



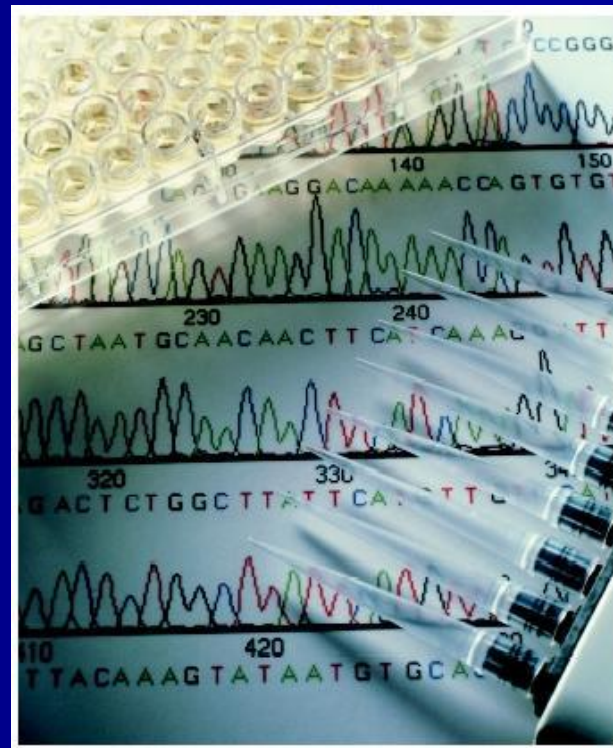
What I do & my vision for the future of beef cattle genomics

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"The mission of the animal genomics and biotechnology extension program is to provide broad, science-based extension programming on the uses of animal biotechnologies in livestock production systems."

<http://animalscience.ucdavis.edu/animalbiotech>





Dairy

High use of AI

Only one breed

Clear selection goal (total net merit)

Large number of high accuracy A.I. sires for training

Extensive, uniform collection of data on traits

Central USDA evaluation (AIPL) receiving all genotypes and recalibrating equations

Obvious way to increase ΔG (decrease age of young progeny test sires)

AI companies funding the genotyping because they get a clear cost savings in terms of young sire program

Beef

Little use of AI

Many breeds, limited resources

Multiple competing selection goals – cow/calf, feedlot, processor – little data sharing between sectors

Relatively few high accuracy sires for training

Few/no records on many important traits (feed efficiency, reproduction)

No centralized “national” cattle evaluation – breed associations perform genetic evaluations

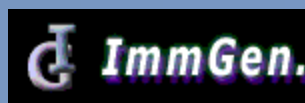
Use young herd sires so less way to decrease generation interval

No clear beneficiary willing to pay!

Crossbreeding is important



Many companies have come and some have gone....



The Power of the IGENITY[®] profile for Angus

The American Angus Association[®] through its subsidiary, Angus Genetics Inc.[®] (AGI), has a vision to provide Angus breeders with the most advanced solutions to their genetic selection and management needs.

1. Dry Matter Intake
2. Birth Weight
3. Mature Height
4. Mature Weight
5. Milk
6. Scrotal Circumference
7. Weaning Weight
8. Yearling Weight
9. Marbling
10. Ribeye Area
11. Fat Thickness
12. Carcass Weight
13. Tenderness
14. Percent Choice (quality grade)
15. Heifer Pregnancy
16. Maternal Calving Ease
17. Direct Calving Ease
18. Docility
19. Average Daily Gain
20. Feed Efficiency
21. Yearling Height

- Arthrogryposis Multiplex (AM)
- Neuropathic Hydrocephalus (NH)
- Bovine Viral Diarrhea – Persistently Infected (BVD PI)
- Coat Color



ANGUS
THE BUSINESS BREED

1. – Calving ease direct
2. – Birth weight
3. – Weaning weight
4. – Yearling weight
5. – Yearling height
6. – Mature weight
7. – Mature height
8. – Dry matter intake
9. – Residual feed intake
10. – Scrotal circumference
11. – Docility
12. – Calving ease maternal
13. – Milking ability
14. – Carcass weight
15. – Fat thickness
16. – Ribeye area
17. – Marbling score
18. – Tenderness

Lead Today with 50K



Pfizer Animal Health
Animal Genetics

50K SNP chip assays
50,000 SNPs spread
throughout genome





American Angus Association performs weekly evaluations with genomic data – recently updated to include new traits



	Igenity	Pfizer
Calving ease (CED)	✓	✓
Growth (BW WW YW Milk)	✓	✓
Residual Average Daily Gain (RADG)	✓	✓
Docility (DOC)	✓	✓
Yearling Scrotal/Height (SC,YH)	✓	✓
Mature Weight (MW)	✓	✓
Carcass (CWT MARB RIB FAT)	✓	✓

