Phase 2 - HackBio Data Contest

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This document contains exploratory data analysis results and their codes for the task given for phase 2 of the HackBio Data Contest.

Decoding and Understanding the dataset

```
#Read the data
sample <- read.table("sample.tsv", header=TRUE)

# Gives us the basic statistical info. Based on the summary statistics, median,
# minimum, maximum and mean were found for all numerical columns especially
# for MAS, Sample Size and P value.

summary(sample)</pre>
```

```
##
       SNPID
                            RSID
                                                 CHR.
                                                                   POS
##
    Length: 25000
                        Length: 25000
                                                   : 1.000
                                                                          67365
    Class :character
                        Class :character
                                            1st Qu.: 4.000
                                                              1st Qu.: 31819538
##
    Mode :character
                        Mode :character
                                            Median : 8.000
                                                              Median: 71068010
##
                                                   : 8.571
                                                                     : 79495862
                                            Mean
                                                              Mean
##
                                            3rd Qu.:13.000
                                                              3rd Qu.:115694815
##
                                            Max.
                                                   :22.000
                                                              Max.
                                                                     :249222450
##
   EFFECT_ALLELE
                        OTHER_ALLELE
                                            EFFECT_ALLELE_FREQ
##
                                                                     BETA
   Length: 25000
                        Length: 25000
                                                   :0.0000
                                                                       :-1.53806
##
                                            Min.
                                                                Min.
                                            1st Qu.:0.0919
                                                                1st Qu.:-0.00539
##
   Class : character
                        Class :character
   Mode :character
                       Mode :character
                                            Median :0.2670
                                                                Median :-0.00005
##
                                                   :0.3389
                                                                      : 0.00030
                                            Mean
                                                                Mean
##
                                            3rd Qu.:0.5470
                                                                3rd Qu.: 0.00515
##
                                            Max.
                                                   :1.0000
                                                                Max.
                                                                       : 1.93485
##
                                                                NA's
                                                                       :194
##
          SE
                             P
                                               N
                                                              ANCESTRY
##
    Min.
           :0.00104
                       Min.
                              :0.0000
                                         Min.
                                                     482
                                                            Length: 25000
##
    1st Qu.:0.00358
                       1st Qu.:0.1163
                                         1st Qu.:
                                                   46408
                                                            Class : character
   Median :0.00654
                       Median :0.3729
                                         Median : 100692
                                                            Mode :character
##
##
    Mean
           :0.01802
                       Mean
                              :0.4087
                                         Mean
                                                : 374821
   3rd Qu.:0.00944
##
                       3rd Qu.:0.6742
                                         3rd Qu.: 264725
##
   Max.
           :1.07000
                              :0.9999
                                         Max.
                                                :1597374
                       Max.
   NA's
           :194
                       NA's
##
                              :194
```

```
# unique super populations present (Ancestrors)
print(superpopulations <- unique(sample$ANCESTRY))</pre>
```

```
## [1] "AFRICAN"
                                  "SOUTH_ASIA" "EAST_ASIA" "HISPANIC"
                    "EUROPEAN"
#find how many chromosomes are present
unique(sample$CHR)
                  4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22
   [1] 1 2 3
# find if NA values present in our sample
print(colSums(is.na(sample)))
##
                SNPID
                                    RSID
                                                         CHR
                                                                            POS
##
                                                           0
                                                                              0
                                        0
##
        EFFECT_ALLELE
                            OTHER_ALLELE EFFECT_ALLELE_FREQ
                                                                            BETA
##
                    0
                                       0
                                                           0
                                                                             194
                                       Ρ
##
                   SE
                                                           N
                                                                       ANCESTRY
##
                  194
                                      194
                                                           0
                                                                              0
```

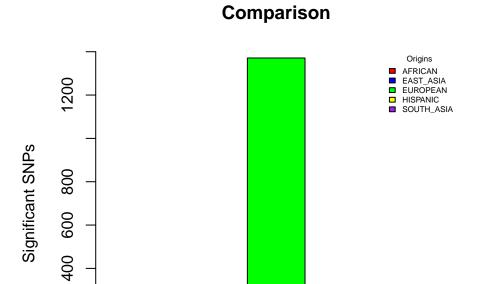
Number of significant SNPs in the total population and even across individual populations

