The Effect of Milk Production on Reproductive Performance in the High Producing and BST Supplemented Dairy Cow.

David McClary, *DVM* Lilly Research Laboratories Greenfield, IN

Introduction

Over the past 35 years numerous studies have measured the relationship between milk production and reproductive performance in dairy cows.¹⁻⁷. Most reported studies indicate a negative relationship between milk yield and key reproductive indices. Hamudikuwanda *et al*⁶ summarized this relationship to be an additional 0.3 days open and .005 services per conception for each 100 lb increase in milk yield for the first 120 days-in-milk. The factors contributing to this inverse relationship between milk production and reproductive performance are discussed.

Production vs Reproduction

Reduced reproductive efficiency in the dairy cow can be related to biological and managerial factors. In most dairy herds management related factors such as failure to detect estrus and faulty insemination technique have a much greater impact on reproductive performance than biological factors. Comparison of reproductive performance of over 40,000 cows from approximately 4,000 herds with annual milk rolling herd averages from 12,000 to 21,000 lbs indicates days open is longer in cows with lower milk production levels⁸. (Table 1) This analysis suggests that reproductive management factors such as heat detection contribute to reduced reproductive performance. Lower producing herds are likely suffering from a number of management deficiencies including deficiencies in reproductive management which contribute to lower production. To evaluate the biological influence of level of milk production or reproduction, management factors must be eliminated and cows must be compared under the same management system.

The negative relationship reported between milk production and reproductive performance is apparently not directly related to genetic improvement. As average milk yield has increased 6500 lb/cow/year since 1950, first service conception rates have declined $(66\% \text{ vs } 45\%)^9$. Yet over the same time period, conception rates in virgin heifers have remained approximately 70%.

The reduction in reproductive efficiency noted in the high producing dairy cow is most likely directly related to TABLE 1 Reproductive Efficiency at Various Milk Production Levels¹

Milk Rolling Herd Ave (1,000s)					
12-13	14-15	16-17	18-19	20-21	
92	128	122	118	126	
142	135	128	126	127	
89	88	86	87	88	
1.9	2.2	2.3	2.2	2.3	
31	40	47	50	54	
	Mil 12-13 92 142 89 1.9 31	Milk Rollin 12-13 14-15 92 128 142 135 89 88 1.9 2.2 31 40	Milk Rolling Herd 12-13 14-15 16-17 92 128 122 142 135 128 89 88 86 1.9 2.2 2.3 31 40 47	Milk Rolling Herd Ave (1 12-13 14-15 16-17 18-19 92 128 122 118 142 135 128 126 89 88 86 87 1.9 2.2 2.3 2.2 31 40 47 50	Milk Rolling Herd Ave (1,000s) 12-13 14-15 16-17 18-19 20-21 92 128 122 118 126 142 135 128 126 127 89 88 86 87 88 1.9 2.2 2.3 2.2 2.3 31 40 47 50 54

¹DHIA Records Raleigh, NC DRPC Nebel RA, Stallings CC: Hoard's Dairyman, 1990. ⁽⁸⁾

the energy balance of early lactation. In early lactation milk production increases rapidly, reaching a peak at four to eight weeks in lactation. Dry matter intake slowly increases during early lactation but does not peak until 10 to 14 weeks into lactation¹⁰. Since a lag exists between peak milk production and maximum dry matter intake, often the cow does not consume enough nutrients to support the high levels of milk production. To support the demands of peak milk production the cow mobilizes body tissue energy stores producing a negative energy balance. Maximum negative energy balance usually occurs at approximately 14 days-in-milk. As dry matter intake continues to increase during early lactation, the cow begins to reduce the magnitude of negative energy balance until an equilibrium between energy intake and energy output is achieved at 56 to 120 days-in-milk¹¹.

During maximum negative energy balance there is little evidence of cyclic ovarian activity. Elevated progesterone concentrations seen throughout most of pregnancy are extremely low following lysis of the corpus luteum of pregnancy. Estrogen concentrations peak near the time of parturition and return to base line immediately postpartum. During the early postpartum period, the frequency of pulsatile GnRH release from the hypothalmus is markedly reduced compared to frequency patterns seen in cyclic cows. Subsequently the pulsatile release of pituitary gonadotropins (LH and FSH) is also reduced. A significant increase in the pulsatile release of GnRH usually starts approximately 14 days post calving. First ovulation in the postpartum dairy cow normally occurs approximately 10 days later or after maximum negative energy balance. Although first ovulation occurs 20 to 24 days-in-milk the range may be quite large (9 to 93 days). Days to first ovulation are likely directly related to the accumulated negative energy balance.

