

# An Integrated Approach To Improve Dairy Cow Fertility



**Victor Cabrera**, Assistant Professor- Management

**Paul Fricke**, Professor - Reproductive Physiology

**Pam Ruegg**, Professor - Physiology and Management

**Randy Shaver**, Professor - Nutrition

**Milo Wiltbank**, Professor -Reproductive Physiology

**Kent Weigel**, Professor and Chair Breeding and Genetics



**University of Wisconsin-Madison**



United States Department of Agriculture  
National Institute of Food and Agriculture



"This project is supported by Agriculture and Food Research Initiative Competitive Grant no. 2010-85122-20612 from the USDA National Institute of Food and Agriculture." Project started March 1<sup>st</sup>, 2010

# Overall Objective

- To improve reproductive efficiency of lactating dairy cows using an interdisciplinary extension and research team that will identify and remove barriers to reproductive success and link outcomes of basic and applied research with an innovative extension delivery program
- Five aims address this overall objective



# SPECIFIC AIM 1

## Prediction of Pregnancy Using Machine Learning Algorithms

### Objective:

To develop a user friendly and intuitive on-farm tool to help farmers make decisions about breeding specific cows.

### Data:

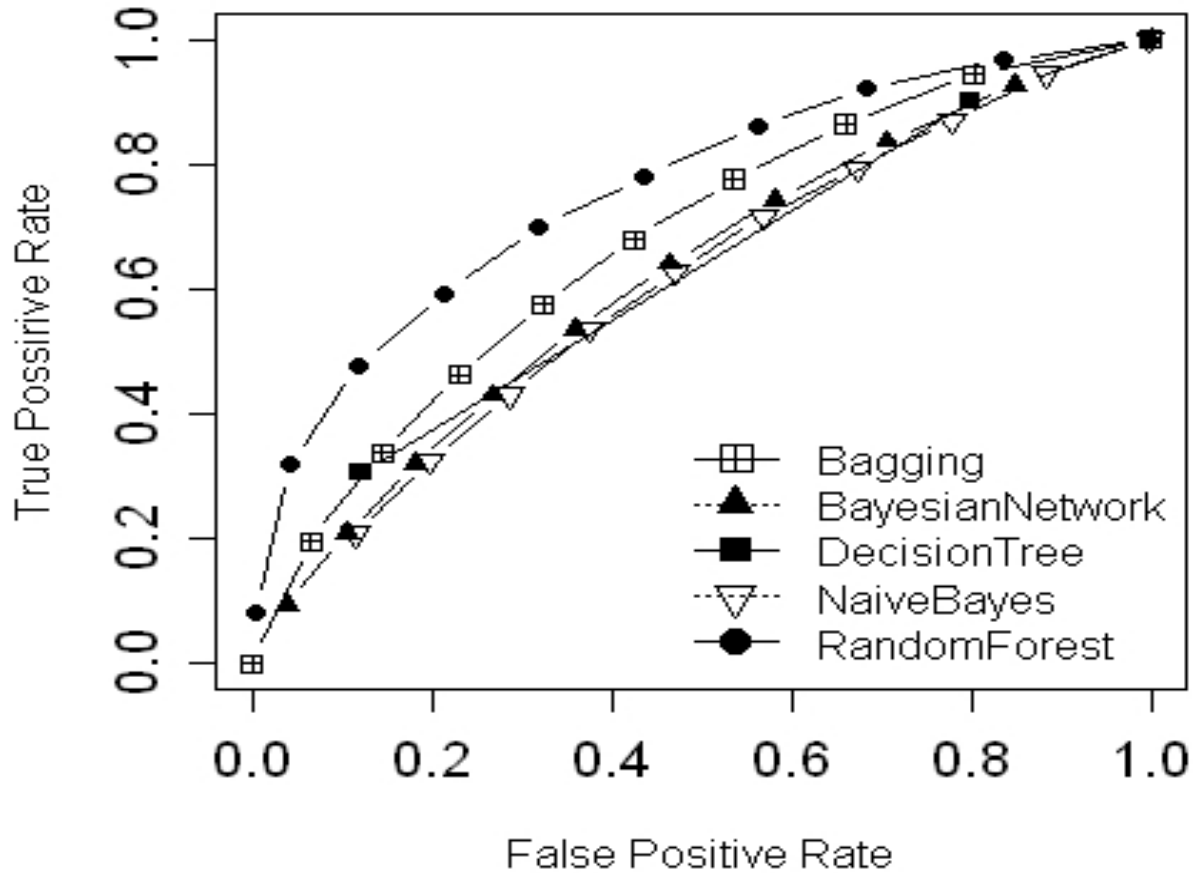
A total of 129,245 breeding records and 28 explanatory variables were available for primiparous cows, and 195,128 breeding records and 31 explanatory variables were available for multiparous cows obtained from 26 dairy farms in the Alta Genetics Advantage Progeny Testing Program (2000- 2010) .

### Machine Learning Algorithms:

Naïve Bayes classifier; Bayesian network; Decision Trees; Bagging

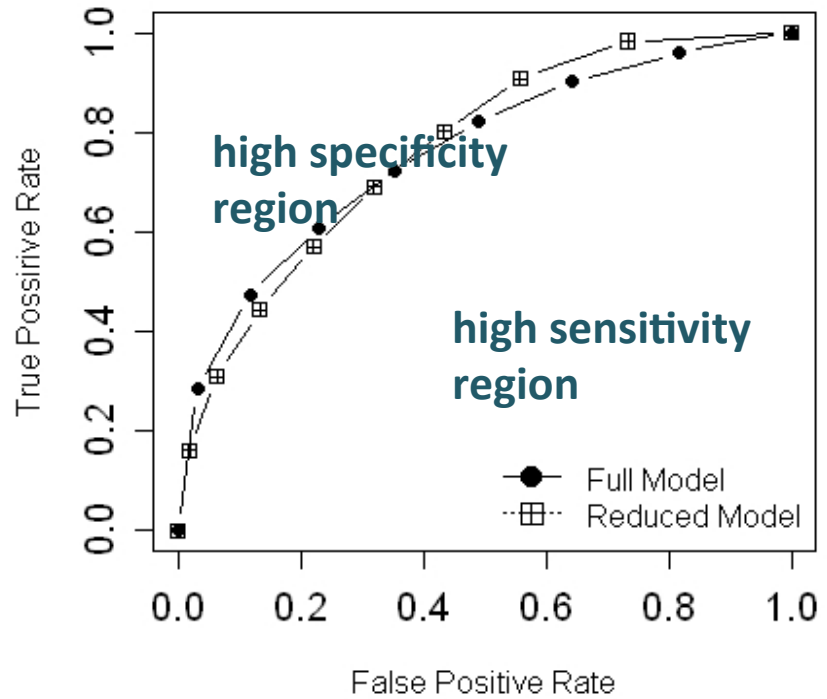


# Results

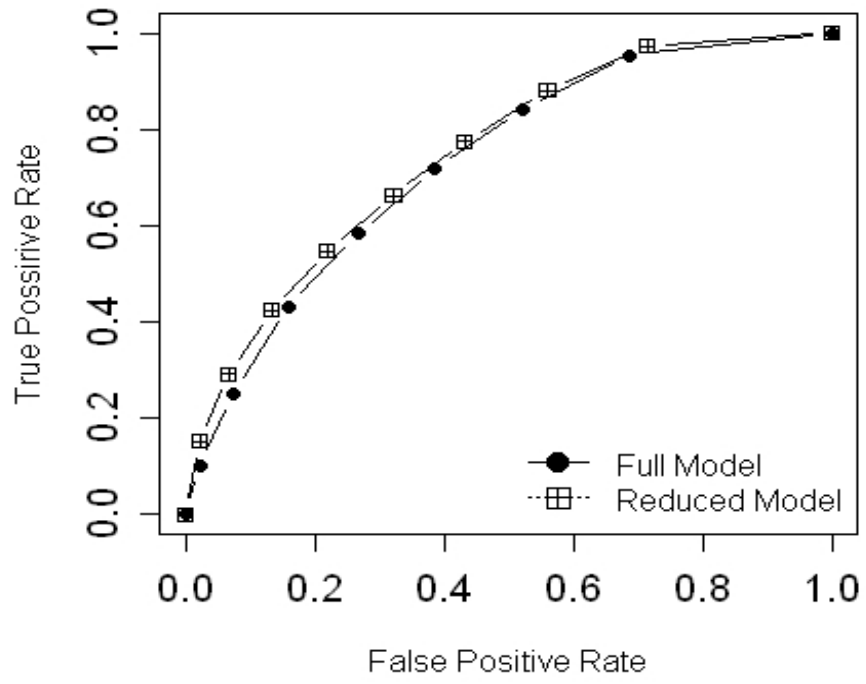


Random Forest Model performed best.

**Explanatory Variables = Incidence of ketosis, number of AIs, herd average conception rate, days in milk at AI, incidence of mastitis.**



**Primiparous cows  
72.3% Correct**



**Multiparous Cows  
73.6%**

**SPECIFIC AIM 2:** Determine the impact of specific nutritional components on reproductive performance of lactating dairy cows.

**Experiment 1:** Evaluate the association of individual components of the diet on fertility traits of dairy cattle.

**Data:** Complete diets were obtained and Dairy Comp 305 backups for fertility and other traits evaluated in 50 dairy farms.

**Results:** Carbohydrate and Methionine

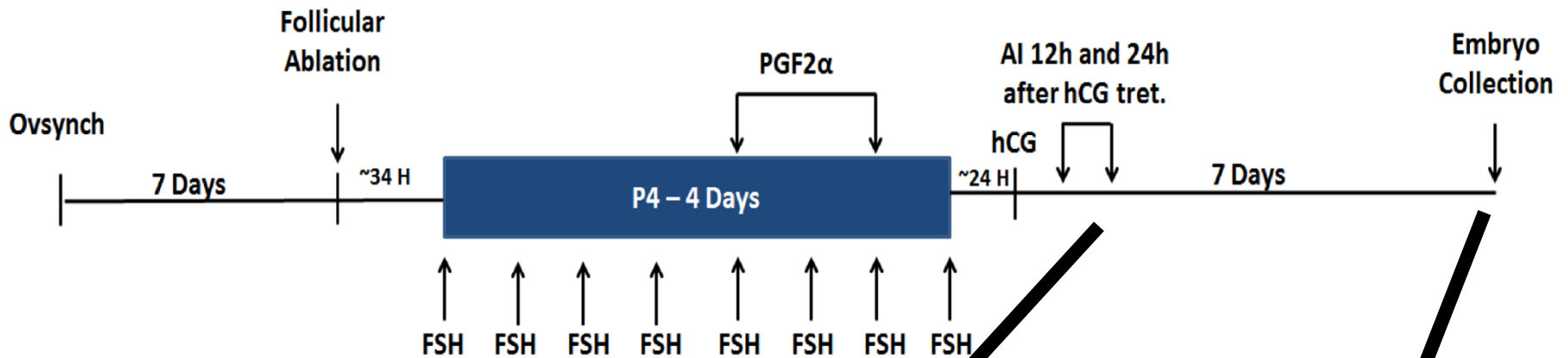
% NFC and 1<sup>st</sup> Service CR:  $r = -0.51$   $P = 0.0002$

