

FAANG
Functional Annotation of Animal Genomes

Examples of FAANG associated projects

- WUR-pigENCODE: Martien Groenen
- Canada and FAANG: Graham Plastow
- Strategy to identify regulatory mutations affecting complex traits: Amanda Chamberlain
- Equine FAANG: Carrie Finno


WU FAANG pilot projects

Martien Groenen, Richard Crooijmans,
Hendrik-Jan Megens, Ole Madsen



Expression and methylation analyses in pig

- RNAseq and miRNA data
 - Improve gene annotation (mRNA)
 - lncRNA
 - miRNA
- RRBS data
 - CpG
- Samples: 3 boars (Large white, Duroc, Pietrain)
 - liver, spleen, testis, brain (frontal lobe), muscle (all)
 - olfactory bulb, lung and lymph node from (LW only)



Co-expression,
correlations

lncRNAs and miRNAs

lncRNAs (Ole Madsen)

- FEELnc (provided by Thomas Derrien, University of Rennes, France)
 - Number of mRNA : 30585
 - Number of lncRNA : 35046
 - Number of lncRNA without interaction with coding gene: **19832**

MiRNAs (Anoop Arya)

- miRDeep2
 - Predicted a total of **1071** precursors.
 - Of these 689 are putative predicted precursors and 382 are known precursors

ChIP-seq

- Histone marks
 - H3K27ac active enhancers and promoters
 - H3K4me3 promoter active genes
 - H3K27me3 silenced genes (active during developmental stages)
 - H3K4me1 active enhancers
- Samples
 - Cell lines (pig intestine, liver)
 - Tissues (start with pig intestine and liver)
 - FR-AgENCODE tissues
- In collaboration with the Molecular Biology Group of the Radboud University, Nijmegen (Henk Stunnenberg)

ChIPseq

- Porcine small intestinal epithelial cell line (IPEC-J2)
 - established from normal intestinal epithelium cells isolated from the jejunum of a neonatal, unsuckled pig
- Liver cell line

