SD
               0.073817
                         0.119798
                                  0.616 0.53777
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
##
      Null deviance: 677.76 on 593 degrees of freedom
## Residual deviance: 453.15 on 554 degrees of freedom
## AIC: 533.15
##
## Number of Fisher Scoring iterations: 5
# Count sig spectra in model
nsig = data.table(summary(MOD)$coeff[-1,"Pr(>|z|)"]) %>%
  subset(V1<0.05) %>% nrow()
print(pasteO(nsig," of 39 spectra significant (p<.05)"))</pre>
## [1] "10 of 39 spectra significant (p<.05)"
# Save model
mod.risk = c(mod.risk,list(t1114=MOD))
# Overall p-value
NLL = glm(data = DAT,formula = D_TRI_CF_ABNORMALITYPR6 ~ 1,family = "binomial")
mod.risk$t1114$p = pchisq(deviance(NLL)-deviance(MOD),
```

## [1] 6.201022e-28

## PC27\_SD -0.1344839

#### 2. International staging system (ISS)

#### 2.1. RUN LOGISTIC REGRESSION

```
DAT = spectra_clinical %>%
 dplyr::select("D_PT_iss",starts_with("PC"))
# Order dependent variable
DAT$ISS = factor(DAT$D_PT_iss, levels = c(1,2,3), ordered = TRUE)
# Ordinal logistic regression model
MOD = polr(data = DAT[,-c("D_PT_iss")], formula = ISS ~ ., Hess = T)
summary(MOD)
## Call:
## polr(formula = ISS ~ ., data = DAT[, -c("D_PT_iss")], Hess = T)
## Coefficients:
##
               Value Std. Error t value
## PC1_SD -0.2749662
                        0.07106 -3.869573
## PC2_SD -0.1333559
                        0.07127 -1.871037
## PC3_SD
          0.3549374
                        0.07179 4.944326
## PC4_SD
          0.3126878
                        0.07203 4.341113
## PC5_SD -0.1743363
                        0.07120 -2.448579
## PC6_SD -0.1477172
                        0.07224 -2.044932
## PC7 SD
          0.0166838
                        0.07019 0.237700
                        0.07054 2.272335
## PC8_SD
           0.1602979
## PC9 SD -0.1729565
                        0.07093 -2.438448
## PC10 SD -0.0107528
                        0.07059 -0.152337
## PC11_SD 0.0517981
                        0.07172 0.722239
## PC12_SD -0.1726008
                        0.07115 - 2.425862
## PC13_SD -0.1743827
                        0.06957 -2.506712
## PC14_SD -0.0209329
                        0.07013 -0.298501
## PC15_SD -0.1563124
                        0.07029 -2.223775
## PC16_SD 0.0820734
                        0.07022 1.168744
## PC17_SD 0.2543831
                        0.07047 3.609659
## PC18_SD -0.0401684
                        0.07203 -0.557635
## PC19_SD 0.2633690
                        0.07220 3.647976
## PC20_SD -0.0066336
                        0.07045 -0.094160
## PC21_SD 0.0927783
                        0.07108 1.305260
## PC22 SD -0.0071847
                        0.07087 -0.101386
## PC23_SD -0.0005706
                        0.07006 -0.008144
## PC24 SD 0.0082876
                        0.07171 0.115568
## PC25_SD 0.1693466
                        0.07045 2.403712
## PC26_SD 0.2382966
                        0.07029 3.390307
```

0.07060 -1.904751

```
## PC28 SD 0.1398285
                        0.07131 1.960725
## PC29_SD 0.1387677 0.07090 1.957315
## PC30 SD 0.0289022
                        0.07145 0.404529
## PC31_SD -0.0464069
                        0.07063 -0.657088
## PC32_SD -0.0366594
                        0.07173 -0.511082
## PC33 SD 0.0126673 0.07093 0.178598
## PC34 SD 0.0243803
                        0.07073 0.344691
                        0.07013 1.921014
## PC35 SD 0.1347160
                        0.07093 -0.513211
## PC36_SD -0.0364027
## PC37_SD -0.0230455
                        0.07037 -0.327481
## PC38_SD 0.1395633
                        0.07135 1.956152
## PC39_SD 0.0207842
                        0.07123 0.291788
## Intercepts:
      Value
              Std. Error t value
## 1|2 -0.7310 0.0834
                         -8.7688
## 2|3 1.0550 0.0886
                         11.9131
##
## Residual Deviance: 1479.279
## AIC: 1561.279
## (21 observations deleted due to missingness)
# Count sig spectra in model
sig = pnorm(abs(coef(summary(MOD))[1:39,"t value"]),lower.tail = FALSE)*2
nsig = data.table(sig) %>% subset(sig<0.05) %>% nrow()
print(pasteO(nsig," of 39 spectra significant (p<.05)"))</pre>
## [1] "15 of 39 spectra significant (p<.05)"
# Save model
mod.risk = c(mod.risk,list(iss=MOD))
# Overall p-value
NLL = polr(data = DAT[,-c("D_PT_iss")], formula = ISS ~ 1, Hess = T)
mod.risk$iss$p = pchisq(deviance(NLL)-deviance(MOD),
                        df.residual(NLL)-df.residual(MOD),
                        lower.tail=FALSE)
mod.risk$iss$p
## [1] 2.999079e-15
Save model results
save(mod.risk,file = "rdata/mod.clinical-risk.rdata")
```