26_testPRS_trainAoU_valAoU_testMETS735_post-imp_PC-AiR

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```
#load libraries
library(data.table)
library(tidyverse)
```

```
## — Attaching core tidyverse packages -
                                                               — tidyverse 2.0.0 —
## ✓ dplyr
              1.1.1
                         ✓ readr
                                     2.1.4
## / forcats 1.0.0
                                    1.5.0

✓ stringr

## ✓ ggplot2 3.4.2

✓ tibble

                                     3.2.1
## ✓ lubridate 1.9.2

✓ tidyr

                                     1.3.0
## ✓ purrr
              1.0.1
## — Conflicts —
                                                         — tidyverse_conflicts() —
                          masks data.table::between()
## * dplyr::between()
                          masks stats::filter()
## * dplyr::filter()
## * dplyr::first()
                          masks data.table::first()
## * lubridate::hour()
                          masks data.table::hour()
## * lubridate::isoweek() masks data.table::isoweek()
## * dplyr::lag()
                          masks stats::lag()
## * dplvr::last()
                          masks data.table::last()
## * lubridate::mday()
                         masks data.table::mday()
## * lubridate::minute() masks data.table::minute()
## * lubridate::month()
                         masks data.table::month()
## * lubridate::quarter() masks data.table::quarter()
## * lubridate::second()
                         masks data.table::second()
## * purrr::transpose()
                          masks data.table::transpose()
## * lubridate::wday()
                          masks data.table::wday()
## * lubridate::week()
                          masks data.table::week()
## * lubridate::yday()
                          masks data.table::yday()
## * lubridate::year()
                          masks data.table::year()
## i Use the conflicted package (<http://conflicted.r-lib.org/>) to force all conflicts to become errors
```

```
#paste function to concatenate filenames/paths
"%&%" = function(a,b) paste(a,b,sep="")
```

```
#read in prs, phenotypes, covariates (file made in 25_compare_AoU_PRS_to_observed_mets735_post-imp.Rmd)
mets = fread("METS735_post-imp_PRSCSx_AoU_scaled_bmi_scores_and_covariates.txt")
#read in mets PCs
pcs = fread("/home/wheelerlab3/2023-09-08_PRSCSx/PRSCSx_testing/mets4prscsx_plink/METS735_PC-AiR_PCs.txt")
mets = left_join(mets,pcs,by=join_by('FID'))
table(mets$country_abbr)
```

```
## GH JA SA US
## 220 176 151 188
```

```
dim(mets)
```

```
## [1] 735 42
```

```
#isa: this is just checking data for outliers
\#qqplot(mets,aes(x=PC1,y=PC2,col=country\ abbr)) + qeom\ point() + scale\ color\ viridis\ d()+ theme\ bw(16)
\#ggplot(mets,aes(x=PC3,y=PC4,col=country\_abbr)) + geom\_point() + scale\_color\_viridis\_d() + theme\_bw(16) \#rm o
utliers?
#plot age distribution by site
\#ggplot(mets,aes(x=age,fill=country\_abbr)) + geom\_histogram() + scale\_fill\_viridis\_d() + facet\_wrap(\sim sex)
#filter out PC3>0.1 and PC3< -0.1
#filter out PC4>0.1 and PC4< -0.1
#filter out age>120
mets_rm_outliers = filter(mets,PC3>(-0.1) & PC3<0.1,PC4>(-0.1) & PC4<0.1,age<120 )
\#ggplot(mets\ rm\ outliers,aes(x=PC1,y=PC2,col=country\ abbr)) + geom\ point() + scale\ color\ viridis\ d()+\ theme
bw(16)
#ggplot(mets_rm_outliers,aes(x=PC3,y=PC4,col=country_abbr)) + geom_point() + scale_color_viridis_d()+ theme_
bw(16)
#plot age distribution by site
#ggplot(mets_rm_outliers,aes(x=age,fill=country_abbr)) + geom_histogram() + scale_fill_viridis_d() + facet_w
rap(~sex)
#test with the outliers and missing age removed
mets = mets_rm_outliers
dim(mets)
```

[1] 714 42

```
#read in AoU weights post-validation
#aou = fread("mean_aou10fold_pop_weights.txt")
#head(aou)
#calc new score in METS using AoU-scores multiplied by AoU-held-out validation weights
#one new ensemble score per person per AoU validation pop (all,afr,amr,eur)
#make matrix of AoU weights
#aou mat=as.matrix(aou[,-1])
#make matrix for new ensemble scores
n=714 #mets sample size (outliers and no-age removed)
p=4 #four AoU validation pops
ens_prs = matrix(nrow=n,ncol=p)
for(i in 1:4){
  #need 1x3 matrix of AoU weights
    aou = fread("AoU_scaled_bmi_phi1e-02_popweights_6-3-24.txt")
    aou_mat=as.matrix(aou[,-1])
    aou_weights = t(as.matrix(aou_mat[,i]))
  #need 3x714 matrix of mets PRS's for matrix multiplication
  mets scores = t(as.matrix(select(mets,AFR,AMR,EUR)))
  #multiply each mets PRS by the AoU weight and take the sum
  # %*% does this via matrix multiplication, makes 1x714 matrix
  new_score = aou_weights %*% mets_scores
  ens prs[,i] = new score
}
colnames(ens_prs) = colnames(aou_mat)
rownames(ens_prs) = mets$FID
head(ens prs)
```

```
## 4327 0.54105447 0.29057426 0.53853805 0.66320015

## 4542 -0.20584607 -0.19253850 -0.16738876 -0.18507024

## 4075 0.13477449 0.06815577 0.13777541 0.17956970

## 4524 0.39122624 0.26503560 0.37040938 0.47389338

## 4629 0.09823187 0.10466854 0.07624379 0.09206169

## 4220 -0.15034532 -0.10508915 -0.14619376 -0.21373481
```

```
#make df to join with mets
ens_prs_df = as.data.frame(ens_prs) |> rownames_to_column("FID")
#make FID character in mets
mets = mutate(mets,FID=as.character(FID))
all_mets = left_join(ens_prs_df,mets,by="FID")
```

raw correlation

```
ensemble_prs_bmi=select(all_mets,ends_with("AoU"),bmi)
cor(ensemble_prs_bmi,use="pairwise")
```

```
## allAoU afrAoU amrAoU eurAoU bmi

## allAoU 1.0000000 0.9327509 0.9945106 0.9879121 0.1622063

## afrAoU 0.9327509 1.0000000 0.8904072 0.8747956 0.1940663

## amrAoU 0.9945106 0.8904072 1.0000000 0.9973052 0.1482495

## eurAoU 0.9879121 0.8747956 0.9973052 1.0000000 0.1406393

## bmi 0.1622063 0.1940663 0.1482495 0.1406393 1.0000000
```

```
cor(ensemble_prs_bmi,use='pairwise',method='spearman')
```

```
## allAoU afrAoU amrAoU eurAoU bmi

## allAoU 1.0000000 0.9338572 0.9944218 0.9867646 0.1432626

## afrAoU 0.9338572 1.0000000 0.8935735 0.8781621 0.1658663

## amrAoU 0.9944218 0.8935735 1.0000000 0.9964309 0.1320622

## eurAoU 0.9867646 0.8781621 0.9964309 1.0000000 0.1259066

## bmi 0.1432626 0.1658663 0.1320622 0.1259066 1.0000000
```

```
#does ranking correspond to sample size in AoU?
#next, split by mets country and compare adjust R2
```

Split by METS site and test allAoU, afrAoU, amrAoU, eurAoU

