Quality control report for kgpexample

H3Agwas QC Pipeline

Mon Jul 11 23:11:20 SAST 2022

1 Introduction

The input file for this analysis was array.orig. {bed,bim,fam}. This data includes:

- 50139 SNPs
- 550 participants

The input files and md5 sums were

array.bed a02ac76feb9712c2ba1b42f8be09d469 array.bim 4ce916cd20501cd825cb3e57f688f9a1 array.fam 9ace90f12d8b222218b42ecf0f4e51e0

Note that some statistics are shown twice – on the raw input data and on the final result, since these statistics are needed or different purposed.

Approach

The pipeline takes an incremental approach to QC, trading extra computation time in order to achieve high quality while removing as few data as possible. Rather than applying all cut-offs at once, we incrementally apply cutoffs (for example, removing really badly genotyped SNPs before checking for heterozygosity will result in fewer individuals failing heterozygosity checks).

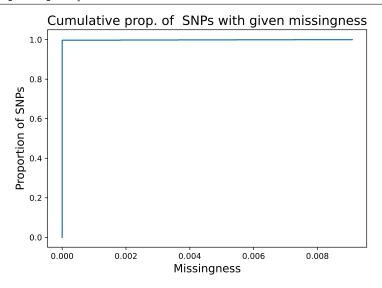
2 QC Phase 0

This phase only removes SNPs which are duplicated (based on SNP name). No other QC is done and so the output of this phase should really be considered as raw data.

- 1. There were 0 duplicate SNPs. The file with them (if any) is called array.dups. Note that duplicate SNPs are determined by the names of the SNPs. SNPs which appear at the same position are probably duplicates but may not be. You can control whether you want to detect these using the parameter remove_on_bp. It is crucial to examine this file to avoid inadvertently removing SNPs. On some chips there are duplicate SNPs at a position you should select what you what want.
- 2. 60 individuals had discordant sex information the full PLINK report can be found in array-nd. sexcheck and an extract of the PLINK report showing only the failed reports can be found can be found in array-nd.badsex, and a more detailed analysis can be found in Section 5.

Figure 1 shows the spread of missingness per SNP across the sample, whereas Figure 2 shows the spread of missingness per individual across the sample. Note that this shows missingness before any filtering or cleaning up of the data.

Figure 1 SNP missingness: For each level of missingness specified on the x axis, the corresponding y-value shows the proportion of SNPs which have missingness less than this. [File is $\{array-nd-snpmiss_plot.pdf\}$]



Minor allele frequency. Table 1 on page 2 shows the minor allele frequency spectrum for the raw data. The number of monomorphic SNPs is shown in the first row. Note that some of the MAFs with very low MAF are actually monomorphic, with the polymorphisms due to genotyping error. Figure 3 on page 4 shows the cumulative distribution of MAF. This can be used to determine an appropriate MAF cut-off.

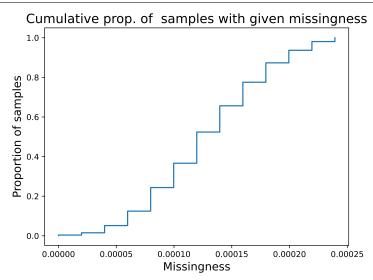
Note that the *minor* allele is determined with respect to the frequency spectrum in this data – 'minor' is not synonym for alternate or non-reference allele, or the allele that has minor frequency in some other data set. Under this definition the MAF is always ≤ 0.5 .

Table 1 Minor Allele Frequency spectrum of the raw data. The number of apparently monommorphic SNPs is shown in the row labelled 0; the other rows show the number of SNPs in the bins shown.

