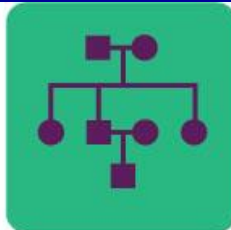







# Integrated Program for Reducing Bovine Respiratory Disease in Beef and Dairy Cattle

Alison Van Eenennaam, Ph.D.

Cooperative Extension Specialist  
Animal Biotechnology and Genomics  
Department of Animal Science  
University of California, Davis  
[alvaneennaam@ucdavis.edu](mailto:alvaneennaam@ucdavis.edu)  
US Bovine Respiratory Disease  
Coordinated Agricultural Project  
<http://www.brdcomplex.org>



Bovine Respiratory Disease Complex  
Coordinated Agriculture Project



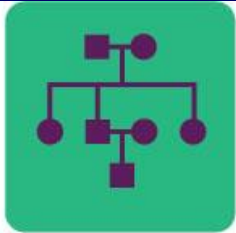
United States  
Department of  
Agriculture

National Institute  
of Food and  
Agriculture

The “Integrated Program for Reducing Bovine Respiratory Disease Complex (BRDC) in Beef and Dairy Cattle” Coordinated Agricultural Project is supported by Agriculture and Food Research Initiative Competitive Grant no. 2011-68004-30367 from the USDA National Institute of Food and Agriculture.



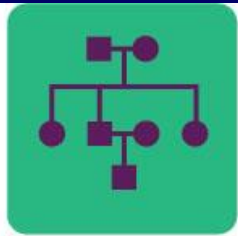
# Overview



Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

- Need for novel approaches for BRD
- What is the BRD Coordinated Agricultural Project
- What is needed to develop accurate genomic/DNA tests
- Work of BRD CAP to address these needs
- What is the relative economic value of BRD
- Implications to beef cattle breeders
- Beef cattle industry structure implications





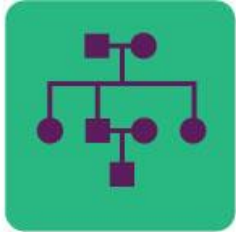
Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

# Background and Rationale

*"Year in and year out, diseases of the respiratory system are a major cause of illness and death in cattle from 6 weeks to two years of age. Sadly, this is as true today as it was 30 years ago despite development of new and improved vaccines, new broad spectrum antibiotics, and increased fundamental knowledge as to the cause of disease"*

- Bovine Respiratory Disease (BRD) has been extensively studied since the 1800s, and yet it remains prevalent
- More effective vaccines have not decreased the morbidity or mortality of BRD
- Mortality has increased as vaccine efficiency has increased
- 1.4% of all US feedlot cattle perish before reaching harvest weight
- **Need to develop new approaches to tackle BRD**

Montgomery, D. 2009. Bovine Respiratory Disease & Diagnostic Veterinary Medicine. Proceedings, The Range Beef Cow Symposium XXI. December 1, 2 and 3 2009, Casper, WY. Pages 1-6.



Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

# BRD Coordinated Agricultural Project

*Long-term goal is to reduce the incidence of BRD in beef and dairy cattle by capitalizing on recent advances in genomics to enable novel genetic approaches to select for cattle that are less susceptible to disease*







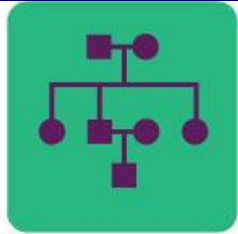
Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

# Potential benefits of genomics are greatest for economically-important traits that:

- Are difficult or expensive to measure
- Cannot be measured until late in life or after the animal is dead
- Are not currently selected for because they are not routinely measured
- Have low heritability

Yep, looks like all of 'em were susceptible





Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

# Disease resistance is a very attractive target trait for genetic improvement

- The presence of genetic variation in resistance to disease, coupled with the increased consumer pressure against the use of drugs, is making genetic solutions to animal health problems increasingly attractive.
- The non-permanent effectiveness of chemical agent (due to development of resistance by the pathogen) further contributes to this interest.

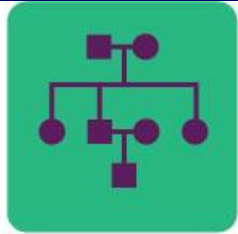
Newman, S. and Ponzoni, R.W. 1994. Experience with economic weights. Proc. 5<sup>th</sup> World Congress on Genetics Applied to Livestock Production. 18:217-223.



# Other animal industries have successfully targeted selection for disease resistance

- In dairy cattle, selection programs have been developed to take advantage of genetic variability in mastitis resistance, despite the fact that the heritability of clinical mastitis is low and mastitis resistance has an adverse correlation with production traits
- Likewise chicken breeders have long used breeding to improve resistance to avian lymphoid leucosis complex and Marek's disease

Stear, M. J., S. C. Bishop, B. A. Mallard, and H. Raadsma. 2001. The sustainability, feasibility and desirability of breeding livestock for disease resistance. *Res Vet Sci* 71: 1-7



Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

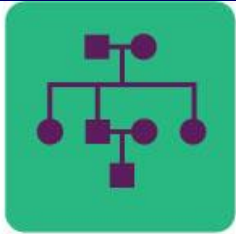
# What is needed to develop DNA-tests for BRD susceptibility?

