Sections

Status: Approved

Basic Information

Additional Info

Basic Information

- Project No. and Title: NE1727 : Influence of ovary, uterus, and embryo on pregnancy success in ruminants (/projects/18329)
- Period Covered: 10/01/2017 to 09/30/2018
- Date of Report: 07/16/2018
- Annual Meeting Dates: 05/13/2018 to 05/15/2018

Participants

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Brief Summary of Minutes of Annual Meeting

Attached file:

NE1727 Minutes 2018.pdf (/system/Sea/minutes_attachments/000/051/298/original/NE1727 Minutes 2018.pdf)

Accomplishments

Identification of the impact that thermal stress has on ovarian signaling is resulting in a mechanistic map being developed, upon which strategies to ameliorate seasonal infertility can be based.

We have identified that dairy cows have an amazing capacity to develop tolerance to chronic lipopolysaccharide exposure and that no effect of this exposure on the growing dominant follicle is observed.

Molecular signaling protein alterations during the follicular and/or luteal phase from lipopolysaccharide or thermal stress treated gilts have been identified.

Determined influence of a physiological stressor, obesity, on the ovarian capacity to respond to a toxic environmental stress.

Identified molecular proteins within the oocyte that contribute to viability of the oocyte.

We have cultured ovarian oocytes to investigate intra-oocyte mechanisms of heat stress induced infertility.

Zinc depletion during preantral development impairs oocyte-somatic cell communication and oocyte growth and competence to complete meiosis.

NR5A2 is a regulator of luteal progesterone production and may be translationally regulated by miRNA.

Changes in the profile of lipids within the CL indicate potential roles for these small molecules as regulators of immune cells, components of intercellular signaling pathways and cell death.

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We have shown that bovine granulosa and theca express CCN1, along with αV , αIIb , $\alpha 6$, $\beta 1$, $\beta 3$ and $\beta 5$ integrin subunits. The suppressive effect of PSI on CCN1 expression in KGN cells was accompanied by a decrease in phosphorylated NF- κ B activity, while the suppressive effect of Calphostin C and PSI on CCN1 expression in HGrC1 cells probably did not involve NF- κ B.

We have determined the effects of Ovsynch manipulation on intraovarian events and pregnancy in multiparous dairy cows.

We have manipulated ERK signaling (via growth factors) in bovine granulosa cells (GCs) to gain insight about its role in immune-mediated apoptosis of GCs and selection of bovine follicles.

Our results provide new information on the property of ZNFO, which will help further elucidation of the molecular mechanisms involved in ZNFO-dependent transcriptional regulation during maternal- embryonic transition.

We determined that feeding rumen-protected methionine to high-producing, multiparous dairy cattle reduced pregnancy loss in multiparous dairy cows.

We determined that high energy diets could reduce oocyte quality, leading to lack of fertilization.

We have defined two other mechanisms that underlie reduced fertility in animals fed high energy diets, namely high progesterone catabolism and reduced embryo quality.

An important practical finding was that loss of body condition score from 21 days before calving until 21 days after calving led to reduced fertility in lactating dairy cows.

We have determined that early plane of nutrition impacts the progression of uterine gland development in heifers, and that this outcome may be mediated by changes in the expression of several local controllers of gland development.

Our recent findings support the contention that maternal obesity modifies a subset of important mediators of embryonic and extraembryonic development in ovine and porcine conceptuses.

We determined that maternal obesity also modifies uterine gene expression during early pregnancy in sheep.

We determined that Bos indicus and Bos taurus cattle differ in their ability to produce viable offspring following nutrient restriction during early pregnancy.

We have discovered that the cytokine, interleukin-6, improves development of the inner cell mass (ICM) in bovine embryos.

We determined that early weaning of beef heifers can have long-term effects on physiology. Some of these changes have implications for lifetime productivity and may be beneficial in some production systems.

We investigated the effects of steroid content of follicle fluid on the cumulus-oocyte complex. Cumulus cell expansion and gene expression differed with estradiol and progesterone concentrations, but in vitro embryo development to the blastocyst stage did not.

