



Disease Resistant Genetics

Steven Larmer, PhD



Disease Resistant Genetics

The only company offering
immune genetics for
greater disease resistance





Immunity+[®]



**2013 USA Dairy Herd Management
Innovation Award**



**2017 Canada Governor
General's Innovation Award**



**2018 UK Royal Dairy Innovation
Award Winner**

Dr. Bonnie Mallard

Department of Pathobiology, University of Guelph

Genetic regulation of the immune system of livestock

22-yr research program Semex supported

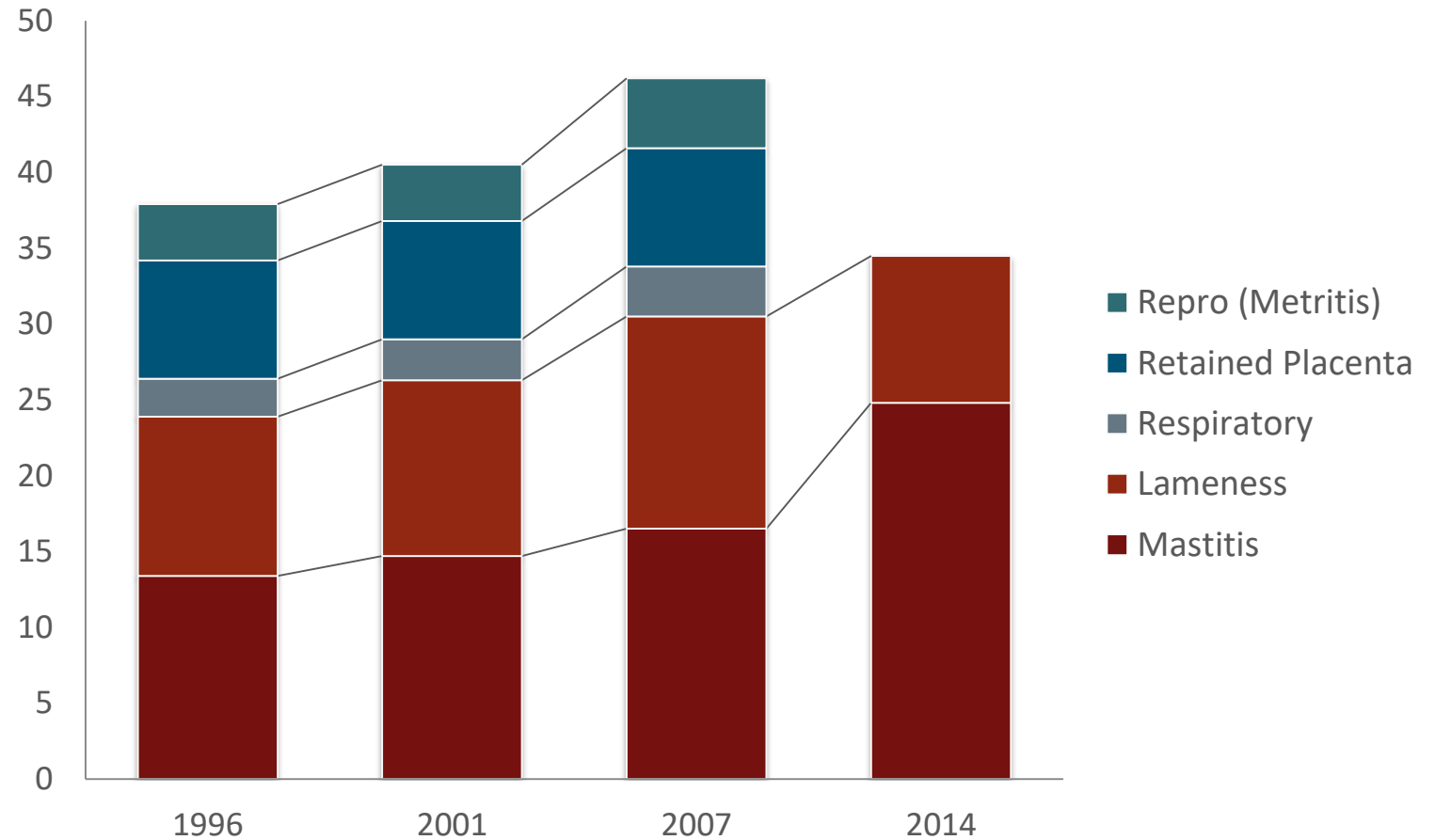
Almost 100 research papers in referred journals on immune response

Several thousand animals tested in research, beta test & commercial herds





Disease Trends



NAHMS Dairy 2014 Part II: Changes in the Dairy Cattle Industry 1996-2014



Past Genetic Selection for Health

- **Few “true” direct health traits**
- **Selecting an end result**
 - Herd Life/Productive Life
 - Daughter Fertility/Daughter Pregnancy Rate
 - Daughter Calving Ease
 - SCS
- **Hope it achieves better health**
- **Low heritability traits**



Need a Better Approach

Selection for Higher Immunity

- It is the ultimate goal
- Even better than direct selection for individual diseases

A starting point for a healthier dairy

Changes the way dairies will select for health



**What is
Immunity?**



Attacks from Air

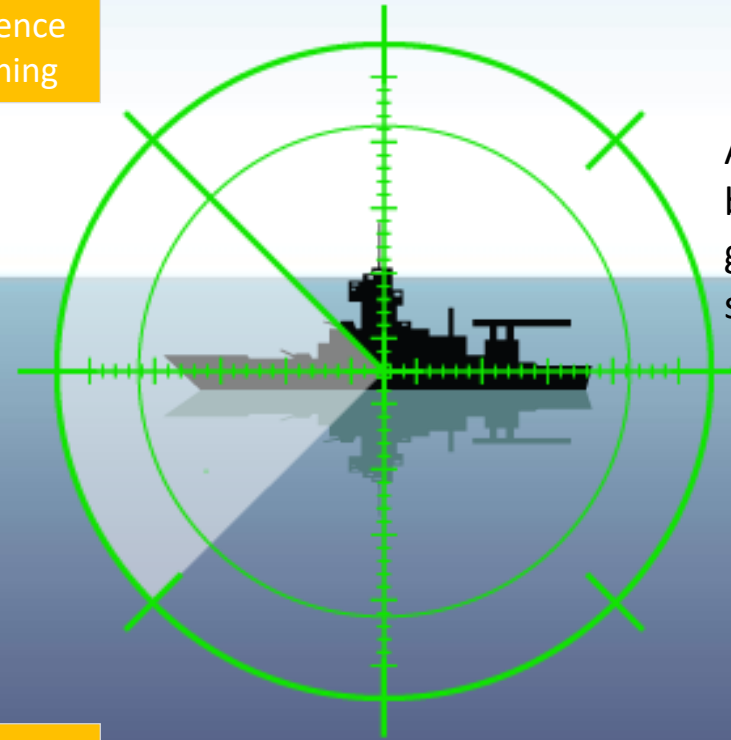


Attacks from Water



Response: Air Defense

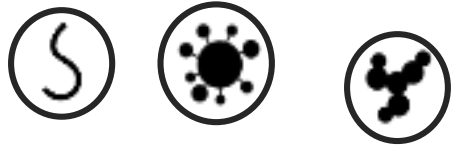
Experience
& training



All actions controlled
by central
government defense
system

Response: Naval Defense

Experience
& training



Bacterial infections

- Mastitis, listeriosis, brucellosis, E. coli scours, bacterial pneumonia, metritis, digital dermatitis

