

Macroalgal extracts improve health status and production performance

As demonstrated in many scientific studies, the onset of infectious pathologies during the first weeks of lactation (60-80% of total pathologies in the dairy herd) is linked to immunosuppression derived from nutritional/metabolic stresses and hormonal changes that take place around calving.

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The increase in non-esterified fatty acids (NEFAs) during the transition period and the increase in steroid hormones (cortisol) at the time of calving have been correlated with two negative impacts on the immune response. Firstly, a decrease in the defence activities of immune cells and secondly, a

reduced ability of immune cells to migrate to infection sites. The lower feed intake experienced during the dry-off period leads to a higher degree of oxidative status which contributes to immune dysfunction. Indeed, a reduction of 50% in the activity of neutrophils is being confirmed in several studies. The glucose requirements of an acute immune response are more than 1kg of glucose within 720 minutes in lactating dairy cows, glucose being the substrate for lactose synthesis.

A compromised barrier function in the gut caused by acidosis and subsequent LPS infiltration, will be followed by local or systemic inflammatory response and a negative impact on the technical performance. A decrease in feed intake during heat stress is only responsible for 50% of reduced milk yield, gut barrier function and lipopolysaccharides (LPS) infiltration will induce an immune response that will consume glucose contributing to

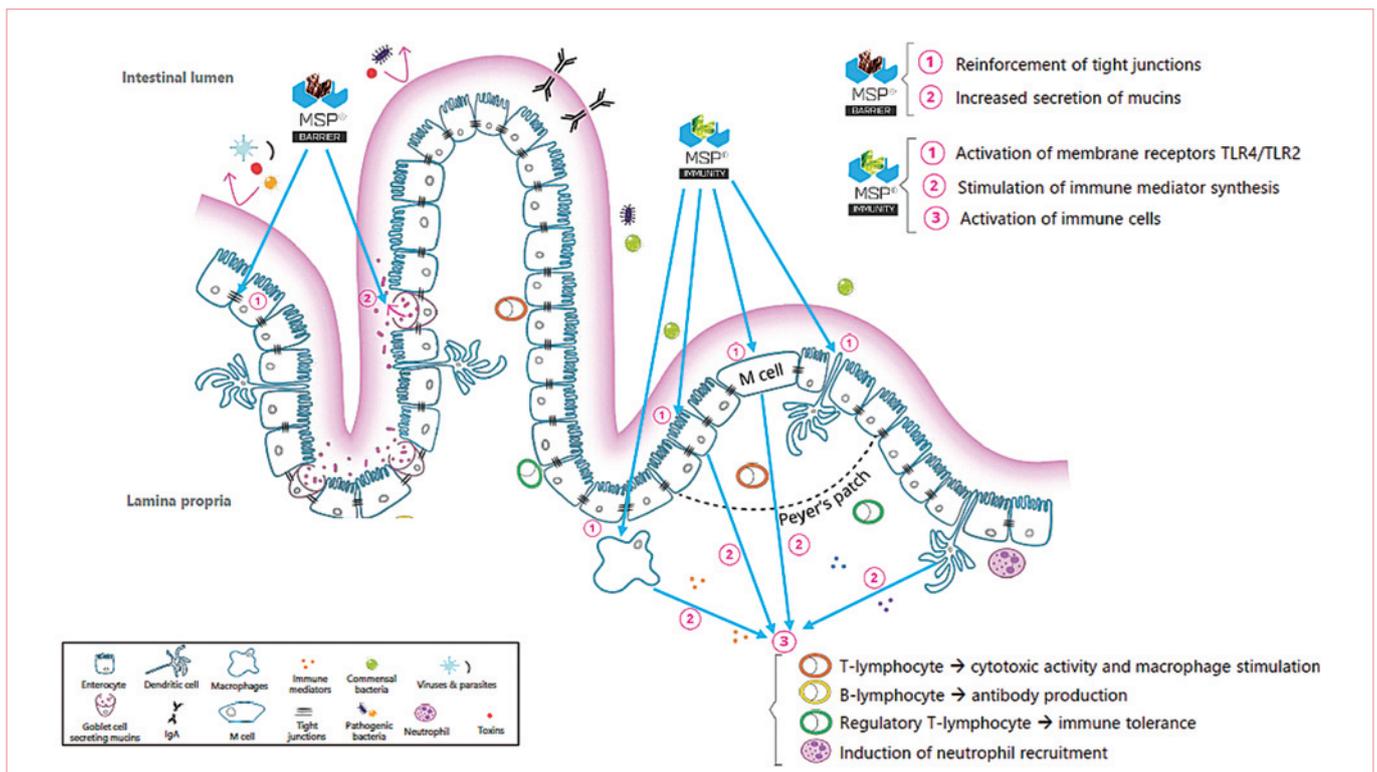
the reduction in the milk yield since lactose is the main osmoregulatory of milk yield. It is paramount to support the immune function and gut barrier function during phases in which cows are exposed to high levels of stressors, among others the dry period, early lactation and in heat stress, in order to reduce or prevent the negative consequences in animal welfare, milk yield and fertility.

Unique structural features of marine macroalgal polysaccharides

Parietal polysaccharides of seaweeds present structural complexity and unique composition that make them very reactive and explain their biological activities towards animals. The complexity and reactivity of seaweed polysaccharides derive from the nature of the sugar units, which are

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Fig. 1. Mode of action of MSP^{IMMUNITY} and MSP^{BARRIER}.



