## Effects of Vitamin E and Selenium on Periparturient Diseases and Fertility in Dairy Cattle

David Morrow, D.V.M., PhD. Department of Large Animal Surgery & Medicine J. W. Thomas, PhD. Department of Animal Sciences Michigan State University East Lansing, Michigan 48824 and R. James Main, D.V.M., Private Practitioner, Lowell, Michigan

Vitamin E and selenium are essential nutrients for cattle. Deficiencies of these two nutrients in cattle are reported to cause white muscle disease, weakness, scours, pneumonia, and unthriftiness in calves and abortions, stillbirths, retained placentas and reduced fertility in cows.

The functions of vitamin E and selenium are interrelated. Vitamin E reduces *in vivo* formation of lipoperoxidases which damage membranes. Selenium is a component of glutathione peroxidase which detoxifies lipoperoxides formed when vitamin E is deficient.

Review of the Literature

The first study to show a relationship between vitamin Eselenium supplementation and retained placenta in dairy cattle was conducted in England.<sup>9</sup> Treatments of 0 or 15 mg of selenium and 680 IU of vitamin E with 15 mg of selenium were injected intramuscularly 28 days prior to the projected calving date. The percent of cows calving in which the placenta had to be manually removed was 27, 0 and 7% respectively, suggesting that the injections of vitamin E and selenium were beneficial in preventing the occurrence of retained placenta.

The first studies in the United States to show a relationship between vitamin E-selenium supplementation and retained placenta in dairy cattle were conducted in Ohio which is a selenium deficient area. The injection of 50 mg selenium and 680 units of vitamin E approximately 21 days prepartum reduced the incidence of retained placentas from 38% in 26 control cows to 0% in 27 treated cows.<sup>5</sup> The plasma selenium values in the treated cows approached 0.1 ppm concentration at parturition. The injection of 50mg selenium and 680 units of vitamin E 20 days prepartum reduced the occurrence of retained placenta from 51% in 80 control cows to 9% in 133 treated cows in a subsequent Ohio field study.<sup>6</sup> Vitamin E-selenium therapy was also effective in reducing the occurrence of retained placenta in a Kentucky study<sup>4</sup> but ineffective in Maryland,<sup>11</sup> Virginia,<sup>3</sup> and New York<sup>2</sup> studies.

An examination of 243 aborted bovine fetuses from British columbia and the State of Washington found that 28% were selenium deficient based on the fluorometric assay of fetal liver.<sup>8</sup> This study suggested that a relationship may exist between selenium deficiency and bovine abortions. By extrapolation, stillbirths, weak calves and other reproductive problems may result from a selenium deficiency in cattle.

In another study, the serum selenium concentration at 14 to 21 days postpartum in 115 Holstein cows was positively correlated with services per conception and days open (P < 0.05).<sup>7</sup>

The objective of the study reported was to determine the effects of vitamin E and selenium injections during the dry period on periparturient disease and fertility in high producing dairy cattle.

#### Material and Methods

All cows due to calve during a 12-month period in the Michigan State 'University dairy herd were allocated at random to a treatment or control group. All animals in the treated group were injected with 50 mg selenium and 680 IU of vitamin  $E^{\pm}$  at 21 days prior to the anticipated parturition. Conventional feeding and management practices were followed. The cows were maintained on a selenium deficient diet and no supplemental vitamin E or selenium was provided. Blood samples were collected to monitor serum selenium values by the fluorometric method.<sup>10</sup>

Criteria evaluated included the following: dystocia (%), milk fever (%), retained placenta (%), metritis (%), displaced abomasum (%), inactive ovaries (%), cystic follicles (%), corpora lutea (%), interval to first estrus (days), interval to first breeding (days), and cows culled or died (%).

a Mu-Se provided by Burns-Biotec, Oakland, California 94621

#### **Results and Discussion**

There were 42 control and 40 treated cows. The mean serum selenium values were 0.018 ug/ml in both control and treated groups. Concentrations below 0.05 ug/ml are generally considered to be deficient by the clinical nutrition laboratory at Michigan State University.

The occurrence of dystocia was decreased only 2% by treatment while the occurrence of milk fever increased 6% (Table 1). There was essentially no change from treatment in the occurrence of displaced abomasum.