Viral & mycobacterial infections

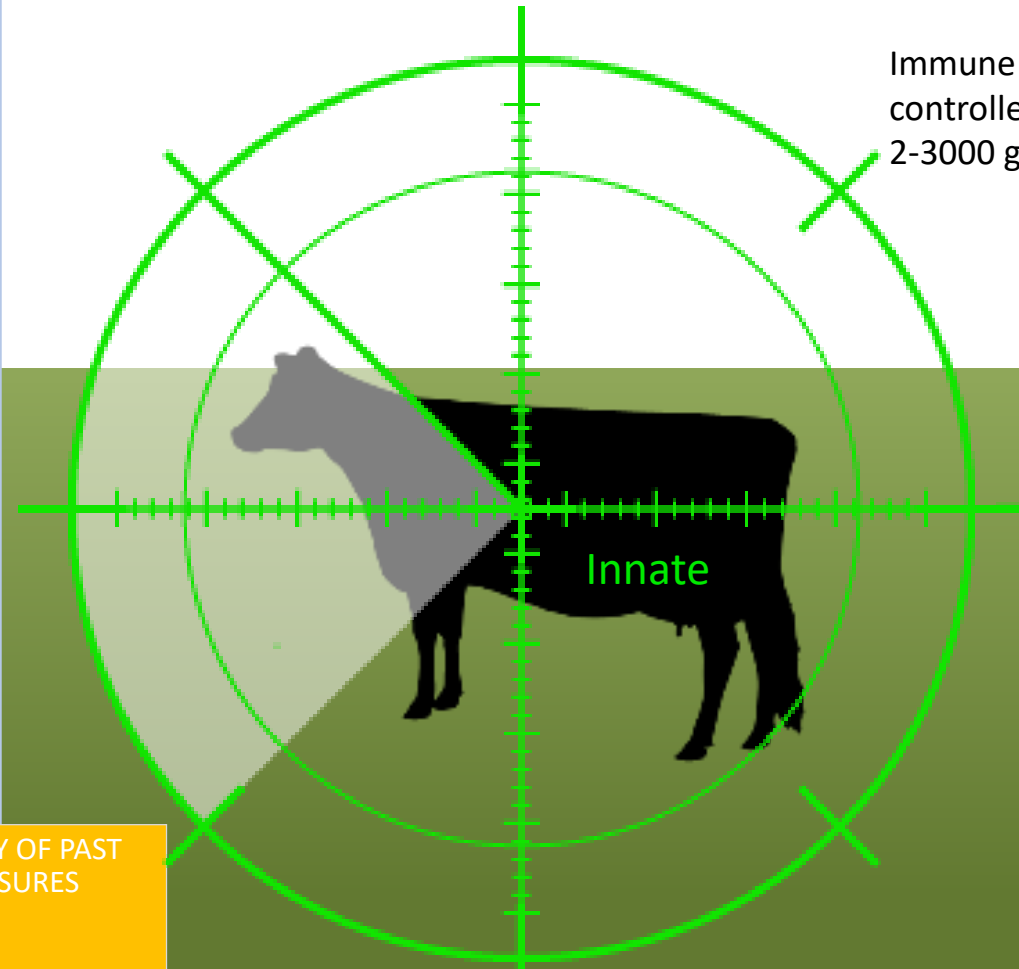
- Viral pneumonia, BVD, IBR, leucosis, foot & mouth, tb, retained placenta, Johne's