Cumulative negative energy balance is related to level of milk production and nutrient intake. While higher producing cows do tend to experience a greater magnitude of negative energy balance¹¹ (Figure 1), reduced nutrient intake can also produce a relative decrease in energy balance. Reduced intake or energy deficient rations may not support high levels of milk production placing increased demands on body tissue nutrient stores.

Numerous disease conditions seen in the postpartum dairy cow can contribute to reduced nutrient intake. Cows with excessive body condition at calving may also suffer reduced intake. Reduced intake often results in excessive mobilization of body fat reserves and a corresponding reduction in reproductive efficiency. Heinonen et al¹² demonstrated a significant correlation between percent body weight loss in early lactation and reproductive efficiency. Cows were categorized as having loss of body weights as >10%, 5-10%, and <5%. Those cows with greater losses of weight had a significant increase in days to first observed estrus (48.3 + / -2.5 vs 42.8 + / -1.3 and 45.7 + / -2.1), days open (89.6 + / -3.2 vs 79.4 + / -1.7 and 78.6 + / -2.5), and first service conception rate (53.1 vs 74.4 and 78.6) as compared to cows losing 5-10% and < 5% body weight, respectively. Butler et al¹³ demonstrated similar results related to loss of body condition score (BCS) during the first 5 weeks of lactation. Cows losing >1.0 BCS regardless of the score at calving had significant reductions in all measured repro-

FIGURE 1 Milk Production and Energy Balance During Lactation



ductive indices. (Table 2) Over-conditioned at calving (BCS > 4) should be avoided to prevent excessive loss in body condition score.

TABLE 2 Body Condition Score1 Loss During First 5Weeks of Lactation vs Reproductive Efficiency

	Body Condition Score Loss			
	<0.5	0.5-1.0	>1.0	
Number	17	64	12	
Days to 1st Ovulation	27+/-2 ^a	31+/-2 ^a	42+/-5 ^b	
Days to 1st Observed Estrus	48+/-6 ^{ab}	$41 + -3^{a}$	62+/-7 ^b	
Days to 1st Service	$68 + / - 4^{a}$	67+/-2 ^a	79+/-5 ^b	
1st Service Conc. Rate (%)	65ª	53ª	17 ^b	
Services/Conc.	18+/-0.4	2.3+/-9.2	2.3+/-0.4	

¹Body Condition Score 1-5 Scale

^{ab}Means With Different Superscripts (P < 0.05) Butler WR et al J Anim Sci, 1981. ⁽¹³⁾

Nutritional Factors Affecting Ovarian Activity in the Postpartum Cow

Although reduced ovarian activity in the postpartum cow is apparently related to the cumulative negative energy balance of the postpartum period, the exact mechanism is complex and not completely understood. The nutritional factors related to ovarian activity are both physical and physiological. Maternalization or suckling apparently decreases the pulsatile release of LH and FSH from the anterior pituitary. While maternalization may be related to postpartum anestrus in the suckled beef cow, it is not an important factor in the dairy cow.

In the postpartum dairy cow, increased utilization of glucose leads to hypoglycemia and hypoinsulinemia which increase fat mibilization. Initially these conditions tend to stimulate appetite. Increased appetite increases the release of opoid compounds (beta-endorphins) within the hypothalamo-hypophyseal unit. Beta-endorphins directly inhibit the pulsatile release of GnRH which subsequently also reduces the pulsatile release of LH. GnRH pulsatility is also apparently reduced by the increased production and utilization of ketones secondary to increased lipolysis. Since insulin has been shown to have a gonadotrophic effect on the ovary¹⁴, hypoinsulinemia in early lactation mav adversely affect responsiveness of ovarian follicles to gonadotropin stimulation.

