Induction of Parturition in Cows with Misoprostol

Atilla Yildiz

Department of Dairy Animal Breeding, College of Sivrice Vocational,

University of Firat, Turkey

Abstract: The purpose of this study, was to determine the efficacy of intravaginal misoprostol for induction of parturition in cows. Twenty-seven multiparous Holstein cows with singleton pregnancies were used in this experiment. As randomly, cows were assigned to 1 of 2 treatment groups (n = 9) and a control group (n = 9). On day 270 of gestation cows in the treatment groups were received 200 μ g or 400 μ g of misoprostol (cytotec®) intravaginally. Intravaginal misoprostol was given every 6 h until the onset of labour. A maximum of 6 doses was administered. The average gestation length in control cows was 281 days. All induced cows (except for one cow administered 400 μ g misoprostol) calved between 24 and 72 h after administered misoprostol. The 400 μ g dosage of misoprostol resulted in induction failures of 11.1%. The rate of placental retention, birth weights, calving difficulty and calf viability did not differ among groups. Vaginal administration of misoprostol is an effective method of inducing parturition in the cow. Misoprostol could be used to produce a predictable calving time.

Key words: Misoprostol, induction of parturition, cow, intravaginal, labour

INTRODUCTION

The induction of parturition in cattle presents great economic incentives and significant economic benefits and advantages as a management tool animal husbandry (Mansell et al., 2006). Induction of parturition may be indicated to terminate pregnancy if desired, to advance the calving date in late conceiving cows, to synchronize the beginning of lactation, to terminate gestations, which have run too far over their time, to control fetal development where gross fetal oversize could be excepted to cause dystocia at full term parturition, to treat of uterine hydrops, chronic traumatic reticulitis, kidney diseases, circulatory incompetence with bronchial pneumonia, cardiac failure, fractures, or other health-related matters, in which salvage of the fetus or the life of the cow are being considered (Barth, 2006; Bazer and First, 1983; Holmes and Garcia, 1999; Shukla et al., 2008). Surgical techniques and pharmacological methods are available for this purpose. Various types and combinations of hormone treatments to induce parturition in cattle have been found applicable to synchronize calving for better and convenient calving management (Barth, 2006). Cortisol and prostaglandins analogues have been used for the on-farm induction of parturition in ruminants (Taverne, 1998). Misoprostol is a synthetic 15 deoxy-16 hydroxy 16-methyl analogue of the naturally occurring prostaglandin E₁. Several studies have found misoprostol effective as an agent for cervical ripening and induction

of labour (Aronsson et al., 2007; El-Sherbiny et al., 2001; Collins, 1990; Zeiman et al., 1997). Misoprostol has a number of advantages for clinical use. It is inexpensive, easy to store and stable at room temperature (Collins, 1990; El-Sherbiny et al., 2001; Goldberg et al., 2001; Sanchez-Ramos et al., 1993). Attempts to the induction of parturition with Misoprostol have been made only in women (El-Sherbiny et al., 2001), goats (Alan and Tasal, 2002) and ewes (Tasal et al., 2001). Similar to those studies, a study in cows had been not defined the beginning of parturition.

This study was undertaken to compare the efficiency and safety of 2 doses of intravaginal misoprostol in achieving induction of parturition.

MATERIALS AND METHODS

Twenty-seven multiparous Holstein cows with singleton pregnancies were used in this experiment. As randomly, cows were assigned to 1 of 2 treatment groups (n = 9) and a control group (n = 9). Cows in control group received no treatment; On day 270 of gestation, cows in the treatment groups were received 200 μ g (Group 1) or 400 μ g (Group 2) of misoprostol (cytotec®) intravaginally. Intravaginal misoprostol was given every 6 h until the onset of labour. A maximum of 6 doses was administered. The tablets were dampened by 3 drops of water immediately prior to insertion in the treatment groups with water. Induction is considered to have failed if cows have

not calved by 72 h after treatment. The number of cows responding, the interval from misoprostol administration to parturition, calving difficulties, calf birth weight, calf vigor, sex ratio of calves born and the incidence of retained placentas were observed and recorded. Calving difficulties were classified as unassisted, manual traction, manual repositioning and traction or caesarean section. A calf was considered of normal vigor if it was able to stand and nurse within 2 h of birth; it was classified as weak if it required any assistance for its survival (Barth *et al.*, 1978). Placentas were considered retained if they were not released by 24 h after calving.