MEMORY OF PAST EXPOSURES

Antibody-mediated adaptive immune response

Immune response is controlled & directed by 2-3000 genes

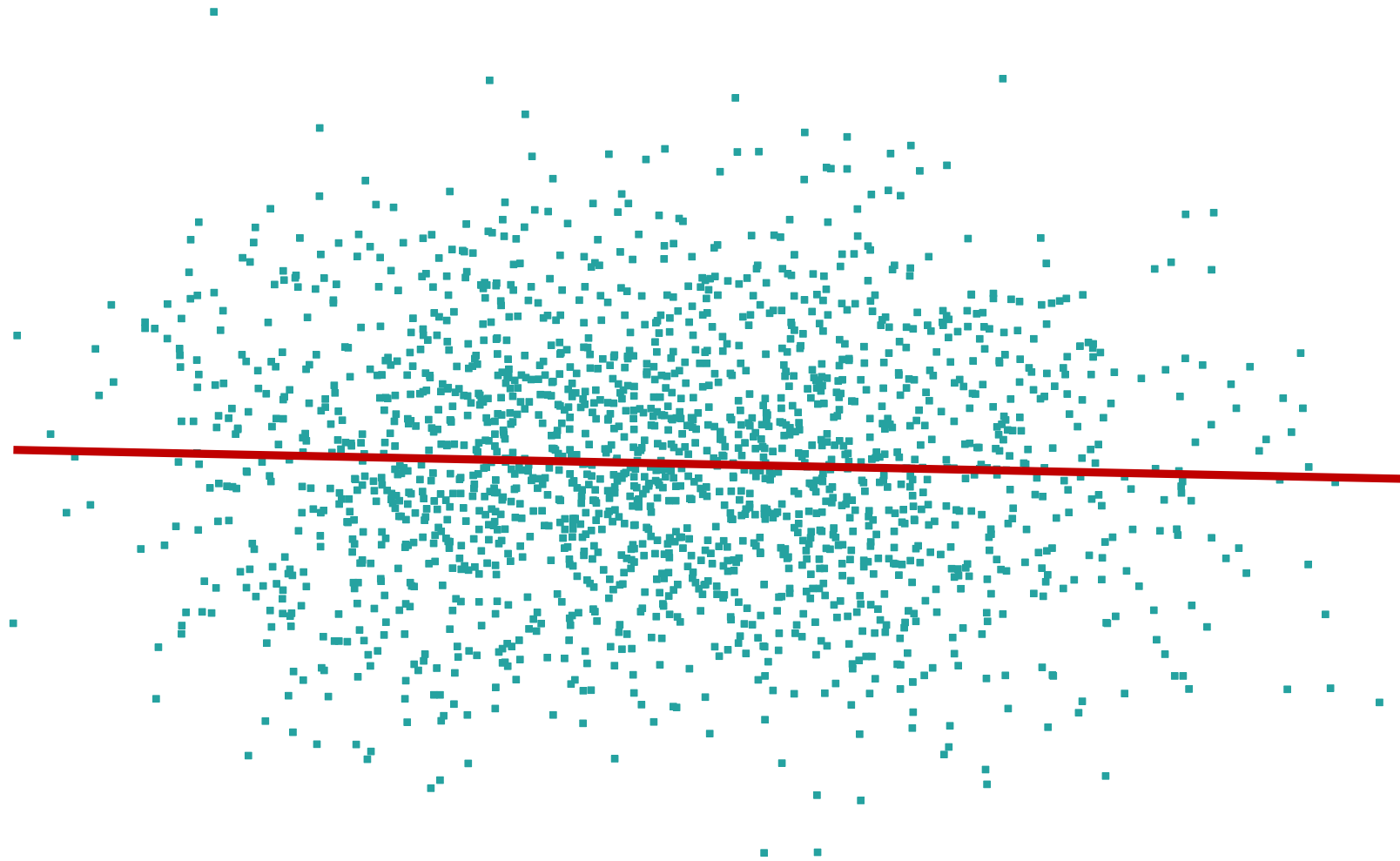


MEMORY OF PAST EXPOSURES

Cell-mediated adaptive immune response



AMIR and CMIR are slightly
NEGATIVELY Correlated