For answering the first question given in the task, "How many SNPs are significant (p-value < 0.01) for variability in height (MAF > 0.01) in all the super populations", the following code and plots conveys the answer.

```
data_color <- c("red", "blue", "green", "yellow", "purple")</pre>
# Filter data- p-value < 0.01 and MAF > 0.01 for each super population
significant_snps <- data.frame()</pre>
for (sp in superpopulations) {
  subset_data <- sample[sample$ANCESTRY == sp, ]</pre>
  # Filter SNPs based on p-value and MAF
  filtered_snps <- subset_data[subset_data$ P < 0.01 &</pre>
                                   subset data$EFFECT ALLELE FREQ > 0.01, ]
  # merge by rows
  significant_snps <- rbind(significant_snps, filtered_snps)</pre>
}
# Count the number of significant SNPs in all super populations
num_of_significant_snps <- nrow(significant_snps)</pre>
num_of_snps <- nrow(sample)</pre>
#or
length(significant_snps$SNPID)
```

[1] 2253

```
#Answer for the 1st question given:
cat("The number of SNPs that are significant (p-value < 0.01) for</pre>
   variability in height (MAF > 0.01) in all the super populations are "
    ,num_of_significant_snps, "\n")
## The number of SNPs that are significant (p-value < 0.01) for
       variability in height (MAF > 0.01) in all the super populations are 2253
cat("Total number of significant SNPs vs Normal: ", num_of_significant_snps, ":", num_of_snps, "\n")
## Total number of significant SNPs vs Normal: 2253 : 25000
# No. of significant SNPs per population:
print( significant_snps_per_super_population <- table(significant_snps$ANCESTRY))</pre>
##
##
     AFRICAN EAST_ASIA EUROPEAN HISPANIC SOUTH_ASIA
                                                       89
##
          294
                     300
                               1371
                                           199
#visual representation of the above data
pop<- c("AFRICAN" , "EAST_ASIA", "EUROPEAN" , "HISPANIC", "SOUTH_ASIA")</pre>
par(mfrow = c (1,1))
barplot(significant_snps_per_super_population, col = c("red", "blue", "green", "yellow", "purple"),
        xlab = "Origins of Populations",
       ylab = "Significant SNPs", xaxt = "n", ylim = c(0,1400),
       main = "Comparison")
legend("topright", legend = pop , fill = c("red", "blue", "green", "yellow", "purple"),
       title = "Origins", xjust = 1, yjust = 0, bty = "n", cex = 0.5)
```



Origins of Populations

Answer - The total number of significant SNPs are 2253 with Europe having the highest - 1371, and South Asia having the lowest - 89.

Question 2

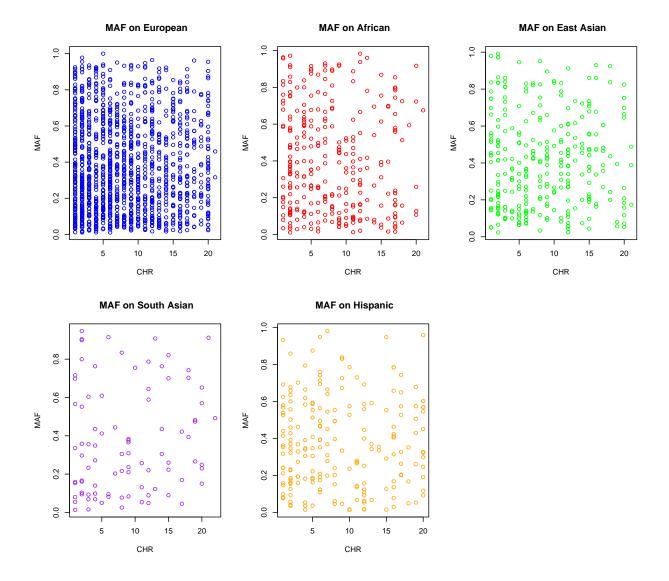
200

Europeans genetic variability can/cannot be found in other super populations

```
#better understanding of the new subset of data
summary(significant_snps)
```