<http://www.angus.org/AGI/GenomicChoice11102011.pdf> (updated 11/18/2011)



ANGUS

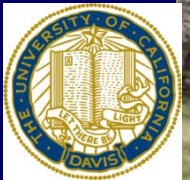
384 SNP

50K SNP



Genetic Correlation (r)/(r ² %)/PE	h ²	Igenity/ Neogen		Pfizer	
Calving Ease Direct	0.20	.47 (22%)	5	.33 (11%)	2
Birth Weight	0.42	.57 (32%)	4	.51 (26%)	3
Weaning Weight	0.20	.45 (20%)	5	.52 (27%)	7
Yearling Weight	0.20	.34 (12%)	2	.64 (41%)	13
Dry Matter Intake	0.31	.45 (20%)	3	.65 (42%)	9
Yearling Height	0.45	.38 (14%)	2	.63 (40%)	5
Yearling Scrotal	0.43	.35 (12%)	1	.65 (42%)	6
Docility	0.37	.29 (8%)	1	.60 (36%)	5
Milk	0.14	.24 (6%)	1	.32 (10%)	3
Mature Weight	0.55	.53 (28%)	2	.58 (34%)	3
Mature Height	0.82	.56 (31%)	1	.56 (31%)	1
Carcass Weight	0.31	.54 (29%)	5	.48 (23%)	3
Carcass Marbling	0.26	.65 (42%)	10	.57 (32%)	7
Carcass Ribeye Area	0.32	.58 (34%)	6	.60 (36%)	6
Carcass Backfat Thick	0.26	.50 (25%)	4	.56 (31%)	7

<http://www.angus.org/AGI/GenomicChoice11102011.pdf> (updated 11/18/2011)





Translational Questions for the Beef Industry




- ? How many phenotypic records are required in the initial experiment estimating the effect of chromosome segments?
- ? How many markers are needed— 384, 9K, 50K, 800K, whole genome?
- ? How does the relationship between the training population and the selection candidate affect accuracy?
- ? How often do chromosome segment effects need to be re-estimated? Every generation?
- ? Do predictions work across breeds?
- ? What is the value generated by the increased accuracy?
- ? Does this technology change breeding program design?

Van Eenennaam, A. L. 2011. Beef translational genomics: Lessons from the literature. Association for the Advancement of Animal Breeding and Genetics. 19: 271-278.

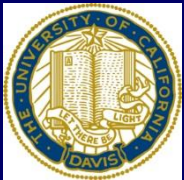


Angus predictions are not very accurate in Red Angus (Data provided by Dorian Garrick)