#### Table 1

Effects of Injecting 50 Mg Selenium and 680 Units of Vitamin E Three Weeks Prepartum on Periparturient Diseases and Fertility in 82 Dairy Cows

| Criteria                   | Control | Treated | P    |
|----------------------------|---------|---------|------|
| Number                     | 42      | 40      |      |
| Dystocia (%)               | 17      | 15      | N.S. |
| Milk Fever (%)             | 2       | 8       | N.S. |
| Retained Placenta (%)      | 31      | 23      | N.S. |
| Metritis (%)               | 38      | 26      | N.S. |
| Displaced Abomasum (%)     | 14      | 13      | N.S. |
| Inactive Ovaries (%)       | 12      | 6       | N.S. |
| Cystic Follicles (%)       | 16      | 20      | N.S. |
| Corpora Lutea (%)          | 72      | 74      | N.S. |
| Interval to first estrus   |         |         |      |
| (days)                     | 44      | 46      | N.S. |
| Interval to first breeding |         |         |      |
| (days)                     | 66      | 69      | N.S. |
| Days open                  | 139     | 144     | N.S. |
| Services/Conception        | 3.4     | 3.5     | N.S. |
| Calf Mortality (%)         | 17      | 22      | N.S. |
| Culled or died (%)         | 26      | 28      | N.S. |

Retained placenta decreased from 31 to 23% however these differences were not significant (P < 0.05).

Ovarian function, measured by the presence of inactive ovaries, cystic follicles, and corpora lutea, was not affected significantly by prepartum treatment with vitamin E and selenium (P < 0.05).

Fertility, measured by the intervals to first estrus and breeding, days open and services per conception, was not significantly affected by the injection of 50 mg selenium and 680 IU of vitamin E 21 days prepartum (P < 0.05).

The occurrence of calf losses during the first 30 days of age and cows culled or died during the subsequent lactation was not affected significantly by treatment (P < 0.05).

The parameters used to measure periparturient disease and fertility in this study were not affected significantly by the injection of 50 mg selenium and 680 units of vitamin E 21 days prepartum in selenium deficeint cows. These parameters are affected by many factors, such as disease, nutrition, season, sanitation, management and heredity which may partially account for the variable responses reported to vitamin E and selenium therapy. It is possible that the amount and/or length of supplementation was inadequate to influence these parameters in this study.

The daily requirement of vitamin E in a dairy cow has not been definitely established<sup>1</sup>: however 300 IU per day appears to be a reasonable amount. It is customary to supplement the ration daily with 50 to 100 IU of vitamin E.

The concentration of selenium in feeds is related to the content in the soil; however, acid soils tend to reduce the uptake of selenium by plants.<sup>1</sup> This factor likely contributes to the variable clinical signs and responses observed in cattle in selenium deficient areas.

The dietary requirement for selenium by cattle is estimated to be approximate 0.1 ppm, depending on the chemical form of selenium and the levels of interfering or enhancing factors in the diet such as vitamin E, sulfur, lipids, proteins and amino acids.<sup>1</sup> The maximum tolerable level of dietary selenium for dairy cattle is estimated to be 2 ppm. Increased amounts of protein and sulfur in the diet help to reduce the potential for toxicity problems. Naturally occurring organic selenium in plants appears to be more toxic than the inorganic form. The range between requirements and toxic levels is 30 to 50-fold.<sup>1</sup> Recommendations for Preventing Selenium Toxicity

1. Add only one selenium supplemented ingredient such as a protein supplement, trace mineral salt, or mineral mix to the diet. When selenium is added from several sources, it is possible to feed more than 2 mg daily. This practice may result in potential toxicity problems.

2. Mix the selenium supplemented ingredient with the ration rather than feed free choice when possible in order to more closely monitor selenium intake.

3. Feed selenium supplements only to the species designated on the feed tag. The species requirements are highly variable.