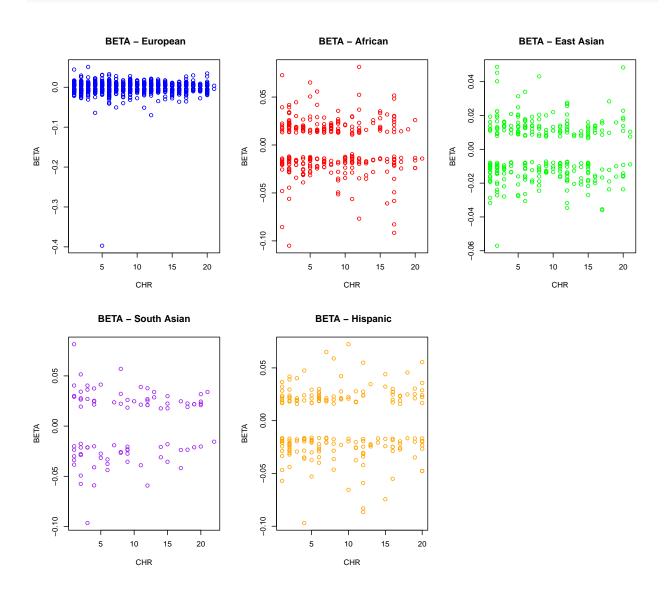
##	SNPID	RSID	CHR	POS
##	Length: 2253	Length: 2253	Min. : 1.000	Min. : 123233
##	Class :character	Class :character	1st Qu.: 3.000	1st Qu.: 34383788
##	Mode :character	Mode :character	Median : 7.000	Median : 70051966
##			Mean : 8.295	Mean : 79655840
##			3rd Qu.:12.000	3rd Qu.:114226535
##			Max. :22.000	Max. :247010734
##	EFFECT_ALLELE	OTHER_ALLELE	EFFECT_ALLELE_FR	EQ BETA
##	Length: 2253	Length:2253	Min. :0.0105	Min. :-0.397104

```
## Class :character
                       Class : character
                                          1st Qu.:0.1590
                                                              1st Qu.:-0.010464
##
   Mode : character Mode : character
                                          Median :0.3350
                                                              Median : -0.003257
##
                                          Mean
                                                 :0.3799
                                                              Mean :-0.001079
##
                                                              3rd Qu.: 0.009352
                                           3rd Qu.:0.5720
##
                                          Max.
                                                 :1.0000
                                                              Max. : 0.081540
##
          SE
                             Р
                                                 N
                                                                ANCESTRY
           :0.001040
                              :0.000e+00
                                                 : 21912
                                                              Length: 2253
## Min.
                       Min.
                                           Min.
                                           1st Qu.: 107061
## 1st Qu.:0.001170
                      1st Qu.:1.210e-07
                                                              Class : character
## Median :0.001790
                       Median :2.433e-04
                                           Median :1573790
                                                              Mode : character
                                                 :1007854
## Mean
          :0.003465
                       Mean
                              :1.840e-03
                                           Mean
## 3rd Qu.:0.004810
                       3rd Qu.:2.841e-03
                                           3rd Qu.:1593869
## Max. :0.137000
                       Max.
                              :9.997e-03
                                           Max.
                                                  :1597373
#subsetting based on populations in the unfiltered dataset
african <- subset(sample, ANCESTRY == "AFRICAN")</pre>
european <- subset(sample, ANCESTRY == "EUROPEAN")</pre>
south_asia <- subset(sample, ANCESTRY == "SOUTH_ASIA")</pre>
east_asia <- subset(sample, ANCESTRY == "EAST_ASIA")</pre>
hispanic <- subset(sample, ANCESTRY == "HISPANIC")
#subsetting based on populations in the filtered dataset
sign_snps_african <- subset(significant_snps, ANCESTRY == "AFRICAN")</pre>
sign_snps_european <- subset(significant_snps, ANCESTRY == "EUROPEAN")</pre>
sign_snps_south_asia <- subset(significant_snps, ANCESTRY == "SOUTH_ASIA")</pre>
sign snps east asia <- subset(significant snps, ANCESTRY == "EAST ASIA")
sign_snps_hispanic <- subset(significant_snps, ANCESTRY == "HISPANIC")</pre>
# remove europe alone
others sign snp <- significant snps [significant snps $ANCESTRY != "EUROPEAN", ]
# since european pop has highest number of filtered significant data (in human height),
# these plots shows how much MAF affects in all chromosomes across all the given populations.
par(mfrow = c (2,3))
plot( sign_snps_european$CHR, sign_snps_european$EFFECT_ALLELE_FREQ, type = "p",
      col = "blue", xlab = "CHR", ylab = "MAF", main = " MAF on European")
plot( sign_snps_african$CHR, sign_snps_african$EFFECT_ALLELE_FREQ, type = "p",
      col = "red", xlab = "CHR", ylab = "MAF", main = " MAF on African")
plot( sign_snps_east_asia$CHR, sign_snps_east_asia$EFFECT_ALLELE_FREQ, type = "p",
      col = "green",xlab = "CHR", ylab = "MAF", main = " MAF on East Asian")
plot( sign_snps_south_asia$CHR, sign_snps_south_asia$EFFECT_ALLELE_FREQ, type = "p",
      col = "purple", xlab = "CHR", ylab = "MAF", main = " MAF on South Asian")
plot( sign snps hispanic$CHR, sign snps hispanic$EFFECT ALLELE FREQ, type = "p",
      col = "orange", xlab = "CHR", ylab = "MAF", main = " MAF on Hispanic")
```