Data are recorded as percentages, means and standard deviation. The time intervals from misoprostol administration to calving and birth weights were compared by analysis of variance. Differences in the number of induction failures, calving ease, rate of placental retention and pregnancy rate were analyzed by chi square (χ^2).

RESULTS AND DISCUSSION

The observations in relation to induction of parturition in cows (mean+SE) are shown in the Table 1.

The normal gestation period of dairy cattle ranges between 278 and 290 days (Adams, 1969). The average gestation length in control cows was 281 days. In both treated groups the mean length of gestation was significantly less (p<0.01) than in the untreated group, however, there was no difference in gestation length between the 2 treated groups (p>0.05). The effect of misoprostol to induction of parturition, as reported in women (Carlan et al., 2001), nanny goat (Alan and Tasal, 2002) and sheep (Tasal et al., 2001), was confirmed in cows. We defined successful induction as parturition occurring within 72 h after administered misoprostol. All induced cows (except for one cow administered 400 µg misoprostol) calved between 24 and 72 h after administered misoprostol. Dams receiving the induction treatment had an insertion-delivery time of >24 h. The overall duration between commencements of treatment to induction of parturition was 38.44±2.78 h in group I and 36.22±3.07 h in group II. The 400 µg dosage of misoprostol resulted in induction failures of 11.1%. In this cow, the cervix did not dilate any further and the cow did not enter the expulsive stage. Therefore, caesarean section was performed. Induction of parturition insignificantly reduced calf birth weights (p>0.01). The low birth weights were a direct result of the 9.0 days shorter gestation length that resulted from the induction treatment. The insignificant decrease in birth weight resulting from induction of parturition was not associated

Table 1: Observations in relation to induction of parturition in cows			
	I	П	Ш
Groups	(200 mcg)	(400 mcg)	(Control)
No. of animals in group	9	9	9
Gestation period up to the first			
treatment (days)	270	270	270
Interval between treatment and			
calving (h) (time+std. error mean)	38.44±2.78	36.22±3.07	-
Induction failure rate (%)	0	11.11	100
Gestation period up to birth (days)) 272	272	281
Retained fetal membranes (%)	33.33	22.22	22.22
Mode of delivery			
Spantaneous vaginal delivery (%)	44.44	55.55	44.44
Manual traction (%)	33.33	22.22	44.44
Positioning and traction (%)	22.22	11.11	11.11
Caesarian (%)	0	11.11	0
State of calf			
Very good (%)	22.22	33.33	33.33
Good (%)	66.66	44.44	55.55
Weak (%)	22.22	11.11	11.11
Mortality (%)	0	0	0
Mean birth weight (kg)	39.5	39.3	41.0
Calf's sex			
Female	5	3	4
Mole	4	6	5

with calf viability or vigor (p>0.05). Although, it was expected that a decreased birth weight might decrease dystocia, that was not the case. Assistance during calving was administered to five of 9 untreated cows (55.5%), 5 of 9 cows in group I (55.5%) and 4 of 9 cows in group II (44.4%) but induction, by itself, did not caused any increase in dystocias. These differences were not significant (p>0.05) among groups. Several of the dystocia was admittedly related to abnormal presentations of the calves. Abnormal presentations occurred in 2 of 9 cows in group I, 1 of 9 cows in group II and 1 of 9 cows in the untreated group. Usually, the head or leg was doubled backwards. Approximately, half of calves born to all groups were delivered unassisted. Meyer et al. (2001) estimated a primiparous and multiparous dystocia rate (score >1) of 28.6 and 10.7%, respectively. Lombard et al. (2007) reported that over 35% of calves required assistance during birthing. It was informed that the proportion of calvings that required assistance in the studied dairies was large, especially for primiparous dams, in which >50% needed assistance at calving in that study. Results of Meyer et al. (2001) suggested a smaller incidence of dystocia than reported in this study (Lombard et al., 2007). In this study, the calf mortality was observed in none in all groups. It could be argued that the calves survived within 24 h of birth because of optimal care. Because dystocia has been shown to have detrimental effects on adaptation of calves to extrauterine life, calves exposed to dystocia required additional attention than calves delivered without assistance (Bellows and Lammoglia, 2000; Berger, 1994; Meyer et al., 2000). The calf mortality may be reduced to some extent by careful observation and prompt assistance. In this study, the calves was routinely given colostrum shortly after birth, provided a dry, clean environment and received care for adverse circumstances. One calf in group I, 2 in group II and 1 in control group were required assistance to nurse.

