

## GEPHE SUMMARY

MC1R ( <a href="https://www.gephebase.org/search-criteria?/and+Gene+Gephebase+^MC1R^#gephebase-summary-title">https://www.gephebase.org/search-criteria?/and+Gene+Gephebase+^MC1R^#gephebase-summary-title</a> )	Gephebase Gene	GP00002302	GepheID
Published	Entry Status	Martin	Main curator

## PHENOTYPIC CHANGE

Morphology ( <a href="https://www.gephebase.org/search-criteria?/and+Trait+Category+^Morphology+^#gephebase-summary-title">https://www.gephebase.org/search-criteria?/and+Trait+Category+^Morphology+^#gephebase-summary-title</a> )	Trait Category		
Coloration (coat) ( <a href="https://www.gephebase.org/search-criteria?/and+Trait+^Coloration+^#gephebase-summary-title">https://www.gephebase.org/search-criteria?/and+Trait+^Coloration+^#gephebase-summary-title</a> )	Trait		
WT coat	Trait State in Taxon A		
cream color in Australian Cattle Dog	Trait State in Taxon B		
Taxon A	Ancestral State		
Domesticated ( <a href="https://www.gephebase.org/search-criteria?/and+Taxonomic+Status+^Domesticated+^#gephebase-summary-title">https://www.gephebase.org/search-criteria?/and+Taxonomic+Status+^Domesticated+^#gephebase-summary-title</a> )	Taxonomic Status		
	Taxon A		Taxon B
Canis lupus ( <a href="https://www.gephebase.org/search-criteria?/and+Taxon+and+Synonyms+^Canis+lupus+^#gephebase-summary-title">https://www.gephebase.org/search-criteria?/and+Taxon+and+Synonyms+^Canis+lupus+^#gephebase-summary-title</a> )	Latin Name	Canis lupus familiaris ( <a href="https://www.gephebase.org/search-criteria?/and+Taxon+and+Synonyms+^Canis+lupus+familiaris+^#gephebase-summary-title">https://www.gephebase.org/search-criteria?/and+Taxon+and+Synonyms+^Canis+lupus+familiaris+^#gephebase-summary-title</a> )	Latin Name
gray wolf	Common Name	dog	Common Name
gray wolf; grey wolf; Canis lupus Linnaeus, 1758	Synonyms	Canis canis; Canis domesticus; Canis familiaris; dog; dogs; Canis familiaris Linnaeus, 1758;	Synonyms
species	Rank	Canis lupus familiaris Linnaeus, 1758	Rank
cellular organisms; Eukaryota; Opisthokonta; Metazoa; Eumetazoa; Bilateria; Deuterostomia; Chordata; Craniata; Vertebrata; Gnathostomata; Teleostomi; Euteleostomi; Sarcopterygii; Dipnotetrapodomorpha; Tetrapoda; Amniota; Mammalia; Theria; Eutheria; Boreoeutheria; Laurasiatheria; Carnivora; Caniformia; Canidae; Canis	Lineage	subspecies	Lineage
Canis () - (Rank: genus) ( <a href="https://www.ncbi.nlm.nih.gov/Taxonomy/Browser/wwwtax.cgi?id=9611">https://www.ncbi.nlm.nih.gov/Taxonomy/Browser/wwwtax.cgi?id=9611</a> )	Parent	Canis lupus (gray wolf) - (Rank: species) ( <a href="https://www.ncbi.nlm.nih.gov/Taxonomy/Browser/wwwtax.cgi?id=9612">https://www.ncbi.nlm.nih.gov/Taxonomy/Browser/wwwtax.cgi?id=9612</a> )	Parent
9612 ( <a href="https://www.ncbi.nlm.nih.gov/Taxonomy/Browser/wwwtax.cgi?id=9612">https://www.ncbi.nlm.nih.gov/Taxonomy/Browser/wwwtax.cgi?id=9612</a> )	NCBI Taxonomy ID	9615 ( <a href="https://www.ncbi.nlm.nih.gov/Taxonomy/Browser/wwwtax.cgi?id=9615">https://www.ncbi.nlm.nih.gov/Taxonomy/Browser/wwwtax.cgi?id=9615</a> )	NCBI Taxonomy ID
No	is Taxon A an Intraspecies?	No	is Taxon B an Intraspecies?