The above graph depicts that European population have the highest spread of MAFs and South Asian populations have the lowest spread of MAFS across all the Chromosomes, majority surpassing the MAF median value of 0.3350.

Relationship between BETA values and Chromosomes across all populations



Performing statistical analysis:

Null hypothesis - Europeans' genetic variability can be found in other super populations

- no significant difference

Alternative hypothesis - Europeans' genetic variability cannot be found in other super populations

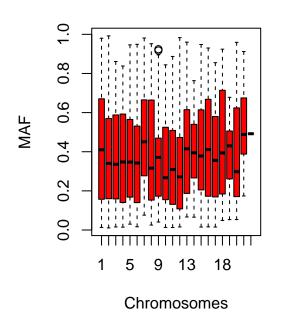
- significant difference

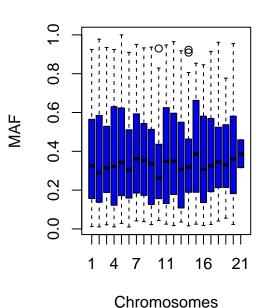
```
#Performing t test
# chosen alpha value 5%, i.e) 0.05%
t.test(sign_snps_european$EFFECT_ALLELE_FREQ, others_sign_snp$EFFECT_ALLELE_FREQ,
      paired = F, var.equal =F )
##
##
   Welch Two Sample t-test
## data: sign_snps_european$EFFECT_ALLELE_FREQ and others_sign_snp$EFFECT_ALLELE_FREQ
## t = -1.7762, df = 1826.2, p-value = 0.07587
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -0.04149409 0.00205483
## sample estimates:
## mean of x mean of y
## 0.3721338 0.3918534
t.test(sign_snps_european$BETA, others_sign_snp$BETA, paired = F, var.equal =F)
##
## Welch Two Sample t-test
## data: sign_snps_european$BETA and others_sign_snp$BETA
## t = 0.74064, df = 1263.5, p-value = 0.4591
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -0.001171384 0.002592225
## sample estimates:
      mean of x
                    mean of y
## -0.0008011157 -0.0015115366
#BETA divided by SE - Effect Size
ES_E <- sign_snps_european$BETA / sign_snps_european$SE
ES_O <- others_sign_snp$BETA / others_sign_snp$SE
t.test(ES_E, ES_O, paired = F, var.equal =F)
##
## Welch Two Sample t-test
```

