

Investigation of codon 136 (A-V) and codon 171 (H-Q-R) polymorphisms for scrapie susceptibility in Makuei Sheep breeds

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Abstract

Prion diseases in livestock can be a potential danger for humans as well. The Makuei sheep breed is one of the most important sources of protein in Iran. In this study, the polymorphism status of codons 136 and 171 of prion protein and their genotypes was investigated in Makuei ecotype sheep. Blood samples were obtained from 60 Makuei sheep and DNA was extracted. The codon 136 (A-V) and codon 171 (H-Q-R) polymorphisms were evaluated by PCR with specific primers. The results showed that allele A (65%) at codon 136 was significantly higher than allele V (35%). AA genotype (46.67%) was also the most common genotype. At codon 171, the Q and R alleles were significantly higher than the H allele. In addition, the allele R was significantly lower than the allele Q. Allele Q (58.33%) and QQ genotypes (40%) were the most common alleles and genotypes. In conclusion, the findings provide strong preliminary lines of evidence that motive large-scale genotyping studies to establish an effective breeding control and successful eradication of scrapie-susceptible genotypes.

Key words: Prion disease, Scrapie, Makuei ecotype, PrP, Genotype

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References

- Baylis, M., & McIntyre, K. M. (2004). Scrapie control under new strain. *Nature*, 432(7019), 810-811.
- Boukouvala, E., Gelasakis, A. I., Kanata, E., Fragkiadaki, E., Giadinis, N. D., Palaska, V., Christoforidou, S., Sklaviadis, T., & Ekateriniadou, L. V. (2020). The association between 171 K polymorphism and resistance against scrapie affection in Greek dairy sheep. *Small Ruminant Research*, 161, 51-56.
- Cassmann, E. D., & Greenlee, J. J. (2020). Pathogenesis, detection, and control of scrapie in sheep. *Am J Vet Res*, 81(7), 600-614. <https://doi.org/10.2460/ajvr.81.7.600>
- Chaudhary, S., & Chaudhary, M. (2013). Scrapie: A neuro degenerative disease in sheep-A review. *Agricultural Reviews*, 34(1), 79-85.
- Curcio, L., Sebastiani, C., Di Lorenzo, P., Lasagna, E., & Biagetti, M. (2016). Review: A review on classical and atypical scrapie in caprine: Prion protein gene polymorphisms and their role in the disease. *Animal*, 10(10), 1585-1593. <https://doi.org/10.1017/s1751731116000653>
- DeSilva, U., Guo, X., Kupfer, D., Fernando, S., Pillai, A., Najar, F., So, S., Fitch, G., & Roe, B. (2003). Allelic variants of ovine prion protein gene (PRNP) in Oklahoma sheep. *Cytogenetic and genome research*, 102(1-4), 89-94.
- Gmür, A., Gaillard, C., & Dolf, G. (2004). Characterization of the prion protein gene (PRNP) region in Swiss sheep breeds. *Journal of Animal Breeding and Genetics*, 121(3), 216-220.
- GOMBOJAV, A., ISHIGURO, N., HORIUCHI, M., SERJMYADAG, D., BYAMBAA, B., & SHINAGAWA, M. (2003). Amino acid polymorphisms of PrP gene in Mongolian sheep. *Journal of veterinary medical science*, 65(1), 75-81.
- Houston, F., Goldmann, W., Foster, J., González, L., Jeffrey, M., & Hunter, N. (2015). Comparative Susceptibility of Sheep of Different Origins, Breeds and PRNP Genotypes to Challenge with Bovine Spongiform Encephalopathy and Scrapie. *PloS one*, 10(11), e0143251-e0143251. <https://doi.org/10.1371/journal.pone.0143251>
- Hunter, N. (2007). Scrapie—Uncertainties, biology and molecular approaches. *Biochimica et Biophysica Acta (BBA)-Molecular Basis of Disease*, 1772(6), 619-628.
- Hussein, M. F., & Al-Mufarrej, S. I. (2004). Prion Diseases: A Review. *Scientific Journal of King Faisal University (Basic and Applied Sciences)*, 5(2), 1425.
- Jeong, M.-J., Kim, Y.-C., & Jeong, B.-H. (2018). Prion-like protein gene (PRND) polymorphisms associated with scrapie susceptibility in Korean native black goats. *PloS one*, 13(10).
- Lan, Z., Li, J., Sun, C., Liu, Y., Zhao, Y., Chi, T., Yu, X., Song, F., & Wang, Z. (2014). Allelic variants of PRNP in 16 Chinese local sheep breeds. *Arch Virol*, 159(8), 2141-2144. <https://doi.org/10.1007/s00705-014-2048-9>
- Mead, S., Lloyd, S., & Collinge, J. (2019). Genetic Factors in Mammalian Prion Diseases. *Annu Rev Genet*, 53, 117-147. <https://doi.org/10.1146/annurev-genet-1202-092352-13>
- Migliore, S., Agnello, S., D'Avola, S., Goldmann, W., Presti, V. D. M. L., & Vitale, M. (2017). A cross-sectional study of PRNP gene in two native Sicilian goat populations in Italy: a relation between prion gene polymorphisms and scrapie incidence. *Journal of genetics*, 96(2), 319-325.
- Pandeya, D. R., Acharya, N., & Hong, S.-T. (2010). The prion and its potentiality. *Biomedical Research-Tokyo*, 21(2).
- Salami, S., Zadeh, R. A., Omrani, M. D., Ramezani, F., & Amniattalab, A. (2011). Allelic frequency and genotypes of prion protein at codon 136 and 171 in Iranian Ghezel sheep breeds. *Prion*, 5(3), 228-231. <https://doi.org/10.4161/pri.5.3.16098>
- Scheckel, C., & Aguzzi, A. (2018). Prions, prionoids and protein misfolding disorders. *Nature Reviews Genetics*, 19(7), 405-418.
- Schulz-Schaeffer, W. J., Wemheuer, W. M., & Wrede, A. (2020). Prion Diseases: Conformational Changes of a Protein Create an Unconventional Infectious Agent. In *Emerging and Reemerging Viral Pathogens* (pp. 479-488). Elsevier.

- Teferedegn, E. Y., Yaman, Y., & Un, C. (2020). Five novel PRNP gene polymorphisms and their potential effect on Scrapie susceptibility in three native Ethiopian sheep breeds. *BMC Vet Res*, 16(1), 122. <https://doi.org/10.1186/s12917-020-02336-0>
- Tongue, S., Pfeiffer, D., Warner, R., Elliott, H., & Vilas, V. D. R. (2006). Estimation of the relative risk of developing clinical scrapie: the role of prion protein (PrP) genotype and selection bias. *Veterinary Record*, 158(2), 43-50.
- Yoshida, N., & Soto, P. (2019). Polymorphisms Modulate Sheep Prion Protein Susceptibility to Misfolding by Altering the Residue Network of Interactions. *Biophysical Journal*, 116(3), 187a.