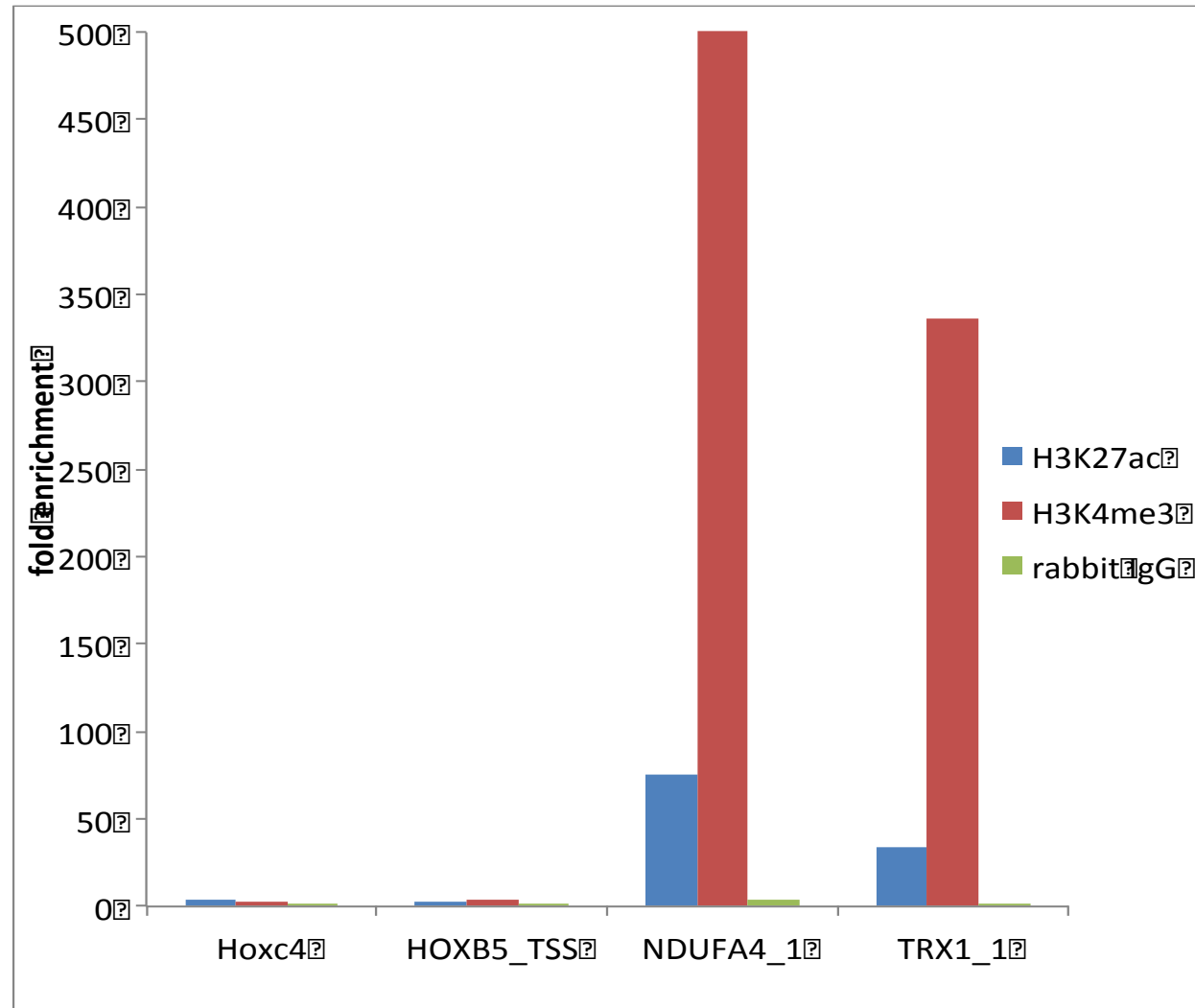
- WGS (30x)
- RNAseq
- RRBS

- Histone marks
 - H3K27ac
 - H3K4me3
 - H3K27me3
 - H3K4me1

- Controls (qPCR)
 - HOXC4
 - HOXB5
 - NDUFA4*
 - TRX1*

expressed (above 100 fpkm) in all our RNA-seq data

ChIPseq



Canada and FAANG

PAG

January 11th 2016



GenomeCanada

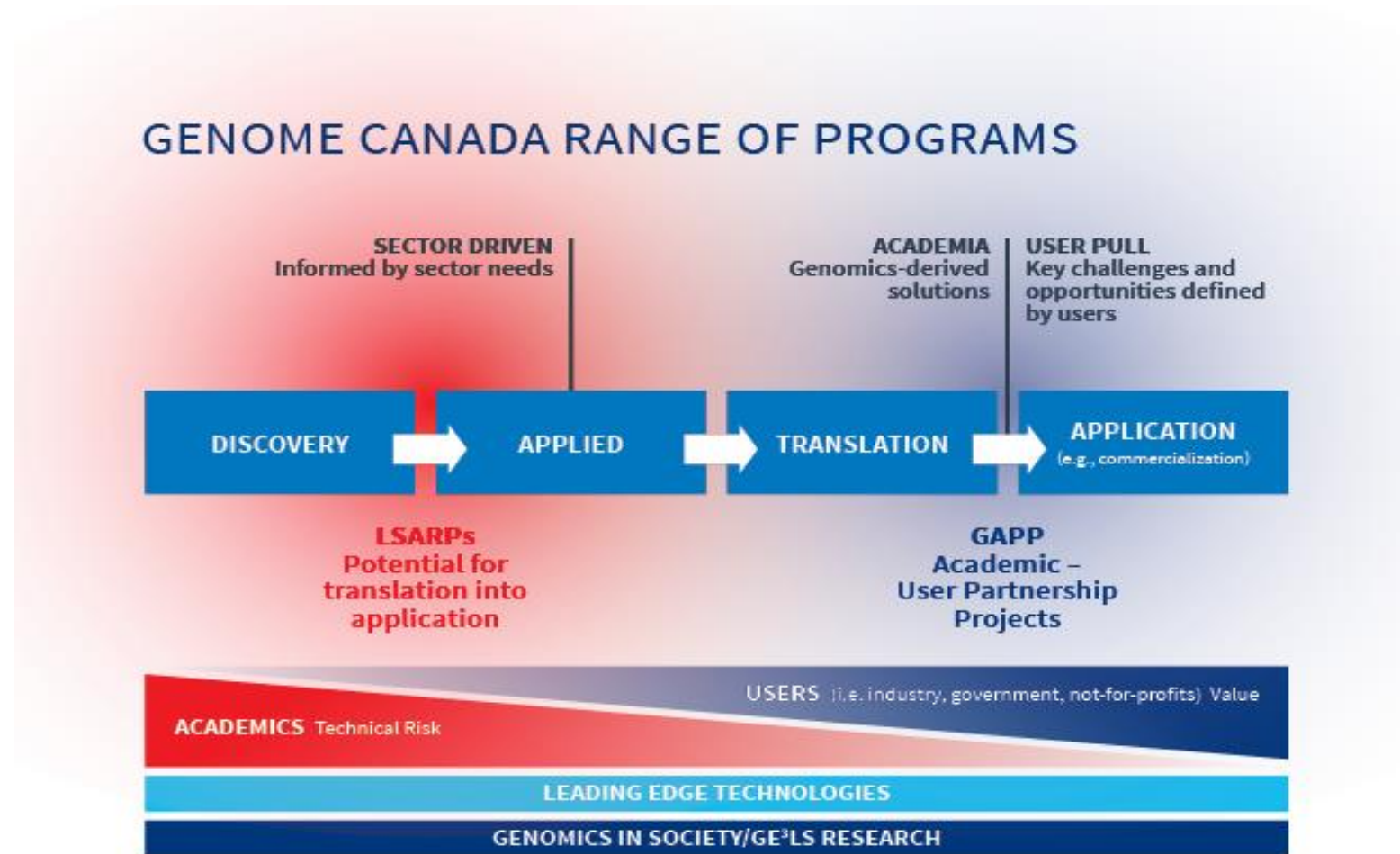
and FAANG

Leading Canada's genomics enterprise



Genome Canada Collaboration Opportunities and Funding for Livestock Genomics, D Bailey W158 10th January

Programs that take discoveries from lab to society



Application of Genomics to Improve Disease Resilience and Sustainability in Pork Production

Project Leader: Michael Dyck (U of Alberta)

Genome Centres: Genome Alberta and Genome Prairie

Project Budget: \$9,801,714

Project Partners:

- ALGP2
- PigGen Canada
- ALMA
- Genome Alberta - PEDV
- Swine Innovation Pork
- Saskatchewan Ministry Agriculture and Food
- INRA
- National Pork Board
- Alltech Inc.
- OGI

ALMA
Alberta Livestock
and Meat Agency Ltd.



Pigs

- Application of genomics to improve disease resilience and sustainability in pork production
- 2016 - 2019
- Industry collaboration with Large White * Landrace F1 barrows in a “natural challenge” – samples from sick/healthy pigs.
(samples can potentially be collected from female litter mates)
- Collection of tissues from LW*LR fetuses (PRRSV challenge plus controls – several time points d84-98 of pregnancy)