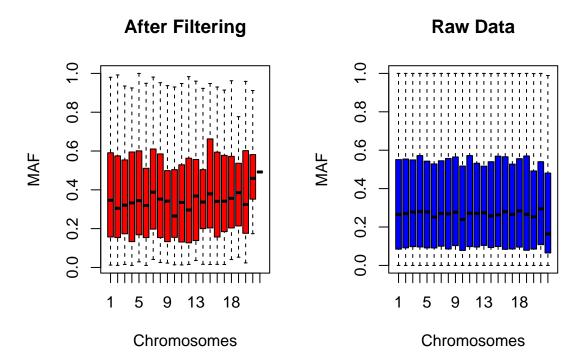
```
##
## data: ES_E and ES_O
## t = -0.94908, df = 2168, p-value = 0.3427
\#\# alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -0.6825874 0.2373659
## sample estimates:
## mean of x mean of y
## -0.3766562 -0.1540455
#in all the above t tests, p-values are greater than alpha, which results
# in the acceptance of the null hypothesis indicating that
# Europeans' genetic variability can be found in other super populations
#t- values are also lesser than 1 in both, indicating difference between
# them is not statistically significant, hence no significant
# difference between the samples.
# ANOVA Tests
aov(BETA ~ ANCESTRY, data = significant_snps)
## Call:
##
     aov(formula = BETA ~ ANCESTRY, data = significant_snps)
##
## Terms:
##
                    ANCESTRY Residuals
## Sum of Squares 0.0014951 0.8932062
## Deg. of Freedom
##
## Residual standard error: 0.01993323
## Estimated effects may be unbalanced
ANOVA_BETA <- aov(BETA ~ ANCESTRY, data = significant_snps)
summary(ANOVA_BETA)
                 Df Sum Sq Mean Sq F value Pr(>F)
                  4 0.0015 0.0003738
## ANCESTRY
                                      0.941 0.439
              2248 0.8932 0.0003973
## Residuals
aov(EFFECT_ALLELE_FREQ ~ ANCESTRY, data = significant_snps)
## Call:
##
      aov(formula = EFFECT_ALLELE_FREQ ~ ANCESTRY, data = significant_snps)
##
## Terms:
                    ANCESTRY Residuals
##
## Sum of Squares
                     0.36848 146.27928
## Deg. of Freedom
                           4
## Residual standard error: 0.2550899
## Estimated effects may be unbalanced
```

Other than Europe

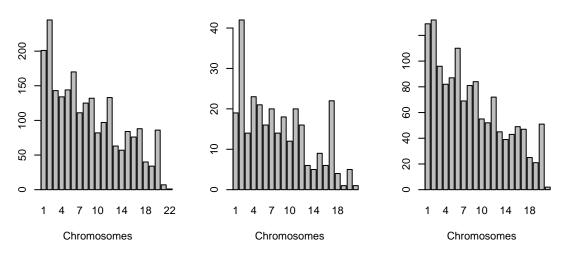
Only Europe



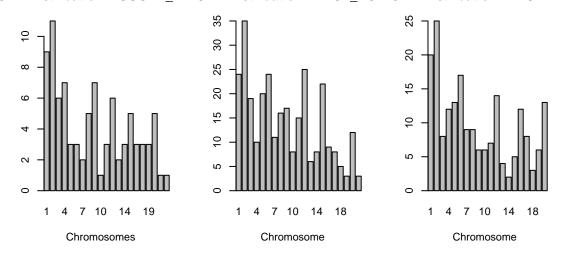




CHR Distribution - Overall CHR Distribution - AFRICA! CHR Distribution - EUROPE!



CHR Distribution - SOUTH_A: CHR Distribution - EAST_AS CHR Distribution - HISPANIC



The plot generated shows that maximum number of data distribution is in the 2nd chromosome. The total range in y-axis is 200, out of which 100 is by European Population.

