



ADDENDUM BBMRI-FUNCTIONAL GENOMICS RP3 USER CODE OF CONDUCT

For the BBMRI-Rainbow Project 3 Functional Genomics (RP3) substantial amounts of genotype and phenotype data will be generated and shared.

Within RP3 existing data contributed and owned by participating biobanks, including GWAS and phenotype data, will be brought together with newly generated transcriptome and methylome data.

The data is only to be used for pre-specified purposes to establish a comprehensive catalogue how genetic variation is impacting DNA methylation and gene expression levels, and how these effects eventually affect phenotypic variation. A description of a planned set of primary analyses according to the grant application is given below. Proposals for follow-up analyses using RP3 data are welcomed and will require approval.

To ensure this data is only accessible to bonafide researchers who obtained proper and explicit permission to conduct such analyses, the following data access and security measures will apply. RP3 data are stored on a centralized ABD secure BBMRI storage facility on the national LifeScience GRID as maintained by SARA. This facility includes tape back-up and fast lightpath connections to Dutch UMCs and GoNL data (RP1+2) are already stored here. To prevent inappropriate use and distribution to the genotype, phenotype, RNA-seq and DNA methylation data, researchers should request and obtain access to the data through the RP3 Data Access Procedure; accept and abide the rules of the Data Access Procedure with regards to security and conduct; request a personal GRID-certificate (enabling a strictly personal authentication procedure and logging of activities on the Grid).

All analyses will take place on the GRID. This implies that is not allowed for researchers to download data and analyze them on their own computing facilities not part of the GRID (e.g. non-GRID clusters and personal computers). When working on a local GRID-cluster, all downloaded data as well as intermediate/output data generated through analyses, must only be accessible by the researcher responsible (i.e. all original and subsequent data will not be readable, writable or executable by anyone other than the researcher). Activities with regards to viewing and downloading data will be monitored at the level of individual researcher to ensure data security.

Planned primary RP3 analyses

- RNA-seq based eQTL catalogue
- 450k-based meQTL catalogue
- DNA methylation vs. gene expression comparisons
- RNA-seq based discovery of exotic mechanisms of post-transcriptional modification (fusion, trans-splicing, editing, alternative poly-adenylation)
- Micro-array expression imputation using RNA-seq
- Next generation DNA methylation in GoNL subset: 'imputation' 450k, epigenetic landscape near insertions/deletions, alternative splicing effects, transmission, allele-specific methylation
- Proof of concept: Functional genomics of GWAS-identified SNPs sets associated with lipids and anthropometry (SNP > Methylation > RNA > Phenotype)