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GENOME-WIDE IDENTIFICATION OF SNPs AND THEIR ANNOTATION IN INDIAN GIR CATTLE

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The study was carried out in Gir cattle for identification of genome wide SNPs and then to annotate the identified high-quality SNPs to the milk production traits. A total of 99,517 SNPs were identified with respect to the *Bos indicus* reference genome. Upon annotation of SNPs identified with respect to *Bos indicus* reference genome, 984 SNPs were found to be associated with 175 candidate genes having to do with milk production traits, notably Acetyl-CoA carboxylase β -gene, which affects milk composition traits by regulation of fatty acid oxidation in the mitochondria; Growth hormone receptor gene which have role in milk yield and its composition traits; LEP gene, involved in energy partitioning and metabolism, were among others. This study provides the first analysis of ddRAD sequences to discover SNPs in Indian Gir cattle breed, aligned to reference genome. The variants mined in this study can be incorporated in existing SNP chips and thus, play an important role in understanding the genetic structure of our cattle to design appropriate breed improvement programmes.

Key words: ddRAD, Gir breed, Milk production traits, SNPs, Zebu cattle.

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ПОВНОГЕНОМНА ІДЕНТИФІКАЦІЯ ОДНОНУКЛЕОТИДНИХ ПОЛІМОРФІЗМІВ І ЇХ МАРКУВАННЯ В ІНДІЙСЬКІЙ ПОРОДІ ХУДОБИ ГІР

Індійську породу великої рогатої худоби Гір було використано у дослідженні, проведенню з метою ідентифікації повногеномних однонуклеотидних поліморфізмів (SNP) та подальшого маркування ідентифікованих високоякісних SNP за ознаками молочної продуктивності. Загалом було ідентифіковано 99 517 SNP стосовно референтного геному *Bos indicus*. Після маркування SNP, ідентифікованих стосовно референтного геному *Bos indicus*, було виявлено, що 984 SNP асоціюються з 175 генами-кандидатами, пов'язаними з ознаками молочної продуктивності, особливо з β -геном ацетил-СоА карбоксилази, що впливає на характеристики складу молока шляхом регуляції окислення жирних кислот у мітохондріях; геном рецептора гормону росту, який відіграє свою роль у надоях молока та характеристиках його складу; геном LEP, залученим до розщеплення енергії і метаболізму, та іншими. У цьому дослідженні вперше проведено аналіз послідовностей ddRAD для виявлення SNP у породі Гір індійської великої рогатої худоби, вирівняних з референтним геномом. Варіанти, розглянуті у цьому дослідженні, можна інкорпорувати в наявні чіпи SNP і, таким чином, вони відіграватимуть важливу роль у розумінні генетичної структури нашої худоби для створення відповідних програм покращення породи.

Ключові слова: ddRAD, порода Гір, характеристики молочної продуктивності, однонуклеотидні поліморфізми (SNP), зебу.

REFERENCES

- Albrechtsen A, Nielsen FC, Nielsen R (2010) Ascertainment Biases in SNP Chips Affect Measures of Population Divergence. Mol Biol Evol 27:2534–2547. <https://doi.org/10.1093/molbev/msq148>
Andrews S (2010) FastQC: a quality control tool for high throughput sequence data

- Ba H, Jia B, Wang G et al (2017) Genome-Wide SNP Discovery and Analysis of Genetic Diversity in Farmed Sika Deer (*Cervus nippon*) in Northeast China Using Double-Digest Restriction Site-Associated DNA Sequencing. *G3 Genes Genom Genet* 7:3169–3176. <https://doi.org/10.1534/g3.117.300082>

Bovine HapMap Consortium, Gibbs RA, Taylor JF et al (2009) Genome-wide survey of SNP variation uncovers the genetic structure of cattle breeds. *Science* 324(5926):528–532. <https://doi.org/10.1126/science.1167936>

Buchanan FC, Fitzsimmons CJ, Van Kessel AG et al (2002) Association of a missense mutation in the bovine leptin gene with carcass fat content and leptin mRNA levels. *Genet Select Evolut* 34. <https://doi.org/10.1186/1297-9686-34-1-105>