Large training/discovery populations with BRD observations and SNP genotypes = used to estimate the value of every chromosome fragment contributing variation BRD susceptibility. This allows for prediction of which chromosome segments regions are important for the trait.

**Prediction equation** = the results of training can then be used to predict the genetic merit of new animals, not contained in the training data set





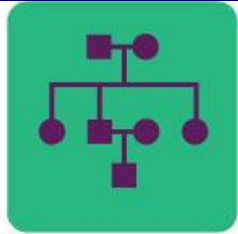


Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

# Need for large discovery populations

- The ready availability of dense single nucleotide polymorphism arrays (i.e. 700 K SNP chips) has given rise to hitherto unforeseen opportunities to dissect host variation and identify possible genes contributing to this variation using genome wide association studies
- To have the power to meaningfully quantify genetic variation or perform a genome scan using a dense SNP chip it is necessary to have datasets comprising observations on several thousands of individuals.

Bishop, S. C., and J. A. Woolliams. 2010. On the genetic interpretation of disease data. Plos One 5: e8940.



Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

# Need for careful “case” definition

- For studies of infectious diseases this usually necessitates utilizing field data because challenge experiments of a sufficient scale will not be possible.
- However, such field data is very ‘noisy’
  - diagnosis of infection or disease may be imprecise; it can be difficult to determine when infection of an individual occurred
  - it is often unclear whether or not apparently healthy individuals have been exposed to the infection
- These factors add environmental noise to the epidemiological data.

Bishop, S. C., and J. A. Woolliams. 2010. On the genetic interpretation of disease data. Plos One 5: e8940.



# Accurate diagnosis (i.e. case definition) of BRD is critical for success of studies

- Traditional methods for detecting morbid cattle include visual appraisal once or twice daily.
- Animals displaying nose or eye discharge, depression, lethargy, emaciated body condition, labored breathing or a combination of these, should be further examined
- Symptomatic animals with a rectal temperature  $\geq 103^{\circ}\text{F}$  are usually considered morbid and given treatment.
- All of these diagnostic systems are **subjective in nature**.
- Confounding factors include the diligence and astuteness of those checking the animals, the variability and severity of the symptoms the animals experience with chronic and acute BRD, and the disposition of the animals





# Accurate diagnosis (i.e. case definition) of BRD is critical for success of studies

- Animals with docile temperaments are more likely to be diagnosed than aggressive or flighty animals
- In one study pulmonary lesions were examined at slaughter in an attempt to confirm the live BRD diagnosis
  - 35% of feedlot steers received treatment for BRD between birth and slaughter, whereas 72% had pulmonary lesions at slaughter.
  - For steers treated for BRD, 78% had pulmonary lesions, whereas 68% of untreated steers also had pulmonary lesions.
- Another study also found that many (29.7%) apparently healthy animals had subclinical disease.
- These studies suggests that the morbidity estimates derived from clinical observations of pen riders likely underestimate the incidence of BRD in feedlot cattle



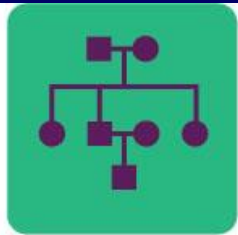


# BRD CAP: BRD field datasets

## Case:control field datasets are being developed for bovine respiratory disease

- 6000 animals – case:control design
  - 2000 dairy calves diagnosed on a collaborating dairy calf rearing ranch (CA)
  - 2000 feedlot cattle diagnosed on a collaborating feedlot (TX)
  - 1000 dairy (NM) and 1000 beef (NV) case:control animals will be used to validate loci associated with BRD in the discovery populations
- All will be genotyped on 700K high density SNP chip



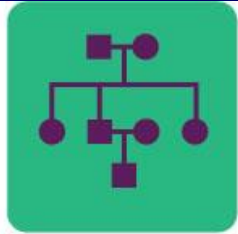


Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

# Year 1: CA Dairy Calf Ranch: 70,000 head capacity



**Jessica Davis, DVM**  
Intern at Veterinary Medicine  
Teaching and Research  
Center, University of  
California, Davis; Tulare  
**Terry Lehenbauer, DVM**  
**Sharif Aly, DVM**  
**Pat Blanchard, DVM**  
California Animal Health and  
Food Safety Laboratory System















Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

# Standardization of BRD Diagnosis

- 1000 case and 1000 control 30-60 day old calves
- Use Dr. Sheila McGuirk's calf respiratory scoring chart
  - Temperature, eyes, ears, nose, +/- cough
  - Additional clinical signs: tachypnea, dyspnea, position of head, appetite
  - Give score and either enroll or not (5 or greater to enroll as case)
- Sample collection
  - Blood for DNA extraction and high density SNP genotyping
  - Nasal swab and deep pharyngeal swab to identify viruses (PCR: IBR, BVD, BRSV, and Corona) and bacteria (*Manheimia haemolytica*, *Pasteurella multocida*, and *Histophilus somni*, and *Mycoplasma* spp.) present in the nasopharyngeal and pharyngeal recesses





