

# How X-Zelit Works to Reduce Milk Fever and Hypocalcemia...

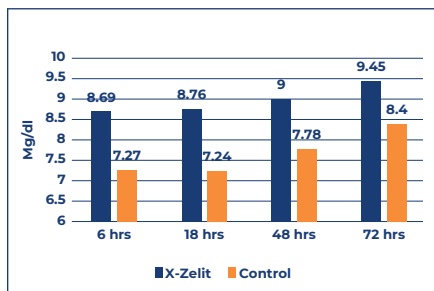
## INTRODUCTION

Traditionally, milk fever prevention programs have attempted to directly alter Ca status of dairy cows at calving. Examples of common approaches include feeding pre-fresh dairy cows diets containing anionic salts, limiting dietary Ca, feeding hyper-dietary Ca and administering Ca boluses.

## IMPROVING BLOOD CALCIUM

New research (Cornell University, UW-Madison, UVM Germany) has discovered that restricting dietary P in pre-fresh cows significantly improves blood Ca status of cows at calving which greatly diminishes hypocalcemia and milk fever. These and other new studies have discovered that modestly restricting dietary P to pre-fresh cows reduces the release of a bone hormone called fibroblast growth factor-23 (FGF23). FGF23 plays a central role in P metabolism regulating the active form of vitamin D and P absorption from the kidneys, intestines and bone mobilization of both P and Ca.

### Blood Calcium Levels



Cornell data, Journal of Dairy Science 2019 (30 cows per treatment)

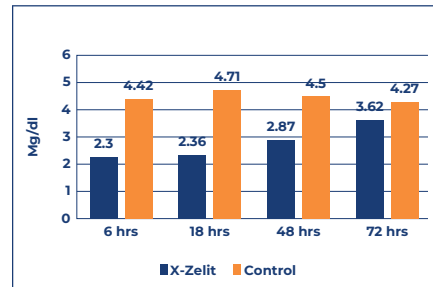
When X-Zelit is fed to pre-fresh cows for 12 – 18 days, in combination with feeding modest levels of dietary P and Ca, enough P is bound to likely cause a reduction in FGF23, yielding a rapid release of both Ca and P from bone thereby

increasing blood Ca. Elevation of blood Ca at calving induced by feeding X-Zelit, a dietary P binder, and subsequent reductions in milk fever and hypocalcemia has been observed in numerous studies and on commercial dairies.

## BLOOD PHOSPHORUS

When X-Zelit is properly fed, it induces a modest dietary P restriction and blood Ca level at calving is significantly improved and with far less variation between cows. Blood Ca improvement is a consistent response observed in studies, with a corresponding lower blood P also observed.