The rate of placental retention did not differ among groups. Average incidence of RFM for normal dairy cows ranges from 4-18% (Han and Kim, 2005). The incidence of retained placenta in this study was 33.3% in group I, 22.2% in group II and 22.2% in group III. It was recorded that the overall incidence of retention of fetal membranes was higher than the normal limits, but the incidence was lower than the reported induction of parturition with prostaglandin F2 alpha and/or corticoids (Allen and Herring, 1976; Murray et al., 1984). Increased occurrence of retained placenta in cows in this study might be due to lack of tone and slow involution or damage to the uterus by mechanical stress resulting from calving difficulty (Klerx and Smolders, 1997). In agreement with other reports (Erb et al., 1985; Pelissier, 1972; Thompson et al., 1983), the dystocia was associated with the incidence of retained placenta in this study. Chassagne et al. (1996) and Joosten et al. (1987) on the relationship between gestation length and the incidence of retained placenta showed that shorter gestation lengths are associated with a higher incidence of retained placenta. Short gestations were not related to an increased risk of developing retained placenta in this study. Sex of the calf had no influence on the retained placenta rate. A similar pattern was found by Muller and Owens (1974). Joosten et al. (1987) founded that the retained placenta for male calves were associated with 0.2% higher rates. It is debatable whether this statistically significant difference is indicative of a real difference or the number of cases analyzed.

CONCLUSION

The rate of placental retention, birth weights, calving difficulty and calf viability did not differ among groups.

The results suggested that misoprostol is a cost-effective and safe alternative by vaginal administration for induction of parturition in the cow. The optimal dose, regime and route of administration needs further investigation.

REFERENCES

Adams, W.M., 1969. The elective induction of labour and parturition in cattle. JAVMA, 154 (3): 261-268. PMID: 5762239.

- Allen, J.G. and J. Herring, 1976. The induction of parturition using dexamethasone in dairy cattle. AVJ., 52 (10): 442-445. DOI: 10.1111/j.1751-0813.1976. tb05386.x.
- Alan, M. and I. Tasal, 2002. Eficacy of prostaglandin F2a and Misoprostol in the Induction of Parturition in Goats. Vet. Rec., 22: 788-789. PMID: 12135076.
- Aronsson, A., C. Fiala, O. Stephansson, F. Granath, B. Watzer, H. Schweer and K. Gemzell-Danielsson, 2007. Pharmacokinetic profiles up to 12 h after administration of vaginal, sublingual and slowrelease oral misoprostol. Hum. Reprod., 22: 1912-1918. PMID: 17488782.
- Barth, A.D., W.M. Adams and J.C. Manns, 1978. Rawlings, Induction of parturition in beef cattle using estrogens in conjunction with Dexamethasone. Can. Vet. J., 19: 175-180. PMID: 698897.
- Barth, A.D., 2006. Induction of parturition in cattle. Large Anim. Vet. Rounds, 6 (2): 1-6. http://www.larounds.ca/crus/laveng 0206.pdf.
- Bazer, F.W. and N.L. First, 1983. Pregnancy and Parturition. J. Anim. Sci., 57: 425-460. PMID: 6352591.
- Bellows, R.A. and M.A. Lammoglia, 2000. Effects of severity of dystocia on cold tolerance and serum concentrations of glucose and cortisol in neonatal beef calves. Theriogenology, 53: 803-813. PMID: 10735045.
- Berger, P.J., 1994. Genetic prediction for calving ease in the United States: Data, models and use by the dairy industry. J. Dairy Sci., 77: 1146-1153. PMID: 8201049.
- Carlan, S.J., S. Bouldin, D. Blust and W.F. O'Brien, 2001. Safety and efficacy of misoprostol orally and vaginally: A Randomized Trial. Obstet. Gynecol., 98 (1): 107-112. PMID: 11430966.
- Chassagne, M., J. Barnouin and B. Faye, 1996. Descriptive epidemiology of placental retention in intensive dairy herds in Brittany. Vet. Res., 27 (4-5): 491-501. PMID: 9026226.
- Collins, P.W., 1990. Misoprostol: Discovery, development and clinical applications. Med. Res. Rev., 10: 149-72. PMID: 2109814.
- El-Sherbiny, M.T., I.H. El-Gharieb and H.A. Gewely, 2001. Vaginal misoprostol for induction of labour: 25 vs. 50 microg dose regimen. Int. J. Gynaecol. Obstet., 72: 25-30. PMID: 11146073.
- Erb, H.N., R.D. Smith, P.A. Oltenacu, C.L. Guard, R.B. Hillman, P.A. Powers, M.C. Smith and M.E. White, 1985. Path model of reproductive disorders and performance, milk fever, mastitis, milk yield and culling in Holstein cows. J. Dairy Sci., 68: 3337-3349. PMID: 4093528.