Calf Health Scoring Criteria			
0	1	2	3
<b>Rectal temperature</b>			
100-100.9	101-101.9	102-102.9	≥103
<b>Cough</b>			
None	Induce single cough	Induced repeated coughs or occasional spontaneous cough	Repeated spontaneous coughs
<b>Nasal discharge</b>			
Normal serous discharge	Small amount of unilateral cloudy discharge	Bilateral, cloudy or excessive mucus discharge	Copious bilateral mucopurulent discharge
			
<b>Eye scores</b>			
Normal	Small amount of ocular discharge	Moderate amount of bilateral discharge	Heavy ocular discharge
			
<b>Ear scores</b>			
Normal	Ear flick or head shake	Slight unilateral droop	Head tilt or bilateral droop
			





Blood collection



Deep  
pharyngeal  
swab  
collection

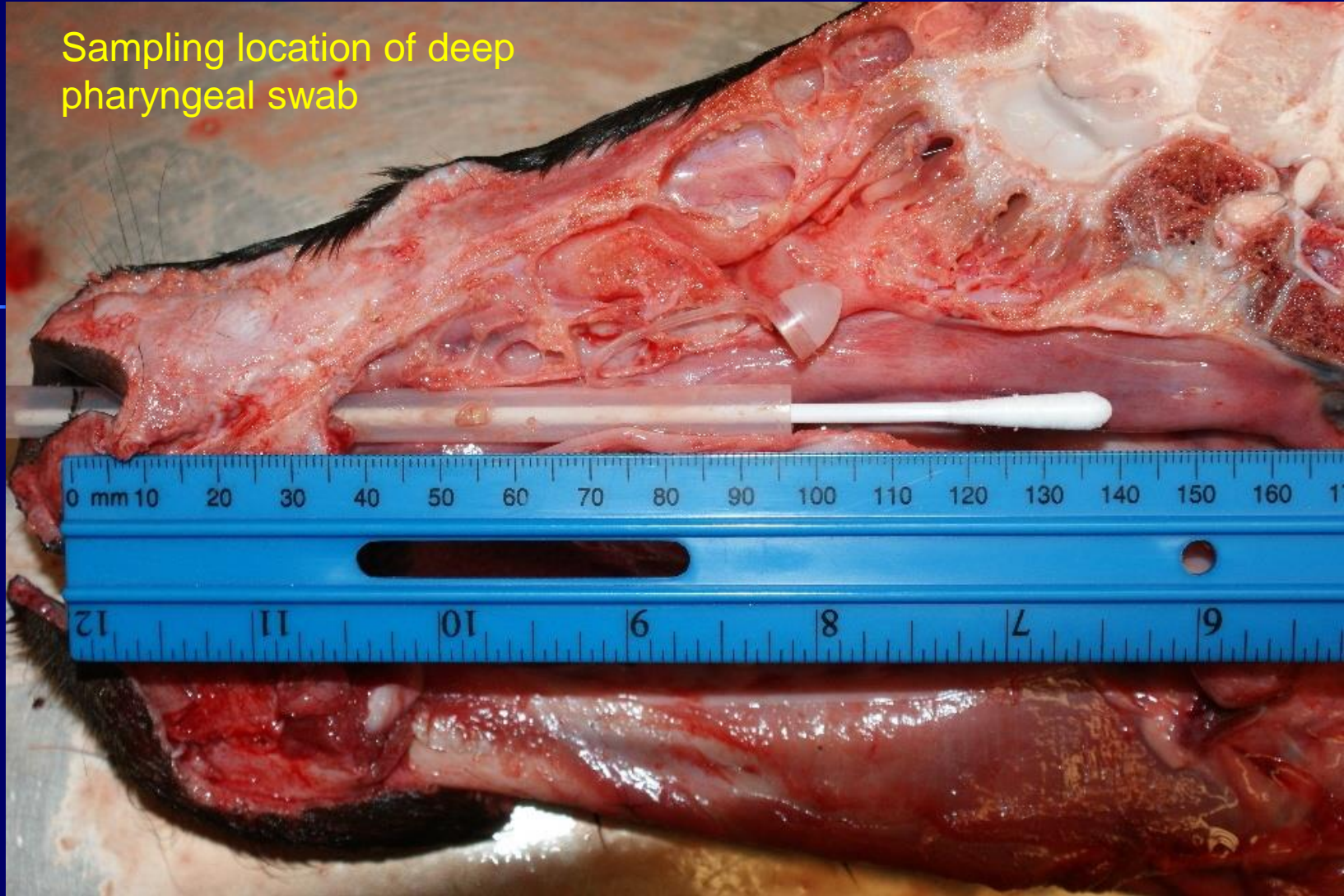


Nasal swab

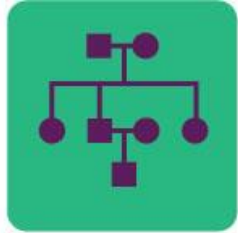




Sampling location of deep  
pharyngeal swab



To culture organisms associated with BRD, pharyngeal swabs offer a less invasive, less stressful and more rapid alternative to bronchoalveolar lavage.



Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

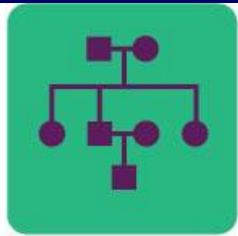
# Controls

- Score control in same way as cases (score of 4 or less)
- Try to select animals in the adjacent hutch, same dairy of origin, and same sex
- Collect samples for control animals in same way as case

**Objective:** Try to identify cases and controls in a relatively constant environment, subjected to the same exposure and stresses, to decrease the environmental “noise” of these field BRD datasets







Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

# Year 2: TX Feedlot Gonzalez, Texas

**Sample collection (1000 case and 1000 controls)  
scheduled to be completed by 3/2013 and analysis of  
genotype data completed by 12/31/2014**



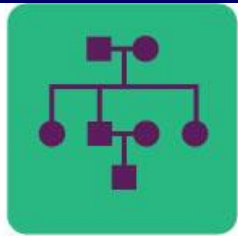
**Cole McQueen, Masters student  
Dr. Noah Cohen  
Dr. Scott Dindot  
Texas A&M University**



# The Future

NEXT EXIT

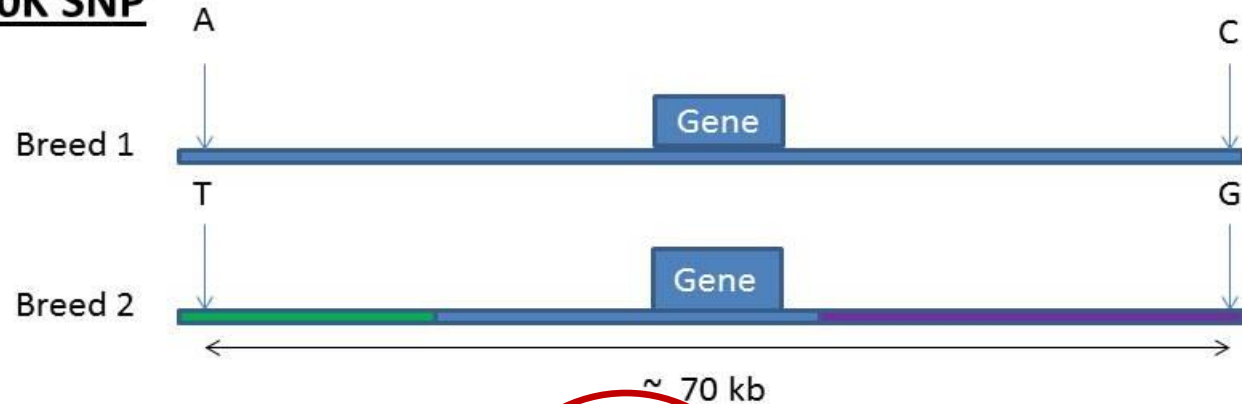




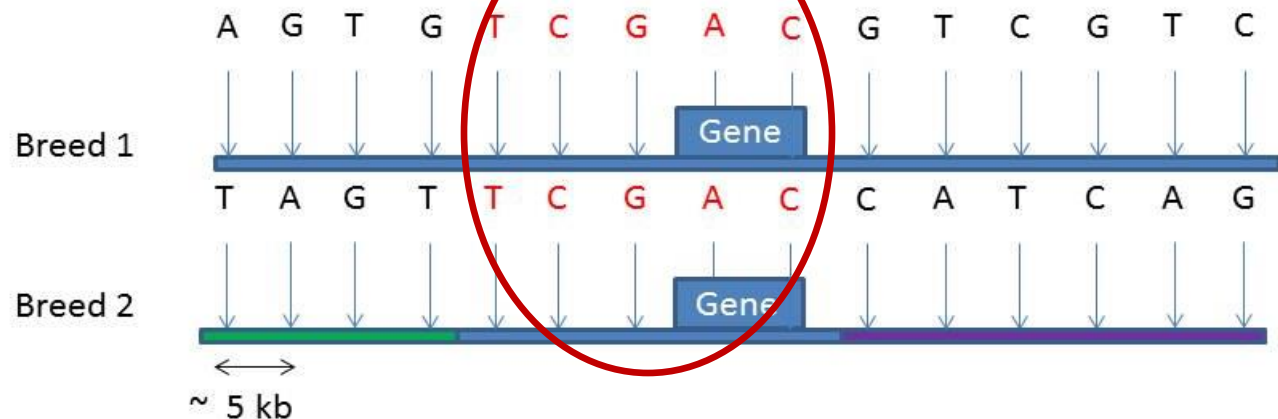
Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

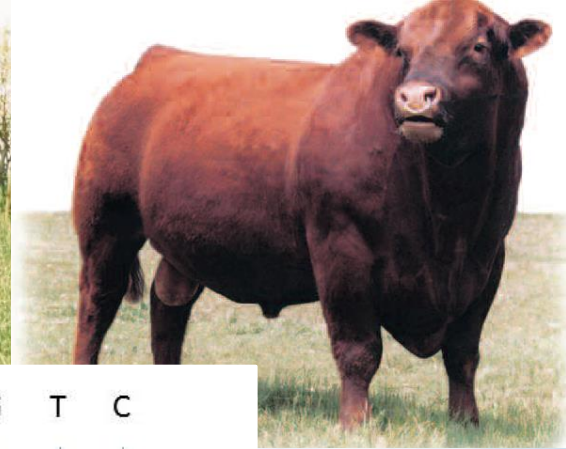
# Hopeful that high density SNP chips will enable tests to work across breeds

## A. 50K SNP



## B. 700K SNP









# What would be the value of selecting for cattle that are less susceptible to BRD?

**OBJECTIVE:** To calculate the weighting (i.e. relative economic value) that should be given to selection for BRD resistance in a multi-trait selection index for Angus terminal sires.