```
#make matrix to store adjusted R2
mets res mat = matrix(nrow=5,ncol=5)
countries = c("GH","SA","JA","US","METS")
for(i in 1:4){
  site = countries[i]
  cat("*****",site,'******\n****************\n')
  data = filter(all_mets,country_abbr==site)
  #baseline model (just covariates)
  cat("*****baseline*****\n")
# prs_pcs = select(data,height,sex,age,starts_with("PC")) #all PCs
  prs_pcs = select(data,bmi,sex,age,PC1,PC2,PC3,PC4,PC5) #5 PCs
  res = summary(lm(scale(bmi)~.,data=prs_pcs))
  mets res mat[i,1] = res$adj.r.squared
  print(res)
  #allAoU ensemble model
  cat("*****allAoU*****\n")
# prs pcs = select(data,allAoU 15k,height,sex,age,starts with("PC"))
  prs_pcs = select(data,allAoU,bmi,sex,age,PC1,PC2,PC3,PC4,PC5) #5 PCs
  res = summary(lm(scale(bmi)~.,data=prs_pcs))
  mets res mat[i,2] = res$adj.r.squared
  print(res)
  #afrAoU ensemble model
  cat("*****afrAoU*****\n")
# prs_pcs = select(data,afrAoU_5k,height,sex,age,starts_with("PC"))
  prs_pcs = select(data,afrAoU,bmi,sex,age,PC1,PC2,PC3,PC4,PC5) #5 PCs
  res = summary(lm(scale(bmi)~.,data=prs pcs))
  mets_res_mat[i,3] = res$adj.r.squared
  print(res)
  #amrAoU ensemble model
  cat("*****amrAoU*****\n")
# prs_pcs = select(data,amrAoU_5k,height,sex,age,starts_with("PC"))
  prs_pcs = select(data,amrAoU,bmi,sex,age,PC1,PC2,PC3,PC4,PC5) #5 PCs
  res = summary(lm(scale(bmi)~.,data=prs_pcs))
  mets_res_mat[i,4] = res$adj.r.squared
  print(res)
  #eurAoU ensemble model
  cat("*****eurAoU*****\n")
# prs pcs = select(data,eurAoU 5k,height,sex,age,starts with("PC"))
  prs_pcs = select(data,eurAoU,bmi,sex,age,PC1,PC2,PC3,PC4,PC5) #5 PCs
  res = summary(lm(scale(bmi)~.,data=prs_pcs))
  mets_res_mat[i,5] = res$adj.r.squared
  print(res)
}
```

```
## ***** GH *****
## *********
## *****baseline****
##
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
               10 Median
##
      Min
                               30
                                      Max
## -1.9983 -0.5295 -0.0682 0.4732 3.9413
##
## Coefficients:
                Estimate Std. Error t value Pr(>|t|)
##
                           0.916931 -1.665
## (Intercept) -1.526540
                                             0.0975 .
                                      6.166 3.64e-09 ***
## sex
                0.831596
                           0.134869
               -0.005602
                           0.008302 -0.675
## age
                                            0.5006
              -45.464098 49.363333 -0.921
## PC1
                                             0.3581
## PC2
               21.617487 14.891308
                                     1.452
                                             0.1481
## PC3
              -50.941247 25.097471 -2.030
                                             0.0437 *
## PC4
              -18.970705 23.067844 -0.822
                                             0.4118
## PC5
               40.938763 24.483498
                                      1.672
                                             0.0960 .
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.9028 on 206 degrees of freedom
    (1 observation deleted due to missingness)
## Multiple R-squared: 0.2117, Adjusted R-squared: 0.1849
## F-statistic: 7.902 on 7 and 206 DF, p-value: 1.748e-08
##
## *****allAoU*****
##
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
##
      Min
               10 Median
                               30
                                      Max
## -1.9764 -0.5517 -0.0526 0.4947 3.8732
##
## Coefficients:
##
                Estimate Std. Error t value Pr(>|t|)
## (Intercept) -1.523340
                           0.918631 -1.658
                                            0.0988 .
## allAoU
                0.156326
                           0.313637
                                      0.498
                                             0.6187
## sex
                0.835473
                           0.135340
                                      6.173 3.53e-09 ***
## age
               -0.005641
                           0.008317 -0.678
                                             0.4984
## PC1
              -44.542471 49.488185 -0.900
                                             0.3691
## PC2
               21.392114 14.925398
                                     1.433
                                             0.1533
## PC3
              -50.673930 25.149098 -2.015
                                             0.0452 *
## PC4
              -19.373116 23.124139 -0.838
                                             0.4031
## PC5
               40.133727 24.581404
                                      1.633
                                             0.1041
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.9045 on 205 degrees of freedom
  (1 observation deleted due to missingness)
## Multiple R-squared: 0.2126, Adjusted R-squared: 0.1819
## F-statistic: 6.92 on 8 and 205 DF, p-value: 4.652e-08
##
## *******
##
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
```

```
##
      Min
                10 Median
                                30
                                       Max
## -1.9597 -0.5526 -0.0608 0.4947
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept) -1.520172
                            0.918909 -1.654
                                               0.0996 .
## afrAoU
                 0.250503
                            0.605060
                                       0.414
                                               0.6793
## sex
                 0.832767
                            0.135171
                                       6.161 3.77e-09 ***
                -0.005679
                            0.008321
                                      -0.683
##
                                               0.4957
  age
## PC1
               -45.230401 49.466131
                                      -0.914
                                               0.3616
## PC2
                21.569462
                           14.921798
                                       1.446
                                               0.1498
## PC3
               -50.476917
                           25.173094 -2.005
                                               0.0463 *
## PC4
               -19.207441 23.121449
                                      -0.831
                                               0.4071
## PC5
                40.569263
                          24.549116
                                       1.653
                                               0.0999 .
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.9047 on 205 degrees of freedom
##
     (1 observation deleted due to missingness)
## Multiple R-squared: 0.2123, Adjusted R-squared: 0.1816
## F-statistic: 6.908 on 8 and 205 DF, p-value: 4.815e-08
  ******amrAoU*****
##
##
## Call:
  lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
##
      Min
                10 Median
                                30
  -1.9821 -0.5513 -0.0528 0.4926 3.8701
##
##
  Coefficients:
##
##
                 Estimate Std. Error t value Pr(>|t|)
##
  (Intercept) -1.524142
                            0.918592
                                     -1.659
## amrAoU
                 0.156229
                            0.305880
                                       0.511
                                               0.6101
                                       6.175 3.49e-09 ***
## sex
                 0.836143
                            0.135405
## age
                -0.005631
                            0.008317 -0.677
                                               0.4991
## PC1
               -44.379365 49.497714 -0.897
                                               0.3710
## PC2
                21.346897 14.927499
                                       1.430
                                               0.1542
## PC3
               -50.719206
                                     -2.017
                                               0.0450 *
                          25.146375
## PC4
              -19.389708
                                               0.4027
                          23.123896 -0.839
## PC5
                40.041463 24.590377
                                       1.628
                                               0.1050
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.9045 on 205 degrees of freedom
     (1 observation deleted due to missingness)
## Multiple R-squared: 0.2127, Adjusted R-squared: 0.182
## F-statistic: 6.922 on 8 and 205 DF, p-value: 4.626e-08
##
## *****eurAoU*****
##
## Call:
  lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
##
                10 Median
                                30
      Min
  -1.9849 -0.5498 -0.0531 0.4889
##
                                   3.8675
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept) -1.523708
                            0.918597 -1.659
                                               0.0987 .
## eurAoU
                 0.121090
                            0.237050
                                       0.511
                                               0.6100
                 0.836500
                            0.135453
                                       6.176 3.48e-09 ***
## sex
```

```
## age
               -0.005637
                           0.008317 -0.678
                                              0.4987
## PC1
              -44.325490 49.502330
                                     -0.895
                                              0.3716
## PC2
               21.321832 14.929316
                                      1.428
                                              0.1548
## PC3
              -50.665866 25.148392
                                     -2.015
                                              0.0452 *
## PC4
              -19.333681
                          23.120258
                                     -0.836
                                              0.4040
## PC5
               40.024902 24.592694
                                      1.628
                                              0.1052
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.9045 on 205 degrees of freedom
    (1 observation deleted due to missingness)
## Multiple R-squared: 0.2127, Adjusted R-squared: 0.182
## F-statistic: 6.922 on 8 and 205 DF, p-value: 4.626e-08
##
## ***** SA *****
## ********
## *****baseline****
##
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
                 10
                     Median
##
       Min
                                   30
                                           Max
## -1.50435 -0.47145 -0.05511 0.39871 2.71724
## Coefficients:
##
                Estimate Std. Error t value Pr(>|t|)
## (Intercept) -1.243236
                           0.767792 - 1.619
                                              0.1077
## sex
                1.228188
                           0.129547
                                      9.481
                                              <2e-16 ***
## age
                0.005897
                           0.009370
                                      0.629
                                              0.5302
## PC1
               12.899772
                           8.481105
                                      1.521
                                              0.1305
## PC2
                0.678094
                           7.362280
                                      0.092
                                              0.9267
## PC3
              -20.789675 23.581687
                                     -0.882
                                              0.3795
## PC4
              -27.715080
                          11.612981
                                     -2.387
                                              0.0184 *
              -44.525934 20.056995 -2.220
## PC5
                                              0.0280 *
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.7766 on 139 degrees of freedom
## Multiple R-squared: 0.4258, Adjusted R-squared: 0.3969
## F-statistic: 14.73 on 7 and 139 DF, p-value: 2.864e-14
##
## *****allAoU*****
##
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
##
                 1Q
                     Median
                                   30
## -1.55400 -0.42619 -0.00025 0.38247 2.61243
##
## Coefficients:
##
                Estimate Std. Error t value Pr(>|t|)
                           0.758216 -1.265 0.20806
## (Intercept) -0.959017
## allAoU
                0.925451
                           0.342009
                                      2.706 0.00767 **
## sex
                1.242697
                           0.126812
                                      9.800 < 2e-16 ***
                0.002764
                           0.009237
                                      0.299 0.76519
  age
## PC1
               15.639477
                           8.356142
                                      1.872 0.06338 .
## PC2
                           7.219191
                                      0.289 0.77282
                2.088236
## PC3
              -18.236641 23.082331 -0.790 0.43084
## PC4
              -28.037789 11.358196 -2.469 0.01479 *
## PC5
              -50.027104 19.720939 -2.537 0.01230 *
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
```