Increasing Feed Efficiency and Reducing Methane emissions through Genomics: A New Promising Goal for the Canadian Dairy Industry

Project Leader: Filippo Miglior (U of Guelph)

Genome Centres: Genome Alberta and Ontario Genomics

Project Budget: \$10,306,910

Project Partners:

- Canadian Dairy Network
- Dairy Producers
- GrowSafe System
- Ontario Ministry of Research and Innovation
- Australia DEDJTR
- UK Scottish Research College
- USDA Beltsville Research Herd
- Qualitas, Switzerland



Réseau laitier canadien



Canadian Dairy Network



Dairy Cattle

- **W146 Increasing Feed Efficiency and Reducing Methane Emissions Using Genomics: An International Approach**
- *Date: Saturday, January 9, 2016, Time: 2:30 PM*
- [Christine F. Baes](#), *Centre for the Genetic Improvement of Livestock, Department of Animal Biosciences, University of Guelph, Guelph, ON, Canada*

- Extremes of feed efficiency and methane emission
- Core tissues
- Core assays

Reverse Vaccinology Approach for the Prevention of Mycobacterial Disease in Cattle

Project Leaders: Andy Potter (VIDO), Robert Hancock (UBC)

Genome Centres: Genome Prairie and Genome BC

Project Budget: \$ 7,358,606

Project Partners:

- University of Saskatchewan
- Genome BC
- AFBI (Agri-Food and Biosciences Institute)



Additional Support

- Federal, provincial and industry support for livestock genomics efforts
- Co-funding and international collaboration are key success factors

Development and Deployment of MBVs/gEPDs for Feed Efficiency and Carcass Traits that Perform in Commercial Beef Cattle