Methionine and 1<sup>st</sup> Service CR:  $r = 0.30$   $P = 0.05$



± Methionine Supplementation (2.43 vs. 1.89 Met %MP)

## Synchronization and superovulation protocol

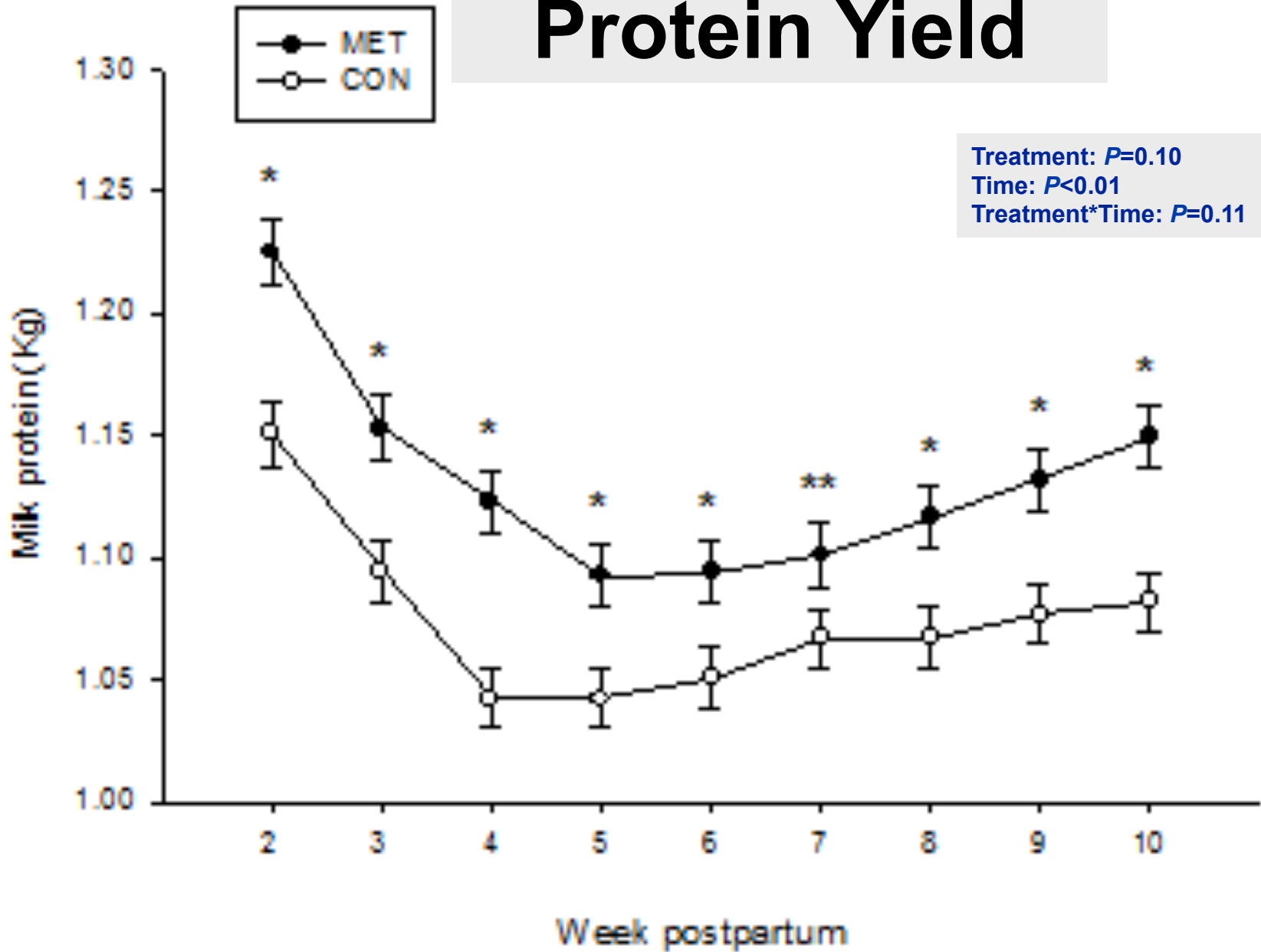


Ovulation  
n = 853

Fertilization  
& Embryo Quality  
n = 571

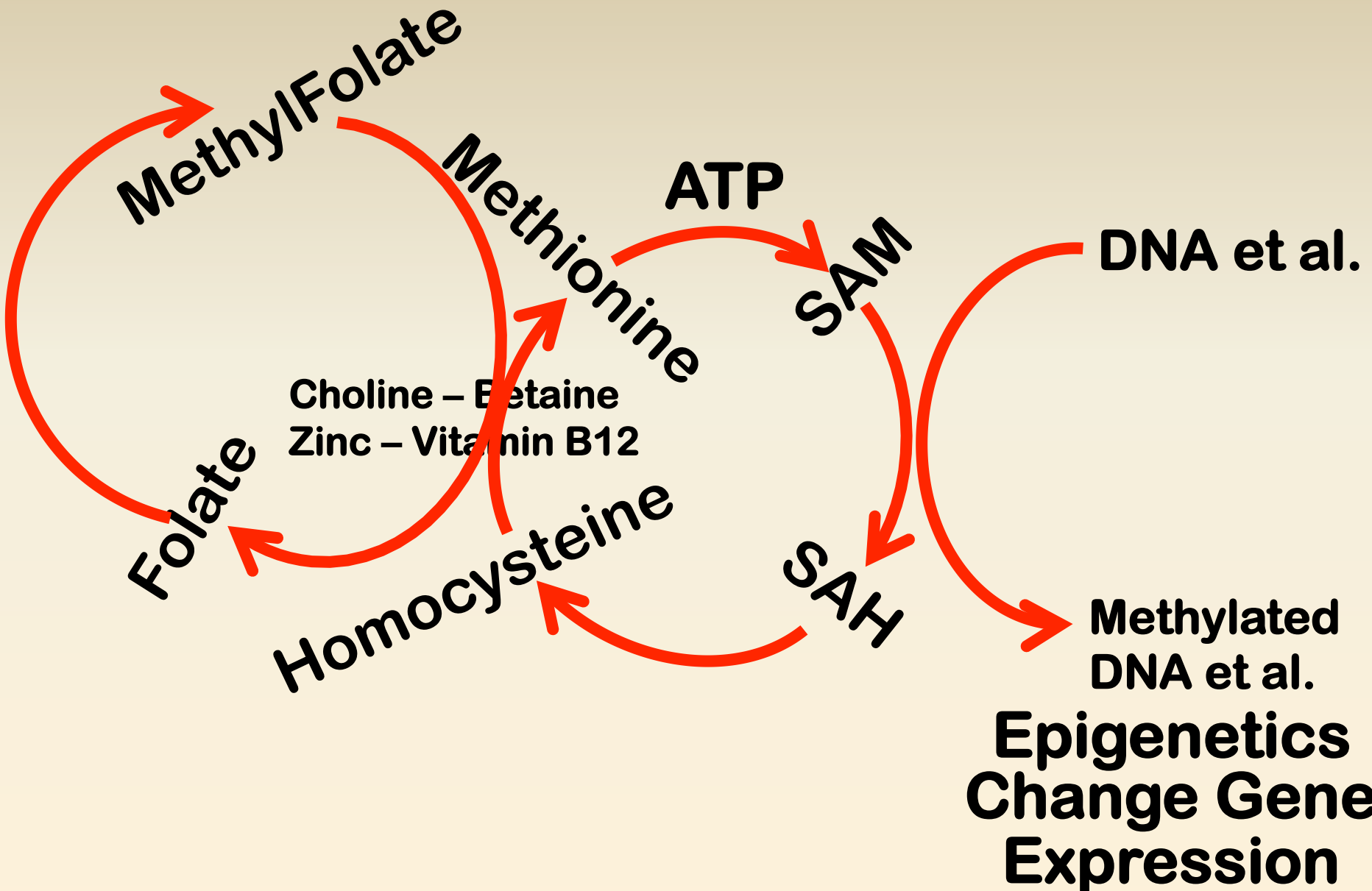


# Protein Yield



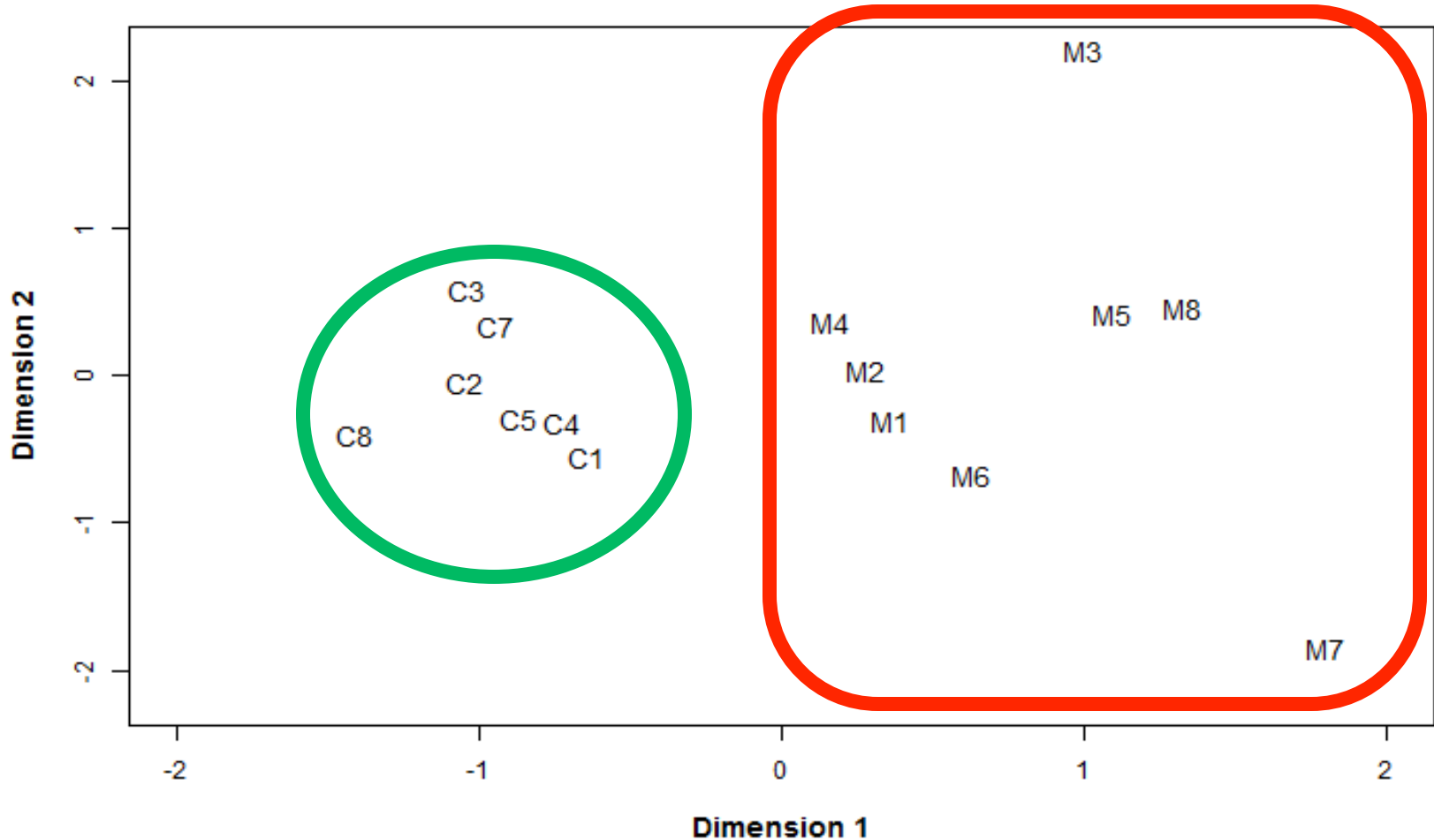
# Embryos of superovulated cows fed MET or CON