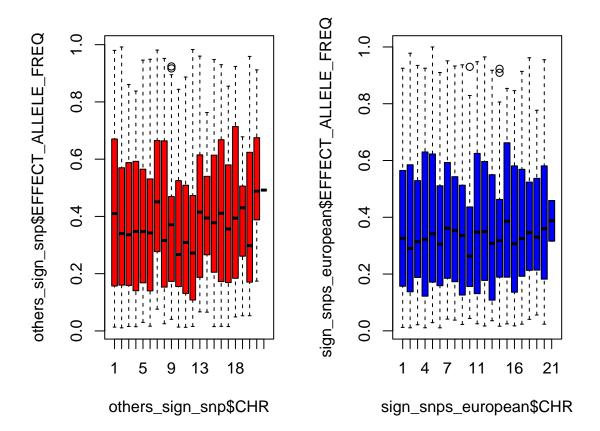
Higher value of MAF is related to Genetic Variability the following code proves that European population has more Minor allele frequency in their data having SNPs with MAFs value greater than the median value

```
#get the median value from here
summary(sign_snps_european$EFFECT_ALLELE_FREQ)
##
     Min. 1st Qu. Median
                            Mean 3rd Qu.
                                            Max.
## 0.0105 0.1580 0.3220 0.3721 0.5635 1.0000
#MAF greater than median in the european population
print( MAF_E <- table(sign_snps_european$EFFECT_ALLELE_FREQ > 0.2670 ))
##
## FALSE TRUE
    573
          798
#get the median value from here
summary(others_sign_snp$EFFECT_ALLELE_FREQ)
##
     Min. 1st Qu. Median
                            Mean 3rd Qu.
                                            Max.
   #how many MAF greater than median in the european population
print( MAF_Others <- table(others_sign_snp$EFFECT_ALLELE_FREQ > 0.3560 ))
##
## FALSE TRUE
    442
total_MAF_SP <- length(others_sign_snp$EFFECT_ALLELE_FREQ)</pre>
total_MAF_E <- length(sign_snps_european$EFFECT_ALLELE_FREQ)</pre>
```

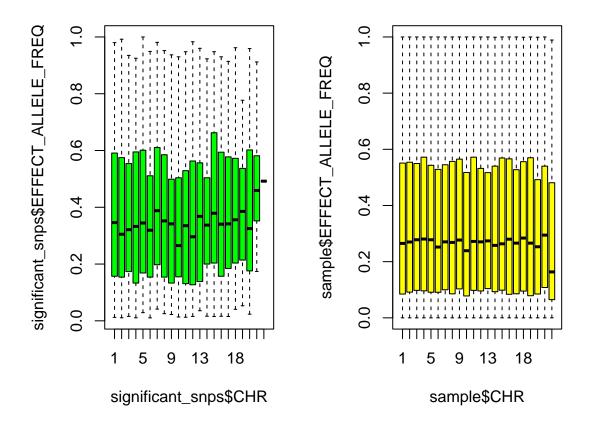
MAFs value greater than thier median value were found to be

higher in the European population

```
# since european pop has highest number of MAF, this shows how much they are significant
par(mfrow = c (1,2))
boxplot(others_sign_snp$EFFECT_ALLELE_FREQ ~ others_sign_snp$CHR , col = "red")
boxplot(sign_snps_european$EFFECT_ALLELE_FREQ ~ sign_snps_european$CHR , col = "blue")
```



```
# normal data MAF vs significant data MAF
par(mfrow = c (1,2))
boxplot(significant_snps$EFFECT_ALLELE_FREQ ~ significant_snps$CHR , col = "green")
boxplot(sample$EFFECT_ALLELE_FREQ ~ sample$CHR , col = "yellow")
```



understanding both unfiltered and filtered datasets:

1st Qu.

31819538

Min.

67365

Median

##

##

```
#Unfiltered: Original (sample)
#to understand the data (mean, min, max, median)
summary(sample$N)
##
                    Median
                               Mean 3rd Qu.
##
                    100692
                                     264725 1597374
             46408
                             374821
summary(sample$P)
##
                               Mean 3rd Qu.
                                                        NA's
      Min. 1st Qu.
                    Median
                                                Max.
    0.0000 0.1163
                    0.3729
                             0.4087
                                     0.6742
                                             0.9999
                                                         194
summary(sample$POS)
```

3rd Qu.

Max.

Mean

71068010 79495862 115694815 249222450

```
summary(sample$BETA)
##
                      Median
       Min. 1st Qu.
                                  Mean 3rd Qu.
                                                    Max.
                                                             NA's
## -1.53806 -0.00539 -0.00005 0.00030 0.00515 1.93485
                                                              194
summary(sample$SE)
      Min. 1st Qu. Median
                             Mean 3rd Qu.
                                                      NA's
                                              Max.
## 0.00104 0.00358 0.00654 0.01802 0.00944 1.07000
                                                       194
#to get the count of each
table(sample$EFFECT_ALLELE)
##
##
           С
      Α
                G
## 6410 6245 6113 6232
table(sample$OTHER_ALLELE)
##
      Α
## 6004 6438 6471 6087
table(sample$CHR)
##
                               6
                                    7
##
                3
                          5
                                      8
                                           9
                                                  10
                                                       11
                                                            12
                                                                 13
                                                                     14
                                                                           15
                                                                                16
## 2084 2185 1632 1630 1407 1675 1384 1470 1330 1455 1230 1232 953 893 746 870
          18
              19
                    20
                         21
                              22
     17
        657
##
   687
             447 745
                        206
                              82
table(sample$ANCESTRY)
##
##
      AFRICAN EAST_ASIA
                           EUROPEAN
                                      HISPANIC SOUTH_ASIA
##
         5000
                    5000
                               5000
                                          5000
                                                     5000
Sample Size Distribution
```