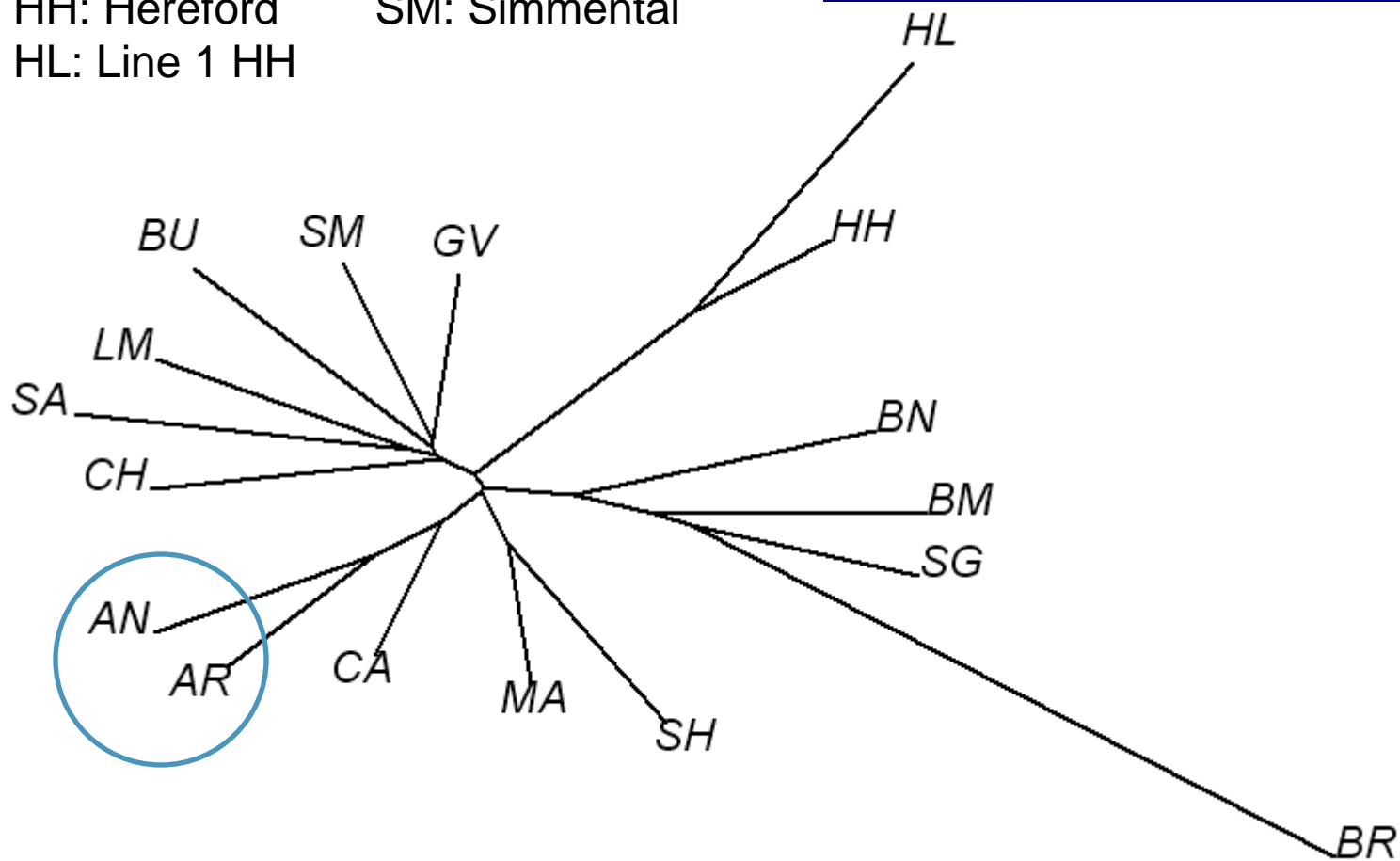
Trait	Validating in American Angus	Validating in Red Angus
BirthWt	0.64	0.27
WeanWt	0.67	0.28
YearlingWt	0.75	0.23
Fat	0.70	0.21
REA	0.75	0.29
Marbling	0.80	0.21
CalvEase (D)	0.69	0.14
CalvEase (M)	0.73	0.18

*Angus = ASREML 5-fold validation Red Angus = correlation
Training on de-regressed EPDs Saatchi et al (GSE)*



AN: Angus	GV: Gelbvieh
BM: Beefmaster	LM: Limousin
BN: Brangus	MA: Maine Anjou
BR: Brahman	RA: Red Angus
BU: Braunvieh	SA: Salers
CA: Chiangus	SG: Santa Gertrudis
CH: Charolais	SH: Shorthorn
HH: Hereford	SM: Simmental
HL: Line 1 HH	

Approximate genetic distance between breeds using data from the 2,000 Bull Project.
Larry Keuhn, USDA MARC
<http://www.nbcec.org/topics/BeefBreeds.pdf>





So what about other breeds?

The following breed associations are working with Dorian Garrick (IA State) to develop their own 50K-based prediction equations



Breed	Breed code	# Training Records
Hereford	HER	1,725
Red Angus	RAN	296
Simmental	SIM	2,853
Brangus	BRG	896
Limousin	LIM	2,319
Gelbvieh	GVH	847
Maine Anjou	RDP	115



Predictions in Some Beef Breeds

(Data provided by Dorian Garrick)

# Records in training	Angus (3,500)	Hereford (800)	Simmental (2,800)	Gelbvieh (847)	Gelbvieh including Angus (1,181)
BirthWt	0.64	0.43	0.65	0.38	0.41
WeanWt	0.67	0.32	0.52	0.31	0.34
YearlingWt	0.75	0.30	0.45	0.21	NC
Milk	0.51	0.22	0.34	0.36	0.34
Fat	0.70	0.40	0.29	NA	NA
REA	0.75	0.36	0.59	0.38	0.48
Marbling	0.80	0.27	0.63	0.54	0.56
CED	0.69	0.43	0.45	NC	0.48
CEM	0.73	0.18	0.32	NC	NC
SC	0.71	0.28	NA	0.50	0.50



Simmental

2,800 records in training



Trait	r	ASA (r ² %)	PE
Calving Ease Direct	0.45	(20%)	5
Birth Weight	0.65	(42%)	6
Weaning Weight	0.52	(27%)	4
Yearling Weight	0.45	(21%)	3
Milk	0.34	(12%)	3
Maternal Calving Ease	0.32	(10%)	3
Stayability	0.58	(34%)	9
Carcass Weight	0.59	(35%)	4
Carcass Marbling	0.63	(40%)	4
Carcass Rib Eye Area	0.59	(35%)	4
Carcass Backfat Thick.	0.29	(8%)	1
Shear Force (Tenderness)	0.53	(28%)	6



**Submit a DNA Sample
Increase EPD accuracy***

If you are interested in knowing
more about your young animals,
ASA's new product can help.

