

**Popular Article** 

# Unleashing the Power of Heat Shock Proteins: Revolutionizing **Livestock Production**

# <sup>1</sup>Avani Singh, <sup>2</sup>Prachi Dhaka, <sup>3</sup> Pavan Kumar Mittal, <sup>4</sup> Sudaksha Saraswat <sup>1, 2 and 3</sup>Department of Veterinary Physiology and Biochemistry, Post Graduate Institute of Veterinary Education and Research, Jamdoli, Jaipur, Rajasthan-302031 <sup>4</sup>Department of Biochemistry, Pandit Deen Dayal Upadhyaya Pashu Chikitsa Vigyan Vishwavidyalaya Evam Go-Anusandhan Sansthan University, Mathura, Uttar Pradesh-281001 https://doi.org/10.5281/zenodo.7983592

# Introduction

Livestock production plays a crucial role in meeting the growing demand for food worldwide. However, the welfare and productivity of farm animals are constantly challenged by environmental stressors, such as heat waves, high humidity, and transportation. In recent years, scientists have been uncovering the extraordinary potential of a group of proteins known as heat shock proteins (HSPs) in mitigating the harmful effects of these stressors. These remarkable proteins have shown promise in revolutionizing livestock production by improving animal health, performance, and overall welfare. In this article, we delve into the fascinating world of heat shock proteins and explore their pivotal role in transforming the industry.

# What are Heat Shock Proteins?

Heat shock proteins are a highly conserved group of molecular chaperones found in almost all living organisms. Heat shock proteins help protect cells from damage by facilitating proper protein folding, preventing the aggregation of misfolded proteins, and promoting cellular repair mechanisms. They are called "heat shock proteins" because they were first discovered as proteins that are rapidly induced in response to heat stress by Ferruccio Ritossa, an Italian geneticist, who made the initial observations in the 1960s. Ritossa was studying the chromosomes of Drosophila



*melanogaster* (fruit flies) using a technique called polytene chromosome staining. During his experiments, Ritossa noticed a peculiar banding pattern on the chromosomes when these flies were exposed to high temperatures. The banding pattern indicated that certain regions of the chromosomes were becoming more active in response to heat stress. In the early 1970s, proteins from these puffed regions were isolated and termed "heat shock proteins" due to their association with heat stress. The name "heat shock proteins" was coined by Paul Tissières, an influential scientist who studied these proteins in bacteria. However, their production can also be triggered by other stressors, such as cold, oxidative stress, infection, inflammation, and even emotional stress.

The primary function of HSPs is to maintain cellular homeostasis and protect cells from damage under stressful conditions. They accomplish this by facilitating proper protein folding, preventing the aggregation of misfolded or damaged proteins, and assisting in the refolding or degradation of non-functional proteins. HSPs also play roles in transporting proteins within cells and ensuring their correct localization.

HSPs are classified into different families based on their molecular weight, such as Hsp70, Hsp90, Hsp60 (also known as chaperonins), and small heat shock proteins (sHSPs). Each family has distinct functions and cellular localization.

**Hsp70**: This family of heat shock proteins is involved in a wide range of cellular processes, including protein folding, protein transport across cellular membranes, and preventing the aggregation of proteins under stress conditions.

**Hsp90**: Hsp90 proteins are essential for the correct folding, stabilization, and activation of many client proteins involved in cell signaling, development, and stress responses. They often work in complex with other proteins called co-chaperones.

**Hsp60** (**Chaperonins**): Chaperonins, such as Hsp60, are large protein complexes that facilitate the folding of newly synthesized proteins or the refolding of denatured proteins. They provide an isolated environment where proteins can fold correctly.

**Small Heat Shock Proteins (sHSPs)**: sHSPs are a diverse group of small proteins that have chaperone-like functions. They can prevent protein aggregation, stabilize partially folded proteins, and assist in their refolding. sHSPs are found in various cellular compartments and are involved in protecting cells from stress-induced damage.



The production of HSPs is regulated by a family of transcription factors known as heat shock factors (HSFs), which are activated upon exposure to stress. These HSFs bind to specific DNA sequences called heat shock elements (HSEs) in the promoters of HSP genes, leading to the transcription and subsequent synthesis of HSPs. The induction of HSPs is a critical cellular response that helps organisms adapt to stressful conditions and maintain cellular integrity. Beyond their protective functions, HSPs are also involved in various cellular processes, including immune response modulation, apoptosis regulation, and cellular signalling pathways.