|                                       | <b>MET</b> | <b>CON</b> |                |
|---------------------------------------|------------|------------|----------------|
| n                                     | 35         | 37         | <b>P-value</b> |
| <b>CL number</b>                      | 17.0 ± 1.3 | 17.7 ± 1.5 | 0.90           |
| <b>Total ova/embryos recovered</b>    | 9.1 ± 1.4  | 6.8 ± 1.0  | 0.18           |
| <b>% Ova/embryos recovered per CL</b> | 49.5 ± 4.9 | 35.8 ± 4.4 | 0.05           |
| <b>Number of fertilized ova</b>       | 6.5 ± 1.1  | 5.5 ± 0.9  | 0.56           |
| <b>% Fertilized ova</b>               | 74.7 ± 5.6 | 82.2 ± 3.8 | 0.27           |
| <b>Number of transferable embryos</b> | 5.0 ± 0.9  | 4.3 ± 0.1  | 0.57           |
| <b>% Transferable embryos</b>         | 56.3 ± 6.5 | 62.5 ± 6.0 | 0.49           |
| <b>Number of degenerate embryos</b>   | 1.5 ± 0.4  | 1.3 ± 0.4  | 0.75           |
| <b>% Degenerate embryos</b>           | 18.5 ± 4.6 | 19.7 ± 4.7 | 0.83           |
| <b>% Degenerate of fertilized ova</b> | 25.1 ± 5.8 | 27.5 ± 6.0 | 0.74           |



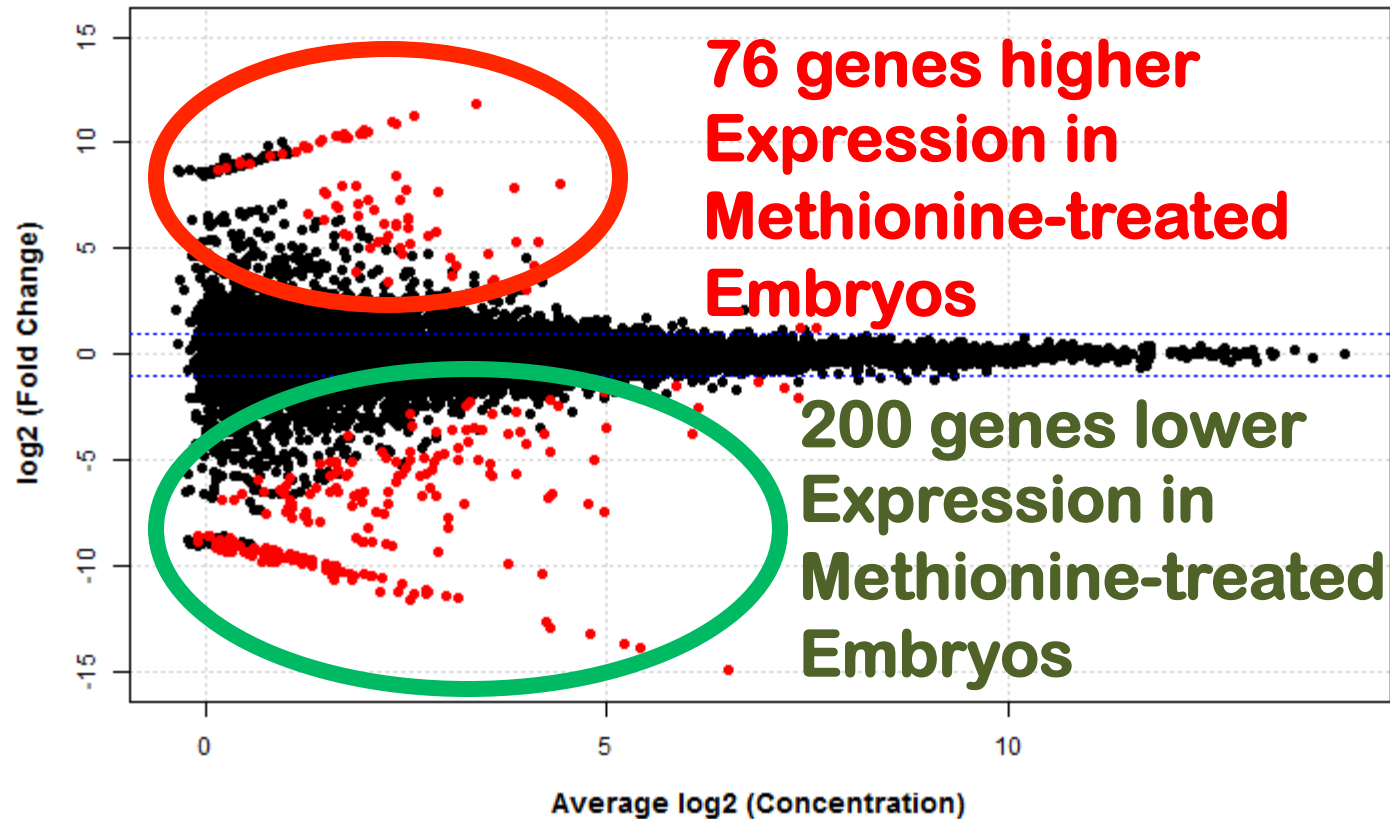
# One Carbon Pathway

# Multidimensional scaling (MDS) plot showing the relative similarities of the samples under study.



# Plot of the log2 fold change against log2 average.

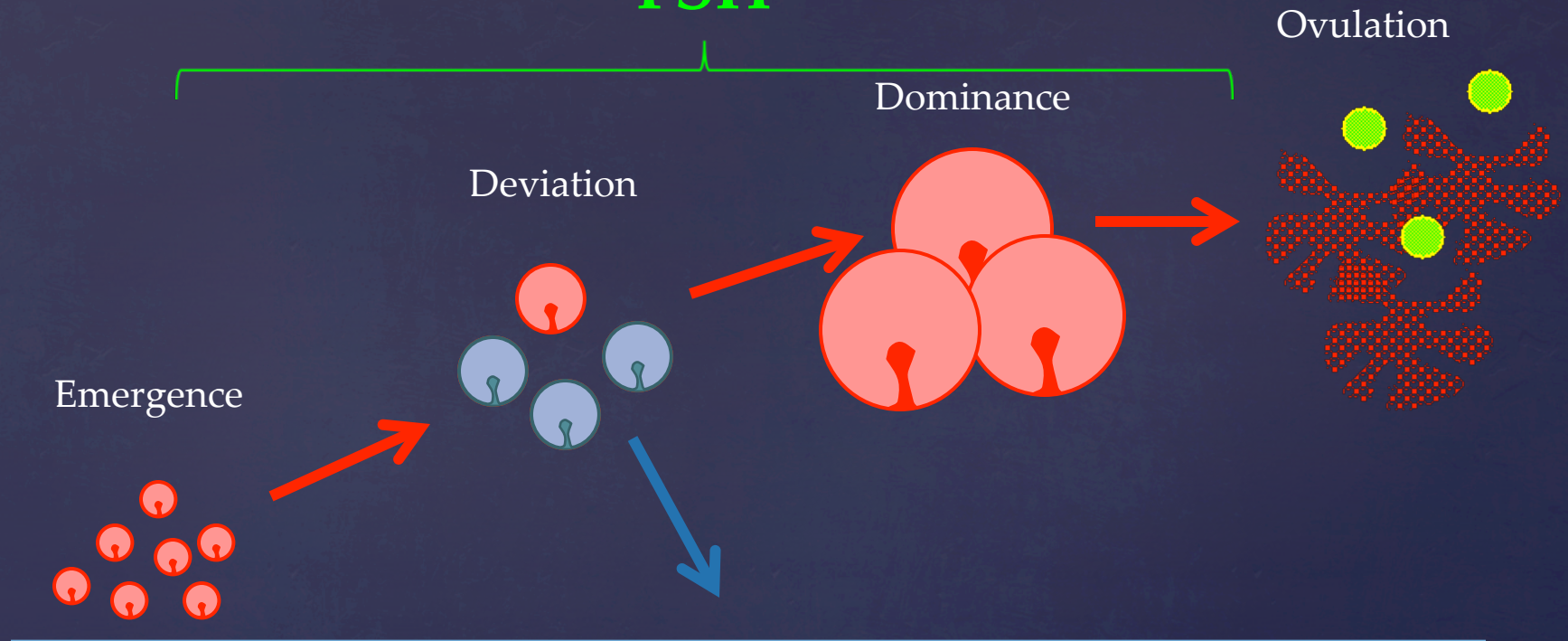
Differential expressed genes are highlighted in red.



| Gene        | Name  | log2 FC | FDR                    |
|-------------|---|---------|------------------------|
| LAPTM5      | Lysosomal protein transmembrane 5                                 | -14.9   | 4.7 x 10 <sup>-9</sup> |
| NKG7        | Natural killer cell group 7 sequence                              | -13.6   | 4.4 x 10 <sup>-8</sup> |
| VIM         | Vimentin  | -13.8   | 1.8 x 10 <sup>-7</sup> |
| TYROBP      | TYRO protein tyrosine kinase binding protein                      | -13.2   | 3.2 x 10 <sup>-6</sup> |
| IFI6        | Interferon, alpha-inducible protein 6                             | -12.6   | 1.5 x 10 <sup>-5</sup> |
| CUFF.2147.1 | Novel transcript unit   | -8.2    | 1.5 x 10 <sup>-5</sup> |
| LOC505451   | Olfactory receptor, family 1, subfamily J, member 2-like          | -13.0   | 1.5 x 10 <sup>-5</sup> |
| SLAMF7      | Signaling lymphocyte-activating molecule family 7 family member 7 | -10.4   | 3.5 x 10 <sup>-5</sup> |
| LOC788199   | Olfactory receptor 6C74-like                                      | -10.4   | 7.6 x 10 <sup>-5</sup> |
| LCP1        | Lymphocyte cytosolic protein 1 (L-plastin)                        | -9.9    | 1.1 x 10 <sup>-4</sup> |