4. Mix feed thoroughly to provide for even distribution of selenium throughout the ration.

5. Read and follow directions closely.

#### References

1. Anon. Nutrient Requirements of Dairy Cattle, 5th rev. ed, National Acad. Sci., Washington, D. C. 18-19, 1978. - 2. Braund, D. G., 1979. Efficacy of Selenium Injection on the Incidence of Retained Placenta and Metritis. Agway R & D Capsules 1:3, 1979. - 3. Gwazdauskas, F. C., Bibb, T. L., McGilliard, M. L., and Lineweaver, J. A. Effects of Prepartum Selenium - Vitamin E Injection on Time for Placenta to pass and on Productive Functions. J. Dairy Sci., 62:978-981, 1979. - 4. Hemken, R. W., Olds, D., Botts, R. L., and Bull, L. S. Selenium Injections Prior to Calving on Prevention of Retained Placenta. Proc. 55th Annual Meeting A.D.S.A., Southern Div. 14., 1979. - 5. Julien, W. E., Conrad, H. R. Jones, J. E., and Moxon, A. L. The Prevention of Retained Placenta with Supplemental Selenium. J. Amer. Dairy Sci. Assoc., 59:1954-1959, 1976. -6. Jullen, W. E., Conrad, H. R., and Moxon, A. L. Selenium and Vitamin E and Incidence of Retained Placenta in Parturient Dairy Cows II. Prevention in Commercial Herds with Prepartum Treatment. J. Dairy Sci., 59:1960-1962, 1976. - 7. Larson, L. L., Marbruck, H. S. and Lowry, S. R. Relationship between Early Postpartum Blood Composition and Reproductive Performance in Dairy Cattle. J. Dairy Sci., 63:283-289, 1980.

「山田市」



Lutaly Sterile Solu

For Inframuscular Use for Estrus Synchronization

MIC 2000

Equix

in Beel Cattle and Non-Lactating Dairy Heiler

Introducing the dawn of a new era in cattle breeding management...

# Lutalyse (dinoprost tromethamine)

## <u>A naturally-occurring prostaglandin</u> for estrus synchronization in beef cattle and non-lactating dairy heifers.

Now with the introduction of Lutalyse you can provide your beef and dairy clients a unique opportunity to make cattle breeding management a realistic possibility without many of the difficulties typically associated with heat detection.

When Lutalyse is injected into normally-cycling beef cattle or non-lactating dairy heifers the prostaglandin stimulates luteolysis. The corpus luteum is naturally regressed and the cattle subsequently exhibit estrus with a closely synchronized ovulation. Prostaglandins are chains of fatty acids that are readily metabolized in the animal's system. Lutalyse is the tromethamine salt of the naturally-occurring prostaglandin F2 alpha. As a naturally-occurring compound, metabolic pathways already exist to handle its metabolism and excretion. Prostaglandin F2 alpha is natural and occurs in all mammals and man.

## Lutalyse can be your key to a total herd management program.

A number of management factors must be practiced by the producer to insure the effectiveness of Lutalyse...because it is not a substitute for poor management. And that's where you can play an important role.

You have a distinct opportunity to provide beef producers and dairymen with a total management program geared around Lutalyse, since this new product is available only to licensed, practicing veterinarians. Such a program would include an evaluation of facilities, nutrition, and herd health.

With your professional expertise and knowledge in these areas, together with Lutalyse, you can become the key to the development of a successful cattle breeding management program for your beef and dairy clients.

Perhaps only once every 20 years is a revolutionary product like Lutalyse introduced. And together, you and Lutalyse can create a new era in cattle breeding management.

Lutalyse from Upjohn ... The Timing Is Right!

### Lutalvse (dinoprost tromethamine)

#### Veterinary

## For intramuscular use for estrus synchronization in beef cattle and non-lactating dairy heilers.

#### DESCRIPTION

This product contains the naturally occurring prostaglandin F2 alpha (dinoprost) as the tromethamine sait. Each mI contains dinoprost tromethamine equivalent to 5 mg dino-prost, also, benzyl a cohol. 9 mg, and water for injection, q. 9 When necessary, pH was adjusted with sodium hydroxide and/or hydrochloric acid. Dinoprost tromethamine is a white or sliphtly off-white crystalline powder that is readily soluble in water at room temperature in concentrations to at least 200 mg/ml.

#### INDICATIONS AND INSTRUCTIONS FOR USE

For Intranuscular Use for Estrus Synchronization in Beef Cattle and Non-Lactating Dairy Heifers Lulalyse (dinoprost tromethamine) Sterile Solution is indicated for its luleolytic effects in beef cattle and in non-lactating dairy heifers. Lulalyse is used to control the timing of estrus and ovulation in estrous cycling cattle that have a corpus luleum

#### WHICH COWS AND HEIFERS WILL RESPOND TO LUTALYSE

Lutalyse is effective only in those normally estrous cycling animals having a corpus luteum, i.e., those which have ovulated at least five days prior to treatment. Lutalyse programs call for two injections 10 to 12 days apart. This avoids the need to consider the animal's precise day of the estrous cycle. Animals in a group situation that are not having estrous cycles will not be harmed by Lutalyse injection.