## GENOTYPIC CHANGE

MC1R	Generic Gene Name	Q01726 ( <a href="http://www.uniprot.org/uniprot/Q01726">http://www.uniprot.org/uniprot/Q01726</a> )	UniProtKB Homo sapiens
CMM5; MSH-R; SHEP2; MSHR	Synonyms	()	GenebankID or UniProtKB
9606.ENSPO0000451605 ( <a href="http://string-db.org/newstring.cgi/show_network_section.pl?identifier=9606.ENSPO0000451605">http://string-db.org/newstring.cgi/show_network_section.pl?identifier=9606.ENSPO0000451605</a> )	String		
Belongs to the G-protein coupled receptor 1 family.	Sequence Similarities		
GO:0008528 : G protein-coupled peptide receptor activity ( <a href="https://www.ebi.ac.uk/QuickGO/term/GO:0008528">https://www.ebi.ac.uk/QuickGO/term/GO:0008528</a> )	GO - Molecular Function		
GO:0004977 : melanocortin receptor activity ( <a href="https://www.ebi.ac.uk/QuickGO/term/GO:0004977">https://www.ebi.ac.uk/QuickGO/term/GO:0004977</a> )			
GO:0004980 : melanocyte-stimulating hormone receptor activity ( <a href="https://www.ebi.ac.uk/QuickGO/term/GO:0004980">https://www.ebi.ac.uk/QuickGO/term/GO:0004980</a> )			

GO:0031625 : ubiquitin protein ligase binding  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0031625>)

#### GO - Biological Process

GO:0007275 : multicellular organism development  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0007275>)  
GO:0045944 : positive regulation of transcription by RNA polymerase II  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0045944>)  
GO:0042438 : melanin biosynthetic process  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0042438>)  
GO:0043473 : pigmentation (<https://www.ebi.ac.uk/QuickGO/term/GO:0043473>)  
GO:0007186 : G protein-coupled receptor signaling pathway  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0007186>)  
GO:0051897 : positive regulation of protein kinase B signaling  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0051897>)  
GO:0019233 : sensory perception of pain  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0019233>)  
GO:0007189 : adenylate cyclase-activating G protein-coupled receptor signaling pathway  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0007189>)  
GO:0035556 : intracellular signal transduction  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0035556>)  
GO:0007187 : G protein-coupled receptor signaling pathway, coupled to cyclic nucleotide second messenger (<https://www.ebi.ac.uk/QuickGO/term/GO:0007187>)  
GO:0032720 : negative regulation of tumor necrosis factor production  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0032720>)  
GO:0010739 : positive regulation of protein kinase A signaling  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0010739>)  
GO:0090037 : positive regulation of protein kinase C signaling  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0090037>)  
GO:0009650 : UV protection (<https://www.ebi.ac.uk/QuickGO/term/GO:0009650>)  
GO:0070914 : UV-damage excision repair  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0070914>)

#### GO - Cellular Component

GO:0005886 : plasma membrane (<https://www.ebi.ac.uk/QuickGO/term/GO:0005886>)  
GO:0005887 : integral component of plasma membrane  
(<https://www.ebi.ac.uk/QuickGO/term/GO:0005887>)

Presumptive Null

No ([https://www.gephebase.org/search-criteria?/and+Presumptive Null+No+Gephebase-summary-title](https://www.gephebase.org/search-criteria?/and+Presumptive+Null+No+Gephebase-summary-title))

Molecular Type

Cis-regulatory ([https://www.gephebase.org/search-criteria?/and+Molecular Type=Cis-regulatory+Gephebase-summary-title](https://www.gephebase.org/search-criteria?/and+Molecular+Type+Cis-regulatory+Gephebase-summary-title))