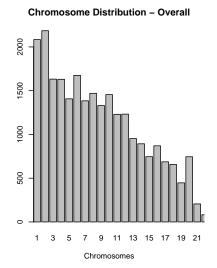
```
# sample size distribution among different origins
par(mfrow = c (2,3))
boxplot(sample$N, notch = TRUE, main = "Plot for sample size- overall")
boxplot(african$N, notch = TRUE, main = "Plot for sample size in Africa")

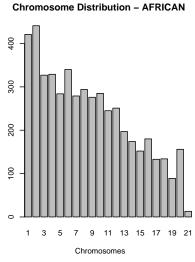
## Warning in (function (z, notch = FALSE, width = NULL, varwidth = FALSE, : some
## notches went outside hinges ('box'): maybe set notch=FALSE
```

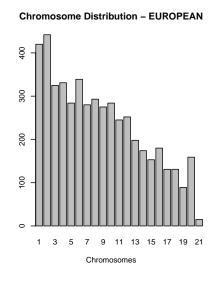
```
boxplot(european$N, notch = TRUE, main = "Plot for sample size in Europe")
boxplot(south_asia$N, notch = TRUE, main = "Plot for sample size in South asia")
## Warning in (function (z, notch = FALSE, width = NULL, varwidth = FALSE, : some
## notches went outside hinges ('box'): maybe set notch=FALSE
boxplot(east_asia$N, notch = TRUE, main = "Plot for sample size in East asia")
boxplot(hispanic$N,notch = TRUE, main = "Plot for sample size in Hispanic")
## Warning in (function (z, notch = FALSE, width = NULL, varwidth = FALSE, : some
## notches went outside hinges ('box'): maybe set notch=FALSE
        Plot for sample size- overall
                                          Plot for sample size in Africa
                                                                             Plot for sample size in Europe
   1000000
                                                                         000000
                                                                         500000
   500000
     Plot for sample size in South asia
                                         Plot for sample size in East asia
                                                                            Plot for sample size in Hispanic
                                                                         50000
   20000
   40000
   30000
   20000
                                      1e+05
```