Van Eenennaam, A. L. and M. D. MacNeil. 2011. **What weighting should be given bovine respiratory disease (BRD) resistance in selection decisions?** Pages 61-68 in Proceedings of the Beef Improvement Federation 43<sup>rd</sup> Annual Research Symposium and Annual Meeting. <http://www.brdcomplex.org/Links/BIF2011.pdf>





# Determining the value of using genomics to improve the accuracy of selection

- Genomic information should not require any fundamental changes to the development of breeding objectives and selection index calculations
- Selection emphasis for new genomically-enabled traits will need to be weighted by effect on profitability relative to other economically-important traits
- Therefore need to determine what is the relative economic value of these new traits in the breeding objective?





# Scenario



- A 1000–cow-calf enterprise with retained ownership
- All progeny of terminal sires were harvested and so no economic value was associated with maternal traits
- Only phenotypes for weaning weight, feedlot average daily gain, feed intake, USDA yield grade, marbling score and BRD incidence (%) contributed to the breeding objective.
- The feedlot phase was divided into three periods. The first period (backgrounding) was terminated at a weight-constant end point of 850 lb.
- The second (growing) and third (finishing) periods were of 50 and 100 days duration, respectively.



# Prices, premiums and discounts used in developing the multi-trait selection index for Angus terminal sires.



<b>Weaned calf weight (lb)</b>	<b>\$/lb</b>		<b>\$/100lb</b>
<b>&lt; 350</b>	1.21	Prime	28.07
<b>351-400</b>	1.15	High Choice	5.53
<b>401-450</b>	1.09	Choice	0
<b>451-500</b>	1.04	Select	-10.20
<b>501-550</b>	1.01	Standard	-20.20
<b>551-600</b>	0.96		
<b>&gt;600</b>	0.92	Yield Grade 1	3.00
<b>Carcass weight (lb)</b>	\$/100 lb	Yield Grade 2	2.00
<b>Base price</b>	155.95	Yield Grade 3	0.00
<b>&lt;550</b>	-15	Quality/Yield Grade	-10.20
<b>&gt;950</b>	-15	Yield Grade 5	-20.20



# Assumptions



- BRD occurred when calves were moved to the feedlot phase at weaning – incidence rate of 10%
- the fixed cost of feedlot phase was unchanged
- a dead calf incurred no feed costs
- there was a 10% mortality from BRD
- there was a 13% reduction in ADG (1.3 lbs/d) for the first phase of feeding (weaning to 850 lbs)
- final yield grade was reduced by 0.1
- the cost to diagnose and treat a BRD calf was \$44





# Results: Decreasing BRD incidence most valuable traits in terminal sire selection index



<b>Trait (unit)</b>	<b>Economic Value (\$)</b>	<b>Genetic SD</b>	<b>Relative economic value (REV)</b>	<b>Relative importance (relative to YG)</b>
<b>BRD incidence (%)</b>	<b>-8424.7</b>	<b>7.94</b>	<b>-66892</b>	<b>37.7</b>
<b>Weaning wt. (lb)</b>	241.4	41.76	10081	<b>5.7</b>
<b>Feed Intake (lb/d)</b>	-5811.8	1.41	-8195	<b>4.6</b>
<b>Feedlot ADG (lb/d)</b>	27654.5	0.24	6637	<b>3.7</b>
<b>Marbling score</b>	8926.0	0.51	4552	<b>2.6</b>
<b>Yield Grade</b>	-5379.2	0.33	-1775	<b>1</b>