- Goldberg, A.B., M.B. Greenberg and P.D. Darney, 2001. Misoprostol and pregnancy. N. Engl. J. Med., 344: 38-47. PMID: 11136959.
- Han, I.K. and I.H. Kim, 2005. Risk factors for retained placenta and the effect of retained placenta on the occurrence of postpartum diseases and subsequent reproductive performance in dairy cows. J. Vet. Sci., 6 (1): 53-59. PMID: 15785124.
- Holmes, C.W. and S.C. Garcia, 1999. Effects of time of calving on the productivity of pasture-based dairy systems: A review. N.Z. J. Agric. Res., 42: 347-362.
- Joosten, I., P. Van Eldik, L. Elving and G.J.W. Van Der Mey, 1987. Factors related to the etiology of retained placenta in cattle. Anim. Reprod. Sci., 14: 251-262.
- Klerx, H.J. and E.A.A. Smolders, 1997. Herd and cow random variation in models of interrelationships between metabolic and reproductive disorders in high yielding multiparous Holstein dairy cattle in The Netherlands. Livest. Reprod. Sci., 52: 21-29. DOI: 10. 1016/S0301-6226(97)00116-4.
- Lombard, J.E., F.B. Garry, S.M. Tomlinson and L.P. Garber, 2007. Impacts of dystocia on health and survival of dairy calves. J. Dairy Sci., 90 (4): 1751-1760. PMID: 17369215.
- Mansell, P.D., A.R. Cameron, D.P. Taylor and J. Malmo, 2006. Induction of parturition in dairy cattle and its effects on health and subsequent lactation and reproductive performance. AVJ, 84 (9): 312-316. DOI: 10.1111/j.1751-0813.2006.00031.x
- Meyer, C.L., P.J. Berger and K.J. Koehler, 2000. Interactions among factors affecting stillbirths in Holstein cattle in the United States. J. Dairy Sci., 83: 2657-2663. PMID: 11104286.
- Meyer, C.L., P.J. Berger, K.J. Koehler, J.R. Thompson and C.G. Sattler, 2001. Phenotypic trends in incidence of stillbirth for Holsteins in the United States. J. Dairy Sci., 84: 515-523. PMID: 11233037.

- Muller, L.D. and I.J. Owens, 1974. Factors associated with the incidence of retained placenta. J. Dairy Sci., 57 (6): 725-728. PMID: 4407513.
- Murray, R.D., W.T. Nutter, S. Wilman and D.B. Harker, 1984. Induction of parturition in cattle using dexamethasone and cloprostenol: Economic performance and disease incidence after treatment. Vet. Rec., 115 (12): 296-300. PMID: 6495587.
- Pelissier, C.L., 1972. Herd breeding problems and their consequences. J. Dairy Sci., 55: 385-391.
- Sanchez-Ramos, L., A.M. Kaunitz, G.O. Del Valle, I. Delke, P.A. Schroeder and D.K. Briones, 1993. Labour induction with the prostaglandin E₁ methyl analogue misoprostol vs. oxytocin: A randomized trial. Obstet. Gynecol., 81: 332-336. PMID: 8437780.
- Shukla, S.P., A. Pandey and S.P. Nema, 2008. Emergency Induction of parturition in buffaloes. Buffalo Bull., 27(1):148-149.http://ibic.lib.ku.ac.th/e-Bulletin/2008-148.htm.
- Tasal, I., M. Alan, E. Saban and Y. Cetin, 2001. Induction of parturition by misoprostol in ewes. Hay. Aras. Derg., 11 (1): 16-19.
- Taverne, M., 1998. Regulation of Parturition: From Theory to Practice. Reprod. Dom. Anim., 33: 247-248. DOI: 10.1111/j.1439-0531.1998.tb01352.x.
- Thompson, J.R., E.J. Pollak and C.L. Pellisier, 1983. Interrelationships of parturition problems, production of subsequent lactation, reproduction and age at first calving. J. Dairy Sci., 66 (5): 1119-1127. PMID: 6875060.
- Zeiman, M., S.K. Fong, N.L. Benowitz, D. Banskter and P.D. Darney, 1997. Absorption kinetics of misoprostol with oral or vaginal administration. Obstet. Gynecol., 90 (1): 88-92. PMID: 9207820.