

Snakes reveal the origin of skin colours

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The skin of corn snakes (Pantherophis guttatus) has an orange base, decorated with red dorsal and lateral spots circled in black. The species can undergo mutations that lead to variations in skin colour, with the lavender corn snake being pink with grey spots. Credit: © UNIGE/ Milinkovitch

The skin color of vertebrates depends on chromatophores—cells found in the superficial layers of the epidermis. A team of specialists in genetic determinism and color evolution in reptiles from the University of Geneva (UNIGE) is studying the wide variety of colors sported by different individuals within the corn snake species. The research, published in the journal *PNAS*, demonstrates that the dull color of the lavender variant of corn snake is caused by the mutation of a gene



involved in forming lysosomes, the "garbage disposal" vesicles of cells. This single mutation is enough to affect every skin color, demonstrating that both the reflective crystals and pigments are stored in lysosomerelated vesicles. The UNIGE study marks a significant step forward in our understanding of the origin of colors and patterns in the skin of vertebrates.

The chromatophores are the cells that determine skin color, thanks to the presence of pigments or crystals that reflect light. There are three types of chromatophores: melanophores, which are responsible for the black or brown color; xanthophores, for red and yellow; and iridophores, with crystals that reflect multiple colors. Mammals only have melanophores, while reptiles and fish carry all three types of chromatophore, meaning they can display a very wide variety of colors and color patterns. The pigments of melanophores are known to be stored in organelles known as LROs or lysosome-related organelles. These are small intra- cellular vesicles that have the same origin as lysosomes, the "garbage disposals" that digest the non-functional molecules in cells. On the other hand, the storage location of the red and yellow pigments and crystals in the other types of chromatophore is unknown.

When snakes turn pink

The skin of <u>corn</u> snakes (Pantherophis guttatus) has an orange base, decorated with red dorsal and lateral spots circled in black. The species can undergo mutations that lead to variations in skin color, with the lavender corn snake being pink with gray spots. The experiments carried out by Athanasia Tzika, a researcher in the Department of Genetics and Evolution in UNIGE's Faculty of Sciences and her doctoral student Asier Ullate-Agote have identified that these altered colors are due to a single mutation pinpointed in the LYST gene, a gene that regulates lysosome trafficking. "It's very long-term work, since snakes only have one litter a year. We had to sequence the entire genome of the corn <u>snake</u> and



identify all the genes within," says Tzika

The liver is key

Mutations in the LYST gene in humans cause the Chediak-Higashi syndrome, which is characterized by albinism, an impaired immune system and an accumulation of enlarged lysosomes. The Geneva team continued its study into corn snakes by analyzing their hepatocytes, the main liver cells in vertebrates, which contain numerous lysosomes. The scientists found that the hepatocytes of lavender corn snakes contain much larger and more aggregated lysosomes. Using electron microscopy, the authors observed that the morphology and arrangement of colored vesicles in all the chromatophores were altered.

The result of evolution

Michel Milinkovitch, a professor in UNIGE's Department of Genetics and Evolution, explains further: "By characterizing the mutant gene, the study has shown for the first time that the different chromatophores were not created from scratch during evolution but that they all entail a basic mechanism involving LROs."

Further studies will provide a better understanding of the mechanisms responsible for the extraordinary variety of skin colors and color patterns in vertebrates, features that play a part in functions as diverse and essential as camouflage, intraspecific communication, and protection against the harmful effects of solar radiation.

More information: Asier Ullate-Agote el al., Genome mapping of a LYST mutation in corn snakes indicates that vertebrate chromatophore vesicles are lysosome-related organelles, *PNAS* (2020). www.pnas.org/cgi/doi/10.1073/pnas.2003724117



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