Project Leader: John Basarab (U of Alberta), Donagh Berry (Teagasc), John Crowley (U of Alberta)

Genome Centre: Genome Alberta

Project Budget: \$4,473,035

Project Partners:

- Genome Alberta
- ALMA
- BIO
- Beefbooster Inc.
- CCHMS
- ICBF
- USDA-ARS



Existing plan

- Collection of FAANG tissues from animals of “high and low” feed efficiency based on EBVs (and GEBVs) includes methane emission
- Potential to expand to carcass and quality traits
- Crossbred steers (defined breed type) and primarily slaughter age
- Project focus on variant identification

New supporting application to Alberta Livestock and Meat Agency

- Lol due 11th January 2016
- *Outline plan*
- Select extreme animals on ebv (or breed Hi/Hi and Lo/Lo) for feed efficiency and methane yield.
- Run a time series at say 4 key points (pre-weaning, first test, puberty, and adult). Last 3 will have own phenotype.
- Collect core FAANG tissues (inc. rumen) from 2 heifers and 2 steers at each time point – 16 samples per trait.
- Angus or Crossbred population (with breed proportions)
- RNAseq and analysis/bioinformatics in Canada, core samples banked and sent (with staff) to expert labs running FAANG core assays (e.g. ChIP-seq assays. DNase-seq or ATAC-seq and Hi-C).

Ontario Ministry of Agriculture

- PI: **Angela Canovas** (P0518 P580)
- Canadian and international partners
Stephen Miller (AgResearch, NZ), Juan Medrano (UC Davis),
Sinead Waters (Teagasc, Ire)
- Focus feed efficiency (and meat quality)
- Cattle

Angus x Simental (UoG Research herd)
Adult (n=14 animals)

Holstein (UoG Research herd)
Adult (n=14 animals)

- Pig
Duroc
Adult, male (n=14 animals)
- Muscle, Adipose, Liver, Gut, Brain

Agriculture and AgriFood Canada

- PI: **Eveline M. Ibeagha-Awemu**
- Dairy and Swine Research and Development Centre, Sherbrooke, Qubec
- Holstein, >2nd lactation (6 cows)
Duodenum, Ileum, Rumen, Colon, Milk fat, Adipose tissue, Skeletal muscle, Liver, Lung, Heart, Kidney, Spleen
- W020 MicroRNAs are Master Regulators of the Bovine Lactation Curve
Date: Tuesday, January 12, 2016 Time: 1:50 PM
- P0087 Long Non-Coding RNA (lncRNA) Is Abundantly Expressed in the Gastro Intestinal Tract of Calves

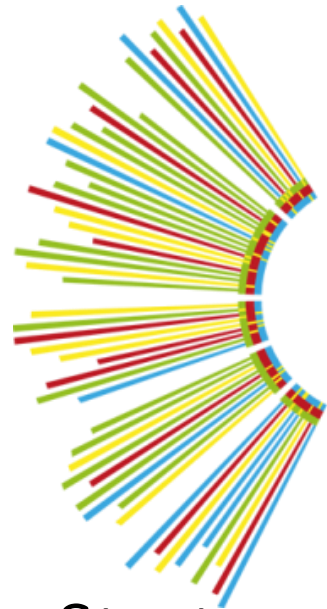
Collaborators

- University of Alberta: Michael Dyck, Leluo Guan, Paul Stothard, John Basarab (AAF)
- University of Saskatchewan: John Harding
- University of Guelph: Angela Canovas, Christine Baes, Filippo Miglior (CDN)
- Agriculture and Agri-Food Canada: Carolyn Fitzsimmons (UofA), ChangXi Li (UofA), Eveline Ibeagha-Awemu
- Roslin Institute: Alan Archibald, Mick Watson
- FAANG ASA – Elisabetta Giuffra, Huaijun Zhou



GenomeCanada

and FAANG



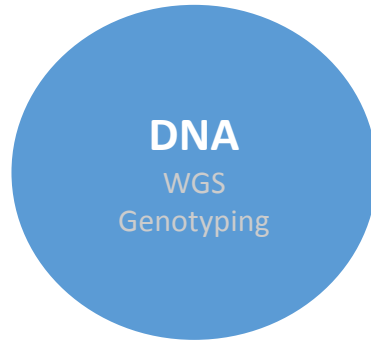
FAANG
Functional Annotation of Animal Genomes



Strategy to identify regulatory mutations affecting
complex traits

Amanda Chamberlain, Mike Goddard, Majid Khansefid, Min Wang, Iona
Macleod, Ben Hayes

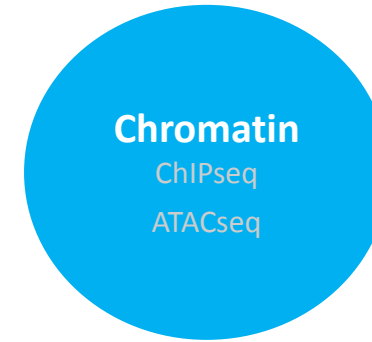
What assays are we using?



