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## **Influence of niacin administration on lipid peroxidation in cows in early lactation**

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**Abstract:** Niacin is a reactive part of NAD and NADP, which are co-enzymes in numerous oxidative and reductive reactions. In early lactation, there is an increased lipid mobilization, accompanied by inflammatory response and oxidative stress. The aim of this study was to examine the effect of niacin administration in cows during early lactation in reducing lipid peroxidation i.e. MDA (malondialdehyde) concentration. Niacin was administered through feed two weeks before and two weeks after calving. Blood samples were taken in the week preceding calving and in the first and second week after calving. MDA concentration was significantly lower in cows receiving niacin compared to the control group during all three samplings: 1.89±0.33:1.42±0.13 (calving, week 0); 2.37±0.41:1.64±0.15 (first week) and 2.6±0.45:1.8±0.17µmol/L (second week). In control cows, no significant correlation between MDA and NEFA concentrations was found compared to cows fed niacin. In cows receiving niacin, the concentration of MDA linearly increased, as did the NEFA concentration. Niacin supplementation in early lactation can decrease lipid peroxidation in cows. Reduced lipid peroxidation in cows can be a consequence of the antilipolytic effect of niacin.

**Keywords:** cows, niacin, lipolysis, oxidative stress.

## Introduction

Niacin is reactive part of NAD and NADP, which are co-enzymes in numerous oxidative and reductive reactions. These enzymes are involved in the mechanism of hydrogen transfer in living cells, including more than 200 reactions of carbohydrates, lipid acids and amino acids. The most important metabolic reactions catalyzed by NAD and NADP are: carbohydrate metabolism (glycolysis – aerobic and anaerobic glucose oxidation, Krebs cycle), lipid metabolism (glycerol synthesis and glycerol degradation, oxidation of fatty acids and synthesis of fatty acids, steroid synthesis), protein metabolism (degradation and synthesis of amino acids, oxidation of carbon chains in the Krebs cycle (Cincović *et al.*, 2018). Pires and Grummer (2007) used abomasal infusion with different concentrations of nicotinic acid (0.6, 30 and 60 mg/kg of body weight) in Holstein cows which received restricted diets. Abomasal infusion was administered as a single bolus dose 48 hours after feed restriction. One hour after the administration of 6 mg nicotinic bolus, plasmatic NEFA concentration reduced from 546  $\mu\text{Eq/l}$  to 208  $\mu\text{Eq/l}$ , and less than 100  $\mu\text{Eq/l}$  3 hours after the infusion of the highest doses of nicotinic acid. The increase of nicotinic acid in the blood can reduce the phosphorylation of hormone-sensitive lipase and prevent increased lipolysis, which reduces plasma NEFA concentrations (Overton and Waldron, 2004; Morey *et al.*, 2011; Kenéz, 2014).

In their review study, Ilkhani *et al.* (2016) showed that niacin administration can decrease lipid peroxidation and reduce MDA concentration in different experimental circumstances. During early lactation, greater lipomobilization occurs, accompanied by inflammatory response and oxidative stress (Cincović, 2016; Cincović and Starič, ed., 2017). The aim of this study was to determine the effect of niacin application in cows during early lactation in reducing lipid peroxidation i.e. the concentration of MDA as a by-product of lipid peroxidation.

## Materials and methods

Cows and management – the experiment included 30 Holstein-Friesian cows in the second and third lactation with no signs of health disorders and with a milk yield of  $7500 \pm 950$  liters. Cows were kept in a free stall system on deep litter. In the transitional period, they were kept in maternity stalls in a tie system. Feed was given in cycles but feeding in maternity stalls after calving was ad libitum. Cows were fed mixed feed that satisfied all their nutritional needs. Water was given ad libitum. Rations for cows in early lactation consisted of: proteins (17.5–19.5% crude proteins, 30–33% degradable proteins, 35–40% non-degradable proteins), carbohydrates (ADF minimum 17–21%, NDF minimum 28–31%, NDF from voluminous feed minimum 18–23%, non-structural carbohydrates 35–42%,

dry matter from voluminous feed minimum 40–45%), energy (NEL 7–7.4 MJ/ kg of meal dry matter), fat per meal 5–7%.