***For more information,
see accompanying article**

\$90/sample

Genotyping performed at
GENESEEEK

Call or email Leoma Wells
406-587-4531
lwells@simmgene.com

THE FUTURE IS HERE!

By Wade Shafer Ph.D.

Approximately a year ago we ran an article in *SimTalk* to update readers about the state of genomic use in our industry. The article closed with the following paragraph: "Unless you have Angus cattle that are evaluated in the American Angus Association's genetic evaluation, currently available DNA tests are of little to no value to you for EPD enhancement; however, don't be dismayed if you don't fit this description. The recent evolution in genomic technology has spurred organizations to take the steps necessary to harvest its value. For example, ASA is presently working on the development of DNA tests specific to our population, as well as the capacity to incorporate test results into ASA's genetic evaluation."

One year later we are pleased to announce that the future has arrived at ASA. We are now offering a DNA test that, when incorporated into our genetic evaluation, can add significant enhancement to lower accuracy EPDs. The result of a multi-year collaborative effort, test development was fueled by ASA members and staff, carcass merit cooperators, USDA, University of Illinois, University of Missouri, Montana State University, GeneSeek and the National Beef Cattle Evaluation Consortium.

Test development was initiated several years ago through requests for donated semen on high accuracy sires. ASA members stepped up to the plate and delivered, with hundreds of DNA samples being collected via donation. Besides older sires, virtually all of the current 500 most heavily used sires were sourced. Samples were also collected on the sires and their offspring fed at the University of Illinois. In total, over 2,700 samples were genotyped.

The genotypes, along with the equivalent of millions of phenotypes (data submitted by breeders and the Carcass Merit Program over the years) were amassed and analyzed by University of Iowa researchers. The analysis resulted in the parameters required for genetic evaluation.

The multi-tiered effort culminated in a very successful outcome. The standard measure of a DNA test used in EPD prediction is the correlation between test results and the traits of interest. Table 1 displays correlations for ASA's test when used on ASA's population. To provide perspective, correlations for Pfizer and Igenty's tests gleaned from the American Angus Association's website are included in the table.

Table 1. Correlations between DNA test results and trait

Trait	Igenty ^a	Pfizer ^a	ASA ^b
CE	0.47	0.33	0.45
BW	0.57	0.51	0.65
WW	0.45	0.52	0.52
YW	0.34	0.64	0.45
Milk	0.24	0.32	0.34
MCE	NA	NA	0.32
Stay	NA	NA	0.58
CW	0.54	0.48	0.59
Mrb	0.65	0.57	0.63
REA	0.58	0.60	0.59
BF	0.50	0.56	0.29
SF	NA	NA	0.53

^aWhen used on the American Angus Association's population

^bWhen used on American Simmental/SimAngus™ population

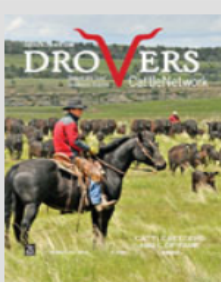
<http://www.simmental.org/site/pdf/register/April%2012%20Register%20The%20Future%20is%20Here.pdf>



Hereford announces genomic-enhanced EPDs (8/20/12)






1,200 records in training


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Hereford introduces genomic-enhanced EPDs

American Hereford Association | Updated: August 20, 2012

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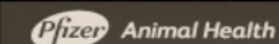
The American Hereford Association (AHA) is the first beef breed to develop and market its own genomic predictions.

The AHA approach is the first of its kind to work with the scientific community and the National Beef Cattle Evaluation Consortium (NBCEC) to build its own training and validation population. This approach is important because AHA now has access to all of the genotypes, phenotypes and pedigrees, which will allow the Association and its members to continue to train and build the Hereford-specific panel.


Today more than 1,200 Herefords have been 50k genotyped and eight sires whole-genome sequenced. This population has been used to train and validate the Hereford-specific panel. Dorian Garrick, Iowa State University Lush Chair in animal breeding and genetics and NBCEC executive director, has led the research and development.

According to Garrick AHA took a pro-active approach testing and validating the tools available to make sure when releasing a Hereford genomic product it was reliable and useful to AHA members. During this process the Association aligned itself with researchers and worked collaboratively with the NCBE, U.S. Department of Agriculture (USDA) Meat Animal Research Center and other global Hereford associations.