	Num SNPs
MAF bin	
0	167
(0.0,0.005]	1740
(0.005,0.01]	2389
(0.01,0.02]	4267
(0.02,0.03]	2905
(0.03,0.04]	1955
(0.04,0.05]	1590
(0.05,0.1]	6133
(0.1,0.15]	5003
(0.15,0.2]	4303
(0.2,0.25]	3854
(0.25,0.3]	3551
(0.3, 0.4]	6351
(0.4, 0.5]	5931

Hardy Weinberg Statistics: Figure 4 shows the cumulative distribution of Hardy-Weinberg p-value for the SNPs in the raw data. This can be used to assess the cost of excluding SNPs with a

Figure 2 Missingness per indvididual: For each level of missingness specified on the x axis, the corresponding y-value shows the proportion of individuals which have missingness less than this. [File is {array-nd-indmiss_plot.pdf}]



particular p-cutoff. We expect the curve the fit tightly to the main diagonal, except for a very small p values (and this deviation may not be observable on a linear plot).

The QQ plot for the HWE scores can be found in Figure 5. The region of deviation from the line of expected versus observed p-values will be more observable here. Note that if there are very small observed p-values in relation to expected values, the expected curve may be very flat — pay attention to the x and y axis coordinates. Since we are plotting on a negative log-scale, note that regions of low probability of deviation from HWE (p-value close to 1) are at the left, and regions of high probability (low p-value) are at the right. The tail of the plot where deviation from the diagonal occurs is likely to be a good cut-off to use for QC.

However, care needs to be taken not exclude SNPs. We are using HWE p-value as a proxy for something having gone wrong with the sample or genotyping, and this is a little crude. In a study with participants from different population groups in a recently admixed group, deviation from HWE is expected and does not indicate problems with QC. Moreover, in a disease study, it is likely that those individuals that are affected, those SNPs that are associated with the condition under study will not in be in HWE. Care needs to be taken – it is easier to handle in a pure case/control study. In a population cross-section study with different conditions being considered, it might be advisable to re-run the QC pipeline for HWE for each study. The current version of the pipeline does not support his more complex analysis, though we plan to extend.

Figure 3 Cumulative frequency of SNPs. For a frequency shown on the x-axis, the corresponding y-value shows the proportion of SNPs with frequency at least this frequency; that is, it shows the proportion of SNPs which will remain if the MAF filter of this x-value is chosen.

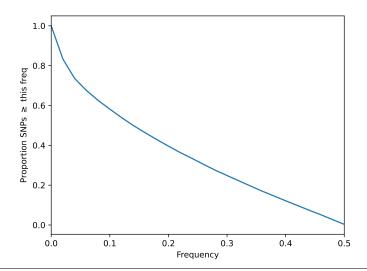


Figure 4 HWE distribution. For an HWE-value shown on the x-axis, the corresponding y-value shows the proportion of SNPs with HWE p-value at least this frequency; that is, it shows the proportion of SNPs which will be removed if the HWE filter of this x-value is chosen. File is array-nd-inithwe.pdf.

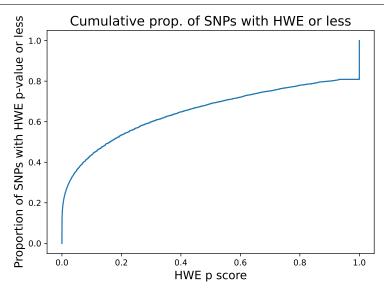
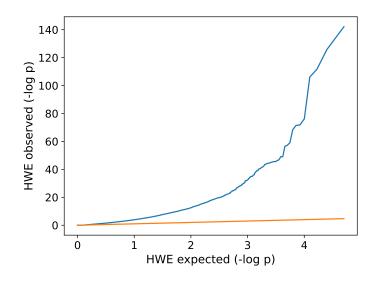


Figure 5 QQ plot for Hardy-Weinberg scores. The graphic can be found in the file array-nd-inithwe-qq.pdf



3 Plate Analysis

No plate analysis can be done because a 10% GenCall score can't be found in the sample sheet or no sample sheet given.

4 QC Phase 1

The details of the final filtering can be found in the Nextflow script. Note that the exact ordering or removal will affect the final results. However, we take a conservative approach.