### Blood Phosphorus Levels

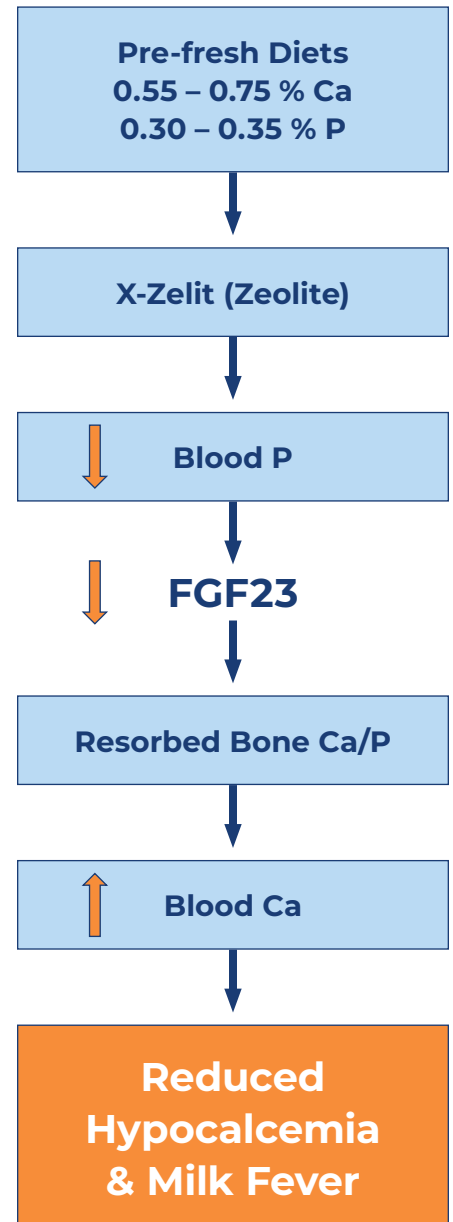


Cornell data, Journal of Dairy Science 2019 (30 cows per treatment)

This brings us to question why is X-Zelit so efficient at binding P? The short answer is because the dairy cow recycles a large fraction of P into saliva to form rumen buffers. This form of P (PO<sub>4</sub>) is abundantly available to be bound. A study at the University of Wisconsin has demonstrated that the amount of P in saliva is reduced when feeding X-Zelit. These phosphorus restriction effects are temporary, with saliva blood P levels recovering rapidly after calving when X-Zelit is no longer fed.

## MECHANISM

Below is a simplified mechanism adapted from Crenshaw et al., 2010. J. Anim. Sci. The effects of FGF23 on kidney function are omitted for brevity.



# KEY RESEARCH

## X-ZELIT RESEARCH

Thilsing-Hansen, T., and Jorgensen, R. J. 2001. Hot Topic: Prevention of parturient paresis and subclinical hypocalcemia in dairy cows by zeolite A administration in the dry period. *J. Dairy Sci.* 84:691 – 693.

Kerwin, A. L., Ryan, C. M., Lenon, B. M., Jakobsen, M., Theilgaard, P., Barbano, D. M., and Overton, T. R. 2019. Effects of feeding synthetic zeolite A during the prepartum period on serum mineral concentration, oxidant status, and performance of multiparous Holstein cows. *J. Dairy Sci.* 102:

Frizzarini, W. S., Monteiro, P. L. J., Campolina, J. P., Vang, A. L., Soudah, O., Lewandowski, L. R., Connelly, M. K., Arriola Apelo, S. I., and Hernandez, L. L. 2024. Mechanisms by which feeding synthetic zeolite A and dietary cation-anion difference diets affect mineral metabolism in multiparous Holstein cows: Part I. *J. Dairy Sci.* 107(7):5204 – 5221.

Frizzarini, W. S., Campolina, J. P., Vang, A. L., Lewandowski, L. R., Teixeira, N. N., Connelly, M. K., Monteiro, P. L. J., and Hernandez, L. L. 2024. Mechanisms by which feeding synthetic zeolite A and dietary cation-anion difference diets affect feed intake, energy metabolism, and milk performance: Part II. *J. Dairy Sci.* 107(7):5222 – 5234.

Frizzarini, W., Diniz, J., Vang, A., Monteiro, P., and Hernandez, L. 2022. Effects of 3 different prepartum diets on dry matter intake, beta-hydroxybutyrate, and mineral concentrations in multiparous Holstein cows. *J. Dairy Sci.* (abstr) Suppl. 1105:350.

Thilsing, T., Larsen, T., Jorgensen, R. J., and Houe, H. 2007. The effect of dietary calcium and phosphorus supplementation in zeolite A treated dry cows on periparturient calcium and phosphorus homeostasis. *J. Vet. Med.* 54:82 – 89.

Pallesen, A., Pallesen, F., Jorgensen, R. J., and Thilsing, T. 2008. Effect of pre-calving zeolite, magnesium and phosphorus supplementation on periparturient serum mineral concentrations. *Vet. J.* 175:234 – 239.

Grabherr, H., Spolders, M., Furl, M., and Flachowsky, G. 2009. Effect of several doses of zeolite A on feed intake, energy metabolism and on mineral metabolism in dairy cows around calving. *J. Anim. Phys. and Anim. Nutr.* 93:221 – 236.

