Exploiting public data

Bas Heijmans
Molecular Epidemiology
Leiden University Medical Center
The Netherlands
bas.heijmans@lumc.nl



Public data

Genomics data is increasingly made publicly available upon publication

- Often requirement of journal
- 'Open science'
- Efficiency of research
- Transparency
- Reproducibility
- More citations



New opportunities for data savvy scientists

Answering a research questions may not require the generation of new data!

(although a combination of new and old is preferred)



Research data (up to many samples)

- Gene Expression Omnibus (GEO)
 https://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GS
 E48472
 E48472
 https://example.com/en/geo/query/acc.cgi?acc=GS
 <a href="https://
- European Genome-phenome Archive (EGA)
 https://www.ebi.ac.uk/ega/datasets/EGAD00001000733
- ARCHS –RNA-seq gene counts from GEO ready for R https://amp.pharm.mssm.edu/archs4/index.html



GWAS summary statistics

- GWAS catalog of genome-wide significant SNPs https://www.ebi.ac.uk/gwas/
- MR-base: summary statistics with focus on Mendelian randomization http://www.mrbase.org/
- UK Biobank GeneAtlas
 http://geneatlas.roslin.ed.ac.uk/



Reference data (few samples)

- Human reference genome
 https://www.ncbi.nlm.nih.gov/projects/genome/guide/human/
- Human reference epigenomes <u>http://epigenomesportal.ca/ihec/</u>
- Cancer genomes (reference & research data)
 https://cancergenome.nih.gov



Biological pathways

- Gene ontology <u>http://www.geneontology.org</u>
- Reactome
 http://www.reactome.org
- StringDB
 <u>https://string-db.org/</u>



Browsers

- Genome browsers like UCSC and ESMBL
 http://www.ensembl.org/index.html
- Omics atlases (effect of SNPs on expression/methylation and much more)
 http://bbmri.researchlumc.nl/atlas/



Public data in practice

Slieker et al. Genome Biology (2016) 17:191 DOI 10.1186/s13059-016-1053-6

Genome Biology

RESEARCH Open Access



Age-related accrual of methylomic variability is linked to fundamental ageing mechanisms



Roderick C. Slieker¹, Maarten van Iterson¹, René Luijk¹, Marian Beekman¹, Daria V. Zhernakova², Matthijs H. Moed¹, Hailiang Mei³, Michiel van Galen⁴, Patrick Deelen², Marc Jan Bonder², Alexandra Zhernakova², André G. Uitterlinden⁵, Ettje F. Tigchelaar², Coen D. A. Stehouwer⁶, Casper G. Schalkwijk⁶, Carla J. H. van der Kallen⁶, Albert Hofman⁷, Diana van Heemst⁸, Eco J. de Geus⁹, Jenny van Dongen⁹, Joris Deelen¹, Leonard H. van den Berg¹⁰, Joyce van Meurs⁵, Rick Jansen¹¹, Peter A. C. 't Hoen⁴, Lude Franke², Cisca Wijmenga², Jan H. Veldink¹⁰, Morris A. Swertz¹², Marleen M. J. van Greevenbroek⁶, Cornelia M. van Duijn¹³, Dorret I. Boomsma⁹, BIOS consortium, P. Eline Slagboom¹ and Bastiaan T. Heijmans^{1*}

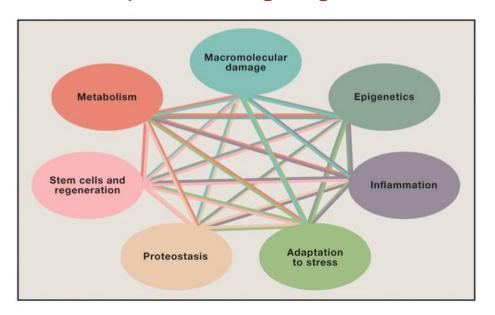


Genomic dysregulation and ageing

Lopes-Otin et al. *Cell* 2013: '9 hallmarks of ageing'

