

DR. LESLIE LYONS Ph.D

Everything you need to know about Genetics...

You can learn from your Cat!

PART THREE

THE GENETIC UNIQUENESS OF BIRMANS AND HOW THEY GOT THEIR GLOVES!

College
of
Veterinary
Medicine,
University
of
Missouri

The Birman, also called 'The Sacred Cat of Burma,' is said to have origins from Burma, the current day Republic of the Union of Myanmar. Although Burma was a British colony during much of the 1800's up to post- World War II, which is during the time in which many cat breeds were developed, the term "Birman" is derived from the French word for Burma – "Birmanie".

GENETIC &/OR DEVELOPMENTAL ORIGINS

Several genetic studies have indicated cats from the Birman breed, consistently, genetically cluster with cats from Southeast Asia, including, Siamese, Korats, Burmese and random bred cats from the region (**Figure 1**)²⁻⁴.

The Birman breed history suggests their development was prominent in France with cross – breeding to Persians to obtain the long hair, particularly after the World Wars, which decimated the breed in Europe.

However the genetic crosses were accomplished, the Birman has retained an overall genetic signature from Southeast Asia (**Figure 1**).

Some breeds, with obvious Southeast Asian traits liked the bobbed tail, specifically the Japanese Bobtail, have not retained their genetic Southeast Asian origins²⁻⁵.

COAT LENGTH

For Birmans, the Siamese points, are clearly a Southeast Asia trait, however, Birmans have the same genetic variant for longhair in the gene *fibroblast growth* factor 5 (FGF5) as Persians, the c.475A>C variant, which is the oldest and most common variant for longhair in all cat breeds^{6,7}.

Other long-haired breeds, such as Ragdolls, Maine Coons and Norwegian Forest cats have the common variant, but also, each of these breeds have different longhair variants that many have been part of their early breed development.

Thus, although the Birman has an influence of a Near eastern trait, the longhair variant in Persians, their overall genetic make-up is mostly Southeast Asian.

The breed history also suggests the Birman breed had to be re-developed from only a few cats that survived the World Wars. Genetic studies also support this history as the Birman has low genetic diversity, only the well-established breeds of Burmese, Singapura and Havana Brown have lower genetic diversity.

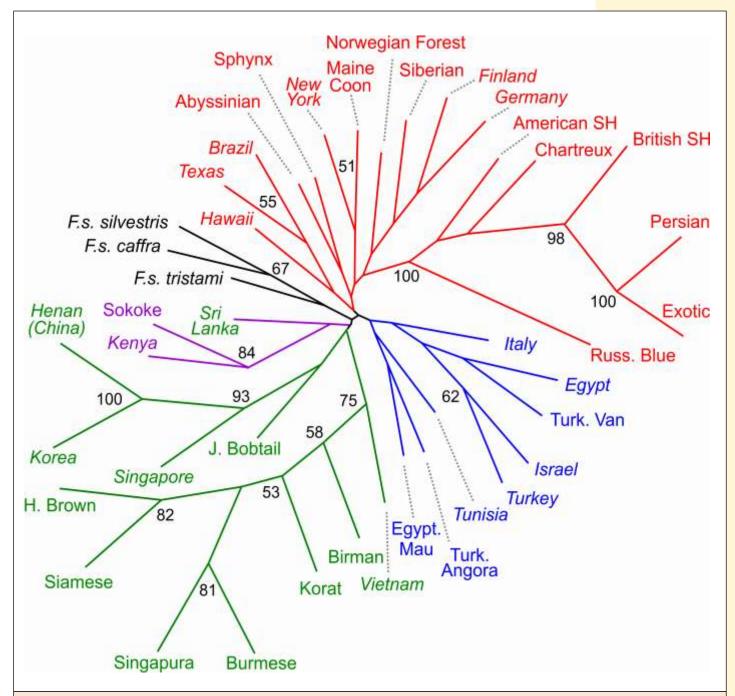


Figure 1. Genetic groupings of cat breeds. Breeds from the Southeastern Asian group together as do Mediterranean and Western European breeds. A number on the line gives some indication of the strength of the grouping. If no number is present, the support for the group is weak, such as with Japanese Bobtail with Southeast Asian cats. Birmans group without Southeast Asian breeds.

LINKAGE DISEQUILIBRIUM

Birmans also have extended "linkage disequilibrium" – more so than any other cat breed^{4,8}. Linkage disequilibrium implies large sections of the Birman genome are the same across the different individuals of the breed, implying, development of the breed is recent (in evolutionary times), or founder effects have been strong, or population bottlenecks have been severe, and or selection for specific traits has been intense.

Since, all Birmans must have points^{9,10}, longhair^{6,7} and gloves¹¹ – these are three traits that are under strong selection in the breed and together with the breed history, helped to explain the extended linkage disequilibrium in the breed.