Evidence of suppressed ovarian activity in the early postpartum dairy cow can be further measured by reduced serum progesterone levels subsequent to early ovulations. Villa-Godoy *et al*¹⁵ noted a direct relationship between serum progesterone concentrations and the relative energy balance in postpartum dairy cows. Apparently excessive caloric deficiency contributes to poor luteal function in the second and third estrus cycles. This relationship likely explains the reduced early postpartum conception rates seen in many high-producing, negative-energy-balance cows.

Effect of BST on Reproductive Efficiency

While numerous studies conducted in the US and other countries have consistently demonstrated increases of 10 to 20% in milk yield in BST-supplemented cows, the effect of BST on reproductive performance is less clear. Some studies have shown no effect on reproductive indices¹⁶⁻²⁶ while others indicate slight increases in days open, calving interval, and/or services-per-conception when BST supplementation was initiated prior to breeding²⁷⁻³². Days-in-milk relative to the initiation of BST supplementation is an important consideration in measuring any negative impact of BST on reproductive efficiency. Treatment during early lactation may contribute to a greater cumulative negative energy balance resulting in a delayed return of cyclic ovarian activity and an increase in days-open.

Data from a series of five trials including 113 non-pregnant cows in early lactation (29-42 days-in-milk) at the initiation of BST treatment were evaluated to determine BST's effect, if any, on reproductive indices³³. Groups consisting of 27, 29, 28, and 29 animals were treated with 0, 320, 640, or 960 mg/28-days BST, respectively. Pregnancy rates were 25 to 27 (93%), 23 of 29 (79%), 24 of 28 (86%), and 24 of 29 (83%) for the respective groups. There was no significant difference nor was there a dose-related pattern in the number of days to first service among the groups. Although increases were noted in the BST-supplemented groups, the differences were not statistically significant. The differences do indicate, however, a trend toward reduced reproductive efficiency in the BST-supplemented cow when BST supplementation is initiated in early lactation. (Table 3)

TABLE 3 FIVE TRIAL SUMMARY

Reproductive Performance of Early Lactation Cows Receiving Somidobove

	Somidobove (mg/28 days)			
Stage	0	320	640	960
No. Cows	- 27	29	28	29
No. Pregnant (%)	25(93)	23(79)	24(86)	24 (83)
Days to 1st Service	83	68	92	78 ´
Days Open	113	120	138	129
No. Calving	23	23	23	23
Calving Interval	393	401	411	399
1st Service Conc. Rate (%)	56	31	36	28
Services/Conception	1.8	2.7	2.5	2.6

Ferguson³⁴ evaluated the reproductive data from a number of BST dose-titration trials in an attempt to measure the effect of treatment prior to pregnancy on reproductive efficiency. Pregnancy rates of 2187 BST-supplemented cows were slightly reduced compared to 1170 nontreated controls (81.2 vs 89.2%). When analyzed on average daily dose equivalent of 0, <10.5, 10.6-25, 26-46, or >46 mg/day BST regardless of the dosage form, daily or sustained release, pregnancy rates tended to be negatively related to dose, (89.5, 87.1, 79.8, 81.4, and 73.8%, respectively). He also noted that pregnancy rates tended to be lower in cows that had below average milk production responses to BST at the various dosage levels.

In the same analysis, Ferguson determined conception rates to be similar in treated and control cows (47.8 vs 48.4%, respectively). Days open were increased 17.2 days in cows receiving >25 mg BST/day compared to controls.

A summary of 21 European BST trials comparing 402 nontreated controls and 483 cows treated with 640 or 960 mg/28 day BST indicated no difference in a number of reproductive indices when BST supplementation was initiated 23 to 130 days-in-milk. A significant increase (p < 0.01) was noted in the multiple birth rate in BST- supplemented cows compared to controls (10.7 vs 4.7%, respectively)³⁵. (Table 4)