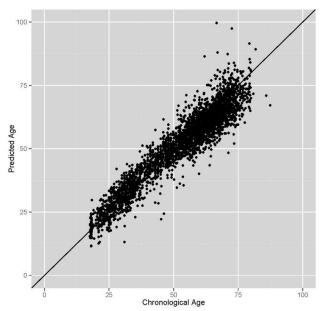


Kennedy et al. *Cell* 2014: '7 pillars of ageing'





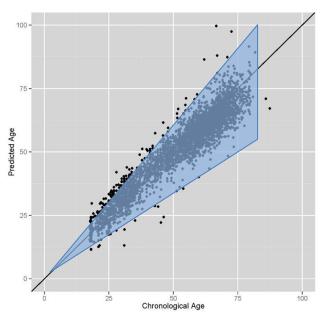
DNA methylation tracks chronological age



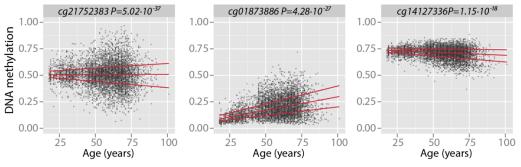
DNAmeth clock of Steve Horvath of 353 CpGs (*Genome Biol* 2013) applied on our own data (N>3000).



Loss of epigenetic control with age



age-related Variably Methylated Positions aVMPs



- Biology remains elusive.
- No link with gene expression.
- We already know chronological age.



Large scale: Biobank-based Integrative Omics Study (BIOS)

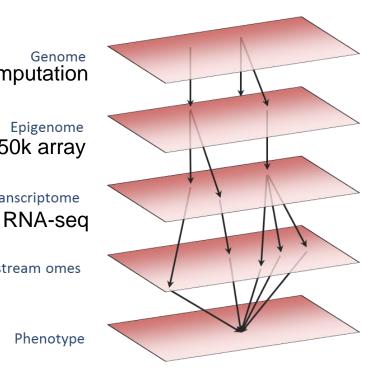
Genome SNP arrays & imputation

> **Epigenome** Illumina 450k array

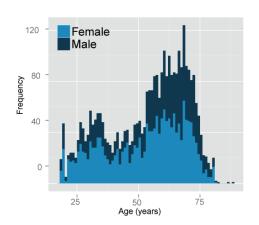
> > Transcriptome

Downstream omes

Phenotype



- Whole blood samples.
- 3295 methylome profiles: 450k array.
- 2044 transcriptome profiles: RNA-seq.







Genomics papers

- 1. Discovery
- 2. Validation
- 3. Interpretation

And often public data helps a lot!



Get a flavour of the possibilities

Slieker et al. Genome Biology (2016) 17:191 DOI 10.1186/s13059-016-1053-6

Genome Biology

RESEARCH Open Access

Age-related accrual of methylomic variability is linked to fundamental ageing mechanisms



Roderick C. Slieker¹, Maarten van Iterson¹, René Luijk¹, Marian Beekman¹, Daria V. Zhernakova², Matthijs H. Moed¹, Hailiang Mei³, Michiel van Galen⁴, Patrick Deelen², Marc Jan Bonder², Alexandra Zhernakova², André G. Uitterlinden⁵, Ettje F. Tigchelaar², Coen D. A. Stehouwer⁶, Casper G. Schalkwijk⁶, Carla J. H. van der Kallen⁶, Albert Hofman⁷, Diana van Heemst⁸, Eco J. de Geus⁹, Jenny van Dongen⁹, Joris Deelen¹, Leonard H. van den Berg¹⁰, Joyce van Meurs⁵, Rick Jansen¹¹, Peter A. C. 't Hoen⁴, Lude Franke², Cisca Wijmenga², Jan H. Veldink¹⁰, Morris A. Swertz¹², Marleen M. J. van Greevenbroek⁶, Cornelia M. van Duijn¹³, Dorret I. Boomsma⁹, BIOS consortium, P. Eline Slagboom¹ and Bastiaan T. Heijmans^{1*}



Reference epigenomes inform on biological function