- 1. Only autosomal SNPs are included
- 2. SNPs and individuals that have been very poorly genotyped (missingness exceeding 10 per cent) are removed.
- 3. Individuals with missingness greater than 0.02 are removed (by filtering out very badly genotyped individuals or SNPs) in the previous steps we may save a few individuals in this step;
- 4. SNP with missingness greater than 0.01 are removed;
- 5. minor allele frequency less than 0.01 (and greater than 1-0.01) are removed;
- 6. HWE adjusted p-value less than 0.008 are removed

Using this approach,

- 0 SNPs that are non-autosomal were removed;
- 0 SNPs were removed due missing genotype threshold constraints;
- 0 individuals were removed due to missing genotype constraints (the list of missing individuals, if any, can be found in the file array-nd-c.irem);
- 0 SNPs were removed as the MAF was too low.
- 10867 SNPs were removed as they were out of the specified Hardy-Weinberg equilibrium.

5 Batch report

The batch quality results shown here (missingness and sex-check failures) is based on the raw input data (other than duplicate SNPs being deleted) as this gives insight into the quality of the raw data differences between batches and sub-batches. The PC analysis is based on the refined data.

5.1 Overall missingness and sex anomaly report

Table 2 on page 6 shows the error rate as shown by phenoqc_ql as found in file pheno.phe.

Table 2 For each group shown, we have: the average percentage missingness in that group (100×100) the sum the number of missing calls over all individuals divided by the total number of genotype calls over all individuals); the percentage of samples in that group with a genotyping error rate above 4 percent; and the percentage of samples in that group that fail the sex check using the standard PLINK parameters on the raw input data.

 phenoqc_ql	Num samples	Missing rate	\% poor	Sex Checkfail
1	493	0.01	0.00	10.55
2	57	0.01	0.00	12.28

Table 3 on page 7 shows the error rate as shown by batch as found in file batch.phe.

5.2 Detailed Sex Check Analysis

This section shows a detailed analysis of sex check anomalies and/or unusual patterns in the X-chromosome. The purpose of this analysis is to help identify trends between sub-groups, as well as possible labelling and sample handling errors. The term *anomaly* or *error* is used to label individuals where the sex of the individual as described in the manifest does not *stricly* match analysis of the X-chromosome and so for QC should be considered further.

In this analysis, we use PLINK to analyse the non-recombining regions of the X-chromosome, and in particular its computation of the inbreeding co-efficient of the X-chromosome. If the F statistic is greater than 0.8, PLINK infers that the sample is male; if it is less than 0.2, it infers that the sample is female.

Reminder: the checking of sex on the raw data before any other QC is shown in Section 1. The rest of this section analyses the data after basic QC on genotype has been done and so differences may be seen

There are two types of apparent anomaly that can happen. Soft anomalies are those cases where an individual is slightly above or below the stated threshholds. These may not be sample handling errors – since the F cut-off values are arbitrary, a too strict F-value may be chosen, or there may be uncommon patterns within the individuals studied. How these samples should be treated will require some thought, but these are not a sign of problems of the experimental protocol, and not by itself probably a sign of problems with genotyping or DNA quality errors. We define a soft anomaly as an indvidual having an F value between 0.2 and 0.8, which is possible but unusual.

Hard errors are cases where the sex in the manifest/fam file is markedly different from what the F-statistic predicts (e.g., the F-statistic says 0.996 and we have this as a female). While there may be very unusual cases were this occurs, many such examples in the data are likely to be a sign of sample handling errors. Great care needs to be taken with this.

A summary of the detailed analysis is shown below. In the output, the file array-nd-c-prune_missing_and_sexcheck.csv is a CSV file that has sample ID, per-individual missingness rate in the raw data (F_MISS is the individual missingness rate not the F-statistic), the phenoqcql status, and whether that sample is hard (H) or soft error (S) for given tolerated per-individual genotyping error rates on the X-chromosome. A '-' indicates that the individual is filtered out at that rate of missingness This can be used to assess whether anomalous sex results are due to poor genotyping rates in individuals.