```
##
## Residual standard error: 0.7595 on 138 degrees of freedom
## Multiple R-squared: 0.4548, Adjusted R-squared: 0.4232
## F-statistic: 14.39 on 8 and 138 DF, p-value: 3.938e-15
## *****afrAoU*****
##
## Call:
##
   lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
##
       Min
                  10
                      Median
                                   30
                                           Max
##
  -1.49779 -0.43746 -0.02989 0.37726 2.55171
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept) -0.999933
                           0.753593 -1.327 0.18674
                           0.612856
                                      2.856 0.00496 **
## afrAoU
                 1.750174
                                      9.662
## sex
                 1.220854
                           0.126363
                                             < 2e-16 ***
## age
                                      0.387
                 0.003549
                           0.009175
                                             0.69949
## PC1
               15.522281
                           8.321717
                                      1.865 0.06427 .
## PC2
                 0.898039
                           7.180212
                                       0.125
                                             0.90065
               -16.048304 23.057046
  PC3
                                     -0.696
                                             0.48758
## PC4
              -27.609663 11.325202 -2.438 0.01604 *
## PC5
               -50.518977
                          19.672118 -2.568 0.01129 *
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.7573 on 138 degrees of freedom
## Multiple R-squared: 0.4579, Adjusted R-squared: 0.4265
## F-statistic: 14.57 on 8 and 138 DF, p-value: 2.707e-15
##
## *******
##
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
##
                  10
                      Median
                                   30
       Min
                                            Max
  -1.56817 -0.41663 0.00402 0.37422 2.63285
##
##
## Coefficients:
##
                Estimate Std. Error t value Pr(>|t|)
## (Intercept) -0.958259
                           0.760126 - 1.261
                                              0.2096
##
  amrAoU
                 0.886304
                            0.339693
                                       2.609
                                              0.0101 *
##
  sex
                 1.247125
                           0.127130
                                      9.810
                                              <2e-16 ***
                0.002689
                           0.009263
                                      0.290
## age
                                              0.7721
## PC1
               15.598605
                           8.373430
                                      1.863
                                              0.0646 .
## PC2
                2.297864
                           7.239801
                                       0.317
                                              0.7514
## PC3
               -18.984669
                          23.114324 -0.821
                                               0.4129
## PC4
                          11.379013 -2.475
               -28.161757
                                              0.0145 *
## PC5
               -49.790105 19.753985 -2.521
                                              0.0129 *
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.7608 on 138 degrees of freedom
## Multiple R-squared: 0.4528, Adjusted R-squared: 0.4211
## F-statistic: 14.28 on 8 and 138 DF, p-value: 4.968e-15
##
## *****eurAoU*****
##
## Call:
  lm(formula = scale(bmi) ~ ., data = prs_pcs)
```

```
## Residuals:
       Min
                  1Q
                      Median
## -1.56778 -0.43466 0.00612 0.37362 2.65237
##
## Coefficients:
##
                Estimate Std. Error t value Pr(>|t|)
## (Intercept) -0.962332
                           0.762156 - 1.263
                                              0.2088
                0.663476
                           0.266131
                                      2.493
                                               0.0138 *
                                       9.792
                1.247775
                           0.127426
                                              <2e-16 ***
##
  sex
                                       0.296
##
  age
                0.002745
                           0.009286
                                               0.7680
## PC1
                15.598548
                           8.396418
                                      1.858
                                              0.0653 .
## PC2
                2.269038
                           7.256050
                                      0.313
                                              0.7550
## PC3
              -19.845135 23.154468
                                     -0.857
                                               0.3929
## PC4
              -28.319798 11.403647 -2.483
                                               0.0142 *
## PC5
              -49.783837 19.803619 -2.514
                                              0.0131 *
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.7624 on 138 degrees of freedom
## Multiple R-squared: 0.4506, Adjusted R-squared: 0.4187
## F-statistic: 14.15 on 8 and 138 DF, p-value: 6.498e-15
##
## ***** JA *****
## *********
## *****baseline****
##
## Call:
  lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
##
                10 Median
  -1.9879 -0.5521 -0.0065 0.4307
                                   3.6803
##
##
  Coefficients:
##
                Estimate Std. Error t value Pr(>|t|)
                          0.745878 - 2.493
## (Intercept) -1.859480
                                              0.0137 *
## sex
               1.158963
                          0.143686
                                     8.066 1.57e-13 ***
                          0.008865
                                    -0.402
                                              0.6884
## age
              -0.003562
## PC1
                4.212483 26.487429
                                     0.159
                                              0.8738
## PC2
               1.099920
                         12.511890
                                     0.088
                                              0.9301
## PC3
                          8.335445
                                     0.214
               1.782624
                                              0.8309
## PC4
              -4.790194
                          8.128785
                                    -0.589
                                              0.5565
## PC5
                0.624282
                          6.119567
                                     0.102
                                              0.9189
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.8585 on 161 degrees of freedom
   (3 observations deleted due to missingness)
## Multiple R-squared: 0.2937, Adjusted R-squared: 0.263
## F-statistic: 9.563 on 7 and 161 DF, p-value: 6.493e-10
##
## ******allAoU*****
##
## Call:
  lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
##
      Min
                10 Median
                                30
                                      Max
##
  -2.0072 -0.4767 0.0016 0.5326 3.6668
##
## Coefficients:
##
                Estimate Std. Error t value Pr(>|t|)
## (Intercept) -1.768022
                          0.744883 - 2.374
                                              0.0188 *
                0.434072
                          0.278128
## allAoU
                                     1.561
                                              0.1206
```

```
## sex
               1.161726
                          0.143060
                                      8.121 1.17e-13 ***
                           0.008852
                                              0.6030
## age
               -0.004613
                                     -0.521
## PC1
                1.907901 26.411420
                                      0.072
                                              0.9425
                0.224771 12.469086
## PC2
                                      0.018
                                              0.9856
## PC3
                0.975943
                           8.314607
                                      0.117
                                              0.9067
## PC4
              -5.455115
                           8.103987
                                    -0.673
                                              0.5018
## PC5
                0.982108
                           6.096774
                                      0.161
                                              0.8722
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.8547 on 160 degrees of freedom
##
    (3 observations deleted due to missingness)
## Multiple R-squared: 0.3043, Adjusted R-squared: 0.2695
## F-statistic: 8.747 on 8 and 160 DF, p-value: 7e-10
##
## *******
##
## Call:
  lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
##
      Min
                10 Median
                                30
  -1.9817 -0.5220 0.0033 0.4982
                                   3.6605
##
## Coefficients:
##
                Estimate Std. Error t value Pr(>|t|)
                           0.742650 -2.443
## (Intercept) -1.814211
                                              0.0157 *
## afrAoU
                0.684689
                           0.422011
                                      1.622
                                              0.1067
## sex
               1.149783
                           0.143075
                                     8.036 1.92e-13 ***
## age
              -0.003902
                          0.008823 -0.442
                                              0.6589
## PC1
                3.537216 26.357454
                                      0.134
                                              0.8934
## PC2
                0.953308 12.449270
                                      0.077
                                              0.9391
                                      0.074
## PC3
               0.615578
                           8.324644
                                              0.9411
## PC4
               -5.148748
                           8.090907
                                     -0.636
                                              0.5254
## PC5
                1.068480
                           6.094931
                                      0.175
                                              0.8611
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.8542 on 160 degrees of freedom
     (3 observations deleted due to missingness)
## Multiple R-squared: 0.3051, Adjusted R-squared: 0.2704
## F-statistic: 8.782 on 8 and 160 DF, p-value: 6.405e-10
##
## *******
##
## Call:
##
  lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
##
      Min
                10 Median
                                30
                                       Max
## -2.0111 -0.4757 0.0122 0.5427 3.6661
##
## Coefficients:
##
                Estimate Std. Error t value Pr(>|t|)
## (Intercept) -1.758602
                        0.745652 -2.358
                                              0.0196 *
## amrAoU
                0.437645
                           0.285007
                                      1.536
                                              0.1266
## sex
                1.165570
                           0.143148
                                     8.142 1.03e-13 ***
## age
               -0.004786
                          0.008864
                                    -0.540
                                              0.5900
## PC1
               1.518216 26.434721
                                              0.9543
                                     0.057
## PC2
               0.051734 12.478141
                                     0.004
                                              0.9967
## PC3
                1.154611
                           8.310583
                                      0.139
                                              0.8897
## PC4
                           8.109597
                                     -0.684
                                              0.4952
               -5.544152
## PC5
                0.970060
                           6.098080
                                      0.159
                                              0.8738
```