Buitenhuis B, Janss LL, Poulsen NA et al (2014) Genome-wide association and biological pathway analysis for milk-fat composition in Danish Holstein and Danish Jersey cattle. *BMC Genomics* 15:1112. <https://doi.org/10.1186/1471-2164-15-1112>

Catchen JM, Amores A, Hohenlohe P et al (2011) Stacks: Building and Genotyping Loci De Novo From Short-Read Sequences. *G3 Genes Ge-nom Genet* 1:171–182. <https://doi.org/10.1534/g3.111.000240>

Chatterjee S, Szustakowski JD, Nanguneri NR et al (2009) Identification of Novel Genes and Pathways Regulating SREBP Transcriptional Activity. *PLoS ONE* 4:e5197. <https://doi.org/10.1371/journal.pone.0005197>

Cingolani P, Platts A, Wang LL et al (2012) A program for annotating and predicting the effects of single nucleotide polymorphisms, SnpEff. *Fly* 6:80–92. <https://doi.org/10.4161/fly.19695>

Clempton AM, Pollott GE, Brickell JS et al (2011) Evidence that leptin genotype is associated with fertility, growth, and milk production in Holstein cows. *J Dairy Sci* 94:3618–3628. <https://doi.org/10.3168/jds.2010-3626>

DaCosta JM, Sorenson MD (2016) ddRAD-seq phylogenetics based on nucleotide, indel, and presence-absence polymorphisms: Analyses of two avian genera with contrasting histories. *Mol Phylogen Evol* 94:122–135. <https://doi.org/10.1016/j.ympev.2015.07.026>

Danecek P, Auton A, Abecasis G et al (2011) The variant call format and VCF tools. *Bioinformatics* 27:2156–2158. <https://doi.org/10.1093/bioinformatics/btr330>

Devadasan MJ, Kumar DR, Vineeth MR et al (2020) Reduced representation approach for identification of genome-wide SNPs and their annotation for economically important traits in Indian Tharparkar cattle. *3 Biotech* 10: <https://doi.org/10.1007/s13205-020-02297-z>

Di Stasio L, Destefanis G, Brugiapaglia A et al (2005) Polymorphism of the GHR gene in cat-tle and relationships with meat production and quality. *Anim Genet* 36:138–140. <https://doi.org/10.1111/j.1365-2052.2005.01244.x>

FAOSTAT. <https://www.fao.org/faostat/en/#home>. Accessed May 25, 2023

Gaur GK, Kaushik SN, Garg RC (2003) The Gir cattle breed of India – characteristics and present status. *Anim Genet Res Inform* 33:21–29. <https://doi.org/10.1017/s1014233900001607>

George L, Alex R, Sukhija N et al (2023) Genetic improvement of economic traits in Murrah buffalo using significant SNPs from genome-wide association study. *Trop Anim Health Product* 55. <https://doi.org/10.1007/s11250-023-03606-3>

Goli RC, Sukhija N, Rathi P et al (2024) Unraveling the genetic tapestry of Indian chicken: A comprehensive study of molecular variations and diversity. *Ecological Genetics and Genomics* 30: 100220. <https://doi.org/10.1016/j.egg.2024.100220>

Gurgul A, Semik E, Pawlina K et al (2014) The application of genome-wide SNP genotyping methods in studies on livestock genomes. *J Appl Genet* 55:197–208. <https://doi.org/10.1007/s13353-014-0202-4>

Han B, Yuan Y, Liang R et al (2019) Genetic Effects of LPIN1 Polymorphisms on Milk Production Traits in Dairy Cattle. *Genes* 10:265. <https://doi.org/10.3390/genes10040265>

India Meteorological Department, Met Center in Ahmedabad, 2023. <https://mausam.imd.gov.in/ahmedabad/> Accessed May 17, 2023

Iqbal N, Liu X, Yang T et al (2019) Genomic variants identified from whole-genome resequencing of indicine cattle breeds from Pakistan. *PLOS ONE* 14:e0215065. <https://doi.org/10.1371/journal.pone.0215065>