Broad-based Defence Against Most Viral & Bacterial Infections

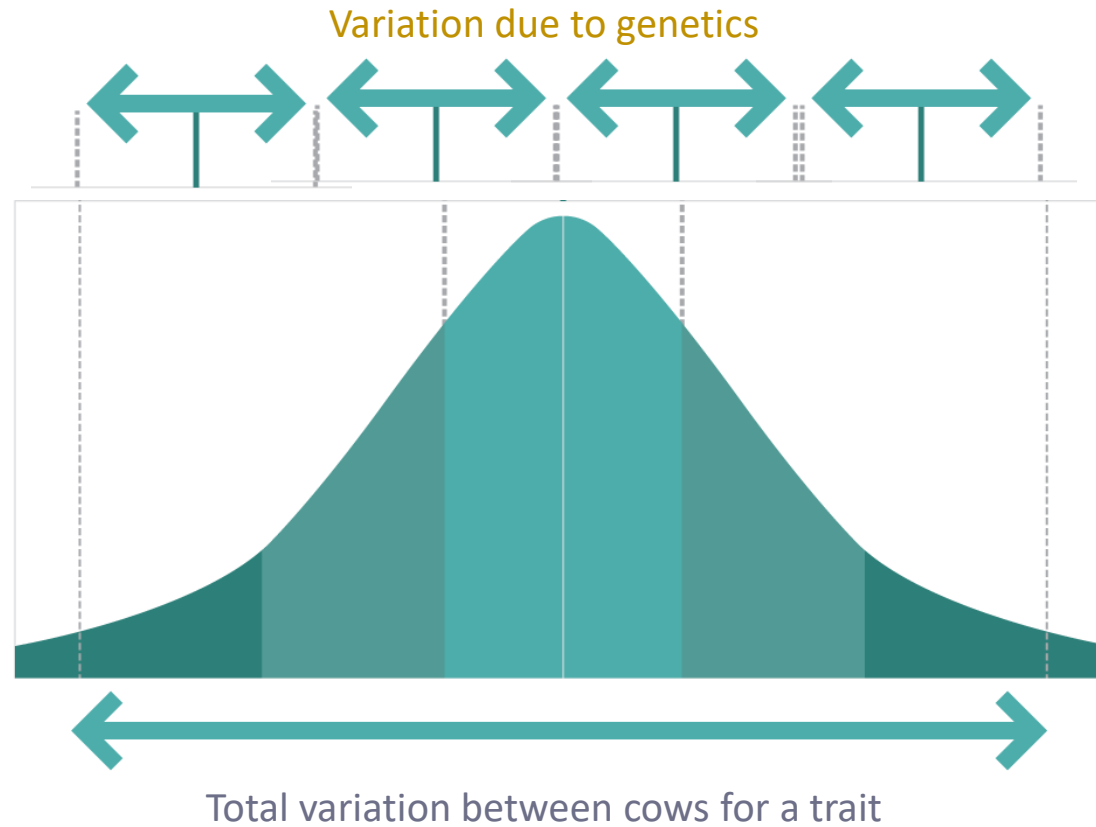


High Immune Response Technology



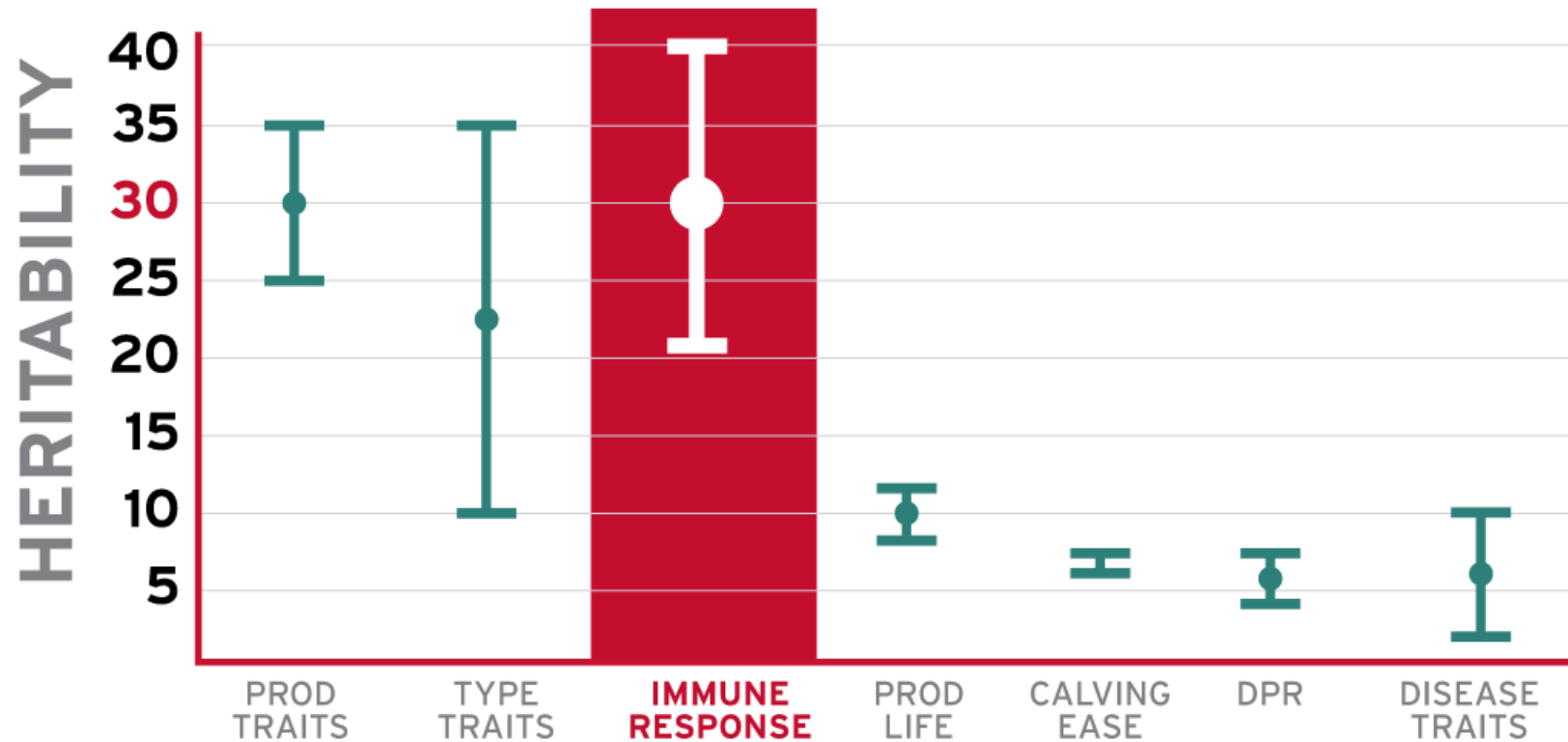
Heritability

% of total variation that's explained by genetics





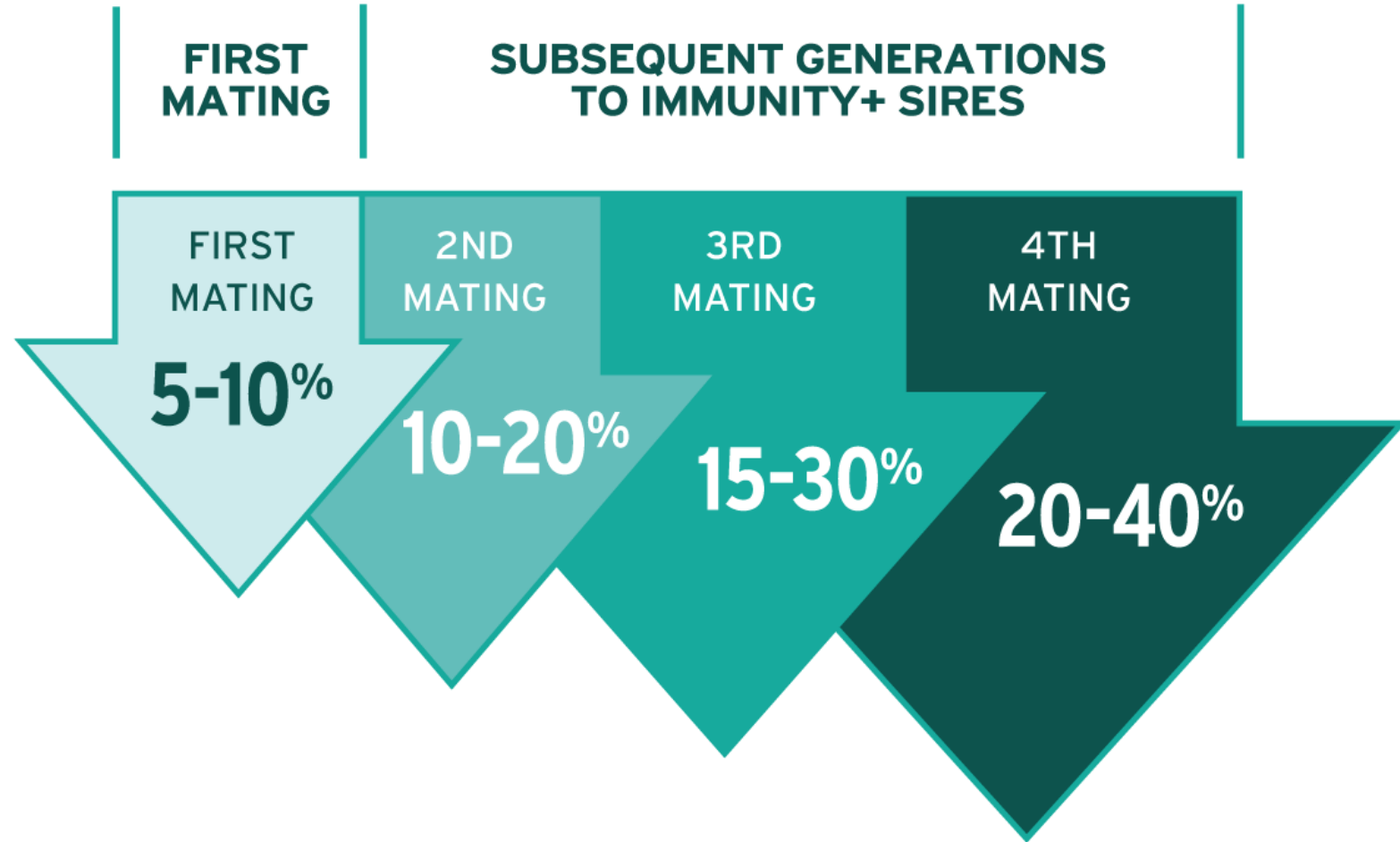
IMMUNE RESPONSE HERITABILITY IS NOW **30%**