# Model

FSH



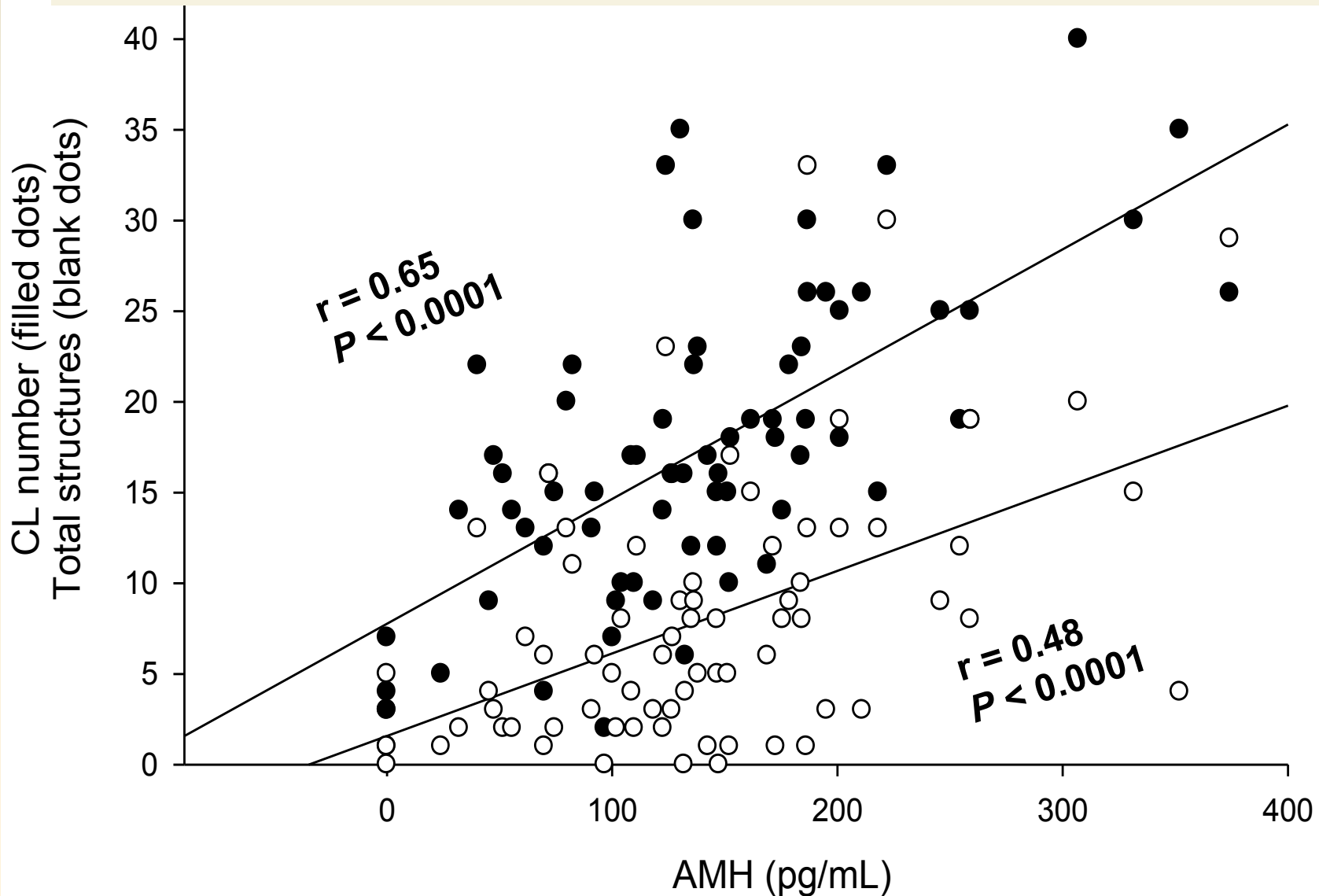
**±Propylene Glycol (↑/↓INSULIN)**

Ovulation Rate: 88.5% vs. 80.5%  $P = 0.002$

Fertilization Rate: 73.7% vs. 63.4%  $P = 0.02$

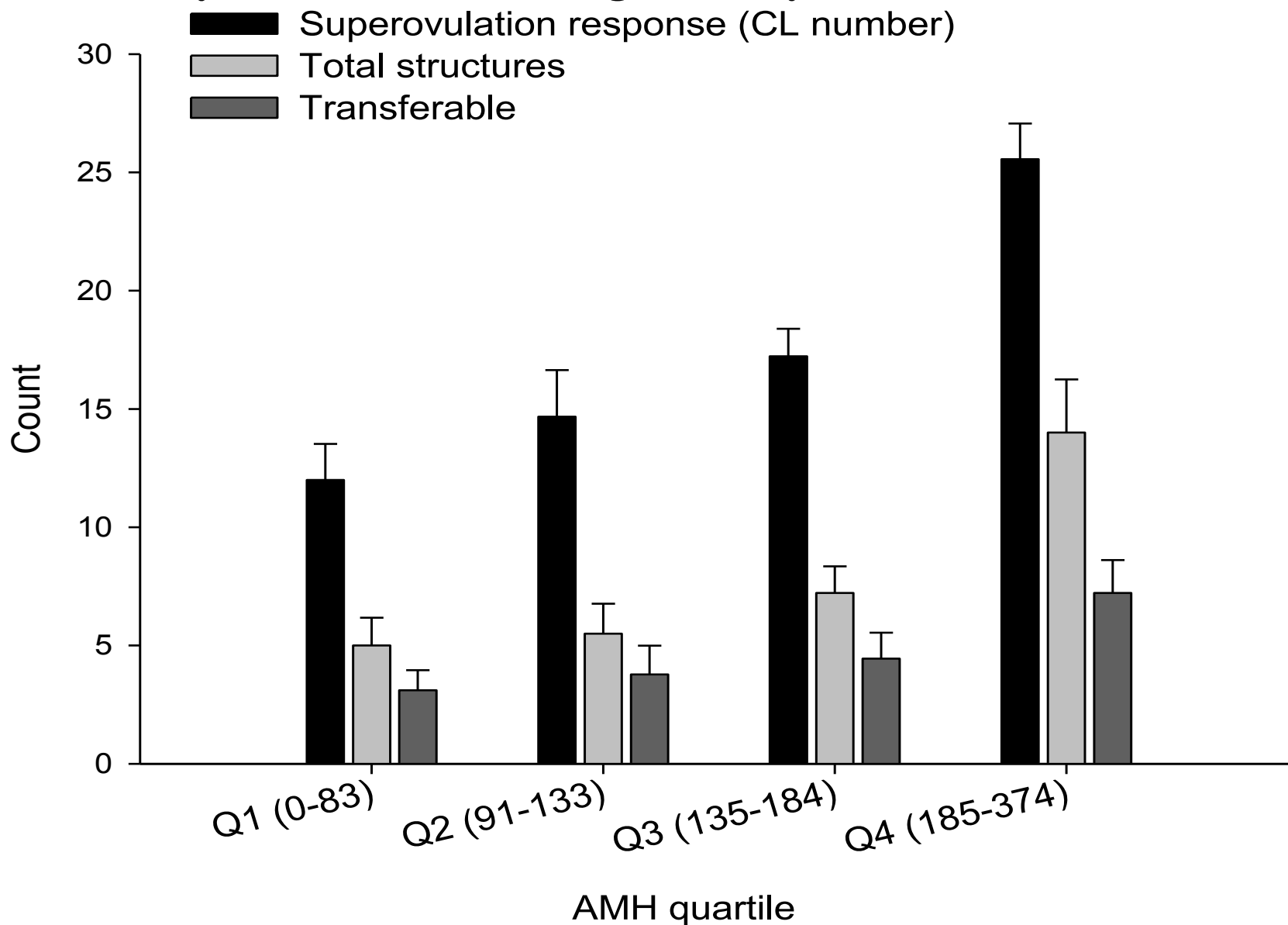
No effect on other embryo measures

# Relationship of AMH to CL number and embryos/oocytes recovered.





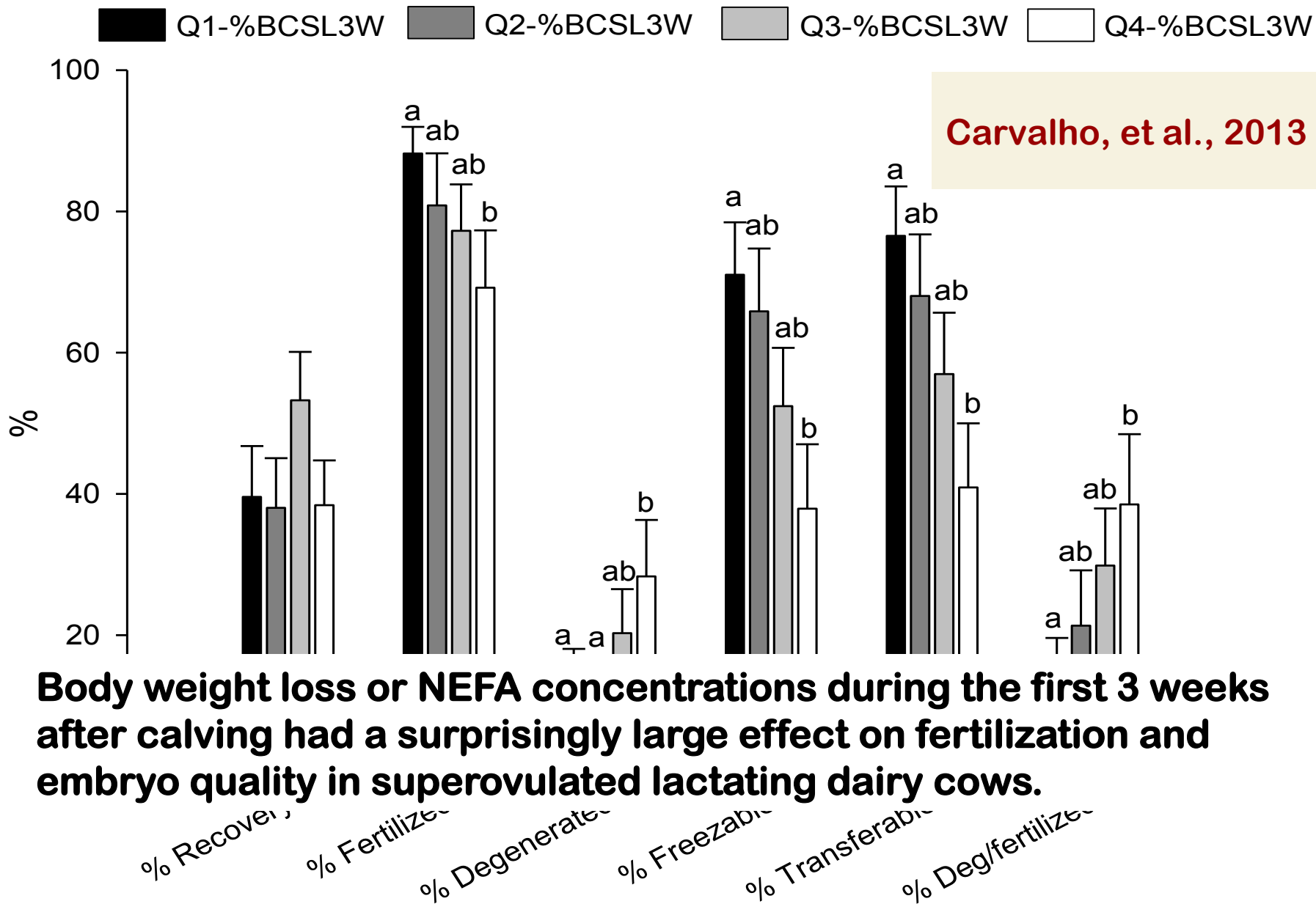
# Circulating AMH is a strong predictor of superovulatory response



# Carvalho, et al., 2013; Effect of postpartum body weight change and circulating NEFA on embryo quality in superovulated dairy cows.