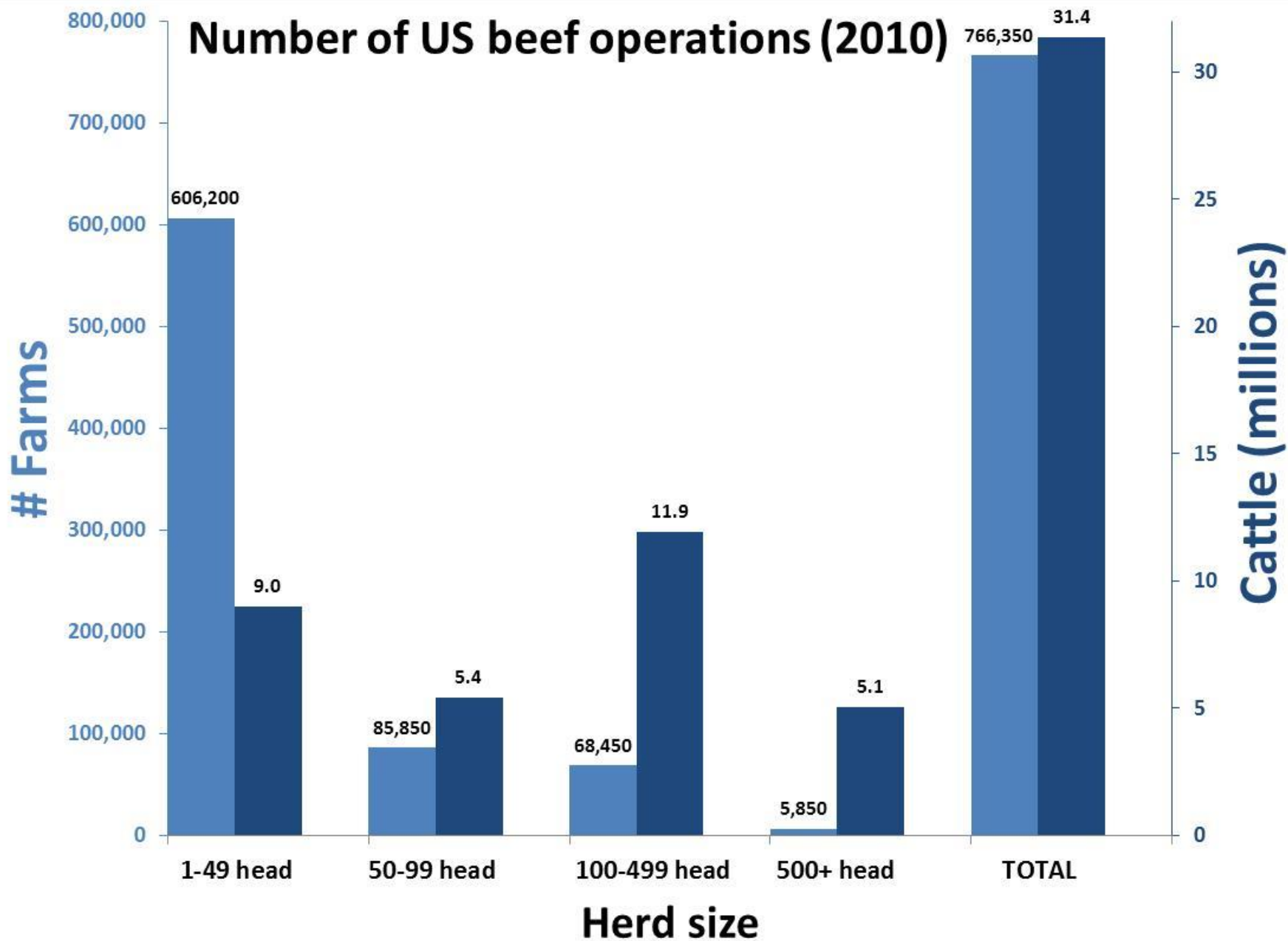


# Ninety percent of US cattle operations have fewer than 100 head, and most sell their cattle at auction prior to feedlot entry

**This relative economic value scenario was developed to maximize the profitability of a vertically-integrated (retained ownership) production system**

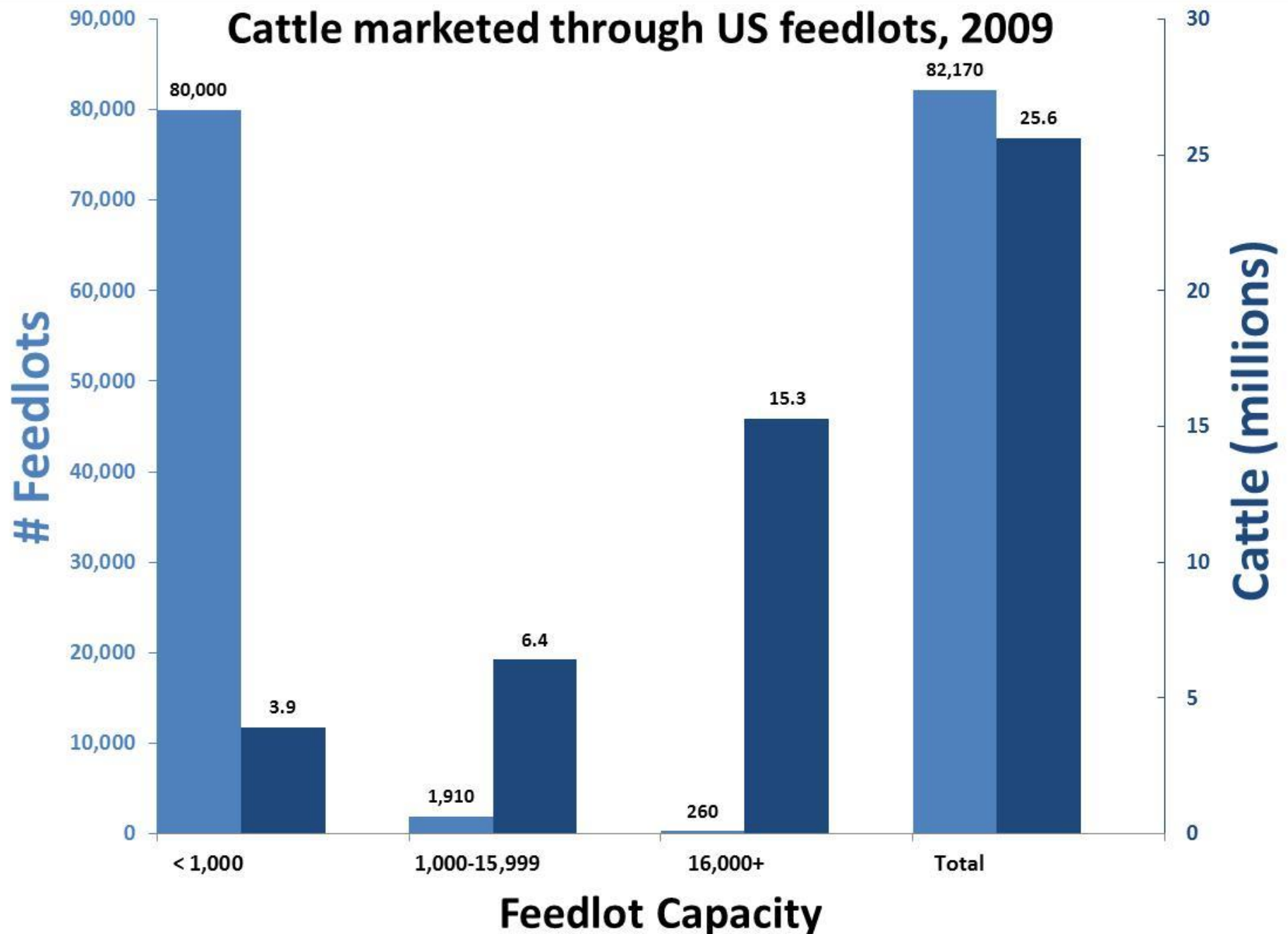
- In reality most producers' financial returns are tied very closely to the number of calves, a function of reproduction, and less to feedlot performance and carcass traits, and even less to bovine respiratory disease incidence.
- To incentivize the inclusion of BRD resistance in selection decisions, a mechanism analogous to a calf preconditioning bonus would be needed to equitably share some of the value derived from reduced feedlot disease incidence and to compensate breeders and producers for reducing selection emphasis on other economically-relevant traits.