We determined that a lack of Ca2+ oscillations is not the main reason of the poor success of ICSI in the bovine. This was demonstrated using both in in vitro matured mouse and bovine oocytes injected with bovine sperm.

We found that bovine sperm heads are highly resistant to sperm head decondensation, which compromises the ability of in vitro maturated oocytes to reprogram the sperm nucleus.

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We identified treatments that abruptly change the metabolic status of bull sperm, enhancing the ability of those sperm upon ICSI to initiate Ca2+ oscillations. This also improved the ability of these sperm to undergo decondensation and support preimplantation embryo development.

We have developed the Short-Resynch protocol. This new reproductive management strategy for re-insemination of lactating dairy cows improves reproductive performance and has other benefits for dairy herd management. Compared to traditional resynchronization strategies widely used on dairy farms, Short-Resynch increased pregnancy rates, increased the fertility of subgroups of sub-fertile cows, while taking advantage of inseminations at detected estrus after a previous insemination.

We demonstrated that the Short-Resynch program resulted in similar pregnancy rate than one of the most effective resynchronization of ovulation programs available for dairy farms (i.e., Resynch-25) thorough more inseminations at detected estrus, a short interval between inseminations, and increased P/AI for sub-fertile cows.

We found that it is possible to collect up to five CL biopsies 48 h apart on the same CL of lactating dairy cows. Five serial CL biopsies from the same CL in lactating dairy cows did not reduce circulating concentrations of P4, CL size, or affect blood flow. Serial CL biopsies did not cause CL regression during the period of biopsy collection.

We determined that a reproductive management strategy designed to increase insemination of cows in estrus after NPD resulted in similar reproductive performance (time to pregnancy) than a typical resynchronization of ovulation protocol used in dairy farms.

Proof of concept studies were performed using in vitro mouse follicles and COCs to understand how TNF and H_2O_2 signaling impact transcriptional and post-transcriptional mRNA synthesis and stability, respectively. We have demonstrated that obesity induces ovarian inflammation which in turn increases the abundance of oocyte-specific transcription factors, oocyte-specific growth factors, and maternal effect genes. Current studies are discriminating between transcriptional and post-transcriptional regulation of these candidate mRNAs in primary follicles, secondary follicles, and cumulus-oocyte complexes by acute treatments with TNF and H_2O_2 . Pathway activation by TNF and H_2O_2 in follicles and COCs are also being elucidated.

We determined that increased abundance of specific polymorphic alleles in the IGF-I and TNFa genes in Holstein cows are favorably associated with increased conception rate to 1st AI and a shorter calving to conception interval during lactation ie. higher fertility.

We found that sperm epigenome (chromatin dynamics) as well as functional genome (macromolecules) are associated with bull fertility, and identified key sperm and seminal plasma metabolites that are associated with bull fertility.

Impacts

1. Low pregnancy rates in beef and dairy cows result in increased costs to producers associated with rebreeding or culling animals, and recouping heifer development costs. Reduced fertility is also a major cause of reduced milk (dairy) and meat (beef) production which impacts the food supply. The focus of the NE1727 multistate project "Influence of ovary, uterus, and embryo on pregnancy success in ruminants", is to understand how altered ovarian function, impaired oocyte quality, and disruptions of conceptus-uterine interactions contribute to infertility, and devise management strategies that will overcome these factors in order to optimize the chances that animals successfully achieve a pregnancy. To achieve this goal, the following three objectives will be performed: (1) Determine the impact of altered ovarian function on reproductive performance; (2) Identify alterations in embryo development and uterine and CL function associated with declining pregnancy establishment; and (3) Identify changes in genetics and reproductive management that lead to improved pregnancy rates. As investigators in NE1727, we will use our collective expertise to understand the mechanisms by which internal and external forces modify these important physiological processes and to use this information about core biological processes to rationally develop new and innovative tools to increase cyclicity and breeding efficiency, and reduce early embryonic failures, thereby improving reproductive efficiency of both dairy and beef cattle operations in the United States.