```
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.8549 on 160 degrees of freedom
## (3 observations deleted due to missingness)
## Multiple R-squared: 0.3039, Adjusted R-squared: 0.2691
## F-statistic: 8.733 on 8 and 160 DF, p-value: 7.251e-10
##
## *****eurAoU*****
##
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
## Residuals:
##
      Min
               10 Median
                               30
                                      Max
## -2.0257 -0.4799 0.0132 0.5386 3.6535
##
## Coefficients:
##
               Estimate Std. Error t value Pr(>|t|)
## (Intercept) -1.753389 0.744831 -2.354
                                             0.0198 *
                                             0.1034
               0.363151
                         0.221697
                                     1.638
## sex
               1.169297
                          0.143080
                                     8.172 8.68e-14 ***
               -0.004924
                          0.008859
                                    -0.556
                                             0.5791
  age
               1.249953 26.412040
## PC1
                                     0.047
                                             0.9623
                                             0.9977
## PC2
                                   -0.003
              -0.035236 12.466271
## PC3
               1.379917
                         8.295855
                                     0.166
                                             0.8681
## PC4
              -5.704319
                          8.105857 -0.704
                                             0.4826
## PC5
               1.078911
                          6.094150
                                     0.177
                                             0.8597
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.854 on 160 degrees of freedom
    (3 observations deleted due to missingness)
## Multiple R-squared: 0.3053, Adjusted R-squared: 0.2706
## F-statistic: 8.791 on 8 and 160 DF, p-value: 6.26e-10
##
## ***** US *****
## *********
## *****baseline****
##
## Call:
  lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
##
               10 Median
                               30
  -1.7578 -0.7416 -0.1056 0.5117 3.6354
##
##
## Coefficients:
##
              Estimate Std. Error t value Pr(>|t|)
                          0.86261 -1.195 0.233593
## (Intercept) -1.03117
## sex
               0.55874
                          0.16589
                                   3.368 0.000936 ***
## age
              -0.01184
                          0.01163 -1.018 0.309992
## PC1
              34.13421
                         32.75799
                                    1.042 0.298886
## PC2
              16.95108
                         15.10588
                                    1.122 0.263381
## PC3
              -1.27532
                         7.48935 -0.170 0.864988
## PC4
              -0.83713
                          7.43781 -0.113 0.910520
              -1.43686
## PC5
                          5.40461 -0.266 0.790670
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.9717 on 170 degrees of freedom
     (2 observations deleted due to missingness)
## Multiple R-squared: 0.09322,
                                   Adjusted R-squared: 0.05588
## F-statistic: 2.497 on 7 and 170 DF, p-value: 0.01821
```

```
## *****allAoU*****
##
## Call:
## lm(formula = scale(bmi) ~ ., data = prs pcs)
## Residuals:
##
      Min
                10 Median
                                30
                                      Max
## -1.6823 -0.6601 -0.0981 0.4919 3.5626
##
## Coefficients:
##
               Estimate Std. Error t value Pr(>|t|)
## (Intercept) -1.21304
                          0.85307 -1.422 0.156877
                0.67560
                          0.27303 2.474 0.014331 *
## allAoU
## sex
               0.56709
                          0.16348
                                   3.469 0.000663 ***
               -0.01092
                          0.01147 -0.952 0.342315
## age
                                   1.152 0.250950
## PC1
               37.20856
                         32.29920
## PC2
               16.75109
                         14.88351
                                   1.125 0.261981
                          7.51690 -0.641 0.522093
## PC3
              -4.82179
## PC4
              -0.74934
                          7.32829 -0.102 0.918678
              -3.97846
                          5.42313 -0.734 0.464203
## PC5
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.9573 on 169 degrees of freedom
    (2 observations deleted due to missingness)
## Multiple R-squared: 0.1249, Adjusted R-squared: 0.0835
## F-statistic: 3.016 on 8 and 169 DF, p-value: 0.003434
##
## *****afrAoU*****
##
## Call:
## lm(formula = scale(bmi) ~ ., data = prs pcs)
##
## Residuals:
                10 Median
                                30
## -1.7389 -0.6276 -0.1126 0.5351 3.3165
##
## Coefficients:
               Estimate Std. Error t value Pr(>|t|)
##
## (Intercept) -1.10089
                          0.84283 -1.306 0.19327
## afrAoU
               1.11370
                          0.36709
                                    3.034 0.00280 **
               0.53773
                          0.16218
                                    3.316 0.00112 **
## sex
               -0.01127
                          0.01136 -0.992 0.32262
## age
## PC1
               34.65434
                         31.99550
                                    1.083 0.28031
               15.14553
                         14.76605
                                    1.026 0.30650
## PC2
## PC3
               -5.07914
                          7.42159 -0.684
                                           0.49468
## PC4
              -0.79988
                          7.26459 -0.110 0.91246
## PC5
              -4.51759
                          5.37551 -0.840 0.40187
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.949 on 169 degrees of freedom
     (2 observations deleted due to missingness)
## Multiple R-squared: 0.1401, Adjusted R-squared: 0.09935
## F-statistic: 3.441 on 8 and 169 DF, p-value: 0.001073
##
## *****amrAoU*****
##
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
##
## Residuals:
##
      Min
                10 Median
                                30
                                      Max
## -1.6741 -0.6679 -0.1113 0.4761 3.6359
```

```
##
## Coefficients:
##
               Estimate Std. Error t value Pr(>|t|)
                           0.85780 -1.430 0.154703
## (Intercept) -1.22623
                                    2.188 0.030018 *
## amrAoU
                0.63294
                           0.28923
                                    3.492 0.000612 ***
                0.57339
                           0.16421
## sex
## age
               -0.01090
                           0.01151 -0.947 0.344946
## PC1
               37.64886
                          32.43872
                                   1.161 0.247435
## PC2
               17,21707
                          14.94080
                                    1.152 0.250804
## PC3
               -4.46812
                          7.54959 -0.592 0.554751
               -0.74440
                           7.35640 -0.101 0.919519
## PC4
## PC5
               -3.58950
                           5.43513 -0.660 0.509880
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.961 on 169 degrees of freedom
     (2 observations deleted due to missingness)
## Multiple R-squared: 0.1182, Adjusted R-squared: 0.07646
## F-statistic: 2.832 on 8 and 169 DF, p-value: 0.005647
##
##
  ******eurAoU*****
##
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
## Residuals:
##
                10 Median
      Min
                                30
                                       Max
  -1.6741 -0.6673 -0.1149 0.4769
##
                                   3.6715
##
## Coefficients:
##
               Estimate Std. Error t value Pr(>|t|)
                           0.85999 -1.422 0.156926
  (Intercept) -1.22274
## eurAoll
                0.46173
                           0.22743
                                   2.030 0.043902 *
  sex
                0.57295
                           0.16454
                                     3.482 0.000633 ***
               -0.01092
                           0.01153 -0.946 0.345295
## age
               37.81286
## PC1
                         32.51182
                                    1.163 0.246449
## PC2
               17.47417
                         14.97128
                                    1.167 0.244782
## PC3
               -4.29433
                          7.56903 -0.567 0.571225
## PC4
               -0.74821
                           7.37057 -0.102 0.919263
## PC5
               -3.27666
                           5.43178 -0.603 0.547159
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.9629 on 169 degrees of freedom
     (2 observations deleted due to missingness)
## Multiple R-squared: 0.1148, Adjusted R-squared: 0.07291
## F-statistic: 2.74 on 8 and 169 DF, p-value: 0.007227
```