39 million variants – 1000 bulls
Ref populations genotyped
GWAS
genomic selection



Gene Atlas Tissue Bank
Experimental populations
eQTL
ASE
Non coding RNA



Gene Atlas Tissue Bank
Experimental populations
Public Data
Enhancers
Promoters
TF binding sites

A catalogue of novel bovine long noncoding RNA across 18 tissues

Lambros T. Koufarotis^{1,2,3*}, Yi-Ping Phoebe Chen¹, Amanda Chamberlain^{2,3}, Christy Vander Jagt^{2,3}, Ben J. Hayes^{1,2,3}

¹ College of Science, Health and Engineering, La Trobe University Bundoora, Melbourne, Victoria, Australia, ² Department of Environment and Primary Industries, AgriBio Bundoora, Melbourne, Victoria, Australia, ³ Dairy Futures Co-operative Research Centre, Melbourne, Victoria, Australia

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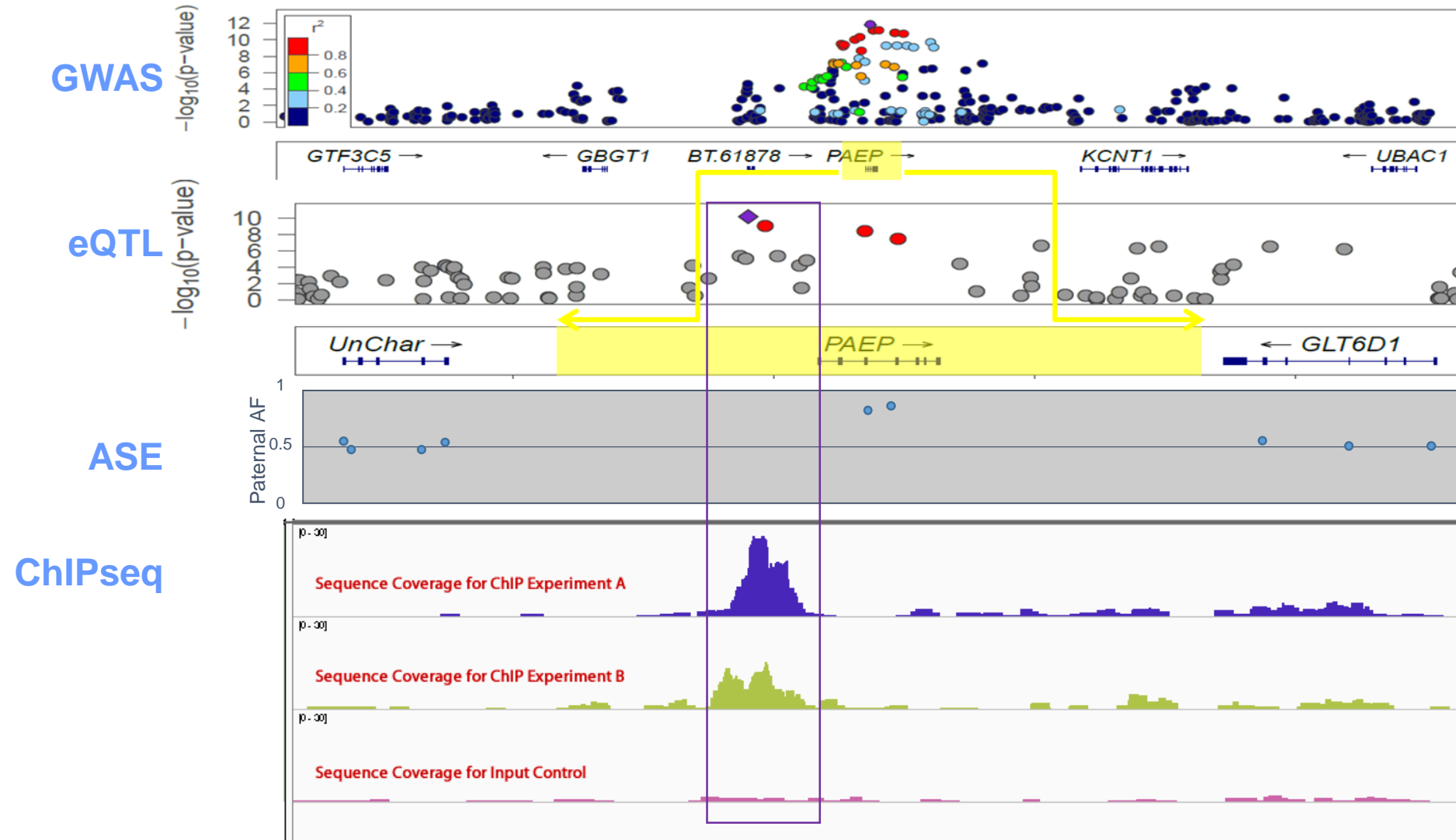


