Genetic Association Course with Application to Sequence and Genotype Data September 26 - 30, 2022

Delbrück Center for Molecular Medicine-Berlin, Germany

Each session will begin with a theoretical introduction followed by practical exercises. The instructors for the course are Suzanne Leal (Columbia University) and Michael Nothnagel (University of Cologne). Please feel free to bring your own data sets for discussion and/or analysis.

The course will be held daily from 9:00 a.m. to 5:00 p.m., except for Wednesday, when the course will end at 1:00 pm to have free time in the afternoon for sightseeing. On Monday, registration will be held from 8:30 to 9:00 am. A dinner at a local restaurant will be held for students and faculty directly following the course on Monday.

MONDAY Morning Lecture

September 26th Introduction to genetic epidemiology, population genetics and statistical testing.

Introduction to PLINK and R; file formats

Computer Exercises;

PLINK and R - manipulating data

Afternoon Lecture

Basic statistical test for the analysis of genotype and sequence data

Introduction to PLINK and R; file formats

Computer Exercises;

PLINK and R – simple tests of association

18:00 -22:00 Dinner at a local restaurant

TUESDAY Morning Lecture

September 27th Linkage disequilibrium (LD), pairwise measures of LD

Data quality control, cleaning genotype data

Computer Exercises

PLINK – Data quality control *Pencil and Paper Exercises*

r2, D', etc.

Afternoon Lecture

Analysis of quantitative and qualitative traits using linear and logistic regression;

confounding and how to control for it in the analysis

Haplotype reconstruction and estimation, testing for associations using haplotype

data

Computer Exercises

UNPHASED – Haplotype and LD estimation & association testing PLINK & R – Logistic and linear regression – adjusting for covariates

WEDNESDAY

Morning Lecture

September 28th

Population substructure/admixture detection and control of confounding due to

population substructure (structure, principal components analysis, etc.)

Computer Exercises

PLINK – Multidimensionality scaling (MDS) and principal components analysis

(PCA)

Afternoon Free for sightseeing

THURSDAY

Morning Lecture

September 29th

Generalized linear mixed models and linear mixed models; Data quality control

for rare variant data obtained from next generation sequencing

Computer Exercises

REGENIE, Variant Association Tools (VAT)-part I

Afternoon Lecture

Complex trait rare variant association analysis of sequence data. population and

family-based aggregate tests.

Computer Exercises
VAT-part II and PSEQ

FRIDAY

Morning Lecture

September 30th

Sample size estimation and power calculations (for Rare Variant Aggregation

Tests) and Genome-Wide Association Studies (GWAS); the multiple testing problem; controlling the family wise error rate (FWER); and permutation and

false discovery rate (FDR)

Cochran-Armitage test for trend power tool, GAS, Genetic Power Calculator

R-permutation, FDR

Computer Exercises

Afternoon Lecture

Imputing genotype data from sequence and genotype data; analyzing imputed

genotype data; polygenic risk scores; detecting gene x gene and gene x

environment interactions

Computer Exercises

PLINK & R -Testing for gene x gene interactions