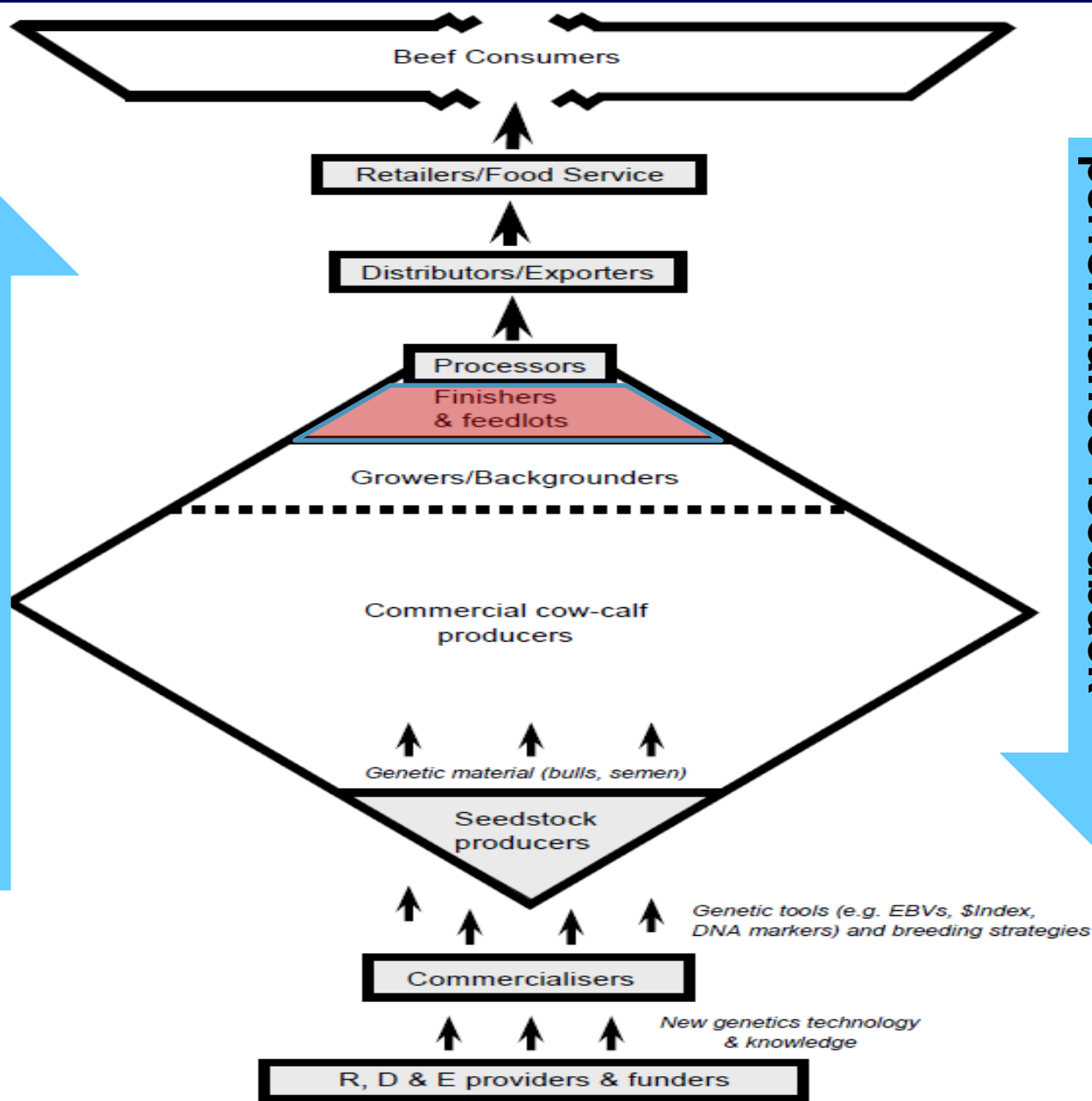


# Cattle marketed through US feedlots, 2009

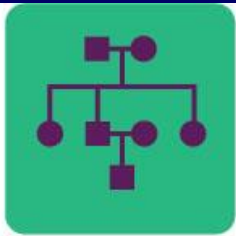




Genetics value proposition



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 .



Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

# BRD Coordinated Agricultural Project



**TEXAS A&M**  
UNIVERSITY

- **Jim Womack, PD**
- Alan Dabney
- Scott Dindot
- Noah Cohen
- Chris Seabury
- Lawrence Falconer
- Lauren Skow
- Gary Snowder

**UCDAVIS**  
UNIVERSITY OF CALIFORNIA

- Laurel Gershwin
- Terry Lehenbauer
- Cassandra Tucker
- **Alison Van Eenennaam**

**Colorado State University**  
United States Department Of Agriculture  
Agricultural Research Service

- Mark Enns



- Jerry Taylor

**WASHINGTON STATE**  
UNIVERSITY

- **Holly Neibergs**
- Shannon Neibergs



- **Milt Thomas**
- Robert Hagevoort
- Tim Ross

## OTHER COLLABORATORS

- Daniel Pomp (NC)
- Shiela McGuirk (WI)
- Adroaldo Zanella (Norway)





Bovine Respiratory Disease Complex  
Coordinated Agriculture Project

# BRD Coordinated Agricultural Project





# CONCLUSIONS



- USDA National Institute of Food and Agriculture has recently funded two large 5-year, multi-institution grants on genomic approaches to feed efficiency and BRD
- Both projects employ high-density genotyping of large numbers of animals
- Both traits are valuable – especially to the feedlot sector
- Need to derive relative economic value of these traits and include them in breeding objectives
- Will beef industry organize increased sharing of feedlot performance data and value to drive investment in using genomics to make genetic improvement in feedlot traits?



# Parting thought....

Breeds/groups that can organize themselves to take advantage of the rapidly-declining cost of genotyping and capture the cumulative supply chain 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.







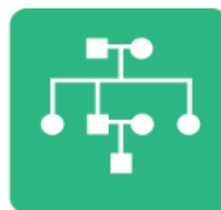
**[www.brdcomplex.org](http://www.brdcomplex.org)**



**United States  
Department of  
Agriculture**

**National Institute  
of Food and  
Agriculture**

The “Integrated Program for Reducing Bovine Respiratory Disease Complex (BRDC) in Beef and Dairy Cattle” Coordinated Agricultural Project is supported by Agriculture and Food Research Initiative Competitive Grant no. 2011-68004-30367 and the DNA value determination project was supported by National Research Initiative competitive grant no. 2009-55205-05057 (“Integrating DNA information into beef cattle production systems”) from the USDA National Institute of Food and Agriculture.



## Bovine Respiratory Disease Complex

### Coordinated Agricultural Project

**BRD Complex**  
Facts

**Prevention**  
What you can do

**Education**  
Teaching Materials

**Links**  
Resources

**About**  
Our Research

We are a collaborative group of researchers whose goal is to reduce the prevalence of bovine respiratory disease complex in beef and dairy cattle for the improvement of animal welfare and profitability. The "Integrated Program for Reducing Bovine Respiratory Disease Complex (BRDC) in Beef and Dairy Cattle" Coordinated Agricultural Project is supported by [Agriculture and Food Research Initiative Competitive Grant no. 2011-68004-30367](#) from the USDA National Institute of Food and Agriculture. Our project is led by [Dr. James Womack of Texas A&M University](#) and includes scientists and educators from the University of California-Davis, Colorado State University, the University of Missouri, New Mexico State University, Washington State University and USDA's Agricultural Research Service.



United States Department of Agriculture  
National Institute of Food and Agriculture

### Researchers

#### Sharif Aly

Assistant Professor  
School of Veterinary Medicine  
University of California Davis  
[saly@ucdavis.edu](mailto:saly@ucdavis.edu)

#### Noah Cohen

Professor  
Department of Large Animal Medicine and Surgery  
Texas A&M University.  
[ncohen@cvm.tamu.edu](mailto:ncohen@cvm.tamu.edu)

**Research Plan (PDF 6MB)**

**Our Researchers**

**Our Advisory Board**

**Calendar of Events (PST)**

**Grant  
Announcements**