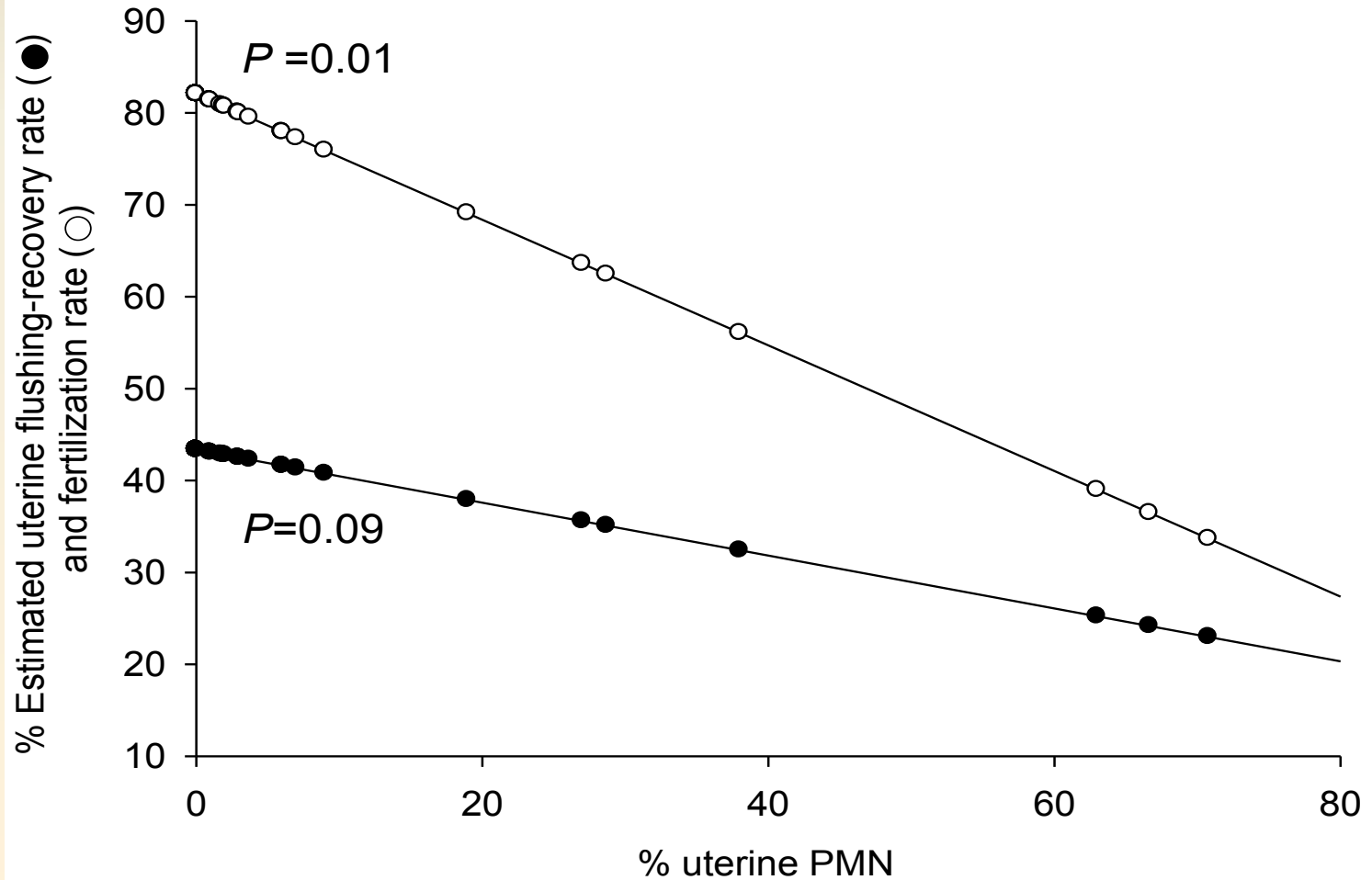
| <b>r<sup>2</sup></b><br><b>P Value</b> | <b>%</b><br>Degenerated          | <b>%</b><br>Fertilized          | <b>% BWC</b><br>3 <sup>rd</sup> week | <b>% BWC</b><br>10 <sup>th</sup> week |
|--|----------------------------------|---------------------------------|--------------------------------------|---------------------------------------|
| NEFA                                   | <b>0.23</b><br><b>0.05</b>       | <b>-0.13</b><br><b>0.27</b>     | <b>-0.44</b><br><b>&lt;0.001</b>     | <b>-0.57</b><br><b>&lt;0.001</b>      |
| % Transferable                         | <b>-0.67</b><br><b>&lt;0.001</b> | <b>0.66</b><br><b>&lt;0.001</b> | <b>0.40</b><br><b>0.001</b>          | <b>0.38</b><br><b>0.002</b>           |
| % Degenerated                          | <b>1.00</b>                      | <b>-0.07</b><br><b>0.56</b>     | <b>-0.25</b><br><b>0.04</b>          | <b>-0.31</b><br><b>0.01</b>           |
| % Fertilized ova                       |                                  | <b>1.00</b>                     | <b>0.28</b><br><b>0.02</b>           | <b>0.21</b><br><b>0.09</b>            |
| % BWC 3 <sup>rd</sup> week             |                                  |                                 | <b>1.00</b>                          | <b>0.71</b><br><b>&lt;0.001</b>       |

Carvalho, et al., 2013



**Body weight loss or NEFA concentrations during the first 3 weeks after calving had a surprisingly large effect on fertilization and embryo quality in superovulated lactating dairy cows.**

# Effect of subclinical endometritis of fertilization



## **SPECIFIC AIM 3**

# **Quantify the impact of mastitis on fertility and pregnancy loss in lactating dairy**

### **Objective:**

1. Determine if the occurrence of pregnancy loss will be greater in cows that experience mastitis.
2. Help to elucidate associations between mastitis case severity, pathogen type, and pregnancy loss.

### **Data:**

Data obtained from 2,800 cows from four large commercial dairy herds which are being enrolled from calving until confirmed pregnant at 150 days post-AI



# FARM 1. IMPACT OF MASTITIS ON REPROD

| PARAMETER                 | NO<br>MASTITIS    | MASTITIS        |                               |                               |
|---------------------------|-------------------|-----------------|-------------------------------|-------------------------------|
|                           |                   | Before AI       | Between<br>AI and 1st<br>Preg | After 1 <sup>st</sup><br>Preg |
| Conception rate           | 37.3<br>(179/480) | 22.9<br>(19/83) | 11.8<br>(4/34)                | 35.9<br>(46/128)              |
| Service per<br>conception | 2.3               | 2.4             | 2.9                           | 2.5                           |
| DIM at first CM           | 0.0               | 34.8            | 96.3                          | 194.6                         |
| Days open                 | 147.5             | 163.2           | 190.9                         | 161.7                         |
| Interbreeding<br>interval | 42.5              | 42.6            | 45.5                          | 41.1                          |

# Ongoing Results

| FARM         | COWS ENROLLED | CLINICAL CASES |
|--------------|---------------|----------------|
| Farm 1       | 739           | 1070           |
| Farm 2       | 747           | 2626           |
| Farm 3       | 636           | 285            |
| Farm 4       | 114           | 70             |
| <b>Total</b> | <b>2,236</b>  | <b>4,051</b>   |

- A total of 2800 cows need to be enrolled
- First TAI pregnancy loss is expected to be 4%
- Cases (pregnancy losses) will be compared against controls (pregnant cows) to determine the effect of mastitis in reproductive performance.

## **SPECIFIC AIM 4**

the impact of reproductive performances using optimization and simulation techniques.

### **Objective 1:**

Examine the use and impact of synchronization programs on reproductive performance and culling rates in dairy herds in WI.

### **Objective 2:**

To assess the economic impact of reproductive performance of dairy cattle under optimal replacement policies using optimization techniques.

### **Data:**

Data was obtained DHI records and benchmarks.



"This project is supported by Agriculture and Food Research Initiative Competitive Grant no. 2010-85122-20612 from the USDA National Institute of Food and Agriculture." Project started March 1<sup>st</sup>, 2010



Souza et al., 2013

## **Impact of timed AI use on reproduction and culling rate in Wisconsin dairy herds.**

Wisconsin farms (n = 200; 600 cows/herd; 12,427 kg/lactation).

Determined in each herd using DC305.