Aberration Type

SNP ([https://www.gephebase.org/search-criteria?/and+Aberration Type=SNP+Gephebase-summary-title](https://www.gephebase.org/search-criteria?/and+Aberration+Type+SNP+Gephebase-summary-title))

Molecular Details of the Mutation

single nucleotide variant within the MITF binding site of the canine MC1R promoter

Experimental Evidence

Association Mapping ([https://www.gephebase.org/search-criteria?/and+Experimental Evidence=Association Mapping+Gephebase-summary-title](https://www.gephebase.org/search-criteria?/and+Experimental+Evidence+Association+Mapping+Gephebase-summary-title))

Main Reference

Two MC1R loss-of-function alleles in cream-coloured Australian Cattle Dogs and white Huskies. (2018) (<https://pubmed.ncbi.nlm.nih.gov/29932470>)

Authors

DÄYarig N; Letko A; Lepori V; Hadji Rasouliha S; Loechel R; Kehl A; HytÄqinen MK; Lohi H; Mauri N; Dietrich J; Wiedmer M; DrÄqgemÄÿaller M; Jagannathan V; Schmutz SM; Leeb T

Abstract

Loss-of-function variants in the MC1R gene cause recessive red or yellow coat-colour phenotypes in many species. The canine MC1R:c.916C>T (p.Arg306Ter) variant is widespread and found in a homozygous state in many uniformly yellow- or red-coloured dogs. We investigated cream-coloured Australian Cattle Dogs whose coat colour could not be explained by this variant. A genome-wide association study with 10 cream and 123 red Australian Cattle Dogs confirmed that the cream locus indeed maps to MC1R. Whole-genome sequencing of cream dogs revealed a single nucleotide variant within the MITF binding site of the canine MC1R promoter. We propose to designate the mutant alleles at MC1R:c.916C>T as e and at the new promoter variant as e . Both alleles segregate in the Australian Cattle Dog breed. When we considered both alleles in combination, we observed perfect association between the MC1R genotypes and the cream coat colour phenotype in a cohort of 10 cases and 324 control dogs. Analysis of the MC1R transcript levels in an e/e compound heterozygous dog confirmed that the transcript levels of the e allele were markedly reduced with respect to the e allele. We further report another MC1R loss-of-function allele in Alaskan and Siberian Huskies caused by a 2-bp deletion in the coding sequence, MC1R:c.816\_817delCT. We propose to term this allele e . Huskies that carry two copies of MC1R loss-of-function alleles have a white coat colour.

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Additional References

## RELATED GEPHE

Related Genes

13 (Agouti (ASIP), GPR22, MFSD12, PMEL17, SLC45A2=MATP, FGF3; FGF4; FGF19; ORAOV1, Kit, Melanophilin (MLPH), Microphthalmia-associated transcription factor, PSMB7, tyrosinase-related protein 1 (TYRP1), beta-defensin 103 (CBD103), RALY (hnRNP associated with lethal yellow)) ([https://www.gephebase.org/search-criteria?/or+Taxon ID=9612+/and+Trait=Coloration/or+Taxon ID=9615+/and+Trait=Coloration/and+groupHaplotypes=true#Gephebase-summary-title](https://www.gephebase.org/search-criteria?/or+Taxon+ID+9612+/and+Trait=Coloration/or+Taxon+ID+9615+/and+Trait=Coloration/and+groupHaplotypes=true#Gephebase-summary-title))

Related Haplotypes

4 ([https://www.gephebase.org/search-criteria?/or+Gene Gephebase=MC1R+/and+Taxon ID=9612+/or+Gene Gephebase=MC1R+/and+Taxon ID=9615+#Gephebase-summary-title](https://www.gephebase.org/search-criteria?/or+Gene+Gephebase+MC1R+/and+Taxon+ID+9612+/or+Gene+Gephebase+MC1R+/and+Taxon+ID+9615+#Gephebase-summary-title))

## EXTERNAL LINKS

## COMMENTS

<https://omia.org/OMIA001199/9615/> @Parallelism