Besides the long hair and the points, what makes a Birman a Birman is their white feet – their gloves. The white feet pattern was first suggested as an autosomal recessive

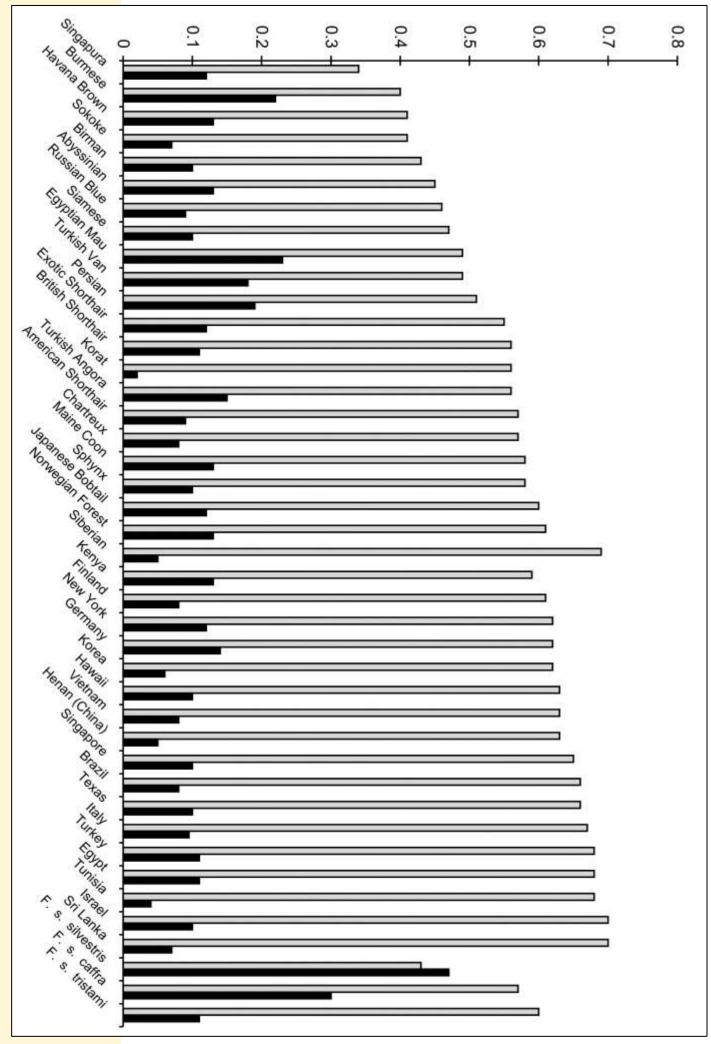


FIGURE 2.

THE GENETIC UNIQUENESS OF BIRMANS AND HOW THEY GOT THEIR GLOVES!

Figure 2. Cat breed genetic diversity. The white bars represent genetic diversity, the dark bars, inbreeding coefficient. A low white bar and a high black bar are indications of poor genetic health. The wild felid species, *Felis silvestris silvestris, caffra* and *tristrami*, had only a few specimens for comparison.

trait in 1991¹². More recently, genetic analyses of a gene called *KIT* (a.k.a. *KIT Proto-Oncogene, Receptor Tyrosine Kinase* or *V-Kit Hardy-Zuckerman 4 Feline Sarcoma Viral Oncogene*) has suggested two side-by-side amino acid changes in the protein are responsible for the gloves in Birmans, as an autosomal recessive allele¹¹

Several other species (horse, cattle, dog) have KIT mutations that influence white spotting as well¹³⁻¹⁵. Most all Birmans are "fixed' or homozygous for this variant, which is very rare to absent in other cat breeds.

Even the Ragdolls do not commonly have this DNA variation and this DNA variation does not predict mittens in Ragdolls. Thus, the Ragdoll breed did not "steal" the glove mutation from Birmans to produce their mitted varieties.

Since the variant is recessive, and because the variant may have been in the random bred population of cats from whence Birmans originated, the DNA variant can be found, but rarely, in other breeds and cat populations.

However, the KIT variant, c.1035_1036delinsCA, is defining for the Birman breed¹¹. Individuals of the breed that are not homozygous for the mutation are usually shown to have recent outcrossing in their ancestry.

BLOOD TYPE ISSUES

Interestingly, the B blood type is more common, but still rare, in different parts of the world, such as Turkey, California, and Australia and more common in British - derived breeds, such as British Shorthairs, Devon and Cornish rex¹⁶⁻¹⁸. Positive selection for the B blood type in itself is unlikely within a cat breeding program.

Cats have naturally occurring antibodies (alloantibodies). In type B cats, the anti-A antibodies are strong agglutinins and hemolysins that lead to the destruction of red blood cells. In contrast, anti-B antibodies in type A cats are weaker agglutinins and hemolysins, leading to less severe destruction of the red blood cells.

When breeding cats, neonatal isoerythrolysis can occur in kittens bearing the A or AB blood group antigen from a mating of B queens to an A or AB tom. Thus, Birman breeders must be vigilant of their cats' blood groups and prevent breedings that lead to neonatal isoerythrolysis and be prepared to intervene with the kitten's ingestion of colostrum within the first few days of life.

SUMMARY

Overall, Birmans have interesting origins and interesting genetics. Birman breeders are encouraged to share cats from around the world to diversify the breed, perhaps obtain some new imports from Southeast Asia if possible.