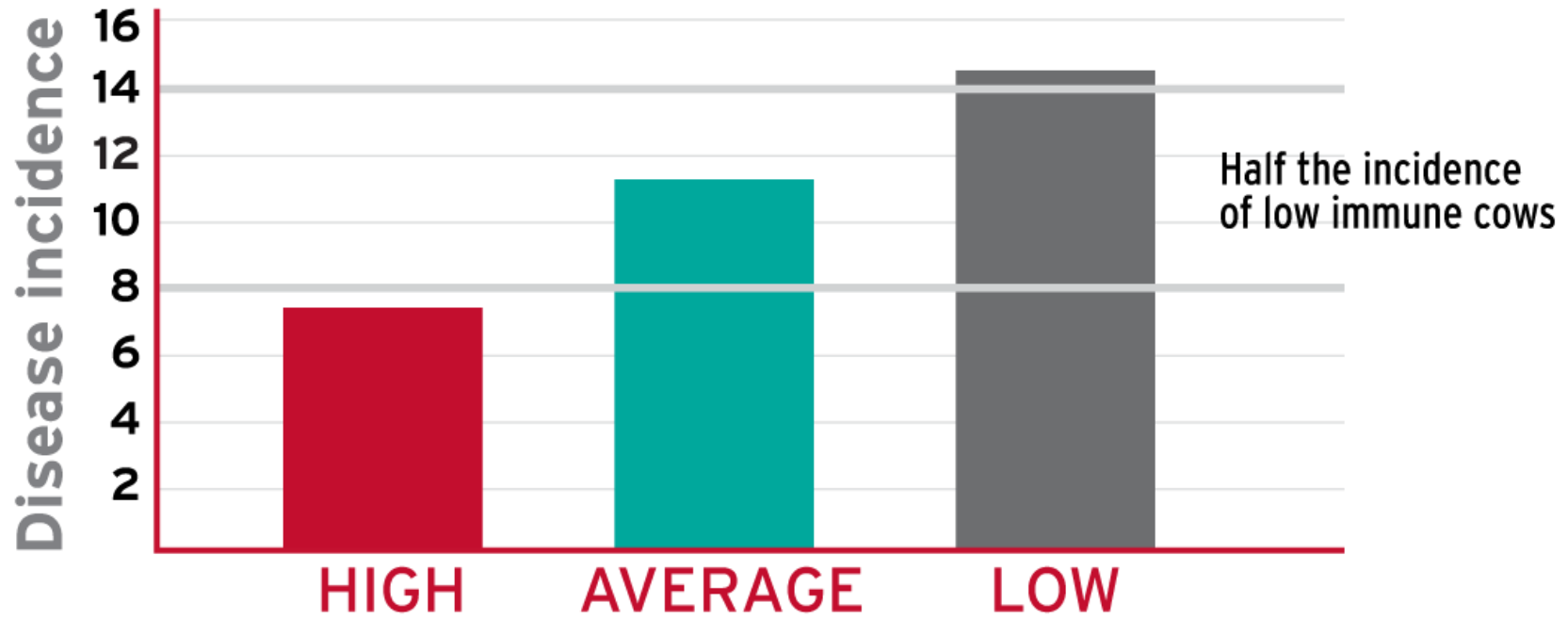
LESS DISEASE

Less disease generation after generation





RESEARCH SHOWS HIGH IMMUNE RESPONSE COWS HAVE LESS DISEASE



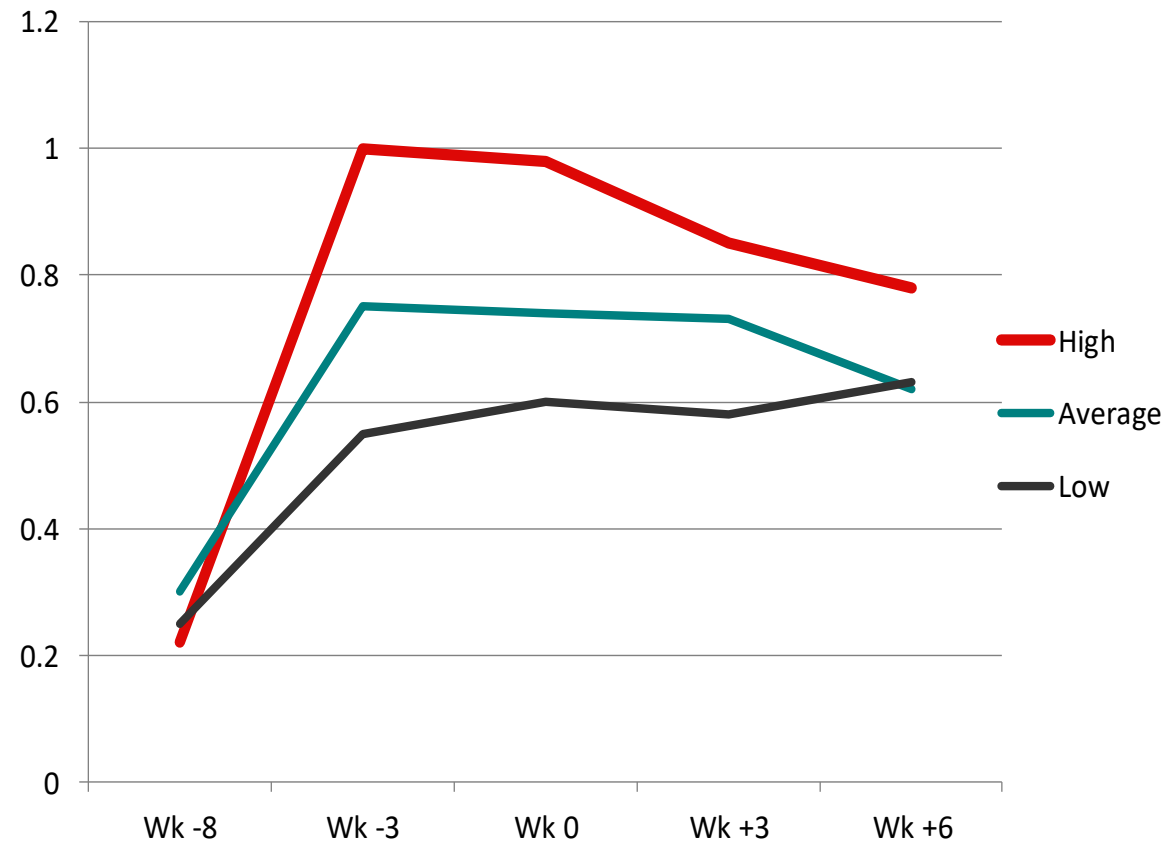
Studies across 64 herds in North America.

Wagter, et al. 2000 J. Dairy Sci. 83:488-498; Thompson-Crispi, et al. 2012. J. Dairy Sci. 95:3888-3893; Thompson-Crispi, et al. 2013. Clin Vacc Immuno. 20:106-112.



Vaccine Response

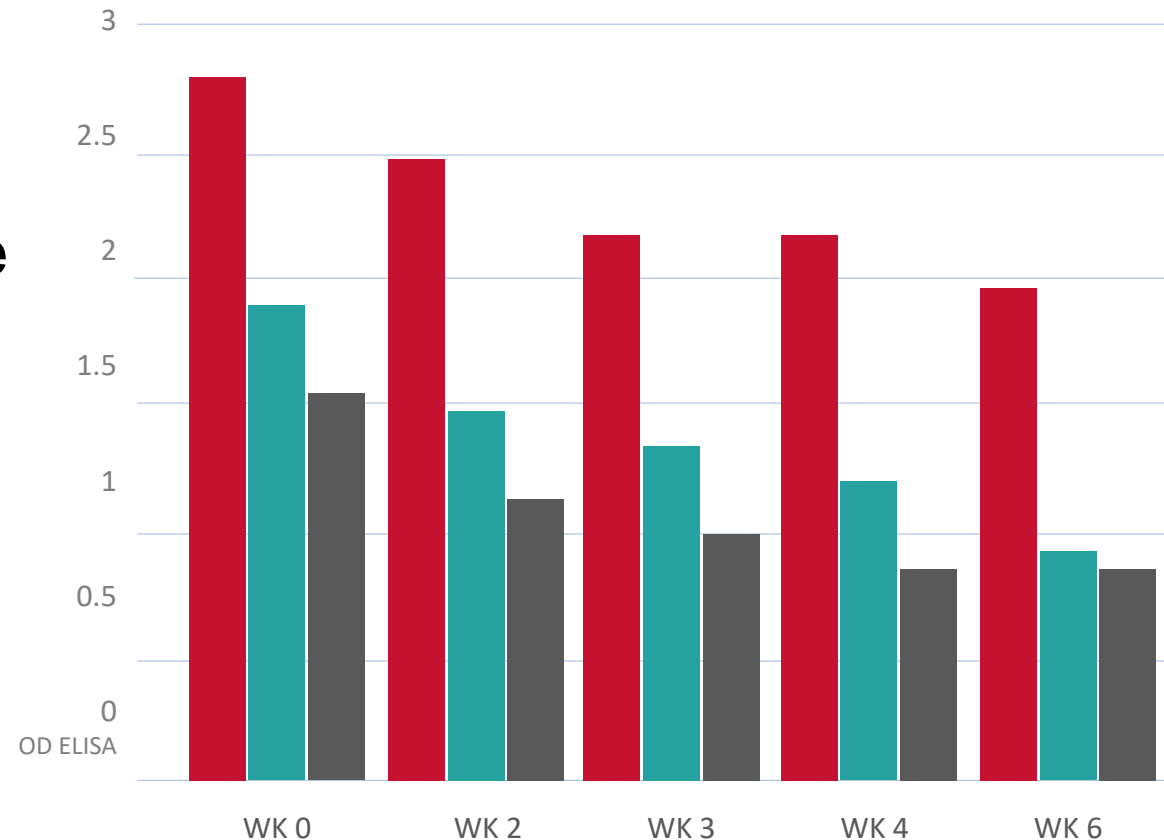
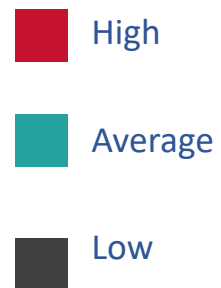
High immune cows respond better to commercial vaccines.





Quality Colostrum

High immune cows have higher quality colostrum with more antibodies.