```
#add all of METS
data = all_mets
i=5
cat("*****baseline*******\n")
```

```
## *****baseline****
```

```
prs_pcs = select(data,bmi,sex,age,PC1,PC2,PC3,PC4,PC5) #5 PCs
res = summary(lm(scale(bmi)~.,data=prs_pcs))
mets_res_mat[i,1] = res$adj.r.squared
print(res)
```

```
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
## Residuals:
##
      Min
               1Q Median
                               30
                                      Max
## -1.7988 -0.5515 -0.1122 0.4226 3.8531
##
## Coefficients:
                Estimate Std. Error t value Pr(>|t|)
##
## (Intercept) -1.4467185 0.2374470 -6.093 1.83e-09 ***
               0.9052514  0.0690480  13.110  < 2e-16 ***
## sex
## age
              -0.0004492 0.0045204 -0.099
                                               0.921
## PC1
               4.3133915 1.0224547
                                     4.219 2.78e-05 ***
## PC2
              -6.3541290 1.2271470 -5.178 2.94e-07 ***
## PC3
              -0.5278048 4.6902408 -0.113
                                               0.910
## PC4
              -2.1507250 4.2151447 -0.510
                                               0.610
## PC5
              -3.8255479 3.2261952 -1.186
                                               0.236
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.865 on 700 degrees of freedom
    (6 observations deleted due to missingness)
## Multiple R-squared: 0.2593, Adjusted R-squared: 0.2518
                  35 on 7 and 700 DF, p-value: < 2.2e-16
```

```
#allAoU ensemble model
cat("******allAoU*******\n")
```

```
## *****allAoU*****
```

```
prs_pcs = select(data,allAoU,bmi,sex,age,PC1,PC2,PC3,PC4,PC5) #5 PCs
res = summary(lm(scale(bmi)~.,data=prs_pcs))
mets_res_mat[i,2] = res$adj.r.squared
print(res)
```

```
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
## Residuals:
##
      Min
                10 Median
                                30
                                      Max
  -1.8934 -0.5553 -0.0968 0.4234 3.8151
##
## Coefficients:
                Estimate Std. Error t value Pr(>|t|)
##
## (Intercept) -1.419683
                          0.234010 -6.067 2.14e-09 ***
                0.660560
                          0.140354
                                     4.706 3.04e-06 ***
## allAoU
## sex
                0.910976
                          0.068039 13.389
                                           < 2e-16 ***
## age
              -0.001361
                          0.004458 -0.305
                                              0.760
                4.536116
                          1.008461
                                     4.498 8.03e-06 ***
##
  PC1
## PC2
              -5.706588
                          1.216822 -4.690 3.29e-06 ***
## PC3
              -2.411084
                          4.638244 -0.520
                                              0.603
              -2.186560
                          4.152880 -0.527
                                              0.599
## PC4
## PC5
              -4.533128
                          3.182087 -1.425
                                               0.155
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.8522 on 699 degrees of freedom
     (6 observations deleted due to missingness)
## Multiple R-squared: 0.282, Adjusted R-squared: 0.2738
## F-statistic: 34.32 on 8 and 699 DF, p-value: < 2.2e-16
```

```
#afrAoU ensemble model
cat("****afrAoU*******\n")
```

```
## *****afrAoU*****
```

```
#prs_pcs = select(data,afrAoU,height,sex,age,starts_with("PC"))
prs_pcs = select(data,afrAoU,bmi,sex,age,PC1,PC2,PC3,PC4,PC5) #5 PCs
res = summary(lm(scale(bmi)~.,data=prs_pcs))
mets_res_mat[i,3] = res$adj.r.squared
print(res)
```

```
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
## Residuals:
##
      Min
               10 Median
                               30
                                      Max
  -1.8492 -0.5552 -0.0827 0.4075 3.5739
##
## Coefficients:
               Estimate Std. Error t value Pr(>|t|)
##
## (Intercept) -1.402013
                         0.233090 -6.015 2.90e-09 ***
                                     5.325 1.36e-07 ***
## afrAoU
               1.154563
                          0.216814
## sex
               0.893611
                          0.067772 13.185 < 2e-16 ***
## age
              -0.001097
                          0.004436 -0.247
                                              0.805
               4.609443
                          1.004582
                                    4.588 5.29e-06 ***
## PC1
## PC2
              -5.809754
                          1.208181 -4.809 1.86e-06 ***
## PC3
                         4.617998 -0.568
                                              0.570
              -2.623937
                                              0.640
## PC4
              -1.935701 4.135313 -0.468
## PC5
              -4.639112
                         3.168628 -1.464
                                              0.144
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.8485 on 699 degrees of freedom
    (6 observations deleted due to missingness)
## Multiple R-squared: 0.2881, Adjusted R-squared: 0.28
## F-statistic: 35.37 on 8 and 699 DF, p-value: < 2.2e-16
```

```
#amrAoU ensemble model
cat("*****amrAoU******\n")
```

```
## ******amrAoU*****
```

```
prs_pcs = select(data,amrAoU,bmi,sex,age,PC1,PC2,PC3,PC4,PC5) #5 PCs
res = summary(lm(scale(bmi)~.,data=prs_pcs))
mets_res_mat[i,4] = res$adj.r.squared
print(res)
```

```
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
## Residuals:
##
      Min
               10 Median
                               30
                                      Max
  -1.8924 -0.5519 -0.0904 0.4275 3.8833
##
## Coefficients:
               Estimate Std. Error t value Pr(>|t|)
##
## (Intercept) -1.425812
                         0.234454 -6.081 1.96e-09 ***
                          0.143166
                                     4.391 1.30e-05 ***
## amrAoU
               0.628698
## sex
               0.915004
                          0.068200 13.417 < 2e-16 ***
## age
              -0.001374
                          0.004467 -0.307
                                              0.759
                                    4.461 9.52e-06 ***
               4.506661
                          1.010316
## PC1
## PC2
              -5.710983
                          1.220248 -4.680 3.44e-06 ***
## PC3
              -2.233858
                         4.646430 -0.481
                                              0.631
              -2.238600
                         4.161198 -0.538
                                              0.591
## PC4
## PC5
              -4.433640
                         3.187877 -1.391
                                              0.165
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.8539 on 699 degrees of freedom
    (6 observations deleted due to missingness)
## Multiple R-squared: 0.2791, Adjusted R-squared: 0.2709
## F-statistic: 33.83 on 8 and 699 DF, p-value: < 2.2e-16
```