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diverse and sometimes rare, like uronic acids, xylose and rhamnose, the variety of glycosidic bonds leading to their branched structure and the presence of sulphate groups. Furthermore, their polyanionic structure and solubility increases their reactivity and facilitates their recognition by host cells. Sulphated polysaccharides are characteristic of macroalgae (they are not found in terrestrial plants, nor fresh water microalgae or yeast cell walls).

Olmix Group, France, has been studying marine biotechnologies for more than 20 years and has focused on the extraction and use of specific macroalgal polysaccharides to support immune and gut barrier functions. Olmix Group marine bioactive ingredient extraction know-how has led to the development of an in-feed product, Algimun, which is based on the combination of two biologically active macroalgal extracts: MSP_{IMMUNITY}, a green algal extract that reinforces innate and adaptive immune responses; and MSP_{BARRIER}, a red algal extract, which enhances the barrier function of the intestinal mucosa (Fig. 1).

MSPs cannot be digested by enzymes in terrestrial animals (the only organisms that can do so are marine micro-organisms), so Algimun will pass the rumen and act as expected on the intestinal mucosa.

Immunomodulating properties of macroalgal polysaccharides

A research project in collaboration with INRA (France) led to the demonstration of the effect of MSP_{IMMUNITY}, on immune mediators' transcription in an in-vitro model (IPEC-1 cell line), including the identification of the metabolic pathways involved in this activation.

Berri et al. (2016) first highlighted that MSP_{IMMUNITY} could positively influence the gene transcription of a broad array of immune mediators involved in defence mechanisms within the innate and the adaptive immune response, among others, the recruitment and activation of antigen-presenting cells, the differentiation and proliferation of lymphocytes B and

lymphocytes T, while inducing immune tolerance thanks to its anti-inflammatory properties. In-vivo scientific studies further confirmed the immunomodulating properties of MSP_{IMMUNITY}, namely by improving the defence activities of monocytes and neutrophils, by favouring the immune transfer through colostrum and milk and by improving the phagocytic activity of macrophages and neutrophils. MSP_{IMMUNITY}, also upregulates the gene expression of immune mediators with anti-inflammatory activities.

Gut barrier function preserving macroalgal polysaccharides

The effect of MSP_{BARRIER} on intestinal barrier function was assessed by measuring the expression of tight junction proteins and mucin-related genes using epithelial cell lines. Results showed that MSP_{BARRIER} upregulates the expression of genes that determine transmembrane (CLND-2) and scaffolding protein (ZO-1) synthesis which are essential for optimal functioning of the tight junction complexes.

Moreover MSP_{BARRIER} upregulates the expression of genes that are necessary for the establishment of the mucus layer (MUC-2 and MUC-4) and the prevention of pathogen colonisation. MSP_{BARRIER} plays an important role in the maintenance of gut mucosa integrity by rendering the tight junctions stronger and promoting a more functional mucus layer. MSP_{BARRIER} strengthens gut integrity as observed by an increase in trans-epithelial electrical resistance (TEER) in IPEC-1 cell line incubated with an E. coli K88 1305.

These findings were corroborated in an in-vivo scientific study. MSP_{BARRIER} reduced the paracellular passage of FITC-dextran (gut permeability biomarker) to the bloodstream in an animal model known to induce specific stress leading to an inflammation of the gut epithelium and consequently a higher degree of permeability.

MSP_{BARRIER} supplementation reinforced the intestinal barrier function of the gut epithelium (significant reduction of the passage of the marker).

Effect of a combination of macroalgal extracts on cow health and performance

The inclusion of Algimun in cow feed during early lactation was evaluated in a study conducted simultaneously in five commercial farms in France. The results obtained were compared with farm data from the previous year (control group) and herd management between the two years remained unchanged in each farm. In total, 500 Holstein breed cows received Algimun during the first 90 days of lactation at a dose of 20g/cow/day.

An increase of 2% in the milk yield and improved milk composition (Fig. 2) were observed in cows receiving Algimun when compared to the control group. In addition, the number of dairy cows treated for clinical mastitis was lower in the Algimun group (n=5) when compared to the control group (n=25), reflecting an improvement of the immune status.

Thanks to this improvement of performance, the addition of Algimun is profitable (ROI= 2:1).

The positive effect of Algimun during early lactation (20g/cow/day) has been confirmed in Italian commercial farms under heat stress periods (summer). Algimun has proven to support the health of the herds as seen by lower somatic cells counts, lower or no drop in milk production and an improved fertility rate in autumn.

The dry period is another risk period in which Algimun (10g/cow/day) use brings benefits too, as seen by an improvement of the colostrum quality: <28% Brix value when compared to the control group's 22% Brix value. In short, Algimun, a macroalgae based solution, can be used as a natural alternative in-feed strategy to support the health status of the gastrointestinal epithelia by reinforcing the barrier function of the digestive mucosa and the immune function. A healthier digestive tract will be less prone to pathologies and more nutrients will be used towards production. ■

References are available from the author on request

Fig. 2. Milk yield during early lactation (left) and milk composition during early lactation (right).¹ Number of farms in which test group results in 2020 were better than control group results in 2019.