Publications

Peer-reviewed journal articles published in 2017 reporting research from this project

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Kvidera, S.K., Horst, E.A., Sanz-Fernandez, M.V., Abuajamieh, M., Ganesan, S., Gorden, P.J., Green, H.B., Schoenberg, K.M., Trout, W.E., Keating, A.F., Baumgard, L.H. 2017. Characterizing effects of feed restriction and glucagon-like peptide 2 administration on biomarkers of inflammation and intestinal morphology. Journal of Dairy Science. 100(11):9402-9417.

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Romereim SM, Summers AF, Pohlmeier WE, Zhang P, Hou X, Talbott HA, Cushman RA, Wood JR, Davis JS, and Cupp AS (2017) Gene expression profiling of bovine ovarian follicular and luteal cells provides insight into cellular identities and functions, *Mol Cell Endocrinol* 439:379-394 PMID: 27693538

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Li, Q., R. Hegge, P.J. Bridges and J. C. Matthews. 2017. Pituitary genomic expression profiles of steers are altered by grazing of high vs. low endophyte-infected tall fescue forages. PLoS One. 12(9): e0184612.

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Wijma, M. M. Pérez, M. Masello, D. G. García, M. L. Stangaferro, and J.O. Giordano. 2017. A resynchronization of ovulation program based on ovarian structures present at nonpregnancy diagnosis reduced time to pregnancy in lactating dairy cows. J. Dairy Sci. 101:1697-1707.

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Lu, Y., Bradley, J.S., McCoski, S.R., Gonzalez, J.M., Ealy, A.D. and Johnson, S.E. 2017. Reduced skeletal muscle fiber size following caloric restriction is associated with calpain-mediated proteolysis and attenuation of IGF-1 signaling. Am. J. Regul. Integr. Comp. Physiol. 312:R806-R815.

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MacGhee, M.E., Bradley, J.S., McCoski, S.R., Reeg, A.M., Ealy, A.D. and Johnson, S.E. 2017. Plane of nutrition affects growth rate, organ size and skeletal muscle satellite cell activity in newborn calves. J. Anim. Physiol. Anim. Nutr. 101:475-483.

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Conference papers and abstracts/presentations:

Yost, E.E., Arzuaga, X., Carlson, L., Keating, A.F., Lehmann, G. 2017. Focusing and refining the evaluation of reproductive endpoints in a systematic review of PCBs. International Symposium on Systematic Review and Meta-Analysis of Laboratory Animal Studies.

Ganesan, S., Nteeba, J. and Keating, A.F. 2017. Impact of glyphosate on ovarian signaling pathways regulating folliculogenesis and steroidogenesis. Society for the Study of Reproduction annual meeting.

Hines, E.A., Romoser, M., Keating, A.F., Baumgard, L.H., Niemi, J., Williams, N.H., Haberl, B., Kerr, B., Touchette, K.T., Ross, J.W. 2017. Supplementation of arginine does not improve gilt reproductive performance under commercial conditions. Society for the Study of Reproduction annual meeting.

Hines, E.A., Romoser, M., Keating, A.F., Baumgard, L.H., Niemi, J., Haberl, B., Kerr, B., Touchette, K., Ross, J.W. 2017. Effect of maternal arginine supplementation on offspring performance of pigs in a commercial production environment. Importance of nutrition and environment on birth weight, muscle growth, health and survival of the neonate workshop meeting.

Baumgard, L.H., Kvidera, S.K., Horst, E.A., Dickson, M.J., Ydstie, J.A., Shouse, C.S., Mayorga, E.J., Al-Qaisi, M., Lei, S., Bidne, K.L., Seibert, J.T., Hall, B.J., Keating, A.F., Ross, J.W., Selsby, J.T., Rhoads, R.P. 2017. Consequences of leaky gut on the immune system, metabolism, physiology and animal performance. American Society of Dairy Science annual meeting.