Wagter & Mallard et al. 2000 JDS 83:488



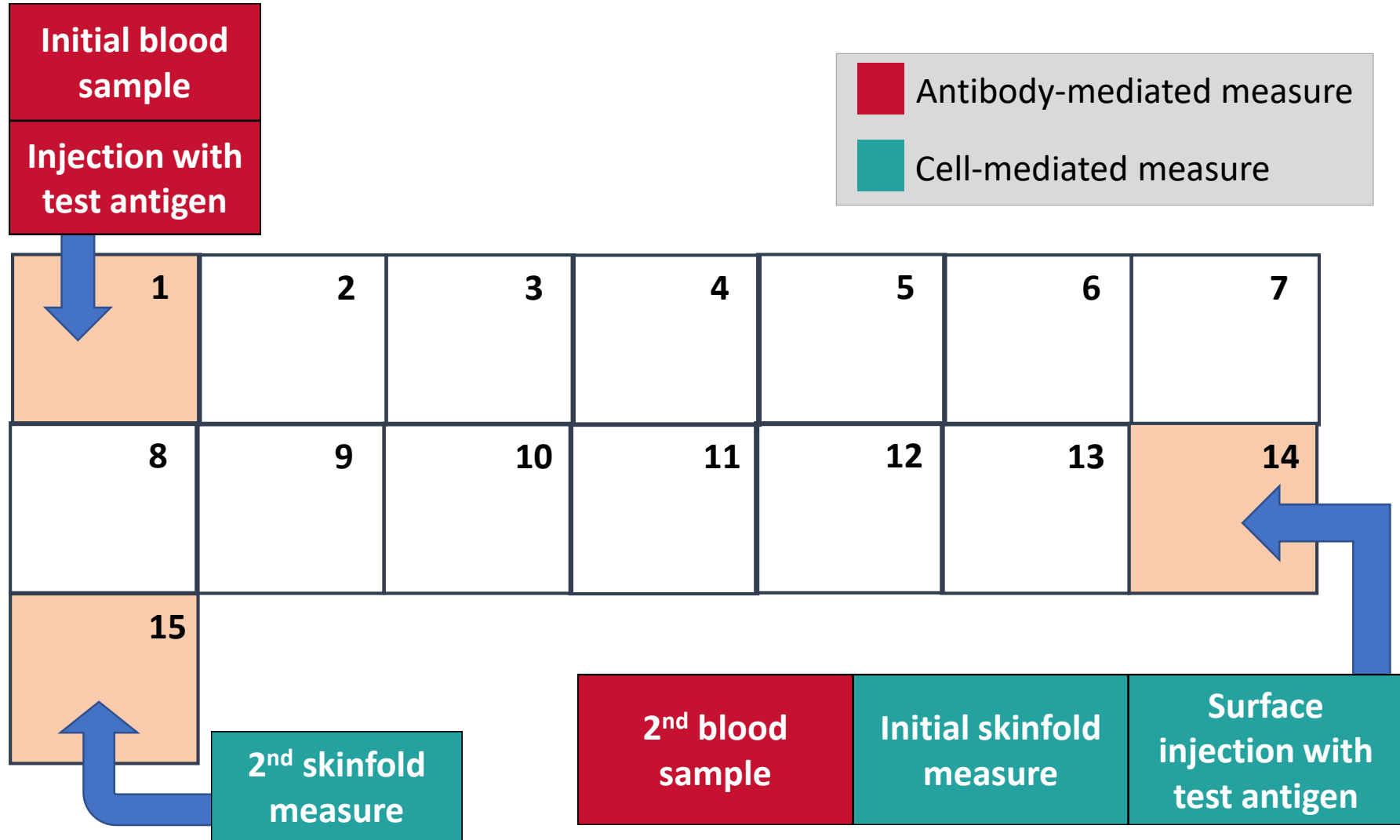
Testing For Immune Response



Two tests performed



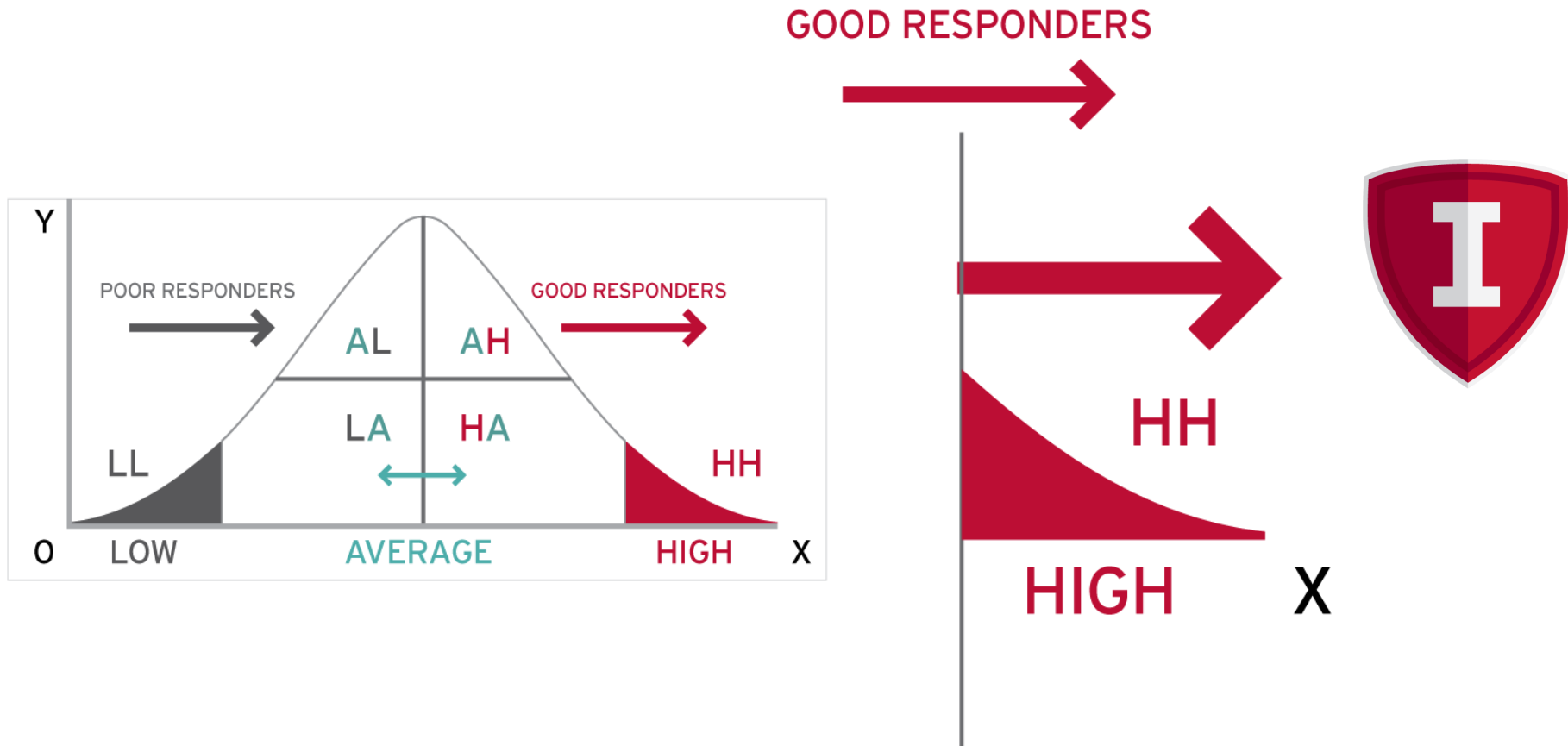
Testing for Immune Response





Bulls Designated as Immunity+

Approximately 10% of sires





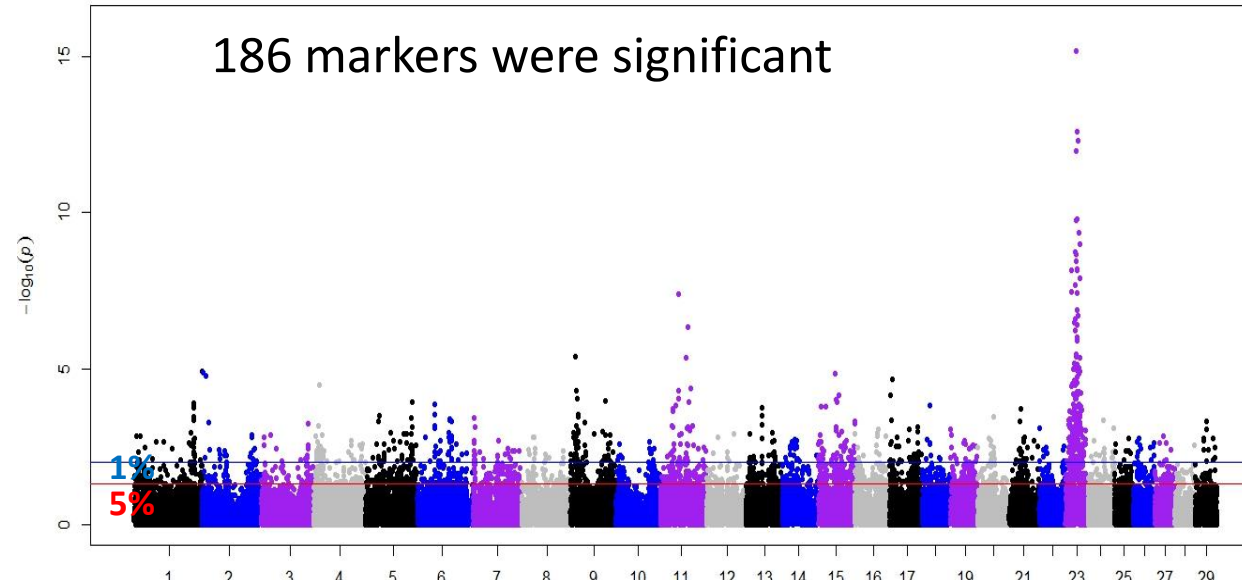
Initial Preliminary Research, AMIR – Genome Association





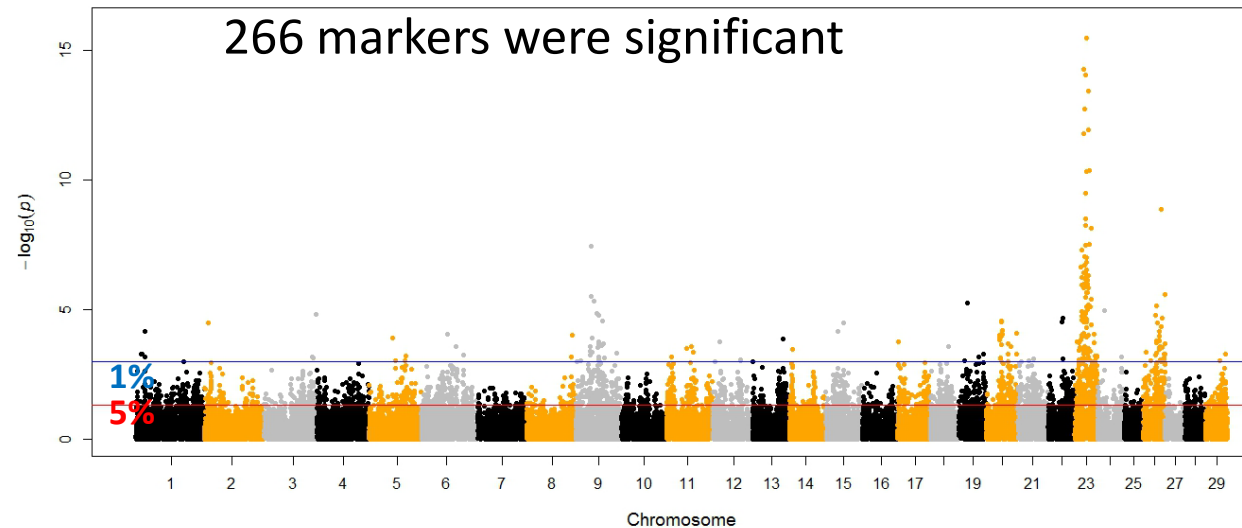
Initial Preliminary Research, AMIR – Genome Association

Female Study



Significant markers with just 163 cows

Male Study



Significant markers with just 631 bulls



Genome Wide Association Studies with Immune Response

- **Chromosome 23** contains the Bovine Major Histocompatibility Complex (BoLA)
 - Complex gene cluster responsible for regulating immunity in cattle
- Studies confirm the HIR measurements are hitting the correct target
- Show great potential for developing a genomic test for Immunity



Proven on Farm™

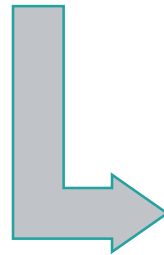




Methods

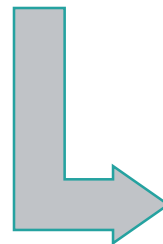
Extract
Health Data

- Cows and heifers are controlled to ensure all groups (Immunity+, non-Immunity+, and Other (Non-Semex)) are of the same average age (Same pathogen exposure)