Table 3 For each group shown, we have: the average percentage missingness in that group ($100 \times 100 \times 100$) the sum the number of missing calls over all individuals divided by the total number of genotype calls over all individuals); the percentage of samples in that group with a genotyping error rate above 4 percent; and the percentage of samples in that group that fail the sex check using the standard PLINK parameters on the raw input data.

batch	Num samples	Missing rate	\% poor	Sex Checkfail
1	175	0.01	0.00	10.86
2	189	0.01	0.00	10.05
3	186	0.01	0.00	11.29

Table 4 of anomalous sex calls. The results are shown for different values *mind* the maximum persample error rate tolerated in the X-chromosome (PLINK parameter). mind=1 means all SNPs included. *Tot* shows the number of individuals included with the given mind value. HErr is the number of hard errors. SAnm is the number of soft apparent anomalies or unusual results.

phenoqc ql	mind=1		mind=0.01		mind=0.03		mind=0.05					
	Tot S	Anm I	HErr	Tot S	Anm I	HErr	Tot S	Anm I	HErr	Tot S	Anm I	HErr
overall	550	27	32	550	27	32	550	27	32	550	27	32
1	493	24	28	493	24	28	493	24	28	493	24	28
2	57	3	4	57	3	4	57	3	4	57	3	4

5.3 Relatedness 5 BATCH REPORT

5.3 Relatedness

Using PLINK, relatedness is computed on using IBD with $\hat{\pi}$ as a proxy. The data used for this analysis was the result of the QC Phase 1 work. The $\hat{\pi}$ of 0.11 is a parameter of the pipeline.

All pairs of individuals with a $\hat{\pi} \geq 0.11$ are examined – we try to remove as few individuals as possible by removing first those who are related to multiple people (e.g. if A is a cousin of B and C, B and C may not be related so it makes sense to remove A rather than B and C).

- There were 15736 pairs of individuals over the cut-off. These can be found in the PLINK report array-nd-c.genome.
- 352 individuals were removed because of relatedness. The list of such individuals can be found in the file array-nd-fail_IBD.txt. (One individual in each pair is removed; any individuals in a pair with relatedness strictly greater than 0.7 are removed.)
- The individuals with $\hat{\pi} \geq 0.45$ can be found in array-nd-c-prune-reltable.csv.

The overall relationship analysis is shown in the Table 5 on page 9. For each group (or if there is only one group, the group as a whole), we show the group and the number of pairs of related individuals. We also show the number of pairs with $\hat{\pi} > 0.95$, which prima facie indicates identical individuals or twins, and the number of pairs of individuals with $\hat{\pi}$ between 0.45 and 0.95 which prima facie indicates relationship of parent/child or siblingship.

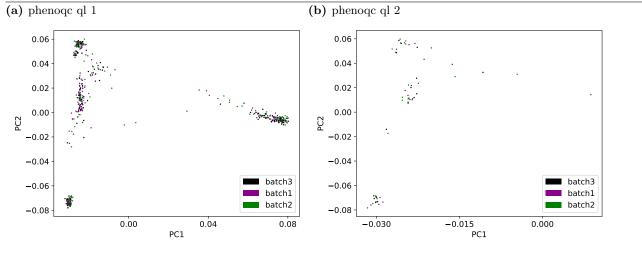
Table 5 Overall breakdown of $\hat{\pi}$ anomalies: for the overall dataset and each sub-group we show the number of total number of pairs with $\hat{\pi} > 0.11$, the number of pairs with $\hat{\pi} > 0.95$ (twins or sample duplication perhaps?), and the number of pairs with $0.95 > \hat{\pi} > 0.45$ (siblings or parent/child?).

phenoqc_ql	Num pairs	$\widehat{\pi} > 0.95$	$\hat{\pi} \in [0.45, 0.95]$	< 0.45
ALL	15736	0	0	15736
1	11671	0	0	11671
2	290	0	0	290
mixed	3775	0	0	3775

5.4 Principal component anlysis

In the PC analysis, the principal components were computed from all samples in the data together, and in the figure(s) we extract the samples for that analysis. The labels and scale of the axes may differ, but the coordinates are the same: the position (0,0) is the same in all sub-figures.