Grabherr, H., Spolders, M., Lebsien, P., Huther, L., Flachowsky, G., Furl, M., and Grun, M. 2009. Effect of zeolite A on rumen fermentation and phosphorus metabolism in dairy cows. *Arch. Of Anim. Nutr.* 63:4, 321 – 336.

Souza Saraiva de Oliveira, R. 2017. An alternative strategy to prevent hypocalcemia by adding zeolite A to the prepartum diet of dairy cows. MS Thesis. Kansas State University, Manhattan, KS.

Jørgensen R. J., Hansen, T., Jensen, M. L., and Thilsing-Hansen, T. 2001. Effect of oral drenching with zinc oxide or synthetic zeolite A on total blood calcium in dairy cows. *J. of Dairy Sci.* 84:609 – 613.

Thilsing-Hansen, T., Jorgensen, R. J., Enemark, J. M. D., and Larsen, T. 2002. The effect of zeolite A supplementation in the dry period on periparturient calcium, phosphorus and magnesium homeostasis. *J. Dairy Sci.*, 85:1855 – 1862.

## DIETARY PHOSPHORUS RESEARCH

Cohrs, I., Wilkens, M. R., and Grunberg, W. 2018. Short communication: Effect of dietary phosphorus deprivation in late gestation and early lactation on the calcium homeostasis of periparturient dairy cows. *J. Dairy Sci.* 101:9591 – 9598.

Grünberg, W., Scherpenisse, P., Dobbelaar, P., Idink, M. J., and Wijnberg, I. D. 2015. The effect of transient, moderate dietary phosphorus deprivation on phosphorus metabolism, muscle content of different phosphorus-containing compounds, and muscle function in dairy cows. *J. Dairy Sci.* 98:5385 – 5400.

Peterson, A. B., Orth, M. W., Goff, J. P., and Beede, D.K. 2005. Periparturient responses of multiparous Holstein cows fed different dietary phosphorus concentrations prepartum. *J. Dairy Sci.* 88:3582 – 3594.

Wachter, S., Cohrs, I., Golbeck, L., Wilkens, M. R., and Grunberg, W. 2022. Effects of restricted dietary phosphorus to dry cows on periparturient calcium status. *J. Dairy Sci.* 105:748 – 760.

Wachter, S., Cohrs, I., Golbeck, L., Scheu, T., Eder, K., and Grunberg, W. Effects of restricted dietary phosphorus supply during the dry period on productivity and metabolism in dairy cows. *J. Dairy Sci.* 105:4370 – 4392.

Julien, W. E., Conrad, H. R., Hibbs, J. W., and Crist, W. L. 1976. Milk fever in dairy cows. VIII. Effect of injected vitamin D3 and calcium and phosphorus intake on incidence. *J. Dairy Sci.* 60:431 – 436.

Anderson, S. T., Kidd, L. J., Benvenuti, M. A., Fletcher, M. T., and Dixon, R. M. 2017. New candidate markers of phosphorus status in beef breeder cows. *Anim. Prod. Sci.* 57:2291 – 2303.

Keanthao, P., Goselink, R. M. A., Schonewille, J. T., Dijkstra, J., and Hendriks, W. H. 2019. Effect of phosphorus intake during the transition period on plasma phosphorus content and hypocalcemia in dairy cows. *Proc. of the 17th Inter. Conf. on Prod. Disease in Farm Anim.* (abstr), P 191.

Barton, B.A., Jorgensen, N. A., DeLuca, H. F., 1987. Impact of prepartum dietary phosphorus intake on calcium homeostasis at parturition. *J. of Dairy Sci.* 70, 1186 – 1191.

## FGF-23 REVIEWS

Crenshaw, T. D., Rortvedt, L. A., and Hassen, Z. 2011. A novel pathway for vitamin D-mediated phosphate homeostasis: Implications for skeleton growth and mineralization. *J. Anim. Sci.* 89:1957 – 1964.

Shimada, T., Hasegawa, H., Yamazaki, Y. 2004. FGF-23 is a potent regulator of vitamin D metabolism and phosphate homeostasis. *J. of Bone and Mineral Res.* 19:429 – 435.