Applied Statistical Methods - Solution 7

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Problem 1: Marker Effects Model

Predict genomic breeding values using a marker effects model. The dataset is available from

```
## https://charlotte-ngs.github.io/asmss2023/data/asm_geno_sim_data.csv
```

Hints

- The variance σ_q^2 of the marker effect is 3.
- The residual variance σ_e^2 is 36
- The sex of each animal can be modelled as a fixed effect

Solution

- Read the data

```
tbl_ex11_p01 <- readr::read_csv(s_ex11_p01_data_path)
```

- Setup mixed model equations to predict marker effects for all the SNP-loci. The model is given as

$$y = Xb + Wq + e$$

where y is the vector of observations, b is the vector of fixed effects and q is the vector of random marker effects for each SNP. The matrices X and W are design matrices. The matrix W is special because it contains the genotype encodings.

From that model the mixed model equations can be specified as

$$\begin{bmatrix} X^TX & X^TW \\ W^TX & W^TW + \lambda_q * I \end{bmatrix} \begin{bmatrix} \hat{b} \\ \hat{q} \end{bmatrix} = \begin{bmatrix} X^Ty \\ W^Ty \end{bmatrix}$$

with $\lambda_q = \sigma_e^2/\sigma_q^2$.

The matrix X

```
mat_X <- model.matrix(lm(P ~ 0 + SEX, data = tbl_ex11_p01))
attr(mat_X, "assign") <- NULL
attr(mat_X, "contrasts") <- NULL
mat_X
```

```
##   SEXf SEXm
## 1    0    1
## 2    1    0
## 3    0    1
## 4    1    0
## 5    1    0
## 6    1    0
## 7    0    1
## 8    0    1
```

The matrix W

```
library(dplyr)
```

```
##
## Attaching package: 'dplyr'
```

```
## The following objects are masked from 'package:stats':
##
##     filter, lag
```

```
## The following objects are masked from 'package:base':
##
##     intersect, setdiff, setequal, union
```

```
tbl_geno_ex11_p01 <- tbl_ex11_p01 %>%
  select(SNP1:SNP100)
mat_W <- as.matrix(tbl_geno_ex11_p01)
mat_W[,1:10]
```

```
##      SNP1 SNP2 SNP3 SNP4 SNP5 SNP6 SNP7 SNP8 SNP9 SNP10
## [1,]    2    1    1    1    0    1    2    0    1     1
## [2,]    2    2    0    1    1    1    2    0    1     2
## [3,]    1    0    0    1    1    2    2    0    1     0
## [4,]    1    2    1    1    2    2    2    0    2     1
## [5,]    0    2    0    2    1    1    2    0    1     0
## [6,]    2    2    0    1    1    1    2    0    2     2
## [7,]    2    1    0    2    1    1    2    0    1     0
## [8,]    2    2    1    1    1    1    2    0    0     1
```

The vector y

```
vec_y <- tbl_ex11_p01$P
vec_y
```

```
## [1] 37.5 18.0 22.4 36.7 33.0 33.1 32.4 18.8
```

The mixed model equations

```
# coefficient matrix
mat_xtx <- crossprod(mat_X)
mat_xtw <- crossprod(mat_X, mat_W)
mat_wtx <- t(mat_xtw)
lambda_q <- sigma_e2 / sigma_q2
mat_ztz_lambda_I <- crossprod(mat_W) + lambda_q * diag(1, nrow = ncol(mat_W))
mat_coef <- rbind(cbind(mat_xtx, mat_xtw),
                  cbind(mat_wtx, mat_ztz_lambda_I))
# right hand side
mat_xty <- crossprod(mat_X, vec_y)
mat_wty <- crossprod(mat_W, vec_y)
mat_rhs <- rbind(mat_xty, mat_wty)
# solution
mat_sol <- solve(mat_coef, mat_rhs)
# partition solutions
vec_sol_fix <- mat_sol[1:2,]
vec_sol_marker <- mat_sol[3:nrow(mat_sol),]
```

The solution for the estimates of the fixed effects are:

```
vec_sol_fix
```

```
##     SEXf     SEXm
## 30.02412 28.31841
```

The solutions for the first few marker effects are

```
vec_sol_marker[1:10]
```

```
##          SNP1          SNP2          SNP3          SNP4          SNP5          SNP6
##  8.637400e-02  1.423242e-01  3.568333e-01  8.887511e-02 -7.332053e-02 -5.900523e-02
##          SNP7          SNP8          SNP9         SNP10
##  5.066427e-15  0.000000e+00  5.476375e-01 -1.618549e-02
```