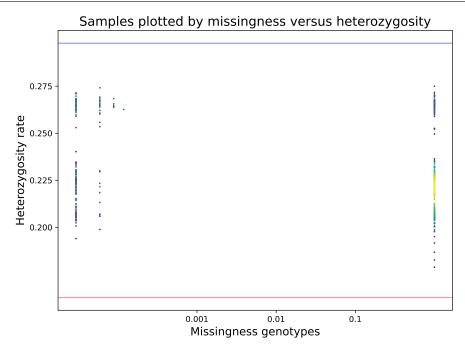
Figure 6 Principal component analysis, using the batch of the samples as the label.



6 Heterozygosity check

Levels of heterozygosity were examined in the data filtered in the previous step. Figure 7 shows plots of heterozygosity versus individual missingness (i.e., the number of SNPs missing per individual). Levels of heterozygosity should be between the ranges given in the configuration file – anything higher may indicate that there is sample contamination, lower may indicate inbreeding. However, each set of data must be treated on its own merits and the analyst must apply their mind the problem. Missingness should be low.

Figure 7 Missingness versus heterozygosity: the lines show the mean heterozygosity plus/minus standard deviations (the brighter the colours, the greater the density). If there is zero missingness, only heterozygosity is shown in a violin plot. [File is {array-nd-c-imiss-vs-het.pdf}]



Individuals out of range heterozygosity were removed. Any indviduals with heterozygosity:

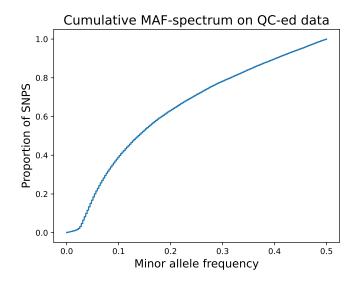
- less than 0.15 are removed. This may indicate inbreeding.
- greater than 0.343 are removed. This may indicate sample contamination.

Overall 0 individuals were removed. These individuals, if any, can be found in the file array-nd-c-fail_het.txt.

7 Minor Allele Frequency Spread

Figure 8 shows the cumulative distribution of minor allele frequency in the data **after** quality control (the figures shown in Section 2 show the MAF before QC. The MAF cut-off should be chosen high enough that one is sure that the variants seen are real (so this would depend on the size of the sample and the quality of the genotyping and whether some of the data is imputed). In this analysis the cut off was 0.01. Again, note that the *minor* allele is determined with respect to the frequency spectrum in this data – 'minor' is not synonym for alternate or non-reference allele, or the allele that has minor frequency in some other data set. Under this definition the MAF is always ≤ 0.5 .

Figure 8 Minor allele frequency distribution [File is {kgpexample-maf_plot.pdf}]



8 Differences between cases and controls

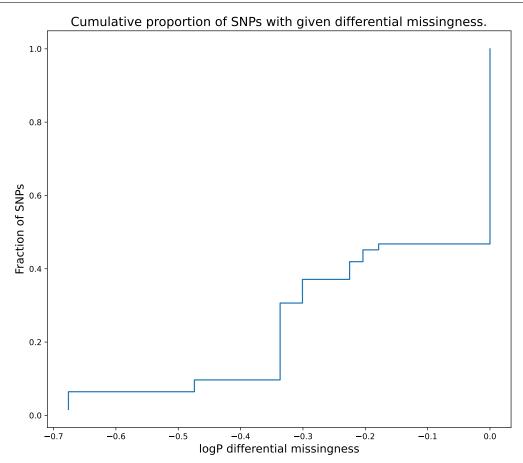
We do not expect there to be large, observable macro-scale differences between cases and controls. Great caution needs to be taken in this case. If the samples are from heterogeneous groups, where the case-control status differs between groups, then there may well statistically significant differences between the cases and controls and a batch analysis should be undertaken.

We compute for each SNP the missingness in the cases, and the missingness in the controls, and the corresponding p-value describing the difference in missingness.

We expect very few SNPs to have highly significant differences. Where many SNPs with very highly significant p-values are found, great care should be taken.