Jaglan K, Ravikumar D, Sukhija N et al (2023) Genomic clues of association between clinical mastitis and SNPs identified by ddRAD sequencing in Murrah buffaloes. *Animal Biotechnology* 34:4538–4546. <https://doi.org/10.1080/10495398.2023.2165937>

Kanaka KK, Sukhija N, Goli RC et al (2023) On the concepts and measures of diversity in the genomics era. *Current Plant Biology* 33:100278. <https://doi.org/10.1016/j.cpb.2023.100278>

Keller I, Bensasson D, Nichols RA (2007) Transition-Transversion Bias Is Not Universal: A Counter Example from Grasshopper Pseudogenes. *PLoS Genetics* 3:e22. <https://doi.org/10.1371/journal.pgen.0030022>

Klein C, Bauersachs S, Ulbrich SE et al (2006) Monozygotic Twin Model Reveals Novel Embryo-

- Induced Transcriptome Changes of Bovine Endometrium in the Preattachment Period1. Biol Reprod 74:253–264. <https://doi.org/10.1095/biolreprod.105.046748>

Xkks S, Reimann E, Lilleoja R et al (2014) Sequencing and annotated analysis of full genome of Holstein breed bull. Mammalian Genome 25:363–373. <https://doi.org/10.1007/s00335-014-9511-5>

Kraus RH, Kerstens HH, Van Hooft P et al (2011) Genome wide SNP discovery, analysis and evaluation in mallard (*Anas platyrhynchos*). BMC Genomics 12. <https://doi.org/10.1186/1471-2164-12-150>

Langmead B, Salzberg SL (2012) Fast gapped-read alignment with Bowtie 2. Nature Methods 9:357–359. <https://doi.org/10.1038/nmeth.1923>

Li H (2011) A statistical framework for SNP calling, mutation discovery, association mapping and population genetical parameter estimation from sequencing data. Bioinformatics 27:2987–2993. <https://doi.org/10.1093/bioinformatics/btr509>

Liefers SC, te Pas MFW, Veerkamp RF, van der Lende T (2002) Associations between Leptin Gene Polymorphisms and Production, Live Weight, Energy Balance, Feed Intake, and Fertility in Holstein Heifers. J Dairy Sci 85:1633–1638. [https://doi.org/10.3168/jds.s0022-0302\(02\)74235-5](https://doi.org/10.3168/jds.s0022-0302(02)74235-5)

Malik AA, Sharma R, Ahlawat S et al (2018) Analysis of genetic relatedness among Indian cattle (*Bos indicus*) using genotyping-by-sequencing markers. Anim Genet 49:242–245. <https://doi.org/10.1111/age.12650>

Malik BS, Ghei GC (1977) Some production characteristics of Gir cattle [dairy cattle, India]. Indian J Anim Sci

Mishra DC, Sikka P, Yadav S et al (2020) Identification and characterization of trait-specific SNPs using ddRAD sequencing in water buffalo. Genomics 112: 3571–3578. <https://doi.org/10.1016/j.ygeno.2020.04.012>

Nayee N, Sahana G, Gajjar S, et al (2018) Suitability of existing commercial single nucleotide polymorphism chips for genomic studies in *Bos indicus* cattle breeds and their *Bos taurus* crosses. J Anim Breed Genet 135:432–441. <https://doi.org/10.1111/jbg.12356>

Patel AB, Subramanian RB, Padh H et al (2017) Identification of Single Nucleotide Polymorphism from Indian *Bubalus bubalis* through Targeted Sequence Capture. Curr Sci 112:1230. <https://doi.org/10.18520/cs/v112/i06/1230-1239>

Peterson BK, Weber JN, Kay EH et al (2012) Double Digest RADseq: An Inexpensive Method for De Novo SNP Discovery and Genotyping in Model and Non-Model Species. PLoS ONE 7:e37135. <https://doi.org/10.1371/journal.pone.0037135>