TABLE 4 SUMMARY OF 21 EUROPEAN BST TRI-ALS

Reproductive Performance of Lactating Cows Receiving Somidobove

	Somidobove	(MG/28 Days)
	0	640/960
No. Cows	402	483
No. Pregnant (%)	327(81.3)	387(80.1)
Pregnant Before Treatment	88	115
Days to 1st Service	71.5	72.4
Days Open	96.9	100.4
Calving Interval	384.7	386.2
1st Service Conc. Rate (%)	45	41.5
Conc. Rate All Services (%)	45.7	44.2
Services/Conception	2.32	2.42
No. Calving (%)	296(90.5)	347(89.6)
No. Abortions	10	8
% Male Calves	51.0	48.2
Male Birth Wt. (KG)	46.3	46.5
Female Birth Wt. (KG)	44.3	43.6
Assisted Birth (%)	72(25.2)	84(23.5)
Milk Fever	31	27
No. Calving (%)	298(90.5)	347(89.6)
Total Calves	310	385
Single Births	282	310
Twin Births (CBTI)	14(4)	36(6)
Multiple Birth Rate %	4.7	10.7*
*p < 0.01		
(35)		

Herrler et al^{36} reported a two fold increase (3.8 + / -1.1 vs 1.9 + / -0.7) in the number of transferable embryos in cows treated with 640 mg/28 day BST five days prior to superovulation treatment with 2500 IU PMSG.

These data suggest BST has an effect on folliculogenesis and ovulation rates in supplemented cows. BST's influence on ovarian function is likely related to the interactions of BST, insulin growth factor-1 (IGF-1), gonadotropins (FSH and LH), and ovarian follicles. Receptors for BST and insulin have been isolated from the bovine ovary. Granulosal cells in the ovary produce IGF-1. IGF-1 production and/or release is increased by BST supplementation. IGF-1 acts synergistically with FSH to; 1) increase granulosal cell division, 2) increase follicular estrogen and progesterone production, and 3) increase granulosal cell LH receptor numbers. Spicer et al³⁷ concluded that treatment with bovine growth hormone-releasing factor (GRF) did not increase IGF-1 concentrations in the ovarian follicle but instead affected the development of large follicles. These factors should be considered when initiating BST supplementation in the postpartum period.

BST will likely have minimal impact on reproductive efficiency in the dairy cow when supplementation is initiated after 100 days-in-milk or after peak lactation and maximum negative energy balance. When BST treatment is initiated earlier in lactation, ie. 30-40 days-in-milk, total lactation milk yield will likely be increased but reproductive efficiency will likely be slightly reduced due to an exacerbated negative energy balance. The reduction in reproductive efficiency will likely be of the same magnitude seen in the genetically-superior, higher-producing cow. If nutrient intake is increased in the early lactation cow the effects of negative energy balance on reproductive efficiency can be minimized. BST supplementation at the time of breeding may increase the incidence of multiple births in the dairy cow.

Summary

Higher-producing dairy cows have reduced reproductive efficiency compared to lower producing herdmates. This relationship is likely to be seen whether increased production is achieved through superior genetics or BST supplementation. Increased days open in the high-producing cow are most likely related to a slight delay in return of

cyclic ovarian activity and reduced first service conception rates. Reduced fertility in the high-producing cow is not directly related to increased milk yield but rather to loss of body weight and body condition. The magnitude of the negative impact of milk production on reproduction depends on the cumulative negative energy balance in early lactation. High-producing cows can achieve optimal reproductive efficiency if they are fed adequate energy and protein to support high levels of milk production in early lactation and replenish adequate body condition in late lactation and the dry period.

While BST-supplemented cows are similar to higherproducing, genetically superior cows in that total milk is increased for the lactation, they may differ in the patterns of milk production over the lactation. Depending on the time of initiation of BST supplementation and the duration of administration, the BST-supplemented cow may have a greater persistency of the lactation curve. Subsequently, the BST-supplemented cow will produce a higher percentage of total milk in the latter part of lactation, changing the rate of profit accumulation. With more milk being produced in late lactation, the importance of maintaining a minimal calving interval may be reduced. The voluntary waiting period prior to breeding may be intentionally extended, thereby reducing the pressures of early postpartum rebreeding when first service conception rates are often low in high-producing cows. Maintaining profitable levels of milk production for longer lactation cycles will allow for later postpartum breeding and reduce culling for reproductive failure. Reduced calvings per year may actually extend herd life by reducing the number of disorders associated with the early postpartum period.

BST will not be indicated for all cows. Even within herds using BST, programs for recommended use will vary depending on a number of factors including milk production, body condition, and reproductive status. The dairy farmer and the health consultant will need to scrutinize individual cows to determine which are most likely to benefit from BST supplementation. To maximize efficiency and profitability, the appropriate stage of lactation in which to use BST must be determined. The bovine practitioner can play a meaningful role in making those management decisions.