The blood type has to be closely managed. The available genetic tests allow Birman breeders to select cats with the long hair variants, gloves, points and blood type to assist the production of health, genetically diverse and beautiful cats.

This is the third
in a series
of articles
based on
feline genetics
by Dr Leslie Lyons
that will be
published in
FELIS HISTORICA
in the

ensuing months

EDITOR:

References

- 1. Hill A. *The Secrets of the Sacred Cat of Burma*. 2006 Trafford Publishing, UK. English translation of Poirier S, Barney G. *Les Secrets du Chat Sacre de Birmanie*.
- 2. Kurushima JD, et al. Variation of cats under domestication: genetic assignment of domestic cats to breeds and worldwide random-bred populations. Anim Genet. 2013 Jun;44(3):311-24. doi: 10.1111/age.12008. Epub 2012 Nov 22.
- 3. Lipinski MJ, et al. The ascent of cat breeds: genetic evaluations of breeds and worldwide random-bred populations. Genomics. 2008 Jan;91(1):12-21. doi: 10.1016/j.ygeno.2007.10.009. Epub 2007 Dec 3.
- 4. Gandolfi B, et al. Applications and efficiencies of the first cat 63K DNA array. Sci Rep. 2018 May 4;8(1):7024. doi: 10.1038/s41598-018-25438-0. Erratum in: Sci Rep. 2018 Jun 4;8(1):8746.
- 5. Lyons LA, *et al*. Whole genome sequencing in cats, identifies new models for blindness in AIPL1 and somite segmentation in HES7. BMC Genomics. 2016 Mar 31;17:265. doi: 10.1186/s12864-016-2595-4.
- 6. Drögemüller C, et al. Mutations within the FGF5 gene are associated with hair length in cats. Anim Genet. 2007 Jun;38(3):218-21. doi: 10.1111/j.1365-2052.2007.01590.x. Epub 2007 Apr 13.
- 7. Kehler JS, et al. Four independent mutations in the feline fibroblast growth factor 5 gene determine the long-haired phenotype in domestic cats. J Hered. 2007 Sep-Oct;98(6):555-66. doi: 10.1093/jhered/esm072. Epub 2007 Sep 1.
- 8. Alhaddad H, *et al.* PLoS One. 2013;8(1):e53537. doi: 10.1371/journal.pone.0053537. Epub 2013 Jan 7.
- Schmidt-Küntzel A, et al. Tyrosinase and tyrosinase related protein 1 alleles specify domestic cat coat color phenotypes of the albino and brown loci. J Hered. 2005 Jul-Aug;96(4):289-301. doi: 10.1093/jhered/esi066. Epub 2005 Apr 27.
- 10. Lyons LA, et al. Tyrosinase mutations associated with Siamese and Burmese patterns in the domestic cat (Felis catus). Anim Genet. 2005 Apr;36(2):119-26. doi: 10.1111/j.1365-2052.2005.01253.x.
- 11. Montague MJ, *et al.* Comparative analysis of the domestic cat genome reveals genetic signatures underlying feline biology and domestication. Proc Natl Acad Sci U S A. 2014 Dec 2;111(48):17230-5. doi: 10.1073/pnas.1410083111. Epub 2014 Nov 10.
- 12. Robinson R. *Genetics for Cat Breeders* 3rd Ed. 1991, Pergamon Press, UK.
- 13. Haase B, et al. Seven novel KIT mutations in horses with white coat colour phenotypes. Anim Genet. 2009 Oct;40(5):623-9. doi: 10.1111/j.1365-2052.2009.01893.x. Epub 2009 May 6.
- 14. Küttel L, et al. A complex structural variant at the KIT locus in cattle with the Pinzgauer spotting pattern. Anim Genet. 2019 Oct;50(5):423-429. doi: 10.1111/age.12821. Epub 2019 Jul 11.
- 15. Wong AK, et al. A de novo mutation in KIT causes white spotting in a subpopulation of German Shepherd dogs. Anim Genet. 2013 Jun;44(3):305-10. doi: 10.1111/age.12006. Epub 2012 Nov 8.
- 16. Bighignoli B, et al. Cytidine monophospho-N-acetylneuraminic acid hydroxylase (CMAH) mutations associated with the domestic cat AB blood group. BMC Genet. 2007 Jun 6;8:27. doi: 10.1186/1471-2156-8-27.
- 17. Giger U, Bucheler J, Patterson DF. Frequency and inheritance of A and B blood types in feline breeds of the United States. J Hered. 1991 Jan- Feb;82(1):15-20. doi: 10.1093/jhered/82.1.15.
- 18. https://eclinpath.com/hemostasis/transfusion-medicine/blood-types/

A BIRMAN BLAST FROM THE PAST!



MRS. WALLER, MRS. HACKMANN, AND MRS. LOHR - WITH THEIR BIRMANS. Mrs. Anneliese Hackmann is holding her Int. Ch. Nadine de Khlaramour.



MODERN DAY BIRMAN KITTENS

Photo: Helmi Flick Cat Photography