Extensive variation between tissues in allele specific expression in an outbred mammal

Amanda J. Chamberlain^{1,2†}, Christy J. Vander Jagt^{1,2†}, Benjamin J. Hayes^{1,2,3*}, Majid Khansefid^{1,2,5}, Leah C. Marett⁴, Catriona A. Millen^{2,5}, Thuy T. T. Nguyen¹ and Michael E. Goddard^{1,5}



Our strategy for identifying regulatory variants



Summary

- Strategy involves using QTL, eQTL, ASE, non-coding RNA, ChIPseq marks, public data to identify regulatory variants and use in genomic prediction.
- Contribution to FAANG:
 - 39 tissues from 2 cows and their foetus'
 - RNAseq
 - ChIPseq (H3K4me1, H3K4me3, H3K27ac, H3K27me3, CTCF)
 - Bioproject PRJNA251439, FAANG tag
- Other data currently available:
 - 1 cow, 18 tissues non-stranded RNAseq, 2 tissues stranded RNAseq, Bioproject PRJNA251439, SRA SRP042639
 - 3 cows, milk and white blood cells stranded RNAseq, Bioproject PRJNA305942, SRA SRP067373



FAANG Associated Projects Update: Equine

Carrie Finno

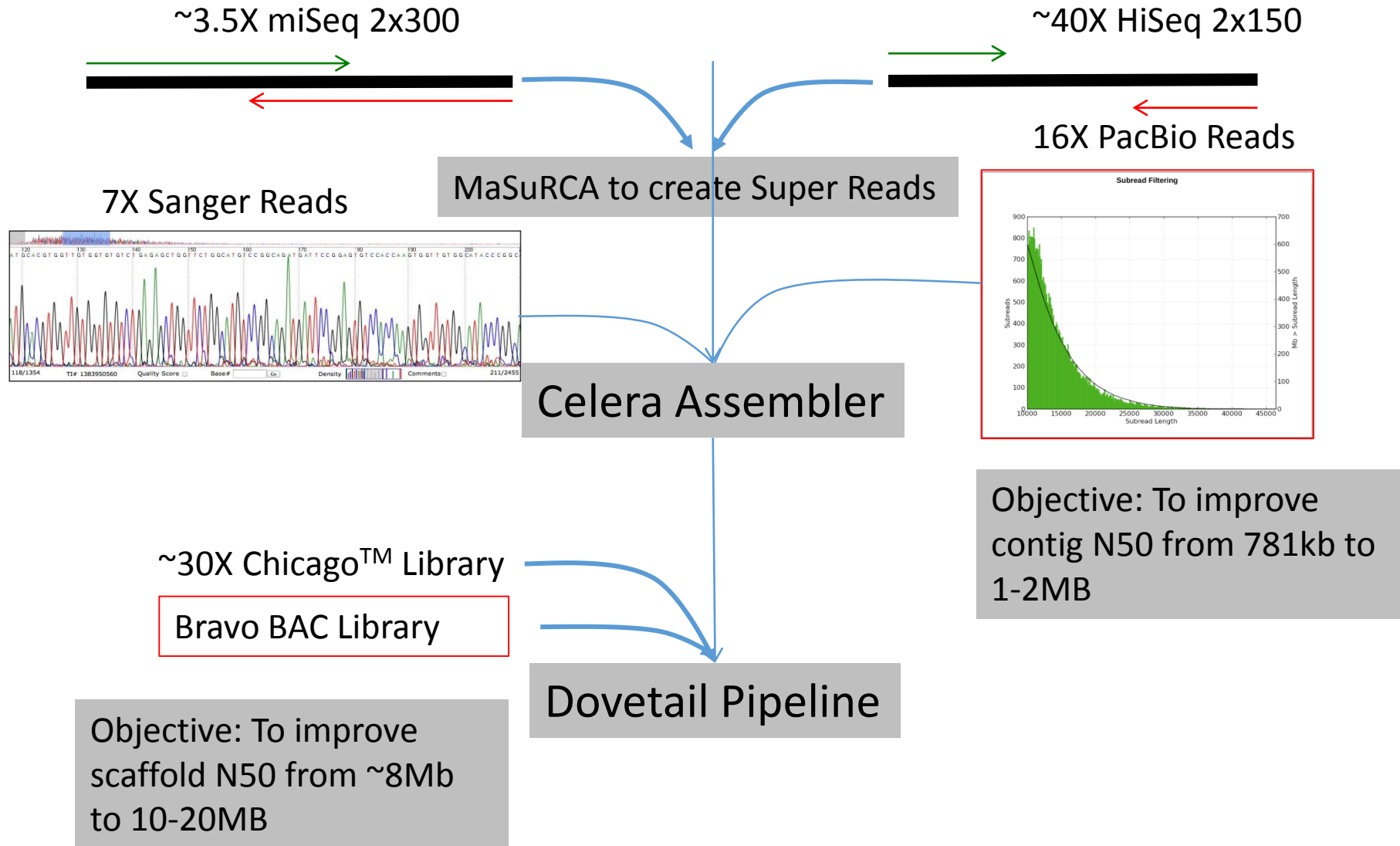
**A Community Effort to Functionally
Annotate the Equine Genome and
Facilitate Genome to Phenome
Analyses**