References

1. Boyd LJ, Seath DM, Olds D: Relationship between level of milk production and breeding efficiency in dairy cattle. J. Anim. Sci. 13:89, 1954. 2. Spalding RW, Everett RW, Foote RH: Fertility in New York artificially inseminated Holstein herds in Dairy Herd Improvement. J. Dairy Sci. 58:718, 1975.. 3. Olds D, Cooper T, Thrift FA: Relationships between milk yield and fertility in dairy cattle. J. Dairy Sci. 62:1140, 1979.. 4. Burger PJ, Shanks RD, Freeman AE, et al: Genetic aspects of milk yield and reproductive performance. J. Dairy Sci. 64:114, 1981. 5. Laben RL, Shanks RD, Burger PJ, et al: Factors affecting milk yield and reproductive performance. J. Dairy Sci. 65:1004, 1982. 6. Hamudikuwanda H, Erb HN: Effects of sixty-day milk yield on postpartum breeding performance in Holstein cows. J. Dairy Sci. 70: 2355, 1987. 7. Lean IJ, Galland JC, Scott JL: Relationships between fertility, peak milk yields and lactational persistency in dairy cows. Theriogenology. 31:1093, 1989. 8. Nebel RA, Stallings, CC: You can have high milk and good reproduction. Hoards Dairyman. Feb. 10, 1990, p. 105. 9. Butler WR, Smith RD: Interrelationships between energy and postpartum reproductive function in dairy cattle. J. Dairy Sci. 72:767, 1989. 10. Nutrient Requirements of Dairy Cattle. Sixth Ed. National Research Council, National Academy Press, Washington, DC 1989, pp 2-52. 11. Erdman RA, Andrew SM: Methods for and estimates of body tissue mobilization in the lactating

dairy cow. In: Meeting the challenges of New Technology, Monsanto Technical Symposium. Syracuse, NY. Oct. 24, 1989, pp 35-44, 12. Heinonen K, Ettala E, Alanko M: Effect of postpartum weight loss on reproductive functions in dairy cows. Acta Vet. Scand. 29:249, 1988. 13. Butler WR, Evert RW, Coppock CE: The relationship between energy balance, milk production, and ovulation in postpartum Holstein cows. J. Anim. Sci. 53:742, 1981. 14. Poretsky L, Kalin NF: The gonadotrophic function of insulin. Endocr. Rev. 8:132, 1987. 15. Villa-Godoy A, Hughes TL, Emery RS, et al: Association between energy balance and luteal function in lactating dairy cows, J. Dairy Sci. 71:1063. 1988. 16. Aguilar AA, Jordan DC, Olsen JD, et al: A short-term study evaluating the galactopoietic effects of the administration of Sometribove (recombinant methionyl bovine somatotropin) in high producing dairy cows milked three times per day. J. Dairy Sci. 71 (Suppl. 1):208 (Abst.), 1988. 17. Cole WJ, Franson SE, Hoffman RG, et al: Response of cows throughout lactation to sometribove, recombinant methionyl bovine somatotropin, in a prolonged release system -- a dose titration study. Part II. Health and reproduction. J. Dairy Sci. 72 (Suppl. 1): 451 (Abst.), 1989. 18. Lamb RC, Anderson MJ, Henderson SL, et al: Production response of Holstein cows to sometribove USAN (recombinant methionyl bovine somatotropin) in a prolonged release system for one lactation. J. Dairy Sci. 71 (Suppl. 1):208 (Abst.), 1988. 19. McDaniel BT, Gallant DM, Fetrow J, et al: Lactational, reproductive and health responses to recombinant bovine somatotropin under field conditions. J. Dairy Sci. 72 (Suppl. 1):429 (Abst.), 1989. 20. Samuels WA, Hard DL. Hintz RL, et al: Long term evaluation of sometribove, USAN (recombinant methionyl bovine somatotropin) treatment in a prolonged release system for lactating dairy cows. J. Dairy Sci. 71 (Suppl. 1):209 (Abst.), 1988. 21. Eppard PJ, Bauman DE, Curtis CR, et al. Effect of 188-day treatment with somatropin on health and reproductive performance of lactating diary cows. J. Dairy Sci. 70:582, 1987. 22. Bunn KB, Jenny BF, Pardue FE, et al: Effect of sustained-release bovine somatotropin (BST) on reproduction and mammary health of dairy cows. J. Dairy Sci. 72 (Suppl. 1):325 (Abst.), 1989. 23. Chalupa W, Baird L, Soderholm C. et al: Responses of dairy cows to somatotropin. J. Dairy Sci. 70 (Suppl. 1):176 (Abst.), 1987. 24. Hansen WP, Otterby DE, Linn JG, et al: Multifarm use of bovine somatotropin (BST) and its effects on lactation, health and reproduction. J. Dairy Sci. 72 (Suppl. 1):429 (Abst.), 1989. 25. Palmquist DL: Response of high-producing dairy cows given daily injections of recombinant bovine somatotropin from D 30-296 of lactation. J. Dairy Sci. 71 (Suppl. 1):206 (Abst.), 1988. 26. Rajmahendran R, Desbbottes S,