Divided into quartiles

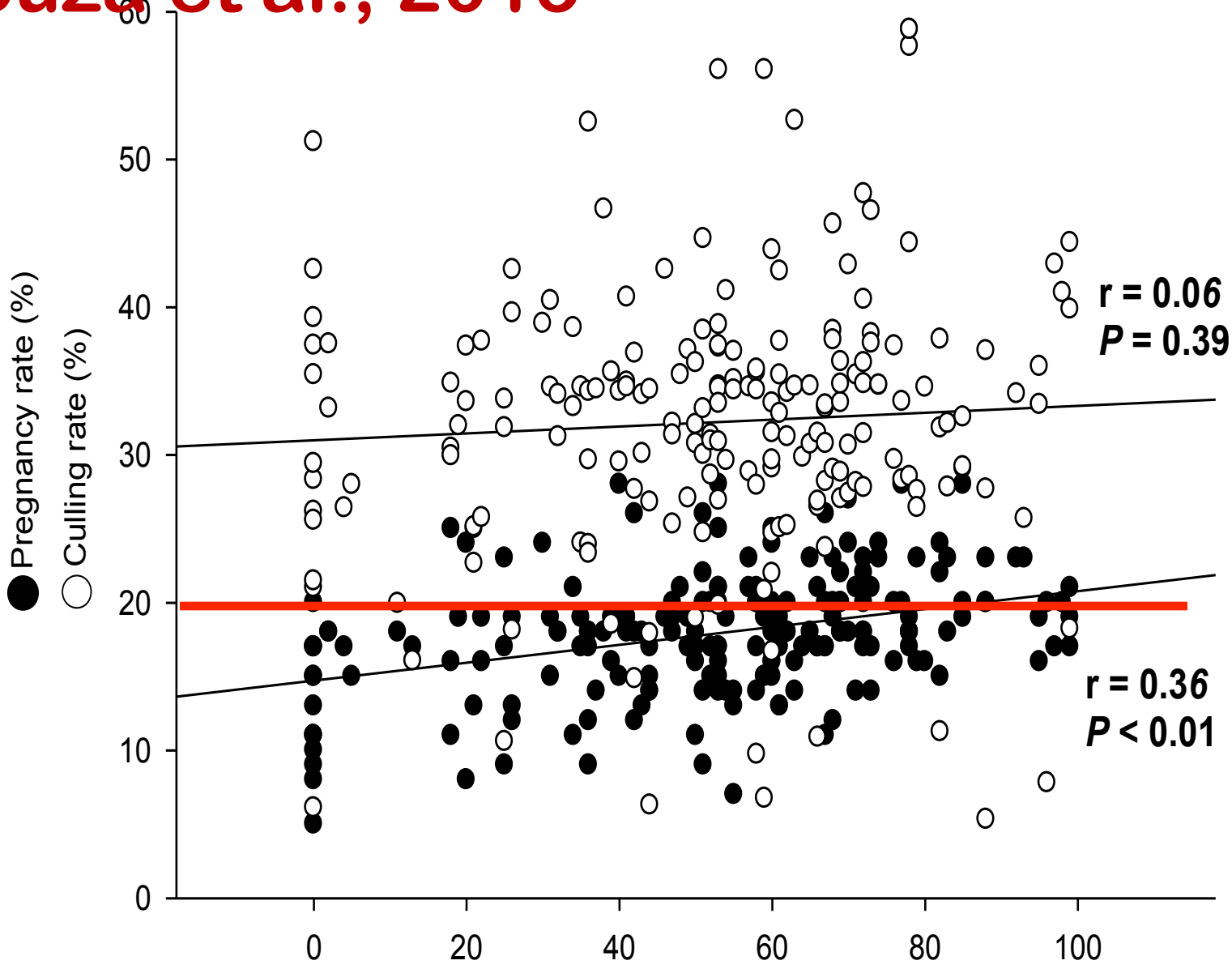
(0-36%; 37-55%; 56-69%; 67-99%)

Average VWP was 57.4 d and % pregnant to 1<sup>st</sup> AI (P/1<sup>st</sup>AI) was 36.6%.

# Souza et al., 2013

| Variable                                | Quartiles - use of timed AI* |                    |                                |                    |
|---|------------------------------|--------------------|--------------------------------|--------------------|
|   | Q1<br>(0% to 36%)            | Q2<br>(37% to 55%) | Q3<br>(56% to 69%)             | Q4<br>(70% to 99%) |
| Average lactating herd size (n)         | 663.5                        | 673.8              | 719.5                          | 582.4              |
| Voluntary waiting period (days)         | 49.6 <sup>a</sup>            | 53.8 <sup>a</sup>  | 58.9 <sup>a</sup> <sup>b</sup> | 67.5 <sup>b</sup>  |
| Conception to 1 <sup>st</sup> AI (%)    | 35.2 <sup>a</sup>            | 35.5 <sup>a</sup>  | 36.1 <sup>a</sup>              | 39.6 <sup>b</sup>  |
| Service rate, VWP set at 50DIM (%)      | 48.7                         | 53.0               | 53.0                           | 49.9               |
| Pregnancy rate, VWP at 50DIM (%)        | 15.9                         | 17.2               | 16.9                           | 17.5               |
| Service rate, actual VWP (%)            | 48.8 <sup>a</sup>            | 54.4 <sup>b</sup>  | 56.9 <sup>b</sup>              | 57.1 <sup>b</sup>  |
| Pregnancy rate, actual VWP (%)          | 15.8 <sup>a</sup>            | 17.7 <sup>b</sup>  | 18.1 <sup>b</sup>              | 19.9 <sup>b</sup>  |
| Culling rate (%)                        | 35.8                         | 37.7               | 35.4                           | 40.5               |
| Days open, pregnant cows (days)         | 139.6 <sup>a</sup>           | 133.3              | 136.2                          | 130.8 <sup>b</sup> |
| Days in milk, all lactating cows (days) | 209.4 <sup>a</sup>           | 201.4              | 205.2                          | 199.6 <sup>b</sup> |

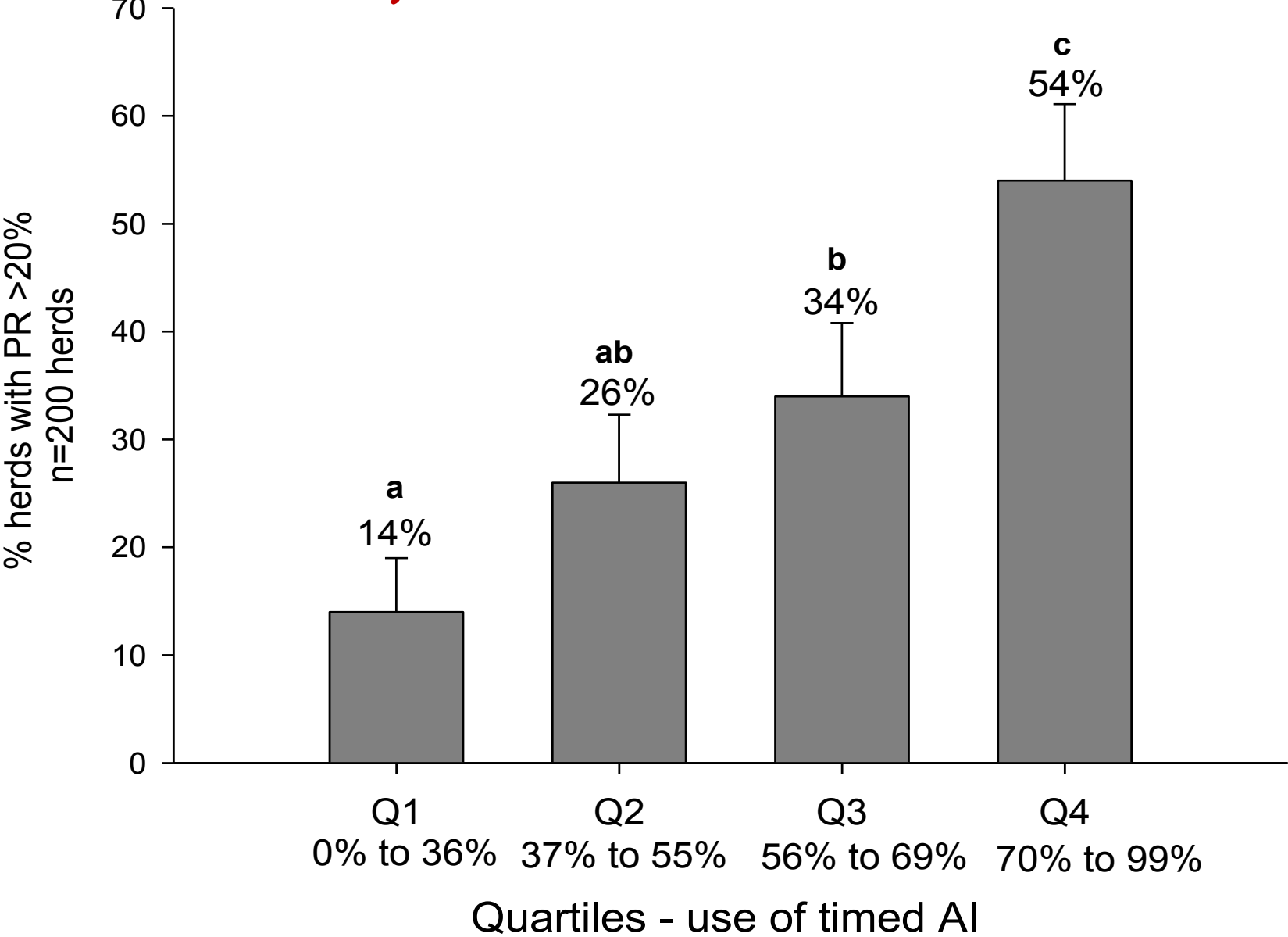
# Souza et al., 2013



Percentage of total AIs performed following timed AI (%)

