

Variants encompassing the Agouti signaling protein gene are associated with dilution of grey shades in Nellore (*Bos indicus*) cattle

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Abstract

Nellore cattle (*Bos indicus*) are well known for their resilience to infectious diseases, survival in low input systems and tolerance to heat. Traits that are often speculated to contribute to heat tolerance are the skin pigmentation and the coat color. Nellore cattle possess dark-black skin, and although their coat color pattern can range widely from solid red to black-spotted, the Brazilian herds were massively selected for white coat. For this type of coat, whereas females are completely white, males can exhibit shades of dark gray in the head, neck, hump and knees. Curiously, some males are also completely white, and they can transmit this phenotype to their progeny in a dominant mode of inheritance. The present study focused on mapping the functional candidate gene responsible for the dominant white phenotype in Nellore cattle. Bulls were evaluated based on visual scores for gray shading in a subjective scale from 1 to 5 by six evaluators, where the smallest score (1) was attributed to all white animals and the largest (5) attributed to animals with black shades. The final score for each animal was decided based on majority voting. A total of 379 animals were evaluated, from which 131 were chosen and included in this study based on extreme scores. These animals were genotyped with the Illumina BovineHD Bead Chip assay, which included 777,000 single nucleotide polymorphism (SNP) markers. After a standard genotype quality control, a genome-wide association analysis (GWAS) was conducted in order to map loci linked to the white phenotype. Considering a Bonferroni-corrected significance level of $\alpha = 1.15 \times 10^{-7}$, we detected highly significant associations ($p = 7.636928 \times 10^{-127}$) mapping to chromosome 13, in the vicinity of the Agouti signaling protein gene (ASIP). Given that ASIP is a well-known color dilution gene, and that the expression pattern of ASIP in the coats of model organisms follows closely the distribution of grey shades found in Nellore cattle, the gene is a promising functional candidate. We are now sequencing the whole genomes of 8 Nellore bulls (4 black and 4 white) for scrutiny of the identified locus in the search for the causal mutation.

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