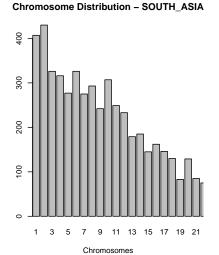
Chromosome distribution in the raw data

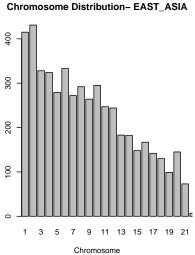
10000

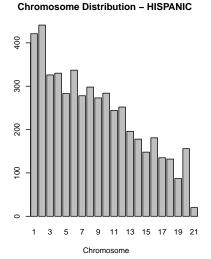






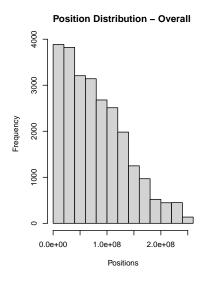


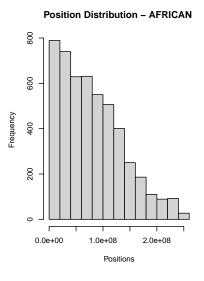


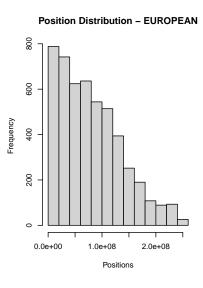


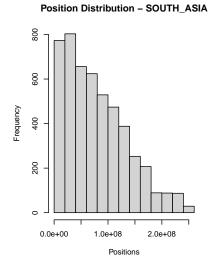
Position distribution

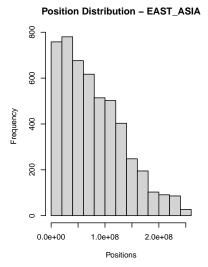
```
# Position distribution across different origins
par(mfrow = c (2,3))
hist(sample$POS, main = "Position Distribution - Overall ", xlab = "Positions")
hist(african$POS, main = "Position Distribution - AFRICAN ", xlab = "Positions")
hist(european$POS, main = "Position Distribution - EUROPEAN", xlab = "Positions")
hist(south_asia$POS, main = "Position Distribution - SOUTH_ASIA", xlab = "Positions")
hist(east_asia$POS, main = "Position Distribution - EAST_ASIA", xlab = "Positions")
hist(hispanic$POS, main = "Position Distribution - HISPANIC ", xlab = "Positions")
```

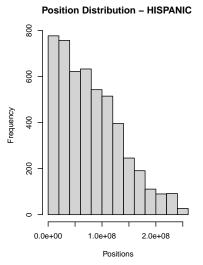












the below code results in the creation of dataframe containing MAFs value greater than 0.95, hence indicating that the following SNPs are highly responsible in Genetic Variation (more common and has been present in the population for a longer time).

```
imp <- subset(significant_snps ,significant_snps$EFFECT_ALLELE_FREQ > 0.95 )
summary(imp)
```

```
##
       SNPID
                            RSID
                                                 CHR
                                                                  POS
                        Length:19
##
    Length: 19
                                                   : 1.000
                                                                        3737495
                                           Min.
                                                             Min.
##
    Class :character
                        Class :character
                                           1st Qu.: 2.000
                                                             1st Qu.: 40990906
##
   Mode :character
                       Mode :character
                                           Median : 5.000
                                                             Median: 64692041
##
                                           Mean
                                                   : 7.211
                                                             Mean
                                                                     : 92057998
##
                                            3rd Qu.:12.000
                                                             3rd Qu.:135323988
                                                   :20.000
##
                                           Max.
                                                             Max.
                                                                     :227503125
##
    EFFECT_ALLELE
                        OTHER_ALLELE
                                           EFFECT_ALLELE_FREQ
                                                                     BETA
                                                   :0.9510
                                                                       :-0.39710
##
   Length:19
                        Length:19
                                           Min.
                                                               Min.
##
    Class : character
                        Class :character
                                           1st Qu.:0.9585
                                                               1st Qu.:-0.03348
##
    Mode :character
                       Mode :character
                                           Median :0.9650
                                                               Median: 0.00745
##
                                           Mean
                                                   :0.9691
                                                               Mean
                                                                       :-0.01870
##
                                           3rd Qu.:0.9790
                                                               3rd Qu.: 0.02245
##
                                                   :1.0000
                                                                       : 0.06510
                                           Max.
                                                               Max.
##
                             Ρ
          SE
                                                                 ANCESTRY
                                                  N
                              :4.100e-08
##
   Min.
           :0.00255
                      Min.
                                           Min.
                                                   : 58371
                                                              Length: 19
##
    1st Qu.:0.00321
                      1st Qu.:2.221e-03
                                           1st Qu.: 104293
                                                              Class : character
##
    Median :0.00983
                      Median :3.520e-03
                                           Median: 262846
                                                              Mode : character
##
   Mean
           :0.01638
                              :3.897e-03
                                                   : 595950
                      Mean
                                           Mean
##
    3rd Qu.:0.01475
                      3rd Qu.:6.866e-03
                                            3rd Qu.:1451252
           :0.13700
                              :8.413e-03
                                                   :1597370
##
    Max.
                      Max.
                                           Max.
# a unique dataframe comtaining MAF values greater than 0.95
MAF_above_0.95 <- data.frame( Origin = imp$ANCESTRY, MAF = imp$EFFECT_ALLELE_FREQ,
imp = imp$SNPID , RSID = imp$RSID, BETA = imp$BETA, sample_size = imp$N )
```

Credits goes to HackBio for the datasets and organizing the data contest

Citation: Yengo, L., Vedantam, S., Marouli, E. et al. A saturated map of
common genetic variants associated with human height. Nature 610, 704–712 (2022).

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Thank you HackBio for the opportunity