In late June AHA transitioned its DNA testing to GeneSeek Inc. located in Lincoln, Neb. With this move, Hereford breeders have the option to utilize a 50k panel to obtain genomic information to enhance the accuracy of Hereford expected progeny differences (EPDs). The genomic information obtained is blended with conventional EPDs to produce genomic-enhanced EPDs (GE-EPDs). GE-EPDs are available on all traits reported by the AHA.






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- USDA feeder cattle market: Yearling steady

<http://www.cattlenetwork.com/e-newsletters/drovers-daily/166794826.html>



Limousin plans to start using DNA in EPDs in December 2012

2,400 (?) records in training

Genomic information is the next step in technology for the beef industry. The North American Limousin Foundation has done extensive work over the last few years in cooperation with Igenity Inc. (now part of GeneSeek Inc.) to develop a Limousin specific 50,000 SNP (pronounced 'snip'; single-nucleotide polymorphism) panel. This test goes beyond the DNA marker tests that were introduced years ago, which included only a small number of markers and were unable to generate values that could be incorporated in National Cattle Evaluations (NCE).

Each SNP alone does not reveal much information. It is only through study of the quantitative effects of a number of critically selected SNPs that generates molecular breeding values (MBVs) for each trait. These MBVs will be factored into the interim EPDs of the animal as well as the NCE based upon the correlation of the MBV to the trait. MBVs will be available for every EPD trait and will be beneficial through increased EPD accuracy of traits that are either difficult or expensive to measure such as Stayability and the carcass traits.

It is important to note that even though a genomic panel yields increased accuracy to EPDs, it is still only one tool in the genetic evaluation toolbox. Weights, measurements, ultrasounds and docility scores will still be vital to build a complete picture of an animal's genetic merit. Those measures will also be crucial to the updating and expanding on the accuracy of the panel and its genetic correlations.

Once animal has had a sample submitted to NALF for the Limousin Genomic Profiler-Igenity® Enabled, the sample will be sent to GeneSeek, Inc. where a panel will be run. NALF will receive the MBVs which will then be factored into the animal's EPDs starting in December 2012 and the NCE soon after. The animal's owner will receive a report with the Igenity 1-10 scores on that animal similar to their earlier Beef Profile report. NALF would recommend that selection decisions be made based on the Genomic-Enhanced EPDs as they are a more accurate tool for determining genetic merit.

http://www.nalf.org/pdf/2012/dna-genomics/Limousin_Genomic_Profiler_and_Genomic-Enhanced_EPDs_7.27.12.pdf

The Future

NEXT EXIT





National Colorado State University-Cornell University-University of Georgia-Iowa State University
Beef Cattle Evaluation
Consortium

NBCEC projects <http://www.nbcec.org>



- Genomic selection in beef cattle (Garrick - IA)
- Single-step approach to incorporating genomic information into EPDs (Misztal - GA)
- Reducing data loss and improving accuracy in National Cattle Evaluation (Bertrand - GA)
- Industry effort to enhance beef cattle selection (aka Weight Trait Project) (Pollak/Spangler - NE)
- Developing selection tools for longevity and reproduction (Pollak – US MARC)
- Genetics of feedlot cattle health (Enns, CO – funded by Pfizer)
- Healthfulness of beef project (Reecy, IA – funded by Pfizer)



USDA funded projects – competitive call for proposals: Part 1 (old CREES proposals)



- **Identification of molecular markers to improve fertility of beef cattle** (Thomas, CO) – finished 12/2011
 - >10,000 DNA samples and phenotypes on heifers
- **BIGS – Bioinformatics to implement genomic selection** (Garrick/Snelling/Golden)/ **Enhanced Bioinformatics to implement genomic selection (BIGS)** (Garrick, Dekkers, Fernando, Reecy, Rothschild) finishes 4/30/2014 See website <http://big.s.ansci.iastate.edu/>
- **Integrating DNA information into beef cattle production systems** (Van Eenennaam, CA) – finishes 12/2013
 - 7,000 DNA samples & weaning weight records, 4500 carcass records