Niacin administration – niacin was administered through feed per os. Rovimix®Niacin was used in a dose that allowed availability in intestines at 6–12 g per day (60–120 g per cow per day). This concentration proved to be the best in previous experiments. Niacin was administered two weeks before and two weeks after calving.

Blood sampling and laboratory analysis – blood samples were taken from the coccygeal vein in cows before morning feeding to avoid the postprandial effect of the meal on metabolic parameters. Blood samples were taken at calving and in the first and second week after calving. Photometric reactions and Rayto (RT1904c) photometer were used. Measurements were conducted following the manufacturer's instructions. Standard kits of Randox (UK) and Pointe scientific (USA) were used.

Statistical analysis – differences in metabolite concentrations between the two groups of cows (cows receiving niacin and control cows) were determined each week. The effect of week and group was determined by ANOVA and LSD test. The correlation between NEFA and MDA in experimental and control cows was determined. The *Statgraphics Centurion* statistical software package was used.

## Results and discussion

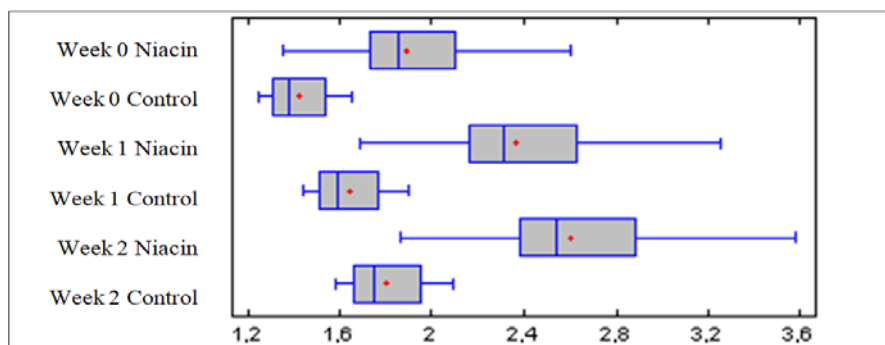
MDA concentration was significantly reduced in cows receiving niacin compared to control cows in all three weeks of the experiment:  $1.89 \pm 0.33$ : $1.42 \pm 0.13$  (calving, week 0);  $2.37 \pm 0.41$ : $1.64 \pm 0.15$  (first week) and  $2.6 \pm 0.45$ : $1.8 \pm 0.17$   $\mu\text{mol/L}$  (second week). ANOVA analysis showed that week of sampling and niacin administration had a very significant effect on MDA concentration in blood samples ( $F=32.46$ ;  $p<0.0001$ ). LSD test showed that cows receiving niacin had lower levels of MDA in all weeks of the experiment. In both groups of cows, a significant increase in MDA was observed in the period from calving to the second week after calving. However, the blood of cows that received niacin showed no significant increase in MDA in the second week compared to the first one.

Experimental cows showed no statistical correlations between MDA and NEFA, unlike control cows, which exhibited a linear increase in NEFA and MDA.

Malondialdehyde (MDA) is a final product of half-saturated fatty acid peroxidation in the cells. The increase in free radicals can cause the hyperproduction of MDA, which is known as a marker of oxidative stress (Gawel *et al.*, 2004). MDA concentration was higher after calving in dairy cows and showed a positive correlation with NEFA and BHB concentrations (Bernabucci

*et al.*, 2005). This experiment showed that week of sampling and niacin administration had a significant influence on MDA concentration in blood samples ( $p < 0.0001$ ). MDA concentration was significantly lower in cows receiving niacin compared to control cows during all three weeks of the experiment. Lower NEFA concentrations in cows receiving niacin were related to lower MDA values during the experiment. Yuan *et al.* (2012) examined the antioxidant level and superoxide dismutase (SOD) in cows during the peripartal period after niacin administration and showed no significant effect of niacin administration. Our results are consistent with the results of scientists who examined the effect of niacin on lipid peroxidation in different animal models. Models that had niacin deficiency showed greater lipid peroxidation. Models that received niacin showed reduced lipid peroxidation (Ghazi-Khansari *et al.* 2005; Cho *et al.*, 2009; Perumal *et al.*, 2005; Atac *et al.*, 2006; Tupe *et al.*, 2011; Doger *et al.*, 2011).