Dickson, M.J., Kvidera, S.K., Horst, E.A., Ydstie, J.A., Bidne, K.L., Wiley, C.E., Gunn, P.J., Keating, A.F., Baumgard, L.H. Chronic lipopolysaccharide infusion has no impact on dominant follicular size but affects 17β-estradiol in lactating dairy cows. American Society of Dairy Science annual meeting.

Kvidera, S.K., Horst, E.A., Sanz Fernandez, M.V., Abuajamieh, M., Ganesa, S., Gordon, P.J., Green, H.B., Schoenberg, K.M., Trout, W.E., Keating, A.F., Baumgard, L.H. 2017. Glucagon-like peptide 2 administration improves biomarkers of inflammation and intestinal morphology in feed restricted lactating Holstein cows. American Society of Dairy Science annual meeting.

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McCain AR, Beede KA, Yates DT, and Wood JR (2017) Maternal and Paternal Obesity Differentially Affect Fetal Growth with Maternal Obesity Associated Growth Restriction Attributed to Decreased 11D-Hydroxysteroid Dehydrogenase Expression Society for the Study of Reproduction, Washington, DC

Timme KA, Xie F, Davis JS, and Wood JR (2017) Ovarian Inflammation and Oxidative Stress Associated with Diet Induced Obesity (DIO) Impacts RNA-Binding Protein Expression and Potentially mRNA Stability in the Murine Ovary and Oocyte. *Society for the Study of Reproduction*, Washington, DC

Romereim SM, Summers AF, Pohlmeier WE, McFee RM, Spuri-Gomes R, Kurz SG, Davis JS, Wood JR, and Cupp AS (2017) A High-Androgen Microenvironment Inhibits Granulosa Cell Proliferation and May Alter Cell Identity. *Society for the Study of Reproduction*, Washington, DC

Abedal-Majed MA, Hart ML, Largen V, Magamage MPS, Kurz SG, Sargent KM, Bergman J, McFee RM, Cushman RA, Davis JS, Wood JR, and Cupp AS (2017) Ovarian Cortex from High A4 Cows Secrete Excess A4 and Exhibits Increased Oxidative Stress, Macrophage Markers and Arrested Follicle Development Which can be Partially Rescued by Angiogenic VEGFA Isoforms. *Society for the Study of Reproduction*, Washington, DC

Nafziger S, Abedal-Majed MA, Tenley S, Summers A, Hart ML, Harsh G, Bergman J, Kurz SG, Wood JR, Cushman RA, and Cupp AS (2017) Endocrine Profiles during Attainment of Puberty may Predict Reproductive Longevity in Heifers. *Society for the Study of Reproduction*, Washington, DC

Wood JR (2017) Maternal Obesity, the Gut Microbiota, and Oocyte mRNAs: Potential Impact on the Developing Embryo and Fetus. 4th World Congress on Reproductive Biology, Naha, Okinawa, Japan (invited speaker)

McCain AR, Beede KA, Yates DT, Shankar K, and Wood JR (2017) Maternal Obesity Results in Fetal Growth Restriction Associated with Reduced Placental Efficiency and an Altered Placental Transcriptome. 14th Annual Gilbert Greenwald Symposium on Reproduction, Kansas City, KS

Timme KA, Xie F, Davis JS, and Wood JR (2017) Ovarian Inflammation and Oxidative Stress Associated with Diet Induced Obesity (DIO) Impacts RNA-Binding Protein Expression and Potentially mRNA Stability in the Murine Ovary and Oocyte. 14th Annual Gilbert Greenwald Symposium on Reproduction, Kansas City, KS

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Active collaboration within the group

NY, PA, NH, VT and WV: Samples collected to investigate associations between fertility outcomes and SNP in candidate genes from dairy cows.

IA and UK: Samples shared to determine the effect of LPS treatment on ovarian inflammation.

PA and WV: Samples shared for analysis of molecular regulators of luteal function.

VT and NY: Samples assayed (NY) for analysis of progesterone concentrations.

NY and WI: Samples collected to investigate the effect of supplementation with rumen-protected methionine on reproductive traits of lactating dairy cows.

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