Figure 9 plots the differences between cases and controls, showing the SNP-wise p-value, unadjusted for multiple testing

Figure 9 The plot shows differential missingness for each (log) level of significance, the number of SNPs with p-value of at least this significance. The flatter the better. [File is array-c-c-diff-snpmiss_plot.pdf]



For removal of SNPs, we compute the p-value adjusted for multiple testing, by performing permutation testing (1000 rounds) using the PLINK mperm maxT option. SNPs are removed from the data set if their adjusted (EMP2) differential missingness p-value is less than 0.05. The SNPs that are removed can be found in the file array-nd-c-c_missing-failed_diffmiss.snps

Figure 10 shows a principal component analysis of the data, identifying the cases and controls. Should the cases and controls cluster differ significantly, something is likely wrong. Moreover should there be any significant clusters or outliers, association testing should take into account stratification. Statistical testing could also be done. Figure 11 shows for each principal component what the corresponding eigenvalue is. The shape of the curve may indicate if there is any structure in the data. Informally, the "broken stick" model sees two processes generating PCs – population structure and

random fluctuation. The eigenvalue is a function of both – the former has a major impact but drops relatively rapidly; the latter has less effect but drops more slowly. If (!) the model is correct then you will see an inflection point in the graph, where the random effects become the leading factor in the change. More formal analysis may be desirable, e.g., using the Tracy-Widom statistic or Velicer's MAP test.

Figure 10 Principal Component Analysis of Cases Versus Controls [File is {array-nd-c-prune-pca.pdf}]

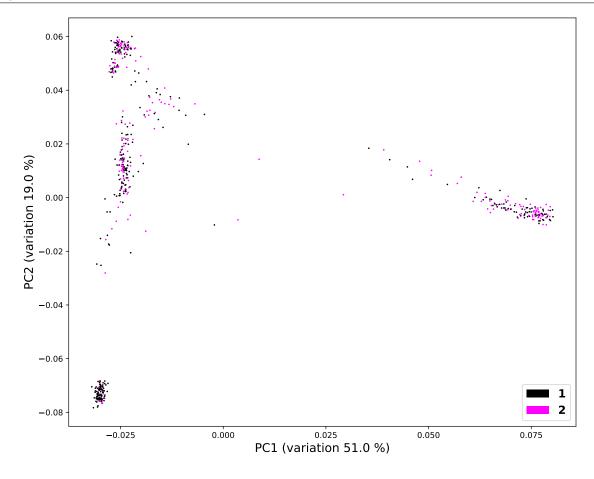
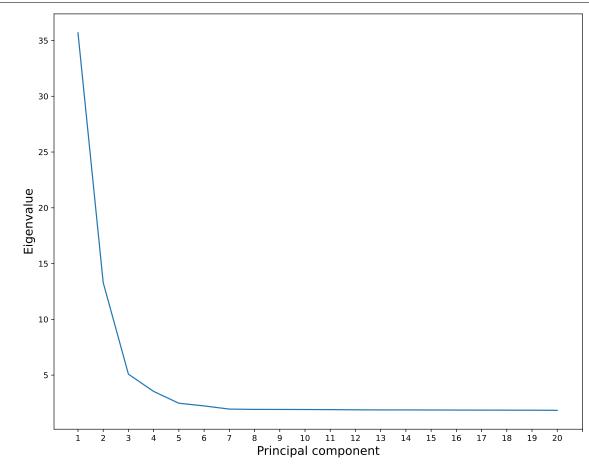


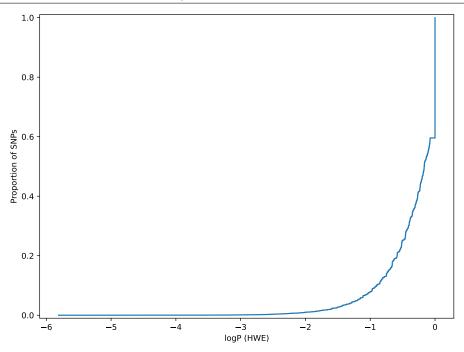
Figure 11 Eigenvalues for each principal component: the shape of the curve gives some indication of how many PCs are important [File is {eigenvalue.pdf}]



9 Hardy-Weinberg Equilibrium

Deviation for Hardy-Weinberg Equilibrium (HWE) may indicate sample contamination. However, this need not apply to cases, nor in a situation where there is admixture. For each SNP, we compute the probability of the null hypothesis (that the deviation from HWE is by chance alone). Figure 12 shows a plot of the corresponding p-value versus the frequency of occurrence.