Fit Poisson
Regression
Model

- Model included effects of parity, herd, age and Immunity+ status (binary)




Calculate
Immunity+
Effect

- Determine odds ratios and relative proportions across groups




Proven on Farm™ – 35 Commercial Dairies (~30,000 Cows, ~75,000 Heifers)

HEALTH EVENT	IMMUNITY+ Prevalence	Non-Immunity+ Prevalence	 Reduction
Mastitis	28.5%	31.7%	10.0%
Persistent Mastitis	4.4%	5.3%	17.0%
Lameness	21.1%	24.0%	12.1%
Miscellaneous Illness	5.9%	6.5%	9.2%
Mortality	4.8%	6.0%	20.0%
Heifer Pneumonia	9.6%	9.8%	2.1%
Heifer Diarrhea	5.4%	5.7%	5.3%
Heifer Mortality	7.7%	9.2%	16.3%



Proven on Farm™ – 35 Commercial Dairies (~30,000 Cows, ~75,000 Heifers)

HEALTH EVENT	IMMUNITY+ Cost/1000 Cows	Non-Immunity+ Cost/1000 Cows	 Reduction
Mastitis ^{1*}	\$63,840	\$71,000	\$7,160
Lameness ^{1*}	\$98,960	\$112,560	\$13,600
Miscellaneous Illness ^{1*}	\$14,390	\$15,860	\$1,470
Mortality ¹	\$105,600	\$132,000	\$26,400
Heifer Pneumonia ²	\$13,240	\$13,520	\$280
Heifer Diarrhea ³	\$5,720	\$6,040	\$320
Heifer Mortality ¹	\$118,580	\$141,680	\$23,100
TOTAL	<i>*Per Lactation</i>		\$72,330