## Heat Shock Proteins and Environmental Stressors

Livestock animals, such as cows, pigs, and poultry, are highly susceptible to environmental stressors. Heat stress, in particular, can lead to reduced feed intake, compromised reproduction, decreased milk production, and even mortality. However, recent studies have demonstrated that heat shock proteins play a vital role in mitigating these adverse effects. When animals are exposed to heat stress, the production of HSPs is triggered, aiding in the stabilization and refolding of damaged proteins, thereby maintaining cellular function. Also due to cold stress, cells undergo physiological and metabolic changes to adapt to the lower temperatures. This can result in disruptions to cellular functions and the misfolding of proteins. However, HSPs help protect cells by stabilizing and refolding proteins, thereby maintaining proper cellular function under cold stress conditions. In addition to their role in protein protection, HSPs also contribute to cellular membranes' stability and fluidity, which can be compromised in cold environments. They help prevent the formation of ice crystals within cells and improve membrane integrity, reducing the risk of cold-induced cell damage.

#### **Enhancing Immune Function**

Beyond their role in protein protection, heat shock proteins have been found to play a crucial role in enhancing the immune response in livestock animals. Heat shock proteins (HSPs) play a crucial role in immune function by modulating various aspects of the immune response. They are involved in both innate and adaptive immunity and contribute to the recognition and clearance of pathogens, as well as the regulation of inflammatory processes. Here are some key roles of HSPs in immune function:

• Antigen Presentation: HSPs are involved in the presentation of antigens to immune cells, such as T cells. They can chaperone and deliver antigens derived from intracellular pathogens or tumour cells



to antigen-presenting cells (APCs) like dendritic cells. This enhances the recognition of antigens by T cells, leading to a more effective immune response.

- **T-Cell Activation**: HSPs can directly interact with T cells and promote their activation and proliferation. HSPs complexed with specific peptides derived from pathogens or tumours can bind to the T cell receptor (TCR) and co-receptors on T cells, triggering immune signalling pathways that initiate T cell activation and differentiation.
- Chaperoning Immune Signalling Molecules: HSPs assist in the proper folding and stabilization of immune signalling molecules, such as Toll-like receptors (TLRs) and cytokines. By ensuring the correct conformation of these molecules, HSPs contribute to their functional activity and signalling cascades involved in immune responses.
- Inflammation Regulation: HSPs can regulate inflammatory processes by modulating the production and release of inflammatory cytokines. They can either promote or suppress inflammation depending on the context. For example, certain HSPs can act as danger signals (also known as damage-associated molecular patterns or DAMPs) when released from damaged or stressed cells, triggering immune responses and inflammation. On the other hand, other HSPs, like Hsp70, have been shown to possess anti-inflammatory properties and can dampen excessive immune responses.
- **Immunomodulatory Effects**: HSPs can modulate the function and differentiation of immune cells. They can influence the maturation of dendritic cells, promote the differentiation of regulatory T cells (Tregs), and regulate the balance between pro-inflammatory and anti-inflammatory immune responses. HSPs may also influence the production and activity of natural killer (NK) cells and macrophages.
- Anti-apoptotic Effects: Some HSPs (27 and 70) have anti-apoptotic properties and can protect immune cells from stress-induced cell death. By preventing cell death, HSPs help maintain a functional immune cell population, ensuring an effective immune response.

The precise mechanisms by which HSPs exert their immune-modulatory effects are still being investigated, and different HSP family members may have distinct roles in immune function. Furthermore, dysregulation of HSPs can have implications for immune-related diseases, such as autoimmune disorders, cancer, and inflammatory conditions.





Understanding the complex interplay between HSPs and the immune system is crucial for developing therapeutic strategies that target HSPs to modulate immune responses, enhance vaccine efficacy, and potentially treat immune-related disorders.

**Improved Reproductive Performance:** Heat shock proteins (HSPs) have been found to play important roles in the reproductive processes of livestock. They contribute to the maintenance of reproductive health, gamete development, embryo development, and overall fertility. Here are some key roles of HSPs in the reproduction of livestock:

- **Spermatogenesis and Oocyte Development**: HSPs are involved in the development and maturation of spermatozoa and oocytes. They assist in the proper folding and assembly of proteins critical for the formation of functional gametes. HSPs also help protect developing sperm and oocytes from stress-induced damage, ensuring their viability and fertility.
- Stress Response and Reproductive Performance: Stressors, such as heat stress or nutritional deficiencies, can negatively impact reproductive performance in livestock. HSPs are induced in response to stress and act as molecular chaperones to mitigate the harmful effects of stress on reproductive organs and cells. They protect against protein misfolding, maintain cellular homeostasis, and prevent oxidative damage, thereby promoting reproductive health and function.
- Embryo Development and Implantation: HSPs are involved in embryonic development and implantation processes. They aid in the proper folding and function of proteins required for embryogenesis. HSPs also play a role in regulating apoptosis (programmed cell death) during embryo development, ensuring the survival and proper development of embryos.
- **Pregnancy and Placental Function**: HSPs are important for maintaining a healthy pregnancy and proper placental function. They assist in the proper folding and trafficking of proteins critical for placental development and hormone production. HSPs also help protect placental cells from stress-induced damage, ensuring their integrity and optimal function.
- Sperm and Embryo Cryopreservation: Cryopreservation of sperm and embryos is commonly used in livestock breeding programs. The freezing and thawing processes can induce stress and damage to the cells. HSPs play a role in protecting cells during cryopreservation by preventing protein denaturation, maintaining membrane integrity, and reducing oxidative stress. They enhance the survival and viability of sperm and embryos after thawing, improving the success of assisted reproductive techniques.



Understanding the involvement of HSPs in reproductive processes is valuable for improving breeding strategies, optimizing reproductive efficiency, and enhancing livestock productivity. Strategies aimed at reducing stress and improving HSP function, such as providing appropriate environmental conditions, managing nutrition, and developing stress-resistant genetic lines, can positively impact reproductive outcomes in livestock production.

## **Maximizing Production Efficiency**

Stress not only affects animal health and reproduction but also reduces productivity and economic efficiency in livestock systems. However, the integration of heat shock proteins into livestock production practices shows promise in maximizing efficiency. By protecting protein structures and maintaining cellular function, HSPs help animals maintain their metabolic rate, nutrient utilization, and growth performance even under stressful conditions. This leads to improved feed conversion efficiency, higher weight gains, increased milk production, and enhanced overall productivity.

## **Future Perspectives**

As our understanding of heat shock proteins deepens, further research is warranted to explore their potential applications in livestock production fully. Scientists are investigating the use of genetic selection to breed animals with an enhanced ability to produce heat shock proteins, as well as the development of nutritional strategies to optimize HSP production. Additionally, advancements in biotechnology may offer avenues for the targeted delivery of heat shock proteins to specific tissues or organs, further improving their protective capabilities.

# Conclusion

Heat shock proteins are emerging as powerful allies in combating the detrimental effects of environmental stressors on livestock animals. By safeguarding cellular function, enhancing immune responses, and improving reproductive performance, HSPs have the potential to revolutionize livestock health and production potential.

## References

Mishra, S.R. and Palai, T.K., 2014. Importance of heat shock protein 70 in livestock-at cellular level. *J Mol Pathophysiol, Apr-Jun, 3*(2).

Feder, M.E. and Hofmann, G.E., 1999. Heat-shock proteins, molecular chaperones, and the stress response: evolutionary and ecological physiology. *Annual review of physiology*, *61*(1), pp.243-282.



- Muralidharan, S. and Mandrekar, P., 2013. Cellular stress response and innate immune signaling: integrating pathways in host defense and inflammation. *Journal of leukocyte biology*, 94(6), pp.1167-1184.
- Li, Z., Menoret, A. and Srivastava, P., 2002. Roles of heat-shock proteins in antigen presentation and cross-presentation. *Current opinion in immunology*, *14*(1), pp.45-51.
- Hauet-Broere, F., Wieten, L., Guichelaar, T., Berlo, S., Van Der Zee, R. and Van Eden, W., 2006. Heat shock proteins induce T cell regulation of chronic inflammation. *Annals of the rheumatic diseases*, 65(suppl 3), pp.iii65-iii68.
- Pockley, A.G., 2003. Heat shock proteins as regulators of the immune response. *The lancet*, 362(9382), pp.469-476.
- Garrido, C., Brunet, M., Didelot, C., Zermati, Y., Schmitt, E. and Kroemer, G., 2006. Heat shock proteins 27 and 70: anti-apoptotic proteins with tumorigenic properties. *Cell cycle*, *5*(22), pp.2592-2601.
- Dix, D.J., 1997. HSP 70 expression and function during gametogenesis. *Cell stress & chaperones*, 2(2), p.73.
- Fraser, L., 2017. Markers for sperm freezability and relevance of transcriptome studies in semen cryopreservation: A. *Theriogenology*, 47.