n=200 herds in WI

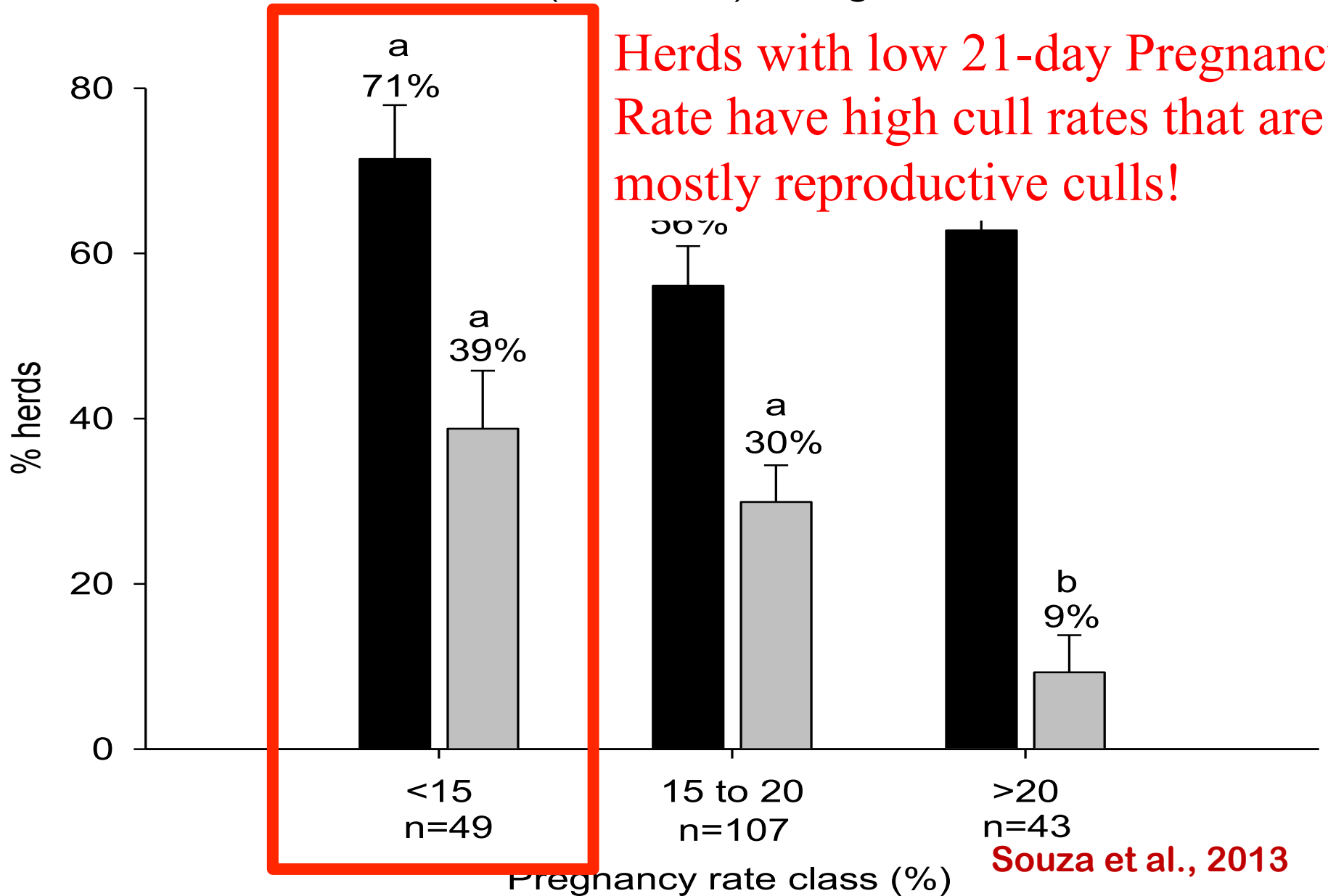
# Souza et al., 2013



# Souza et al., 2013

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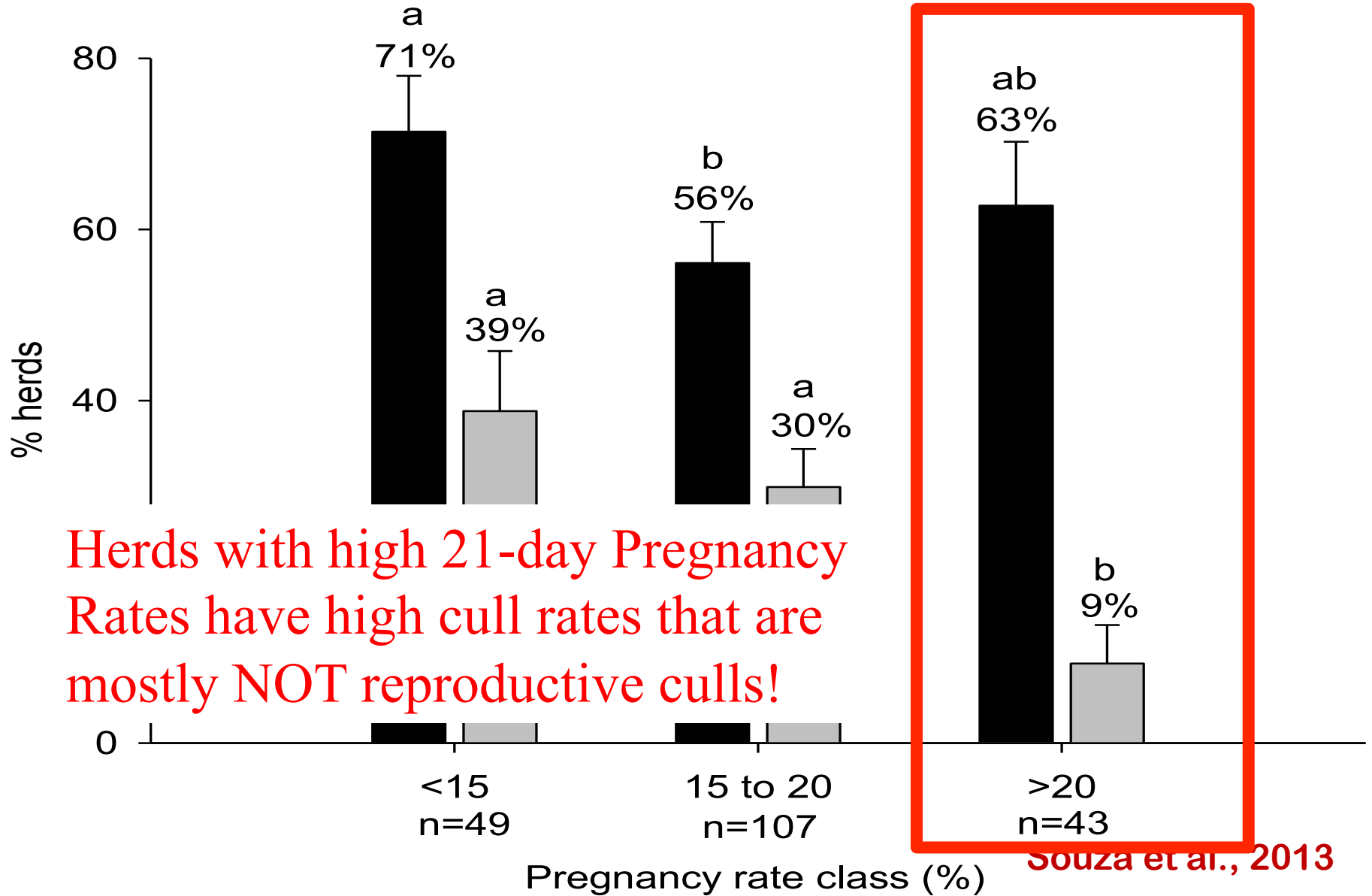
■ Herds with culling rate > 30%  
■ Herds with late (>330 DIM) culling rate > 30% of total culls



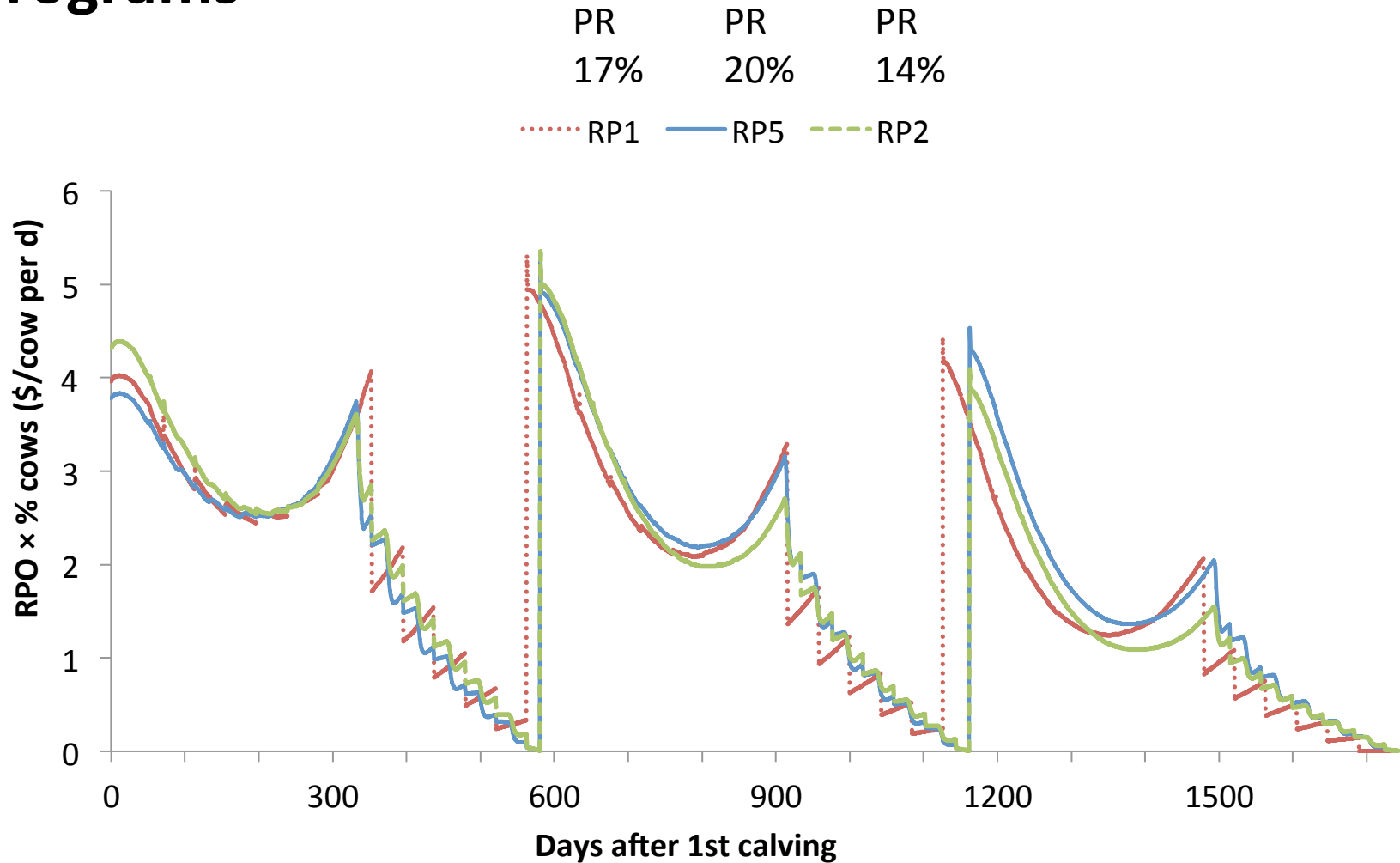
Herds with low 21-day Pregnancy Rate have high cull rates that are mostly reproductive culls!

Souza et al., 2013

■ Herds with culling rate > 30%  
■ Herds with late (>330 DIM) culling rate > 30% of total culls



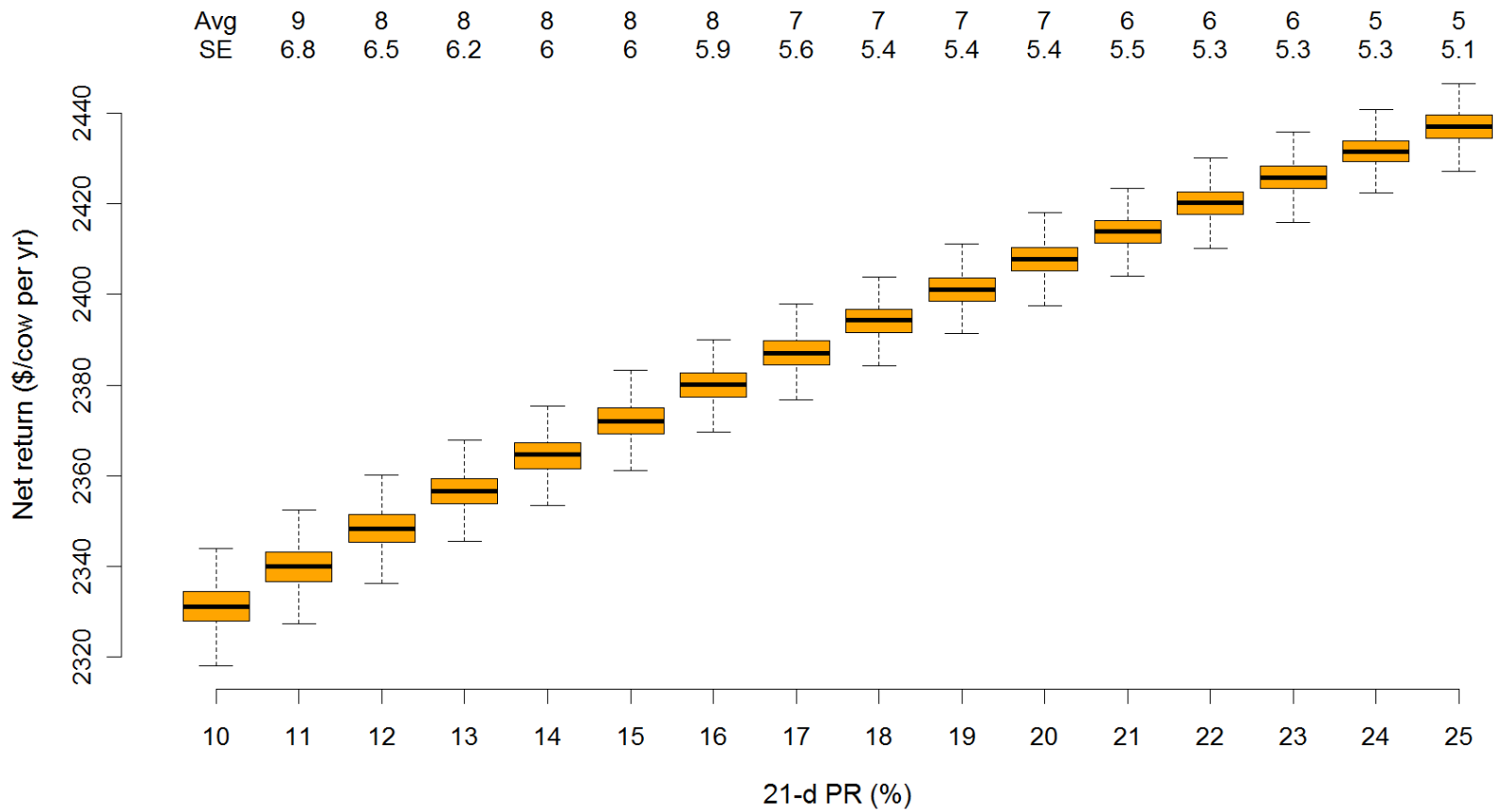
# Sources of differences between reproductive programs



