

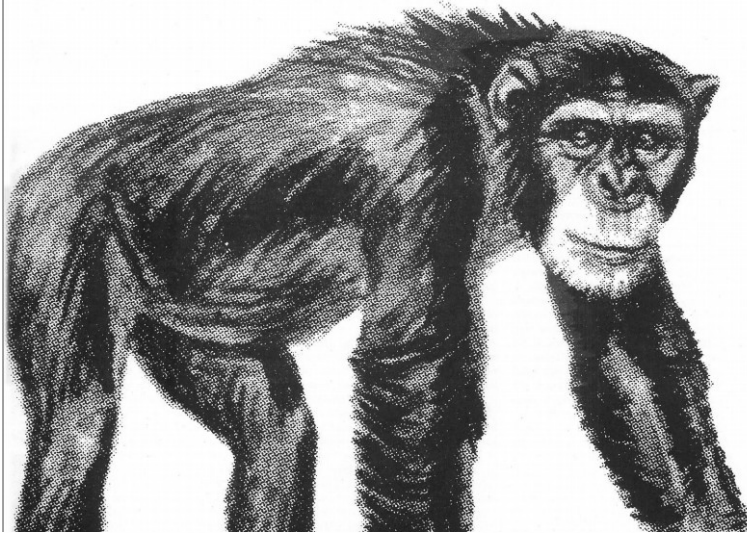
The contribution of non-coding mutations to human phenotypes and disease is unknown

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SCIENCE

Evolution at Two Levels in Humans and Chimpanzees

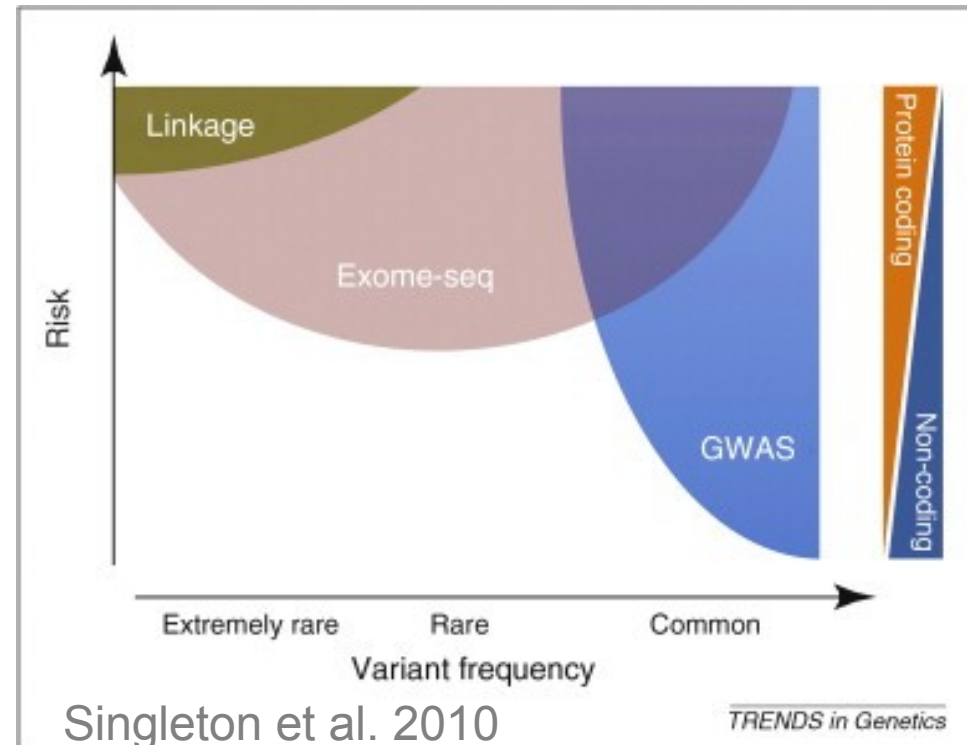
Mary-Claire King and A. C. Wilson



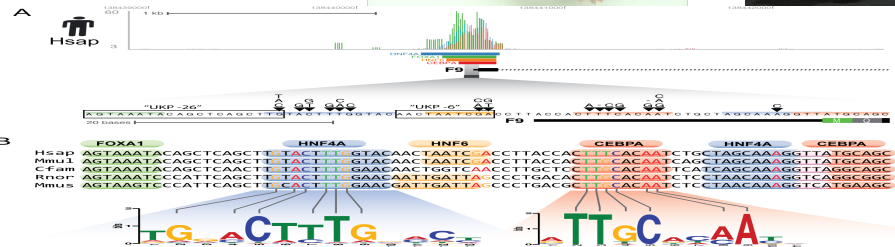
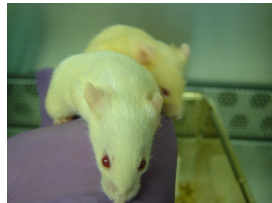
Evolution at Two Levels in Humans and Chimpanzees

Their macromolecules are so alike that regulatory mutations may account for their biological differences.

Mary-Claire King and A. C. Wilson



Wilson Lab – Understanding human gene regulation and disease with comparative genomics



1) Generating/analyzing “big” datasets to assign function to the non-coding genome

- Hemostasis (F. Gagnon, S. Mital, and C. Hayward labs)
- Puberty (M. Palmert and A. Goldenberg labs)
- Early zebrafish heart development (I. Scott lab)
- Genome biology of topoisomerases

2) Testing regulatory DNA function in human cells and model organisms

Wilson Lab – Understanding human gene regulation and disease with comparative genomics

1) Generate “big” datasets to assign function to the non-coding genome

- Protein-DNA and epigenomic assays
 - ChIP-seq, ChIP-exo ...
- Gene expression
 - RNA-seq
 - high-throughput qPCR (Fluidigm BioMark with Rossant lab),
 - Single cell mRNA sequencing (Fluidigm C1 system)

2) Analyze “big” datasets to assign function to the non-coding genome

- detect differential DNA occupancy, gene expression
- Compare NGS results between species
- Integrative analyses to assign function to the non-coding genome

3) Test regulatory DNA function in human cells and model organisms

- Targeted sequencing/cloning/reporter assays of non-coding DNA
 - Fluidigm Access Array microfluidic PCR
- CRISPR/Cas9 genome editing to delete/modify non-coding DNA
- Gene reporter assays in zebrafish (with Ian Scott’s Lab)