Many factors contribute to success and failure of reproduction management, and these factors are important also when time of breeding is to be regulated with Lutalyse. Some of these factors are:

- 1. Physical facilities must be adequate to allow cattle handling without being detrimental
- Priystan aumites most or adequate prior to and during the breeding season as this to the animal;
  Nutritional status must be adequate prior to and during the breeding season as this has a direct effect on conception and the initiation of estrus in heiders or return of estrous cycles in cows following calving;
  Cattle must be ready to breed—they must be estrous cycling and must be healthy.
  Estrus must be detected accurately if timed Al is not employed.
  Semen of high fertility must be used.
  Semen must be inseminated properly.

A successful A I, program can employ Lulalyse effectively, but a poor A I, program will continue to be poor when Lulalyse is employed unless other management deficiencies are remedied linst.

#### USE PROGRAMS ARE:

- Program I—Estrus Observation

- Inject 5 ml Lutalyse intramuscularly (25 mg dinoprost).
  Repeat the injection 10 to 12 days after the first injection, then,
  Observe for estrus after the second injection, and
  Inseminate at the usual time relative to detection of each estrus following the second
- injection. 5. If the catile are estrous cycling estrus is expected to occur 2 to 5 days after second injection. Cattle that do not become pregnant to that breeding will be expected to return to estrus between days 21 and 27 after the second injection

#### Program II—Timed Al

- Inject 5 ml Lutalyse (25 mg dinoprost) intramuscularly. Repeat the injection 10 to 12 days after the first injection, then, Inseminate about 80 hours after the second Lutalyse injection without estrus detection
- or observation Cattle that do not become pregnant to that breeding will be expected to return to estrus between 21 to 27 days after the second injection

Experimental data have demonstrated that pregnancy rates at 2 to 5 days after second injection in Program I and Program II, were markedly greater than pregnancy rates for contemporary controls. However, due primarily to the mechanics of Program 2 there was an increase in services per conception.

#### WARNINGS

Not for human use. Women of child-bearing age asthmatics, and persons with bronchial and other respira-tory problems should exercise extreme caution when handling this product. In the early stages, women may be unaware of their pregnancies. Dinoprost fromethamine is readily absorbed through the skin and can cause abortion and/or bronchiospasms. Direct con-tact with the skin should, therefore, be avoided. Accidental spillage on the skin should be washed off immediately with soap and water.

#### PRECAUTION

Do not administer to pregnant cows, as abortion may result

Do not administer intravenously (I.V.), as this route might potentiate adverse reactions.

#### ADVERSE REACTIONS

The most frequently observed side effect is increased rectal temperature at a 5x or 10x overdose. However, rectal temperature change has been transient in all cases observed and has not been delimental to the animal
 Limited salviation has been reported in some instances.
 Intravenous administration might increase heart rate.

#### DOSAGE AND ADMINISTRATION

Lulalyse (dinoprost tromethamine) is supplied at a concentration of 5 mg dinoprost per ml. Lutalyse is luteolytic in cattle at 25 mg (5 ml) administered intramuscularly. As with any multidose viail practice asoptic techniques in withdrawing each dose. Adequately clean and disinfect the vial closure prior to entry with a sterile needle.

#### HOW SUPPLIED

Lutatyse (dinoprost tromethamine) Sterile Solution is available in 10 ml vials

#### CAUTION

Federal (U.S.A.) law restricts this drug to use by or on the order of a licensed veterinarian

from the strong red line of



Veterinary Products, Kalamazoo, Michigan 49001

- 8. Taylor, R. F., Puls, R., and MacDonald, K. R. Bovine Abortions Associated with Selenium Deficiency in Western Canada, Amer. Assn. Vet. Lab. Diag. 22 Ann. Proc., 77-84, 1979. - 9. Trinder, N., Woodhouse, C. K., and Renton, C. P. The Effect of Vitamin E and Selenium on the Incidence of Retained Placenta in Dairy Cows. Vet. Rec. 85:550, 1969. -10. Whetter, P. A., and Ullrey, D. E. Improved Fluormetric Method for Determining Selenium. J. Assoc. Off. Anal. Chem., 61:927-930, 1978. -11. Williams, W. F., Yver, Dr. R., Diefenderfer, D. L., Douglas, L. W. and Vandersall, J. H. Influence of Prepartum Selenium-Vitamin E on Retained Placenta in Dairy Cattle. J. An. Sci., 45:326. Supp. 1., 1977.

This paper was presented at the XIII AABP Annual Convention, Toronto, Canada, November 19-22, 1980. (See Panel Discussion, p. 182-183 in the Proceedings).