- A. S. Kalantari and V. E. Cabrera (2012)

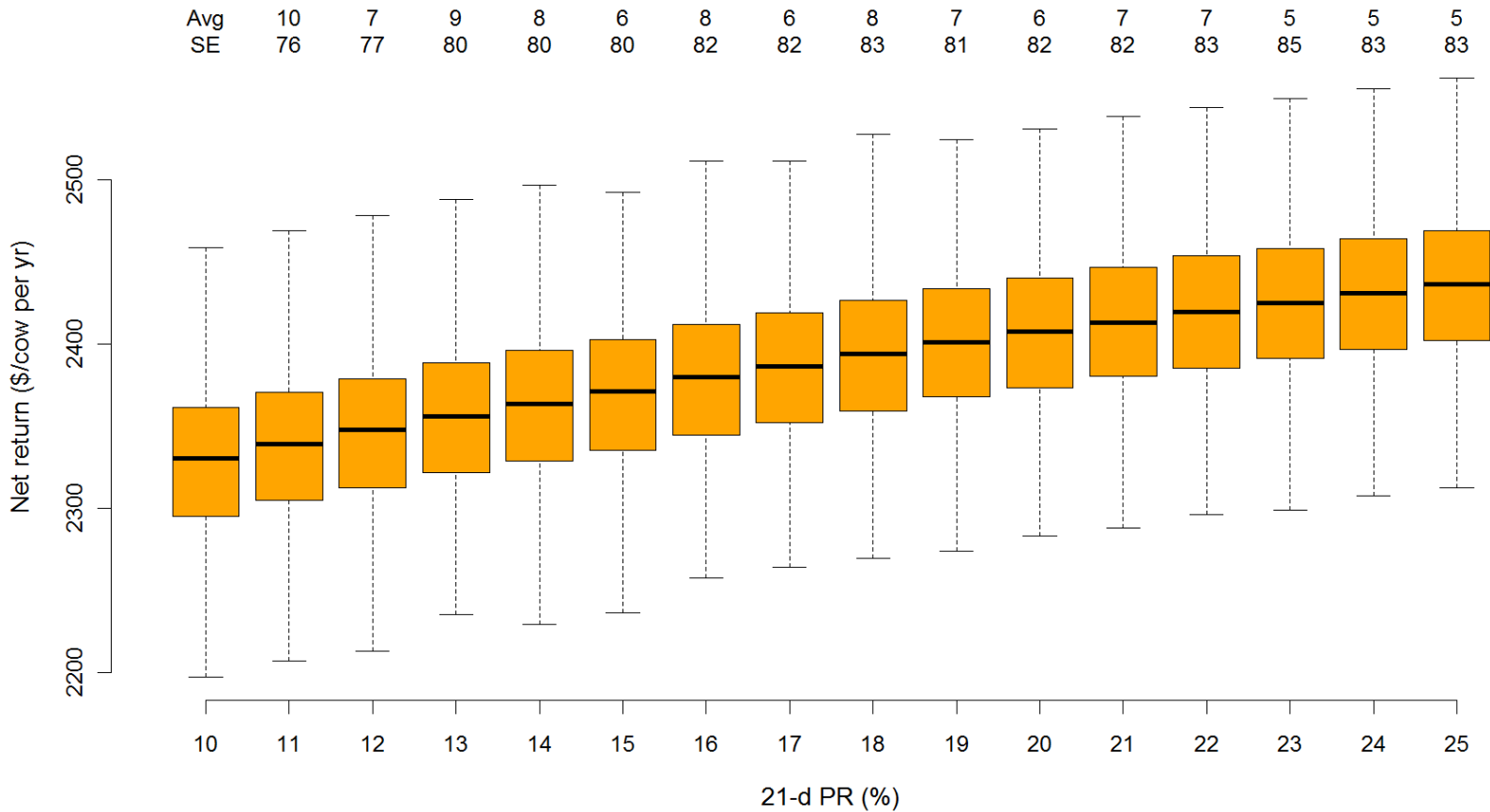


# Variation in the net return without Milk production variation with 2,000 Replicates



- A. S. Kalantari and V. E. Cabrera (2013; in review)

# Variation in the net return including Milk production variation with 2,000 Replicates



# SPECIFIC AIM 5

## Repro Money Program



### Objective:

To improve reproductive efficiency and profitability of the dairy farms by means of farmer-directed team-base program called Repro Money

### Data:

Data obtained from farm records of all farms that enrolled and completed the program (n=13) between Fall 2010 and Summer 2012 was analyzed before and after participating in the program to evaluate the program's impact on enhancing reproductive performance and herd profitability.



"This project is supported by Agriculture and Food Research Initiative Competitive Grant no. 2010-85122-20612 from the USDA National Institute of Food and Agriculture." Project started March 1<sup>st</sup>, 2010

| <b>OUTCOME</b>        | <b>BEFORE PROGRAM</b> | <b>AFTER PROGRAM</b> | <b>IMPROVEMENT</b> | <b>P-value</b> |
|-----------------------|-----------------------|----------------------|--------------------|----------------|
| 21-d Preg Rate (%)    | 19                    | 22                   | 3                  | 0.0083         |
| Insemination Risk (%) | 53                    | 56                   | 5                  | 0.0003         |
| Conception Rate (%)   | 37                    | 42                   | 3                  | 0.0016         |
| Cows AI 21d           | 61                    | 79                   | 17                 | 0.0141         |

**Economic Gain = \$55/cow per year  
(0 to \$278)**

# Repro Money Extension Efforts



- Webpage for program news & updates
- 27 dairy farms currently enrolled and ongoing
- National meetings poster presentations ( ADSA, DCRC, AABP)
- State-wide yearly workshop showcasing our results:
  - Winter 2013: 22 workshops
  - More than 500 participants
    - Farmers, industry representatives, technical college students, industry professionals.

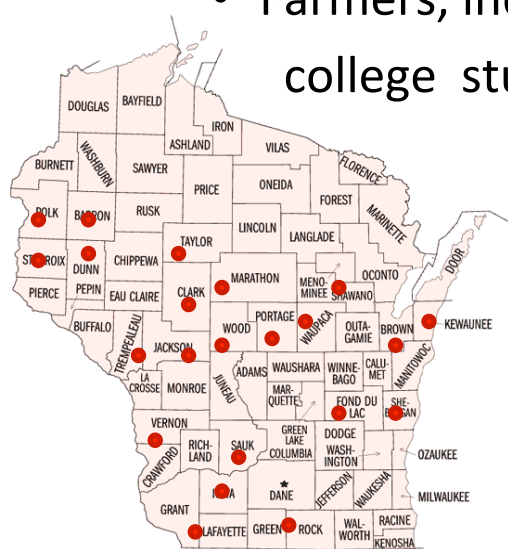


Figure 1: Workshop locations

**2013 Repro Money Road Show Series**  
**Reproducing Profitability Workshop**

**SPEAKERS & TOPICS**

**Dr. Victor Cabrera**  
 "The economic value of a dairy cow, the value of a pregnancy, the cost of a pregnancy loss and their relationship with reproduction performance"

**Dr. Paul Fricke**  
 "Update on reproduction research at UW Madison"

**Dr. Pamela Ruegg**  
 "Subclinical mastitis-the silent disease that can reduce reproductive performance"

**Dr. Pat Hoffman**  
 "Dairy heifer management in the age of genomics"

**Dr. Amy Stanton**  
 "Eliminating road blocks to getting your heifers pregnant: The impact of early calf health and rearing on fertility and time to first calving"

**Dr. Randy Shaver**  
 "What's new with nutrition and reproduction in dairy cows?"

**Dr. Connie Cordoba**  
 "How the Repro Money Program can increase your net income by enhancing your herd's reproductive efficiency"

This solution-oriented workshop will provide you with practical ideas delivered by UW experts who will share proven methods and decision-making tools to help you better manage your reproductive program.

**Contact your county agent for details**

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Accelerated Genetics

Pfizer Animal Health

ANIMART  
 Dairy & Livestock Solutions

NorthStar Cooperative

Figure 2: Workshop flyer

# An Integrated Approach To Improve Dairy Cow Fertility



**Specific Aim 1: Predicting Fertility**

**Saleh Shahinfar, Ph.D. Student**

**Kent Weigel, Professor and Chair Breeding and Genetics**



**Specific Aim 2: Nutrition-Reproduction**

**Alexandre Souza, Post Doctoral fellow**

**Milo Wiltbank, Professor -Reproductive Physiology**

**Randy Shaver, Professor - Nutrition**



**Specific Aim 3: Mastitis-Reproduction**

**Maria José Fuenzalida Valenzuela, Ph.D. Student;**

**Paul Fricke, Professor - Reproductive Physiology**

**Pam Ruegg, Professor - Physiology and Management**

**Carol Hulland, Milk Quality**



**Specific Aim 4: Economics of Reproduction**

**Afshin Kalantari, Ph.D. Student**

**Victor Cabrera, Assistant Professor- Management**

**Specific Aim 5: Repro Money Extension Program**

**Victor Cabrera, Assistant Professor- Management**

**Connie Cordoba, M.V, M.S.; Extension Specialist**



United States Department of Agriculture  
National Institute of Food and Agriculture