```
#eurAoU ensemble model
cat("*****eurAoU******\n")
```

```
## *****eurAoU*****
```

```
prs_pcs = select(data,eurAoU,bmi,sex,age,PC1,PC2,PC3,PC4,PC5) #5 PCs
res = summary(lm(scale(bmi)~.,data=prs_pcs))
mets_res_mat[i,5] = res$adj.r.squared
print(res)
```

METS

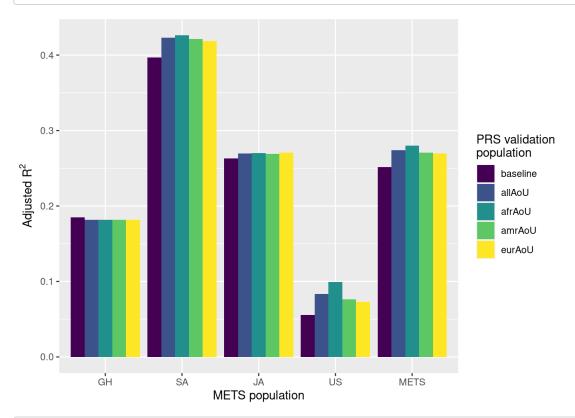
```
## Call:
## lm(formula = scale(bmi) ~ ., data = prs_pcs)
## Residuals:
##
      Min
               10 Median
                               30
                                     Max
## -1.8929 -0.5461 -0.0863 0.4263 3.9174
##
## Coefficients:
##
               Estimate Std. Error t value Pr(>|t|)
                         0.234684 -6.089 1.87e-09 ***
## (Intercept) -1.428994
                                    4.220 2.76e-05 ***
## eurAoU
               0.472044
                          0.111850
               ## sex
## age
              -0.001361 0.004472 -0.304
                                             0.761
                                    4.448 1.01e-05 ***
## PC1
               4.498332
                         1.011344
              -5.698516
                         1.222581 -4.661 3.77e-06 ***
## PC2
              -2.085313 4.649584 -0.448
## PC3
                                             0.654
              -2.226844 4.165462 -0.535
## PC4
                                             0.593
## PC5
              -4.274405
                         3.189912 -1.340
                                             0.181
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.8548 on 699 degrees of freedom
    (6 observations deleted due to missingness)
## Multiple R-squared: 0.2777, Adjusted R-squared: 0.2694
## F-statistic: 33.59 on 8 and 699 DF, p-value: < 2.2e-16
rownames(mets res mat) = countries
colnames(mets_res_mat) = c("baseline","allAoU","afrAoU","amrAoU","eurAoU")
mets_res_mat
##
         baseline
                      allAoU
                                 afrAoU
                                           amrAoU
                                                     eurAoU
       0.18489007 0.18190534 0.18159822 0.1819549 0.18195519
## GH
## SA
       0.39692830 0.42316403 0.42645324 0.4211148 0.41873716
       0.26297826 0.26949275 0.27037567 0.2691426 0.27060388
## .JA
       0.05588161 0.08350011 0.09934733 0.0764644 0.07290634
## US
## METS 0.25184835 0.27379047 0.27998757 0.2708930 0.26939468
table(all_mets$country_abbr)
##
## GH JA SA US
## 215 172 147 180
mets_res_mat/mets_res_mat[,1]
##
       baseline
                   allAoU
                             afrAoU
                                       amrAoU
                                                eurAoU
              1 0.9838567 0.9821956 0.9841249 0.9841263
## GH
## SA
              1 1.0660969 1.0743836 1.0609343 1.0549441
## JA
              1 1.0247720 1.0281294 1.0234406 1.0289972
## IIS
              1 1.4942325 1.7778181 1.3683286 1.3046572
```

```
table(all_mets$country_abbr)
```

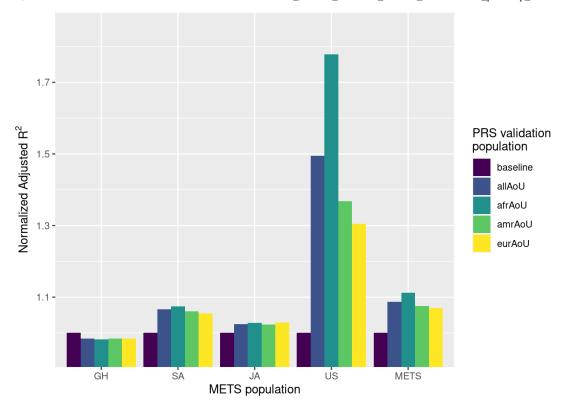
1 1.0871243 1.1117308 1.0756194 1.0696702

```
## ## GH JA SA US
## 215 172 147 180
```

```
mets_res_df = rownames_to_column(as.data.frame(mets_res_mat),"site")
mets_res_df_long = mets_res_df |> pivot_longer(!site, names_to = "PRS_val_pop", values_to = "Adj_R2")
ggplot(mets_res_df_long,aes(x=factor(site,levels=c("GH","SA","JA","US","METS")),y=Adj_R2,fill=factor(PRS_val_pop,levels=c("baseline","allAoU","afrAoU","eurAoU","eurAoU")))) +
    geom_bar(position="dodge", stat="identity") +
    scale_fill_viridis_d(name="PRS validation\npopulation") +
    labs(x="METS population",y=bquote("Adjusted"~R^2))
```



```
#normalize by baseline
norm_mets_res = rownames_to_column(as.data.frame(mets_res_mat/mets_res_mat[,1]),"site")
norm_mets_long = norm_mets_res |> pivot_longer(!site, names_to = "PRS_val_pop", values_to = "Adj_R2")
ggplot(norm_mets_long,aes(x=factor(site,levels=c("GH","SA","JA","US","METS")),y=Adj_R2,fill=factor(PRS_val_p
op,levels=c("baseline","allAoU","afrAoU","amrAoU","eurAoU")))) +
    geom_bar(position="dodge", stat="identity") +
    scale_fill_viridis_d(name="PRS validation\npopulation") +
    labs(x="METS population",y=bquote("Normalized Adjusted"~R^2)) +
    coord_cartesian(ylim=c(0.95,1.85))
```



looking at normalized by baseline: makes sense that US is dramatically better because it's closest to aou pops. looking at raw adj r2: i do NOT know

scale across all METS first

```
all_mets=mutate(all_mets,scale_bmi = scale(bmi))
#make matrix to store adjusted R2
mets_res_mat = matrix(nrow=4,ncol=5)
countries = c("GH","SA","JA","US")
for(i in 1:4){
  site = countries[i]
  data = filter(all_mets,country_abbr==site)
  #baseline model (just covariates)
  prs_pcs = select(data,scale_bmi,sex,age,PC1,PC2,PC3,PC4,PC5)
  res = summary(lm(scale_bmi~.,data=prs_pcs))
  mets_res_mat[i,1] = res$adj.r.squared
  #allAoU ensemble model
  prs_pcs = select(data,allAoU,scale_bmi,sex,age,PC1,PC2,PC3,PC4,PC5)
  res = summary(lm(scale_bmi~.,data=prs_pcs))
  mets_res_mat[i,2] = res$adj.r.squared
  #afrAoU ensemble model
  prs_pcs = select(data,afrAoU,scale_bmi,sex,age,PC1,PC2,PC3,PC4,PC5)
  res = summary(lm(scale_bmi~.,data=prs_pcs))
  mets_res_mat[i,3] = res$adj.r.squared
  #amrAoU ensemble model
  prs_pcs = select(data,amrAoU,scale_bmi,sex,age,PC1,PC2,PC3,PC4,PC5)
  res = summary(lm(scale_bmi~.,data=prs_pcs))
  mets_res_mat[i,4] = res$adj.r.squared
  #eurAoU ensemble model
  prs_pcs = select(data,eurAoU,scale_bmi,sex,age,PC1,PC2,PC3,PC4,PC5)
  res = summary(lm(scale_bmi~.,data=prs_pcs))
  mets res mat[i,5] = res$adj.r.squared
}
rownames(mets_res_mat) = countries
colnames(mets res mat) = c("baseline","allAoU","afrAoU","amrAoU","eurAoU")
mets_res_mat
```

```
## baseline allAoU afrAoU amrAoU eurAoU

## GH 0.18489007 0.18190534 0.18159822 0.1819549 0.18195519

## SA 0.39692830 0.42316403 0.42645324 0.4211148 0.41873716

## JA 0.26297826 0.26949275 0.27037567 0.2691426 0.27060388

## US 0.05588161 0.08350011 0.09934733 0.0764644 0.07290634
```

```
mets_res_mat/mets_res_mat[,1]
```

```
table(all_mets$country_abbr)
```

```
##
## GH JA SA US
## 215 172 147 180
```

raw correlation b/t PRS and height

```
#used these numbers (afrAoU) for ASHG abstract
all_mets |> group_by(country_abbr) |> summarize(cor=cor(scale(bmi), allAoU, use="pairwise",method='spearman'))
```

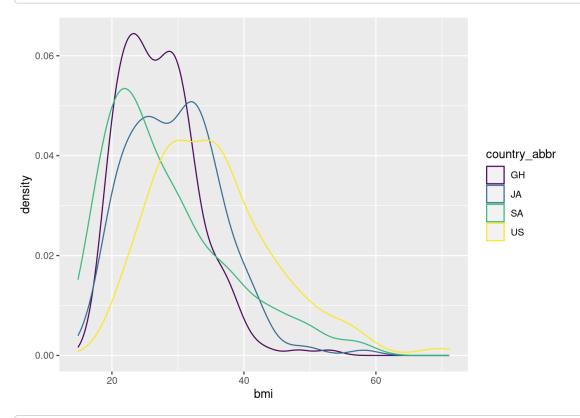
```
all_mets |> group_by(country_abbr) |> summarize(cor=cor(scale(bmi), afrAoU, use="pairwise",method='spearma
n'))
```