- Compute predicted genomic breeding values based on the estimated marker effects. The predicted genomic breeding values are obtained by the matrix-multiplication of matrix W times the vector of the estimated marker effects.

```
mat_mem_gbv <- crossprod(t(mat_W), vec_sol_marker)
mat_mem_gbv
```

```
##            [,1]
## [1,]  5.8994159
## [2,] -7.0191825
## [3,] -2.8080245
## [4,]  4.2735526
## [5,]  3.1400904
## [6,]  0.3090506
## [7,]  2.8508736
## [8,] -8.1159149
```

Problem 2: Breeding Value Based Model

Use the same dataset as in Problem 1 to predict genomic breeding values based on a breeding-value model. The dataset is available from

```
## https://charlotte-ngs.github.io/asmss2023/data/asm_geno_sim_data.csv
```

Hints

- The genomic variance σ_u^2 of the marker effect is 9.
- The residual variance σ_e^2 is 36
- The sex of each animal can be modelled as a fixed effect
- Use the following function to compute the genomic relationship matrix G based on the matrix of genotypes

```
computeMatGrm <- function(pmatData) {
  matData <- pmatData
  # check the coding, if matData is -1, 0, 1 coded, then add 1 to get to 0, 1, 2 coding
  if (min(matData) < 0) matData <- matData + 1
  # Allele frequencies, column vector of P and sum of frequency products
  freq <- apply(matData, 2, mean) / 2
  P <- 2 * (freq - 0.5)
  sumpq <- sum(freq*(1-freq))
  # Changing the coding from (0,1,2) to (-1,0,1) and subtract matrix P
  Z <- matData - 1 - matrix(P, nrow = nrow(matData),
                            ncol = ncol(matData),
                            byrow = TRUE)
  # Z%*%Zt is replaced by tcrossprod(Z)
  return(tcrossprod(Z)/(2*sumpq))
}
```

- If the genomic relationship matrix G which is computed by the function above cannot be inverted, add $0.05 * I$ to G which results in G^* and use G^* as genomic relationship matrix.

Solution

- Read the data

```
tbl_ex11_p02 <- readr::read_csv(s_ex11_p02_data_path)
```

- Compute the inverse genomic relationship matrix using the given function for the genomic relationship matrix. The genomic relationship matrix G is computed using the above given function with the matrix W from the marker effect model as an argument.

The matrix W

```
library(dplyr)
tbl_geno_ex11_p02 <- tbl_ex11_p02 %>%
  select(SNP1:SNP100)
mat_W <- as.matrix(tbl_geno_ex11_p01)
```

The genomic relationship matrix G

```
mat_G <- computeMatGrm(pmatData = mat_W)
mat_G
```

```
##             [,1]       [,2]        [,3]        [,4]        [,5]       [,6]        [,7]       [,8]
## [1,]  1.0766407 -0.1128958  0.11473153 -0.39192290  0.10371730 -0.3038091 -0.18265259 -0.3038091
## [2,] -0.1128958  0.6067003 -0.42863699 -0.20100964 -0.32216613  0.3570445 -0.28545204  0.3864158
## [3,]  0.1147315 -0.4286370  1.00321248  0.02661771  0.05231758 -0.4433226  0.14777421 -0.4726939
## [4,] -0.3919229 -0.2010096  0.02661771  0.66544286  0.04497476  0.0192749  0.02294631 -0.1863240
## [5,]  0.1037173 -0.3221661  0.05231758  0.04497476  0.80495640 -0.3662230 -0.01009637 -0.3074805
## [6,] -0.3038091  0.3570445 -0.44332263  0.01927490 -0.36622304  0.7535567 -0.24139514  0.2248738
## [7,] -0.1826526 -0.2854520  0.14777421  0.02294631 -0.01009637 -0.2413951  0.79027077 -0.2413951
## [8,] -0.3038091  0.3864158 -0.47269390 -0.18632400 -0.30748050  0.2248738 -0.24139514  0.9004130
```

We have to check whether G can be inverted. This is done by computing the rank of the matrix

```
Matrix::rankMatrix(mat_G)
```

```
## [1] 7
## attr(,"method")
## [1] "tolNorm2"
## attr(,"useGrad")
## [1] FALSE
## attr(,"tol")
## [1] 1.776357e-15
```

This shows that matrix G does not have full column rank. Hence we add $0.05 * I$ to get to matrix G^*.

```
mat_G_star <- mat_G + 0.05 * diag(1, nrow = nrow(mat_G))
Matrix::rankMatrix(mat_G_star)
```

```
## [1] 8
## attr(,"method")
## [1] "tolNorm2"
## attr(,"useGrad")
## [1] FALSE
## attr(,"tol")
## [1] 1.776357e-15
```

Matrix G^* can be used as genomic relationship matrix.