Figure 12 The plot shows for each level of significance, the number of SNPs with H WE p-value [File is {array-nd-c-c-unaff-hwe_plot.pdf}]



10 Final results

The quality control procedures as described below were applied to the data. The final, cleaned result contains:

- 33791 SNPs
- 178 participants

The final output files are

- kgpexample.bed,
- kgpexample.bim, and
- kgpexample.fam.

The ouput files' md5 sums are shown below

kgpexample.bed d29a18e7c5077bb82d073f0d7cdbd292 kgpexample.bim 60636744e6b8790ddcef6a90d5efb615 kgpexample.fam 0a5ee2230114243254f3d03ae6206653

Table 6 Docker Images Used				
Nextflow process	Docker Image			
default	quay.io/h3abionet_org/py3plink			

11 Technical details

The analysis and report was produced by the h3aGWAS pipeline (http://github.com/h3abionet/h3agwas) produced by the Pan-African Bioinformatics Network for H3Africa (http://www.h3abionet.org).

The following tools were used:

- PLINK v1.90b6.16 64-bit (17 Feb 2020) [Chang et al 2015]
- 22.04.3 [Di Tommaso et al, 2017]
- A local copy of the workflow was used
- The command line below was called [NB: if the command line is long, the linebreak may break oddly after a hyphen or dash so take care.]

```
nextflow run /home/jeantristan/Travail/git//h3agwas/qc/main.nf --input_dir data/
array_plk --input_pat array --output_dir qc --output kgpexample --phenotype data/
pheno/pheno_test.all --pheno_col phenoqc_ql --case_control data/pheno/pheno_test.
all --case_control_col Sex --batch data/pheno/pheno_test.all --batch_col batch -
profile slurmSingularity -resume
```

- The profile slurmSingularity was used: the docker images used are found in Table 6
- The full configuration can be found in the appendix.

12 References

- Chang CC, Chow CC, Tellier LC, Vattikuti S, Purcell SM, and Lee JJ. (2015). Second-generation PLINK: rising to the challenge of larger and richer datasets. *GigaScience*, 4(1), 1-16. http://doi.org/10.1186/s13742-015-0047-8
- Di Tommaso P, Chatzou M, Prieto Barja P, Palumbo P, Notredame C. Nextflow enables reproducible computational workflows. *Nature Biotechnology*, 35, 316-319, 2017. Nextflow can be downloaded from https://www.nextflow.io/