```
all_mets |> group_by(country_abbr) |> summarize(cor=cor(scale(bmi), eurAoU, use="pairwise",method='spearma
n'))
```

plot BMI and PRS distributions

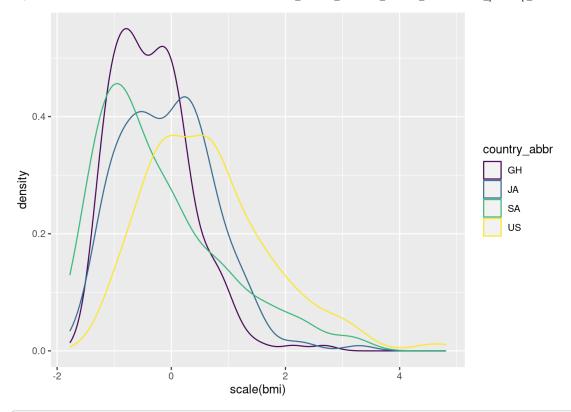
```
#height by site
ggplot(all_mets,aes(x=bmi,col=country_abbr)) + geom_density() + scale_color_viridis_d()
```

Warning: Removed 6 rows containing non-finite values (`stat_density()`).

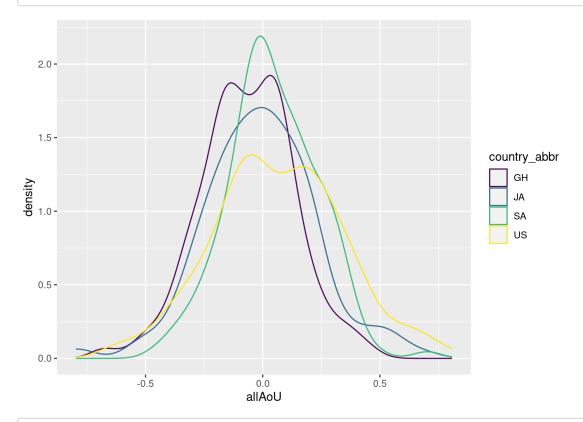


```
#scaled height by site
ggplot(all_mets,aes(x=scale(bmi),col=country_abbr)) + geom_density() + scale_color_viridis_d()
```

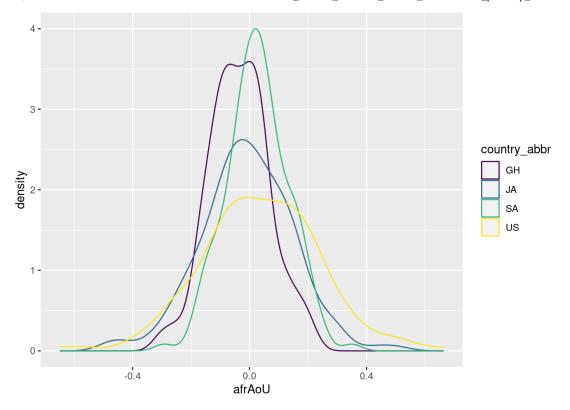
Warning: Removed 6 rows containing non-finite values (`stat_density()`).



#allAoU PRS
ggplot(all_mets,aes(x=allAoU,col=country_abbr)) + geom_density() + scale_color_viridis_d()

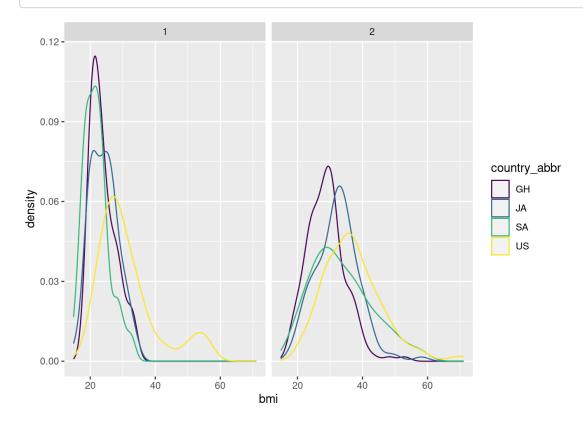


#afrAoU PRS
ggplot(all_mets,aes(x=afrAoU,col=country_abbr)) + geom_density() + scale_color_viridis_d()



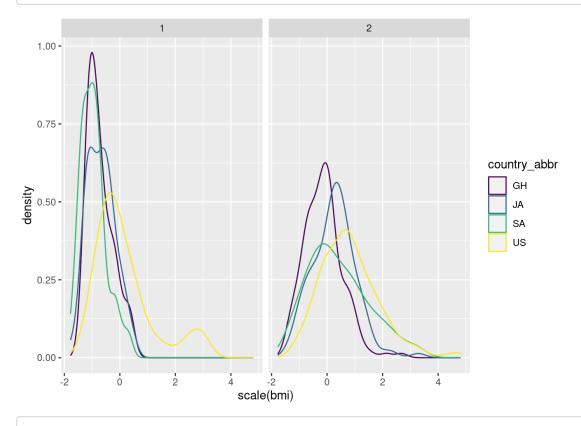
#facet by sex
#height by site
ggplot(all_mets,aes(x=bmi,col=country_abbr)) + geom_density() + scale_color_viridis_d() + facet_wrap(~sex)

Warning: Removed 6 rows containing non-finite values (`stat_density()`).

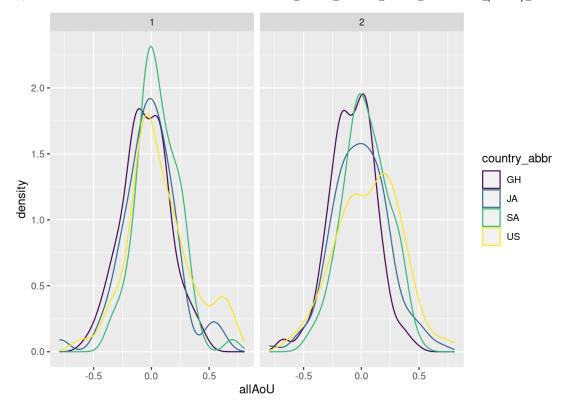


#scaled height by site
ggplot(all_mets,aes(x=scale(bmi),col=country_abbr)) + geom_density() + scale_color_viridis_d() + facet_wrap
(~sex)

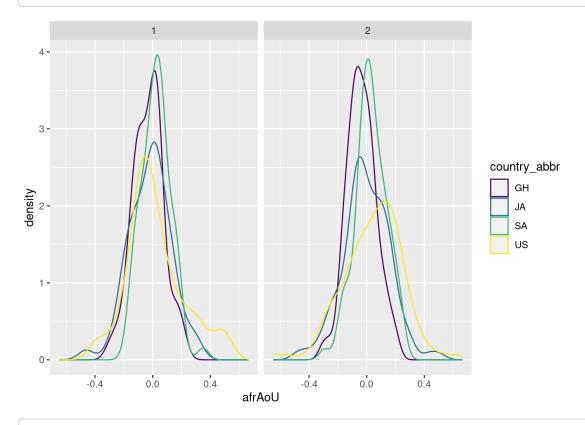
Warning: Removed 6 rows containing non-finite values (`stat_density()`).



#allAoU PRS
ggplot(all_mets,aes(x=allAoU,col=country_abbr)) + geom_density() + scale_color_viridis_d() + facet_wrap(~se
x)



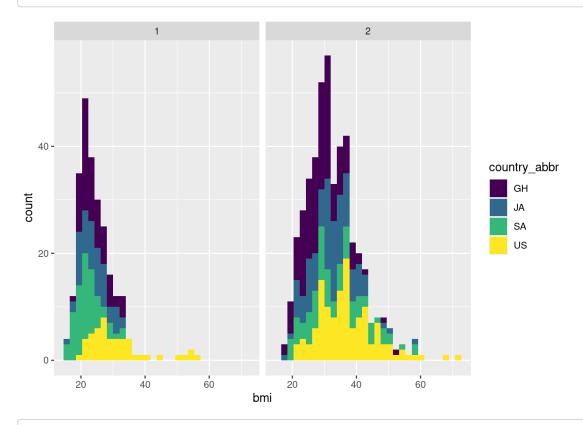
#afrAoU PRS
ggplot(all_mets,aes(x=afrAoU,col=country_abbr)) + geom_density() + scale_color_viridis_d() + facet_wrap(~se
x)