*Graphic 1.* MDA concentration in cows receiving niacin and control cows in the period of 0–2 weeks after calving



*Graphic 2.* Correlation between NEFA and MDA in niacin and control cows in all experimental weeks

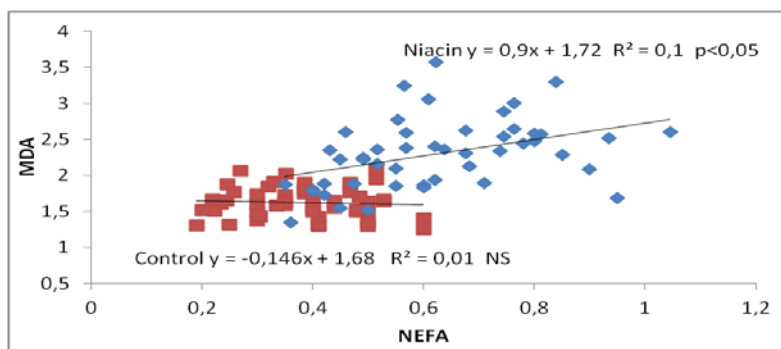


Table 1. LSD test of MDA value differences between different experimental groups for ANOVA analysis

<i>Contrast</i>	<i>Sig.</i>	<i>Difference</i>
Week 0 Control - Week 0 Niacin	p<0.01	0.467333
Week 0 Control - Week 1 Control	p<0.01	-0.474
Week 0 Control - Week 1 Niacin	p<0.01	0.252
Week 0 Control - Week 2 Control	p<0.01	-0.710667
Week 0 Control - Week 2 Niacin	NS	0.09
Week 0 Niacin - Week 1 Control	p<0.01	-0.941333
Week 0 Niacin - Week 1 Niacin	NS	-0.215333
Week 0 Niacin - Week 2 Control	p<0.01	-1.178
Week 0 Niacin - Week 2 Niacin	p<0.05	-0.377333
Week 1 Control - Week 1 Niacin	p<0.01	0.726
Week 1 Control - Week 2 Control	p<0.05	-0.236667
Week 1 Control - Week 2 Niacin	p<0.01	0.564
Week 1 Niacin - Week 2 Control	p<0.01	-0.962667
Week 1 Niacin - Week 2 Niacin	NS	-0.162
Week 2 Control - Week 2 Niacin	p<0.01	0.800667

### Conclusion

Niacin administration during early lactation causes reduction in lipid peroxidation in cows. Reduced lipid peroxidation in cows could be a consequence of the antilipolytic effect of niacin.

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## UTICAJ APLIKACIJE NIACINA NA LIPIDNU PEROKSIDACIJU KOD KRAVA U RANOJ LAKTACIJI

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### Rezime

Niacin je reaktivna polovina NAD i NADP, koji su koenzimi u velikom broju oksidativnih i reduktivnih reakcija. U periodu rane laktacije postoji povećana lipidna mobilizacija, inflamatorni odgovor i oksidativni stres. Cilj ovog rada je da se ispita da li aplikacija niacina kod krava u ranoj laktaciji dovodi do smanjenja lipidne peroksidacije odnosno koncentracije MDA. Niacin je aplikovan putem hrane u periodu dve nedelje pre i dve nedelje posle teljenja. Krv je uzeta u nedelji teljenja a potom i u prvoj i drugoj nedelji posle teljenja. Koncentracija MDA je bila značajno niža kod krava koje su primale niacin u odnosu na kontrolnu grupu tokom sve tri nedelje ispitivanja:  $1,89 \pm 0,33; 1,42 \pm 0,13$  (teljenje, nulta nedelja);  $2,37 \pm 0,41; 1,64 \pm 0,15$  (prva nedelja) i  $2,6 \pm 0,45; 1,8 \pm 0,17 \mu\text{mol/L}$  (druga nedelja). Kod krava ogleadne grupe nema statistički značajne korelacije između vrednosti MDA i NEFA, za razliku od kontrolne grupe koja nije primala niacin gde je koncentracija MDA linearno rasla sa porastom koncentracije NEFA. Upotreba niacina u ranoj laktaciji dovodi do smanjenja lipidne peroksidacije kod krava. Smanjena lipidna peroksidacija kod krava može biti posledica antilipolitičkog delovanja niacina.

**Ključne reči:** krave, niacin, lipoliza, oksidativni stres.