A Configuration

A.1 config

```
singularity.cacheDir = "/home/jeantristan/.singularity/"
A.2
     nextflow
plugins {
  id 'nf-azure'
py3Image = "quay.io/h3abionet_org/py3plink"
gemmaImage="quay.io/h3abionet_org/py3plink"
latexImage="quay.io/h3abionet_org/h3agwas-texlive"
py2fastImage="quay.io/h3abionet_org/py2fastlmm"
swarmPort = '2376'
queue = 'batch'
manifest {
   homePage = 'http://github.com/h3abionet/h3agwas'
    description = 'GWAS Pipeline for H3Africa'
   mainScript = "main.nf"
}
aws {
accessKey='******
secretKey='******
    region
           ='us-east-1'
/*cloud {
    // imageId = "ami-710b9108"
                                    // specify your AMI id here
    instanceType = "m4.xlarge"
    subnetId = "null"
                     = "null"
    sharedStorageId
    sharedStorageMount = "/mnt/shared"
    bootStorageSize = "20GB" // Size of disk for images spawned
        instanceStorageMount = "" // Set a common mount point for images
        instanceStorageDevice = "" // Set a common block device for images
    autoscale {
        enabled = true
        maxInstances = 1
        terminateWhenIdle = true
    }
params {
    // Directories
                            = "/$PWD"
    work_dir
    input_dir
                            = "sample"
    // Can use S3 too
    //input_dir
                              = "s3://h3abionet/sample"
    input_pat
                            = "sampleA"
    output_dir
                            = "${params.work_dir}/output"
    scripts
                            = "${params.work_dir}/scripts"
                            = "out"
    output
                        = 95
    max_forks
```

```
high_ld_regions_fname = ""
    sexinfo_available = true
    cut_het_high
                      = 0.343
                        = 0.15
    cut_het_low
                        = "0.05"
    cut_diff_miss
                         = "0.01"
    cut_maf
                         = "0.02"
    cut_mind
                         = 0.01
    cut_geno
    cut_hwe
                         = 0.008
   pi_hat
                         = 0.11
    super_pi_hat = 0.7
                        = 0.8 // default for F-sex check -- >= means male
    f_lo_male
    f_hi_female
                        = 0.2 // \ll means female
                        = 0.4 // 10% Gen Call confidence -- 0.4 is very low
    gc10
    case_control = "${params.input_dir}/sample.phe"
    case_control_col
                       = "PHE"
    phenotype = "0"
    pheno_col = ""
    batch = "0"
    batch_col = 0
    idpat
                        = 0 // or (\w+)-DNA_(\w+)_.* or .*_(.*)
                        = "1750MB"
   plink_mem_req
    other_mem_req
                        = "1750MB"
                        = '12h'
    big_time
    sharedStorageMount
                        = "/mnt/shared"
   max_plink_cores
}
profiles {
    // For execution on a local machine, no containerization. -- Default
    standard {
       process.executor = 'local'
    }
    awsbatch {
        process.executor = "awsbatch"
 aws.region ='us-east-1'
        aws.uploadStorageClass = 'ONEZONE_IA'
        process.queue = 'h3a-00'
   }
    azurebatch {
       process.executor = 'azurebatch'
    slurm {
      process.executor = 'slurm'
      process.queue = queue
    // Execute pipeline with Docker locally
    docker {
       docker.remove
                          = true
```

```
docker.registry
                           = 'quay.io'
        docker.enabled
                           = true
                           = 'auto'
        docker.temp
        docker.fixOwnership= true
docker.runOptions = '-u $(id -u):$(id -g) --rm'
        docker.process.executor = 'local'
    // Execute pipeline with Docker Swarm setup
    dockerSwarm {
        docker.remove
                           = true
        docker.runOptions = '--rm'
        docker.registry = 'quay.io'
        docker.enabled
                           = true
                           = 'auto'
        docker.temp
        docker.fixOwnership= true
        docker.process.executor = 'local'
        docker.engineOptions = "-H :$swarmPort"
    }
    // For execution on a PBS scheduler, no containerization.
        process.executor = 'pbs'
        process.queue = queue
    }
    // For execution on a SLURM scheduler, no containerization.
    slurm {
        process.executor = 'slurm'
        process.queue = queue
    }
    slurmSingularity {
        singularity.cacheDir = "${HOME}/.singularity"
        process.executor = 'slurm'
        singularity.autoMounts = true
        singularity.enabled = true
        singularity.runOption = "--cleanenv"
        process.queue = queue
    }
    singularity {
        singularity.cacheDir = "${HOME}/.singularity"
        singularity.autoMounts = true
        singularity.enabled = true
     }
}
process {
    withLabel:bigMem {
```

A.2 nextflow A CONFIGURATION

```
memory = '8GB'
    }
    container=py3Image
    withLabel: latex {
          container = latexImage
    }
    withLabel: py2fast {
          container = py2fastImage
    }
}
\verb|timeline| \{
    \verb"enabled=true"
    file = "nextflow_reports/timeline.html"
report {
    enabled = true
    file = "nextflow_reports/report.html"
}
```