```
#histograms
#height by site
ggplot(all_mets,aes(x=bmi,fill=country_abbr)) + geom_histogram() + scale_fill_viridis_d() + facet_wrap(~sex)
```

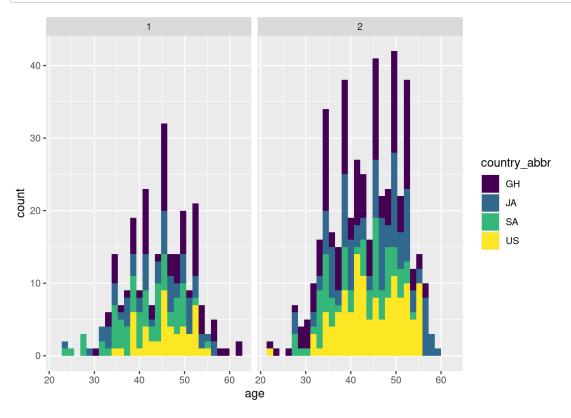
`stat_bin()` using `bins = 30`. Pick better value with `binwidth`.

Warning: Removed 6 rows containing non-finite values (`stat_bin()`).



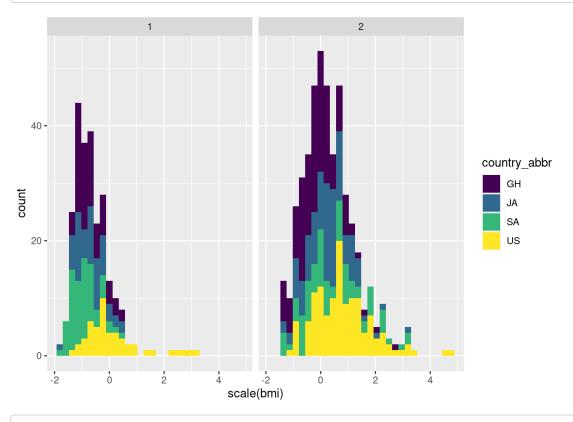
#age by site
ggplot(all_mets,aes(x=age,fill=country_abbr)) + geom_histogram() + scale_fill_viridis_d() + facet_wrap(~sex)

`stat_bin()` using `bins = 30`. Pick better value with `binwidth`.



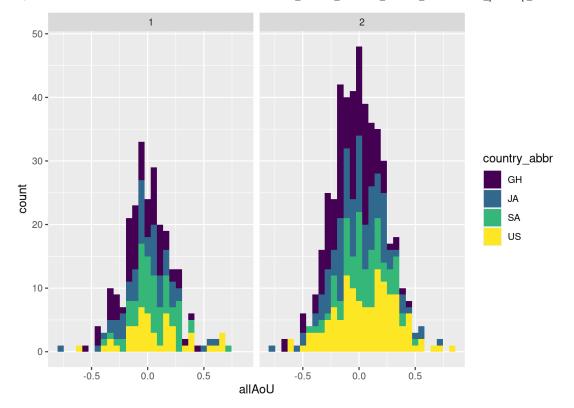
`stat_bin()` using `bins = 30`. Pick better value with `binwidth`.

Warning: Removed 6 rows containing non-finite values (`stat_bin()`).



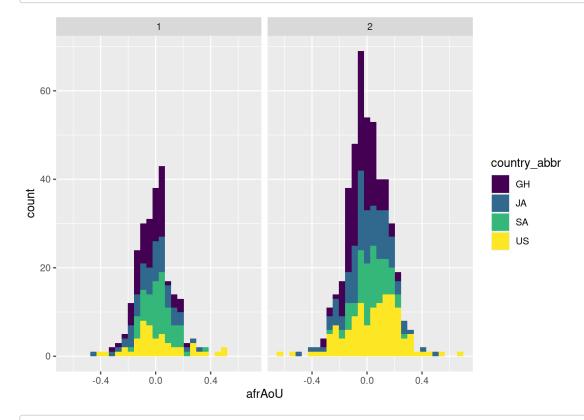
#allAoU PRS
ggplot(all_mets,aes(x=allAoU,fill=country_abbr)) + geom_histogram() + scale_fill_viridis_d() + facet_wrap(~s
ex)

`stat_bin()` using `bins = 30`. Pick better value with `binwidth`.



#afrAoU PRS
ggplot(all_mets,aes(x=afrAoU,fill=country_abbr)) + geom_histogram() + scale_fill_viridis_d() + facet_wrap(~s
ex)

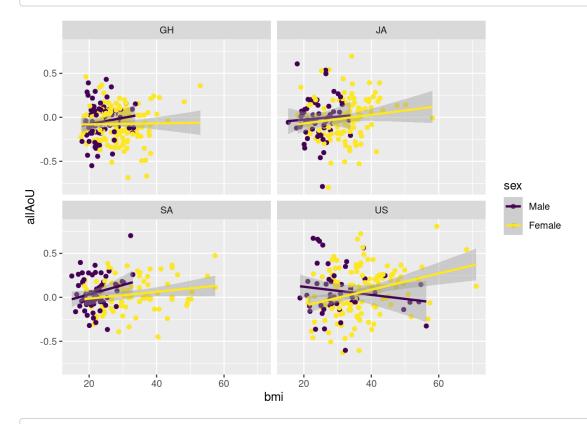




```
## `geom_smooth()` using formula = 'y \sim x'
```

Warning: Removed 6 rows containing non-finite values (`stat_smooth()`).

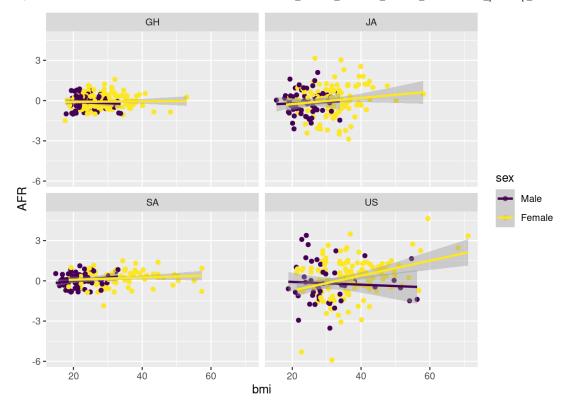
Warning: Removed 6 rows containing missing values (`geom_point()`).



ggplot(all_mets,aes(x=bmi,y=AFR,col=factor(sex,labels=c("Male","Female")))) + geom_point() + geom_smooth(met hod='lm') + facet_wrap(~country_abbr) + guides(col=guide_legend(title="sex"))+ scale_color_viridis_d()

```
## `geom_smooth()` using formula = 'y \sim x'
```

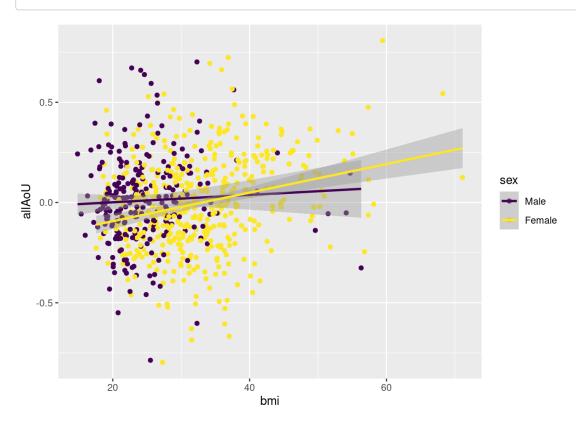
Warning: Removed 6 rows containing non-finite values (`stat_smooth()`).
Removed 6 rows containing missing values (`geom_point()`).



ggplot(all_mets,aes(x=bmi,y=allAoU,col=factor(sex,labels=c("Male","Female")))) + geom_point() + geom_smooth
(method='lm') + guides(col=guide_legend(title="sex"))+ scale_color_viridis_d()

```
## geom_smooth() using formula = y \sim x'
```

Warning: Removed 6 rows containing non-finite values (`stat_smooth()`).
Removed 6 rows containing missing values (`geom_point()`).



ggplot(all_mets,aes(x=bmi,y=afrAoU,col=factor(sex,labels=c("Male","Female")))) + geom_point() + geom_smooth
(method='lm') + guides(col=guide_legend(title="sex"))+ scale_color_viridis_d()

```
## geom_smooth() using formula = y \sim x'
```

```
## Warning: Removed 6 rows containing non-finite values (`stat_smooth()`).
## Removed 6 rows containing missing values (`geom_point()`).
```