Pimentel ECG, Bauersachs S, Tietze M et al (2010) Exploration of relationships between production and fertility traits in dairy cattle via association studies of SNPs within candidate genes derived by expression profiling. Anim Genet 42:251–262. <https://doi.org/10.1111/j.1365-2052.2010.02148.x>

Porto-Neto LR, Sonstegard TS, Liu GE et al (2013) Genomic divergence of zebu and taurine cattle identified through high-density SNP genotyping. BMC Genom 14:876. <https://doi.org/10.1186/1471-2164-14-876>

Raven L-A, Cocks BG, Goddard ME et al (2014) Genetic variants in mammary development, prolactin signalling and involution pathways explain considerable variation in bovine milk production and milk composition. Genet Select Evolut 46:29. <https://doi.org/10.1186/1297-9686-46-29>

Ravi Kumar D, Joel Devadasan M, Surya T et al (2020) Genomic diversity and selection sweeps identified in Indian swamp buffaloes reveals its uniqueness with riverine buffaloes. Genomics 112:2385–2392. <https://doi.org/10.1016/j.ygeno.2020.01.010>

Schmieder R, Edwards R (2011) Quality control and preprocessing of metagenomic datasets. Bioinformatics 27:863–864. <https://doi.org/10.1093/bioinformatics/btr026>

Sivalingam J, Nirajan SK, Yadav DK et al (2024) Phenotypic and genetic characterization of unexplored, potential cattle population of Madhya Pradesh. Trop Anim Health Product 56. <https://doi.org/10.1007/s11250-024-03946-8>

Sivalingam J, Vineeth MR, Surya T et al (2020) Genomic divergence reveals unique populations among Indian Yaks. Scientific Reports 10. <https://doi.org/10.1038/s41598-020-59887-3>

Stothard P, Choi J-W, Basu U et al (2011) Whole genome resequencing of black Angus and Holstein cattle for SNP and CNV discovery. BMC Genomics 12. <https://doi.org/10.1186/1471-2164-12-559>

Sukhija N, Goli RC, Sukhija M et al (2024b) Evolutionary stamps for adaptation traced in *Cervus nippon* genome using reduced representation sequencing. Conserv Genet Res 16:135–146. <https://doi.org/10.1007/s12686-023-01339-x>

Sukhija N, Kanaka KK, Purohit PB et al (2023) Mendelism: Connecting the Dots Across Centuries. Cytol Genet 57:500–516. <https://doi.org/10.3103/s0095452723050067>

Sukhija N, Malik AA, Devadasan JM et al (2024a) Genome-wide selection signatures address trait specific candidate genes in cattle indigenous to arid regions of India. Anim Biotechnol 35:. <https://doi.org/10.1080/10495398.2023.2290521>

Surya T, Vineeth MR, Sivalingam J et al (2019) Genome-wide identification and annotation of SNPs in

- Bubalus bubalis*. Genomics 111:1695–1698. <https://doi.org/10.1016/j.ygeno.2018.11.021>

Viale E, Tiezzi F, Maretto F et al (2017) Association of candidate gene polymorphisms with milk technological traits, yield, composition, and somatic cell score in Italian Holstein-Friesian sires. J Dairy Sci 100:7271–7281. <https://doi.org/10.3168/jds.2017-12666>

Vineeth MR, Surya T, Sivalingam J et al (2019) Genome-wide discovery of SNPs in candidate genes related to production and fertility traits in Sahiwal cattle. Trop Anim Health Product 52:1707–1715. <https://doi.org/10.1007/s11250-019-02180-x>

Wang H, Jiang L, Liu X et al (2013) A Post-GWAS Replication Study Confirming the PTK2 Gene Associated with Milk Production Traits in Chinese Holstein. PLoS ONE 8:e83625. <https://doi.org/10.1371/journal.pone.0083625>

Zwane AA, Schnabel RD, Hoff J et al (2019) Genome-Wide SNP Discovery in Indigenous Cattle Breeds of South Africa. Frontiers in Genetics 10. <https://doi.org/10.3389/fgene.2019.00273>

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