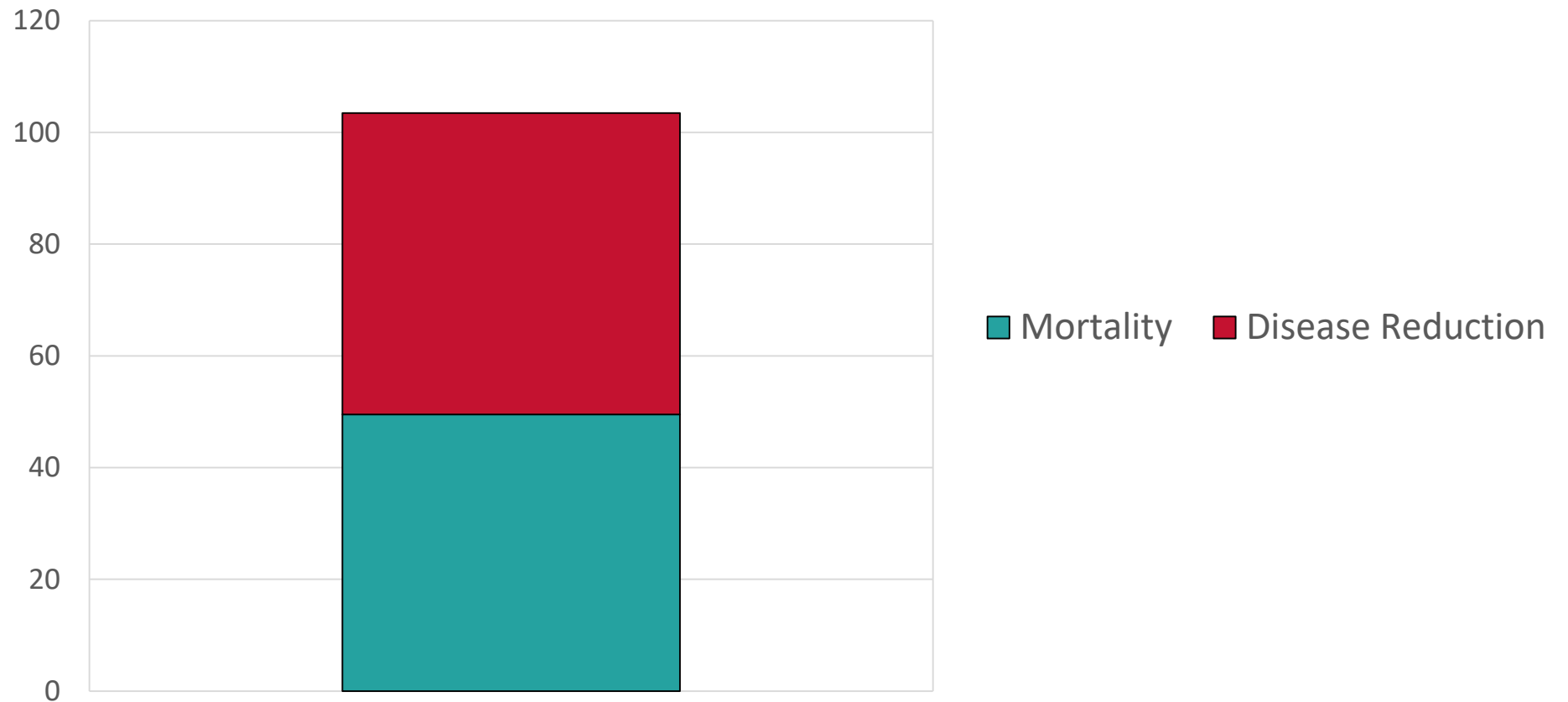
¹Guard, C. *The costs of common diseases of dairy cattle – CVC in San Diego Proceedings*

²*Pneumonia in heifer calves costs more than \$100/head – Vet Times*

³Mohd, N et al. *Estimating the costs of rearing young dairy cattle... - Preventative Vet Med*



Total Economic Benefit



Total Lifetime Savings of \$103,450 per 1,000 cows without considering increased colostrum quality and vaccine response



HIGH QUALITY COLOSTRUM WITH GENETICS

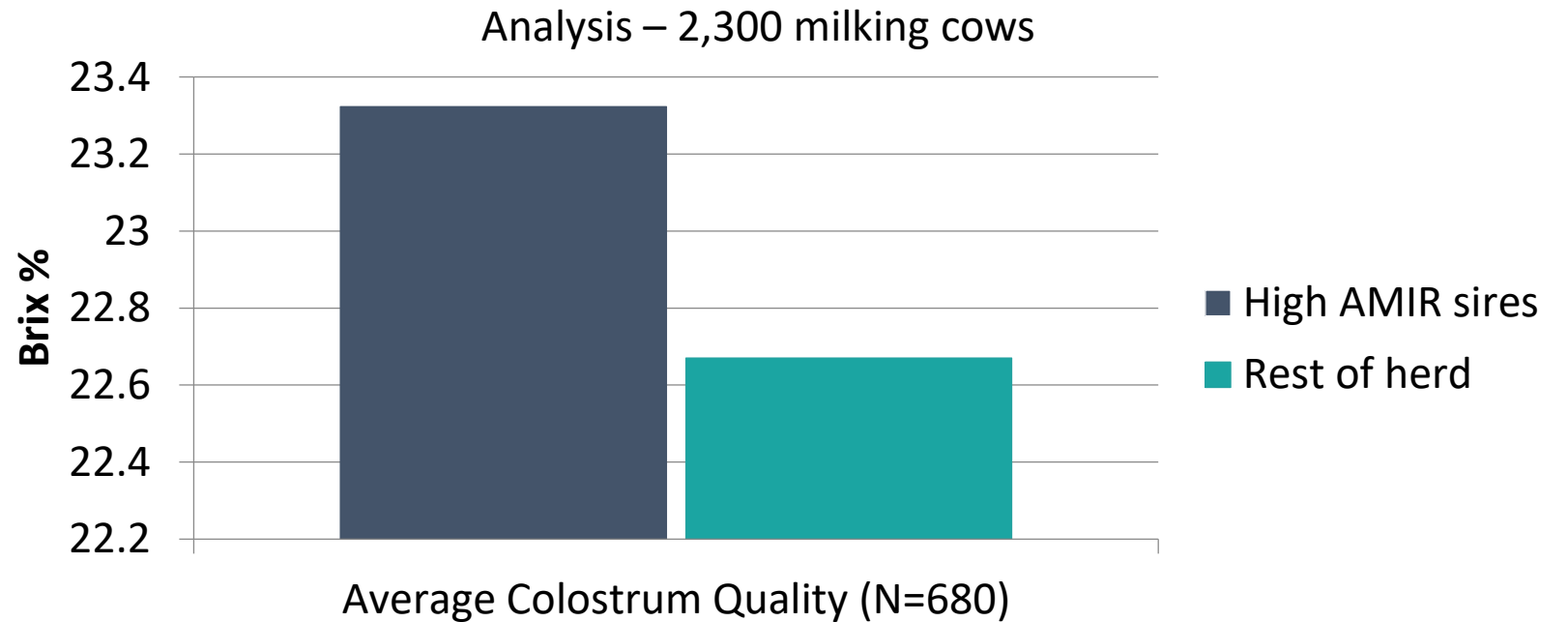
PROVEN RESULTS

MORE ANTIBODIES

A HEALTHY START



Proven on Farm™ – Colostrum (2016)



Correlation between colostrum quality and overall immune response: **0.55**

High AMIR sires <22% = **0%**

Rest of herd = **27%**



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