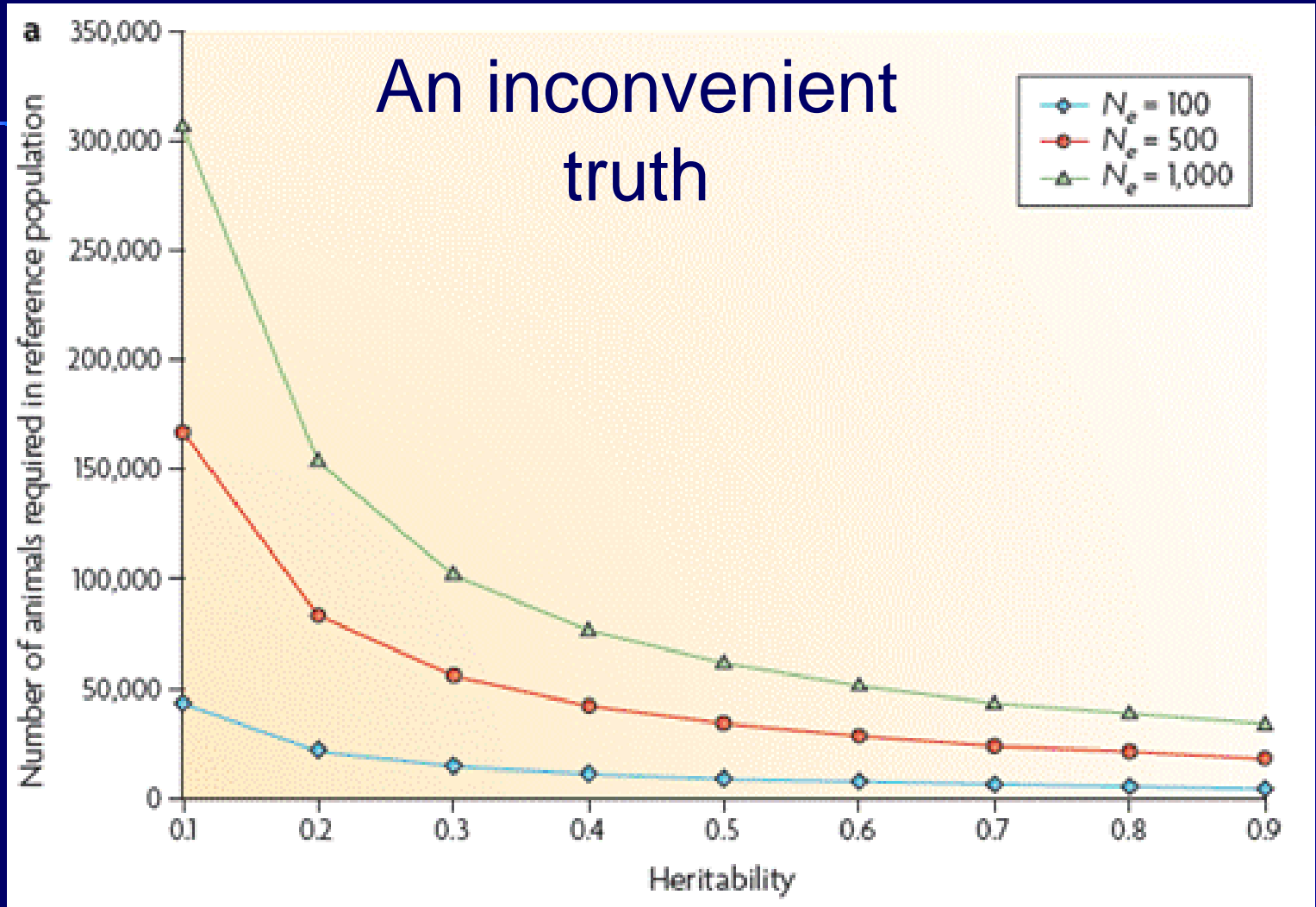
USDA funded projects – competitive call for proposals: Part 2 (new NIFA AFRI proposals)



- **National program for genetic improvement of feed efficiency in beef cattle**
(Taylor, MO) – finishes 4/2016 (<http://www.beefefficiency.org>)
 - Genotype ~ 2,400 head on HD chips; 7000 records FE records
 - \$5 million, 5 year project; April 2011 – April 2016
- **Integrated program for reducing bovine respiratory disease (BRD) in beef and dairy cattle** (Womack, TX) – finishes 4/2016 (<http://www.brdcomplex.org>)
 - Collection and HD genotypes on 6,000 BRD case-control animals
 - \$10 million, 5 year project; April 2011 – April 2016
- **Identification and management of alleles impairing heifer fertility while optimizing genetic gain in Angus cattle**
(Patterson, Taylor, et al. MO; Van Eenennaam, CA) – finishes 12/17
 - Sequence up to 200 cattle from up to 10 different beef breeds
 - \$3 million, 5 year project; Jan 2013 – December 2017



Effect of population size and heritability on the number of animals required in the training population (for an accuracy of 0.7)



Goddard, M. E., and B. J. Hayes. 2009. Mapping genes for complex traits in domestic animals and their use in breeding programmes. *Nature Reviews Genetics* 10: 381-391.