Shelford JA, et al: Effect of recombinant bovine somatotropin (rbSt) on milk production and reproductive performance of dairy cows. J. Dairy Sci. 72 (Suppl. 1):444 (Abst.), 1989. 27. Cole WJ, Eppard PJ, Lanza GM, et al: Response of lactating dairy cows to multiple injections of sometribove, USAN (recombinant methionyl bovine somatotropin) in a prolonged release system. Part II. Health and reproduction. J. Dairy Sci. 71(Suppl. 1):184 (Abst.) 1988.

28. Huber JT, Willman S, Marcus K, et al: Effect of sometribove (SB), USAN (recombinant methionyl bovine somatotropin) injected in lactating cows at 14-day intervals on milk yields, milk composition and health. J. Dairy Sci. 71 (Suppl. 1):207 (Abst.), 1988. 29. Elvinger F, Head HH, Wilcox CJ, et al: Effects of administration of bovine somatotropin on lactation milk yield and composition. J. Dairy Sci. 70 (Suppl. 1):121 (Abst.), 1987. 30. Burton JH, McBride BW, Bateman K, et al: Recombinant bovine somatotropin: effects on production and reproduction in lactating cows. J. Dairy Sci. 70 (Suppl. 1):175 (Abst.), 1987. 31. Cleale RM, Rehman JD, Rob EJ, et al: On-farm lactational and reproductive responses to daily injections of recombinant bovine somatotropin. J. Dairy Sci. 72 (Suppl. 1):429 (Abst.), 1989. 32. Hard DL, Cole WJ, Franson SE, et al: Effect of long term sometribove, USAN (recombinant methionyl bovine somatotropin), treatment in a prolonged release system on milk yield, animal health and reproductive performance-pooled across four sites. J. Dairy Sci. 71 (Suppl. 1):210 (Abst), 1988. 33. McClary DG: The impact of BST on animal health and reproduction. In: Bovine Somatotropin: Proceedings of Symposia. Elanco Products Co. and Lilly Research Labs. Columbus, OH, Oct. 16, 1989; St. Paul, MN, Oct. 26, 1989, pp 5-11. 34. Ferguson JD: Interaction between milk yield and reproduction in dairy cows. In: Meeting the challenges of New Technology, Monsanto Technical Symposium. Syracuse, NY. Oct. 24, 1989, pp 35-44. 35. Wilkinson JID, McGuffey RK: Recent developments in somatotropin research. Fourth International Symposium on Modern Cattle Production, German Cattle Breeders Association (DGL), Giessen, FRG, Nov. 24-26, 1989. 36. Herrler A, Farries E, Neimann H: A trial to stimulate insulin like growth factor 1 levels to improve superovulatory response in dairy cows. Theriogenology. 33:248, 1990. 37. Spicer LJ, Enright WJ: Concentrations of insulin-like growth factor-I (IGF-I) in follicular fluid (FFL) of preovulatory bovine ovarian follicles: Effect of daily injections of a potent growth hormone (GH)-releasing factor (GRF) analog and/or thyrotropin-releasing hormone (TRH). J. Dairy Sci. 72 (Suppl. 1):346 (Abst.), 1989.