Untitled

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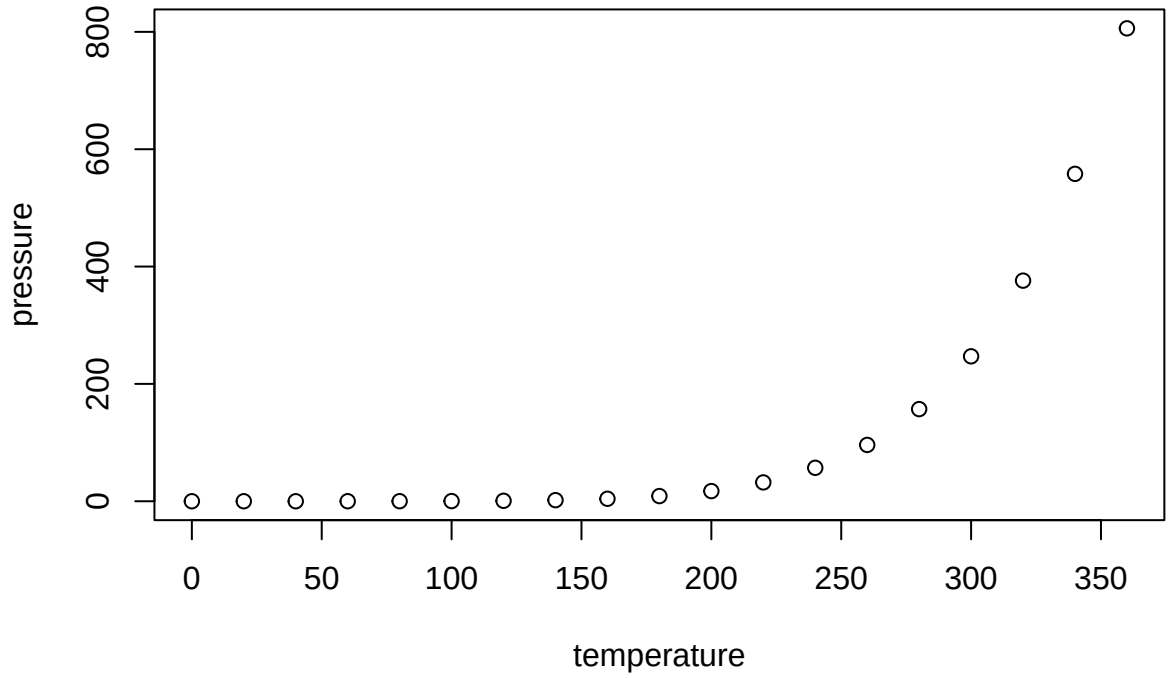
R Markdown

This is an R Markdown document. Markdown is a simple formatting syntax for authoring HTML, PDF, and MS Word documents. For more details on using R Markdown see http://rmarkdown.rstudio.com.

When you click the **Knit** button a document will be generated that includes both content as well as the output of any embedded R code chunks within the document. You can embed an R code chunk like this:

```
summary(cars)
```

```
##      speed           dist       
##  Min.   : 4.0   Min.   :  2.00  
##  1st Qu.:12.0   1st Qu.: 26.00  
##  Median :15.0   Median : 36.00  
##  Mean   :15.4   Mean   : 42.98  
##  3rd Qu.:19.0   3rd Qu.: 56.00  
##  Max.   :25.0   Max.   :120.00
```