US cattle numbers (x 10⁶)

US Beef operations 766,350
Million Cows 31.4
Average herd size 41

Breeder

1.1

Seedstock Cows

**Commercial
cow/calf
producer**

35.7

Commercial Cows + replacements

Feedlot

13.6 (on feed at any one time)
25.6 (cattle fed per year in 2009)

Processing

43.2

Retailer

Other Beef Animals (calves, steers, heifers and bulls)




Extracting DNA multiple times in different labs for different applications makes about as much sense as simultaneously paying to have access to all of the following communication devices





TECHNOLOGICAL CHALLENGE: How do we collect phenotypes and DNA on all animals and make genotype information available to all sectors?

ONLY THESE SECTORS PRODUCE NEW ANIMALS



Use	Seedstock	Commercial	Feedlot	Processor
DNA-assisted selection	XXXX	X	XXXX	XXXX
Parentage	XX	X		
Recessive allele testing	XX	X		
Control of Inbreeding	XX	X		
Mate selection	XX	X		
DNA-assisted management/purchasing		X	XX	XX
Product differentiation				XXXX
Traceability				XX

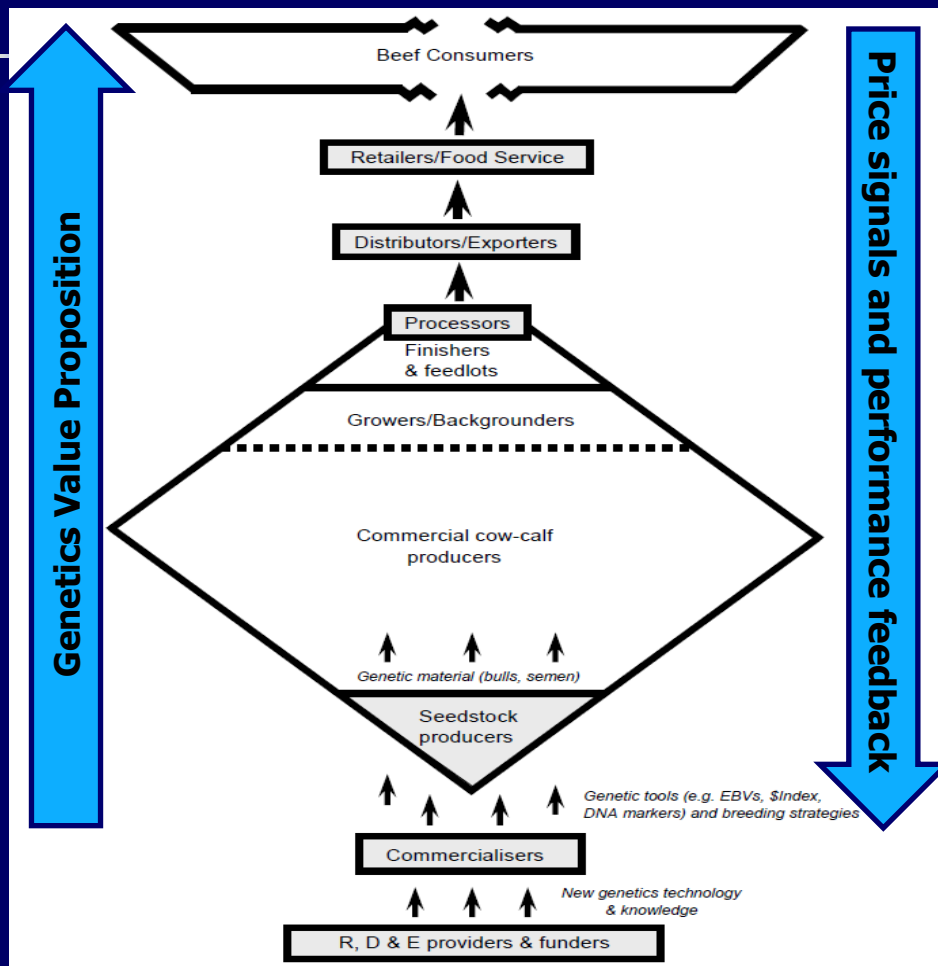


Ideally cattle would be genotyped ONCE early in life and genotypes shared with downstream production sectors to derive the maximum value from the fixed DNA collection and extraction costs