- Setup mixed model equations to predict genomic breeding values. The breeding value based model is given by

$$y = Xb + Zu + e$$

where y is the vector of observations, b is the vector of fixed effects and u is the vector of random genomic breeding values. The matrices X and Z are design matrices.

The mixed model equations are

$$\begin{bmatrix} X^TX & X^TZ \\ Z^TX & Z^TZ + \lambda_g * (G^*)^{-1} \end{bmatrix} \begin{bmatrix} \hat{b} \\ \hat{u} \end{bmatrix} = \begin{bmatrix} X^Ty \\ Z^Ty \end{bmatrix}$$

with $\lambda_u = \sigma_e^2/\sigma_u^2$.

The matrix X

```
mat_X <- model.matrix(lm(P ~ 0 + SEX, data = tbl_ex11_p02))
attr(mat_X, "assign") <- NULL
attr(mat_X, "contrasts") <- NULL
colnames(mat_X) <- NULL
mat_X
```

```
##   [,1] [,2]
## 1    0    1
## 2    1    0
## 3    0    1
## 4    1    0
## 5    1    0
## 6    1    0
## 7    0    1
## 8    0    1
```

The matrix Z

```
# model matrix from data
mat_Z <- model.matrix(lm(P ~ 0 + as.factor(ID), data = tbl_ex11_p02))
attr(mat_Z, "assign") <- NULL
attr(mat_Z, "contrasts") <- NULL
colnames(mat_Z) <- NULL
mat_Z
```

```
##   [,1] [,2] [,3] [,4] [,5] [,6] [,7] [,8]
## 1    1    0    0    0    0    0    0    0
## 2    0    1    0    0    0    0    0    0
## 3    0    0    1    0    0    0    0    0
## 4    0    0    0    1    0    0    0    0
## 5    0    0    0    0    1    0    0    0
## 6    0    0    0    0    0    1    0    0
## 7    0    0    0    0    0    0    1    0
## 8    0    0    0    0    0    0    0    1
```

The vector y

```
vec_y <- tbl_ex11_p02$P
vec_y
```

```
## [1] 37.5 18.0 22.4 36.7 33.0 33.1 32.4 18.8
```

The mixed model equations are

```r
# coefficient matrix
mat_xtx <- crossprod(mat_X)
mat_xtz <- crossprod(mat_X, mat_Z)
mat_ztx <- t(mat_xtz)
lambda_u <- sigma_e2 / sigma_u2
mat_ztz_u_inv_lambda <- crossprod(mat_Z) + lambda_u * mat_G_star
mat_coef <- rbind(cbind(mat_xtx, mat_xtz), cbind(mat_ztx, mat_ztz_u_inv_lambda))
# right hand side
mat_xty <- crossprod(mat_X, vec_y)
mat_zty <- crossprod(mat_Z, vec_y)
mat_rhs <- rbind(mat_xty, mat_zty)
# solution
mat_sol <- solve(mat_coef, mat_rhs)
# partition the solution
vec_sol_fix <- mat_sol[1:2,]
vec_sol_gbv <- mat_sol[3:nrow(mat_sol),]
```

The solution for the estimated fixed effects are

```r
vec_sol_fix
```

```
## [1] 30.33937 27.63563
```

The predicted genomic breeding values are

```r
vec_sol_gbv
```

```
## [1]  2.5835105 -4.0500796 -2.2763349  1.7142930 -0.3002899  2.0786012  1.0235856 -0.7732859
```

Comparing order of animals according to predicted genomic breeding values from Problem 1 and Problem 2:

- marker effect model

```r
order(mat_mem_gbv[,1], decreasing = TRUE)
```

```
## [1] 1 4 5 7 6 3 2 8
```

- breeding value based model

```r
order(vec_sol_gbv, decreasing = TRUE)
```

```
## [1] 1 6 4 7 5 8 3 2
```