Equine Genome

- 2007 Equine Genome-6.8X Coverage
 - EquCab2.0 (Wade *et al.*, 2009)
- Towards EquCab3.0
 - Kalbfleisch, MacLeod, Orlando *et al.* W293)
 - -Morris Animal Foundation D15EQ-019



On-going Effort

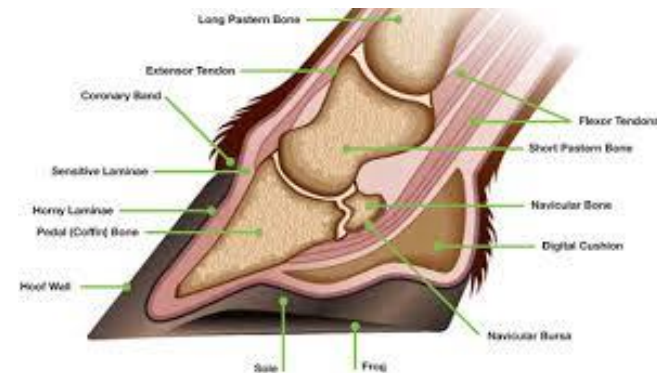


Structural Annotation

- RNA-seq
 - 1) Gene structure ~69K transcripts from pool of 43 tissues (Thoroughbred)
 - Hestand *et al.*, 2015 PLoS One 10(6):e0124375
 - 2) Eight tissue-specific transcription profiles (multiple breeds): embryo (ICM and TE), cerebellum, brainstem, spinal cord, retina, skeletal muscle, and skin.
 - Mansour *et al.* (P0563)
 - Publically available resource- UCSC track hubs
 - Tissue Specificity
 - Improve annotation: Extension of UTRs & Identification of novel transcripts including ncRNA
 - Correcting potential errors

Equine Functional Annotation: Proposed

- Samples: 2 Adult Thoroughbred horses (female and male)
- 50 tissues to be collected
- 8 prioritized tissues for assays: Skeletal muscle, liver, ovary/testis, cerebral cortex, spleen, heart, lung, and laminae
- Assays: WGS, RNA-Seq, microRNA-Seq, ChIP-seq (H3K4me3, H3K4me1, H3K27me3, H3K27ac, CTCF) and DNase-Seq
- Funding:
 - USDA Species Coordinator funds for sample collection
 - Submitted grant pending (Finno, Petersen, Bellone)





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