Cattle industry Sector	Type of DNA product // DNA information access required	Cost?? (\$US)	Mobile Device// Data Access Plan
Nucleus seedstock/AI bulls	Full genome sequence	\$250	ipad
Seedstock/bull multiplier	HD 770 K genotype	\$50	iphone
Registered females and stock bulls for commercial sector	50K genotype + parentage + single gene traits/recessives	\$25	Talk and text smart phone
Commercial cattle – Marker-assisted management (MAM), replacement heifer selection	Imputation LD chip + parentage + single gene traits/recessives	\$10	Prepaid cellular phone
Feedlot cattle purchasing, sorting and marker-assisted management (MAM)	Access genotypes from supplier (subset of LD imputation chip).	<\$1	Pay as you go contract
Traceability for voluntary labelling e.g. Angus beef	Access genotypes from supplier (subset of LD imputation chip).	<\$1	Friends and family plan
Traceability for disease outbreak/contaminated meat	Access genotypes from supplier (subset of LD imputation chip).	<\$1	Emergency only phone (911 calls)

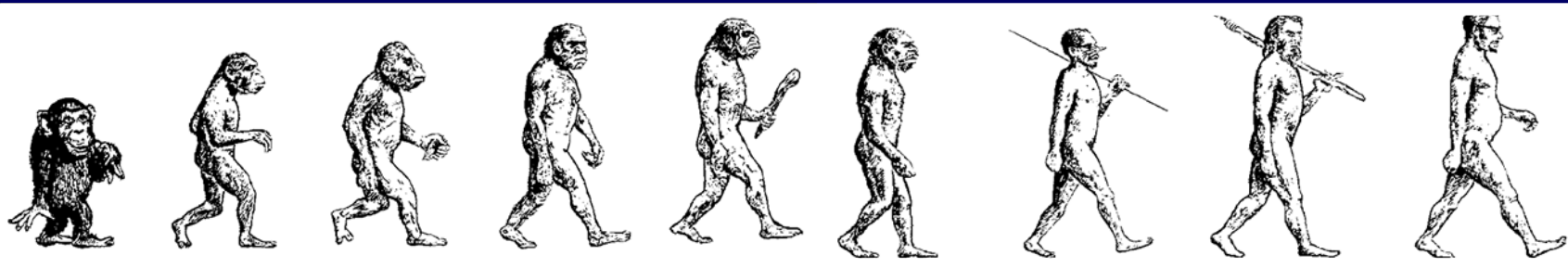


Industry structure may evolve to enable the exchange of information and value between the different sectors.



For widespread technology adoption, breeders need to be adequately rewarded for making DNA investments and selection decisions for traits that benefit the different sectors of the beef industry.

Parnell, P.F. 2007. Effective value chain partnerships are essential for rapid adoption of beef genetics technology. Association for the Advancement of Animal Breeding and Genetics. 18. 167-174 .



2003

2008

2013

2020

- single marker/
single trait
- reported
genotypes
- single
marker
accounted
for small
amount of
genetic
variation
- limited
adoption
- technology
oversold

- multimarker tests
for a few traits
reported in a
variety of formats
- no tie between
DNA test results
and national
genetic evaluation
or breed
associations
- tests accounted
for < 10% additive
genetic variation
- limited validation
- technology not in
a form producers
could use

- panels with
thousands of markers
for many traits
- results reported in
units of the trait
- incorporation of
DNA information into
national genetic
evaluation
- DNA-based
evaluations improve
accuracy of EPDs
- large numbers of
genotyped
populations are
available for
calibration/validation

- universal marker
panel used by
worldwide beef cattle
community
- seamless submission
of genotype data to
national genetic
evaluation/breed
associations
- testing cost is low
- DNA information
used for traceability,
parentage, genetic
defects, selection,
marker-assisted
management, product
differentiation





Ramifications of genomics to beef industry



- The benefits of genomics are best captured in well-structured industries that are already making significant genetic progress
- May encourage more vertical integration to collect phenotypes to enable predictions for economically relevant traits for all sectors
- May see genetic evaluations developed for novel traits – if large enough phenotyped populations can be amassed
- Feedlots stand to gain the most from collecting phenotypes and sharing data back with suppliers
- Will beef follow the pig/poultry model of developing a single breeding objective optimized for industry good?



Concluding thought....



Breeds/groups that can organize themselves and **technologically** and **structurally** to seamlessly obtain and marry entire supply chain phenotypes and genotypes and take advantage of the rapidly-declining cost of genotyping to capture the cumulative value derived from using genomic information for multiple purposes (selection, parentage, genetic defects, marker-assisted management, product differentiation, traceability) will be ideally positioned to fully realize the nascent potential of genomic information.

Van Eenennaam, A. L., and D. J. Drake. 2012. Where in the beef cattle supply chain might DNA tests generate value? *Animal Production Science*. 52(3) 185-196.



Questions?