Select top predictive proteins

```
top_n <- 20

load('../data/biomarker-clean.RData')

library(dplyr)

dat <- biomarker_clean
dat$group <- factor(dat$group)

proteins <- setdiff(names(dat), c('group','ados'))

tt_res <- sapply(proteins, function(p){
    x <- dat[[p]]
    grp <- dat$group
    ok <- !is.na(x) & !is.na(grp)
    if(sum(ok) < 3) return(NA)
    t <- try(t.test(x[ok] ~ grp[ok]), silent=TRUE)
    if(inherits(t,'try-error')) return(NA)
    t$p.value
})
tt_df <- tibble::tibble(protein = proteins,
                                            pvalue = as.numeric(tt_res)) %>%
```

```
    arrange(pvalue) %>%
    slice_head(n = top_n)

if(!requireNamespace('randomForest', quietly = TRUE)) {
    stop("Package 'randomForest' required. Please install it before running this chunk.")
}
rf_dat <- dat %>% dplyr::select(dplyr::all_of(proteins))
rf_resp <- dat$group
rf_fit <- randomForest::randomForest(x = rf_dat, y = rf_resp, ntree = 1000, importance = TRUE)
rf_imp_mat <- randomForest::importance(rf_fit, type = 2)
rf_imp_val <- if(is.matrix(rf_imp_mat)) rf_imp_mat[,1] else rf_imp_mat
rf_df <- tibble::tibble(protein = names(rf_imp_val), importance = as.numeric(rf_imp_val)) %>%
    arrange(desc(importance)) %>%
    slice_head(n = top_n)

if(!requireNamespace('glmnet', quietly = TRUE)) {
    stop("Package 'glmnet' required. Please install it before running this chunk.")
}
X <- as.matrix(rf_dat)
y <- as.numeric(dat$group) - 1
cv <- glmnet::cv.glmnet(X, y, family = 'binomial', alpha = 1, nfolds = 5)
# Get coefficients using the generic coef() function
coef_min <- as.matrix(coef(cv, s = 'lambda.min'))
# Remove intercept (first row) and get coefficients
coefs <- coef_min[-1,1]
lasso_df <- tibble::tibble(protein = proteins, coef = as.numeric(coefs)) %>%
    mutate(abscoef = abs(coef)) %>%
    arrange(desc(abscoef)) %>%
    slice_head(n = top_n)

tt_top <- tt_df$protein
rf_top <- rf_df$protein
lasso_top <- lasso_df$protein

cat(glue::glue("Selected top {top_n} proteins by each method:\n"))
```

```
## Selected top 20 proteins by each method:
```

```
cat("- t-test (by p-value):\n")
```

```
## - t-test (by p-value):
```

```
print(tt_top)
```

```
##  [1] "DERM"            "RELT"            "FSTL1"           "C1QR1"
##  [5] "Calcineurin"     "CXCL16, soluble" "IgD"             "MRC2"
##  [9] "PTN"             "Cadherin-5"      "MAPK2"           "TGF-b R III"
## [13] "DAF"             "MIA"             "Notch 1"         "gp130, soluble"
## [17] "MMP-2"           "ALCAM"           "ROR1"            "MATN2"
```

```
cat('\n- Random Forest (importance):\n')
```

```
##
## - Random Forest (importance):
```

```
print(rf_top)
```

```
##  [1] "DERM"        "MAPK14"      "IgD"         "TSP4"        "RELT"
##  [6] "FSTL1"       "Notch 1"     "TGF-b R III" "ERBB1"       "eIF-4H"
## [11] "PTN"         "MAPK2"       "M2-PK"       "MMP-2"       "TrkC"
## [16] "CSK"         "SRCN1"       "Cadherin-5"  "SOST"        "IGFBP-1"
```

```
cat('\n- LASSO (coef magnitude):\n')
```

```
##
## - LASSO (coef magnitude):
```

```
print(lasso_top)
```

```
##  [1] "IgD"                        "DERM"
##  [3] "14-3-3 protein zeta/delta" "Epo"
##  [5] "MAPK2"                      "ENTP5"
##  [7] "Protein S"                  "FSTL1"
##  [9] "IL-17 RC"                   "SRCN1"
## [11] "CD59"                       "IL-6 sRa"
## [13] "ITI heavy chain H4"         "TWEAKR"
## [15] "PAI-1"                      "PYY"
## [17] "CSRP3"                      "FAM3D"
## [19] "hnRNP K"                    "HGFA"
```

```
unique_proteins <- unique(c(tt_top, rf_top, lasso_top))
cat('\nTotal unique proteins across methods: ', length(unique_proteins), '\n')
```

```
##
## Total unique proteins across methods:  45
```

```
selected_lists <- list(tt = tt_df, rf = rf_df, lasso = lasso_df)

selected_lists
```

```
## $tt
## # A tibble: 20 x 2
##    protein               pvalue
##    <chr>                  <dbl>
##  1 DERM            0.00000000827
##  2 RELT            0.0000000782
##  3 FSTL1           0.000000466
##  4 C1QR1           0.000000479
##  5 Calcineurin     0.000000537
```

```
##  6 CXCL16, soluble 0.000000875
##  7 IgD             0.000000933
##  8 MRC2            0.00000103
##  9 PTN             0.00000135
## 10 Cadherin-5      0.00000175
## 11 MAPK2           0.00000204
## 12 TGF-b R III     0.00000330
## 13 DAF             0.00000397
## 14 MIA             0.00000483
## 15 Notch 1         0.00000500
## 16 gp130, soluble  0.00000530
## 17 MMP-2           0.00000552
## 18 ALCAM           0.00000664
## 19 ROR1            0.00000786
## 20 MATN2           0.00000799
##
## $rf
## # A tibble: 20 x 2
##    protein     importance
##    <chr>            <dbl>
##  1 DERM             0.991
##  2 MAPK14           0.656
##  3 IgD              0.613
##  4 TSP4             0.519
##  5 RELT             0.478
##  6 FSTL1            0.461
##  7 Notch 1          0.427
##  8 TGF-b R III      0.411
##  9 ERBB1            0.375
## 10 eIF-4H           0.358
## 11 PTN              0.355
## 12 MAPK2            0.343
## 13 M2-PK            0.336
## 14 MMP-2            0.322
## 15 TrkC             0.312
## 16 CSK              0.285
## 17 SRCN1            0.281
## 18 Cadherin-5       0.275
## 19 SOST             0.273
## 20 IGFBP-1          0.267
##
## $lasso
## # A tibble: 20 x 3
##    protein                     coef abscoef
##    <chr>                      <dbl>   <dbl>
##  1 IgD                        0.563   0.563
##  2 DERM                       0.525   0.525
##  3 14-3-3 protein zeta/delta  0.319   0.319
##  4 Epo                        0.276   0.276
##  5 MAPK2                      0.272   0.272
##  6 ENTP5                     -0.266   0.266
##  7 Protein S                  0.231   0.231
##  8 FSTL1                      0.229   0.229
##  9 IL-17 RC                  -0.229   0.229
```

```
## 10 SRCN1                     0.216   0.216
## 11 CD59                     -0.214   0.214
## 12 IL-6 sRa                 -0.165   0.165
## 13 ITI heavy chain H4        0.154   0.154
## 14 TWEAKR                   -0.147   0.147
## 15 PAI-1                    -0.145   0.145
## 16 PYY                       0.140   0.140
## 17 CSRP3                    -0.139   0.139
## 18 FAM3D                    -0.124   0.124
## 19 hnRNP K                   0.119   0.119
## 20 HGFA                      0.111   0.111
```