

Two genes explain variation in color and behavior in the wall lizard

March 1 2019



A common wall lizard individual exhibiting orange coloration. The presence of orange is explained by variation in the SPR gene, which is a gene in a highly conserved pathway that is linked to the production of neurotransmitters like serotonin and dopamine. Credit: Guillem Pérez i de Lanuza



How are reptiles capable of generating such a diversity of bright colors? And how is it possible that within a single population of the same species, different individuals exhibit strikingly different coloration patterns? In a new paper published in the journal *PNAS* an international team of scientists, led by researchers from CIBIO/InBIO (University of Porto) and Uppsala University, reveal two genes implicated in yellow to red pigmentation in reptiles, and demonstrate that these pigmentation genes also affect behavior and other traits.

The study focused on the European wall lizard, which is common across most of Southern Europe.

"While for a casual observer it may seem like an ordinary lizard, with its camouflaged back blending perfectly with rocky surroundings, in most populations of this species the belly and throat show a remarkable diversity in color, being either white, bright orange/red or bright yellow. Strikingly, lizards of different colors differ in other characteristics, such as size, fighting ability and reproductive strategies. They also preferentially mate with individuals of their own color," said Pedro Andrade, Ph.D. student at the University of Porto and lead author of the paper.

The researchers found that the genomes of differently colored lizards are virtually identical – except for very specific portions of the DNA sequence. "In nature, when individuals from different populations or species differ in multiple characteristics, the expectation is that many genes are involved. So, for us it was a big surprise to see that in these lizards, the color differences and associated characteristics are largely explained by small regions of the genome that contain only two genes," said senior author Miguel Carneiro from CIBIO, University of Porto.

Orange/red hues are explained by the SPR gene, a vital gene in the pterin synthesis pathway, while yellow hues are explained by the BCO2 gene,



which is involved in the metabolism of carotenoids. "The characterization of the molecular basis of the color variation is an important step because it allows us to understand how it has originated and exactly which metabolic pathways are involved," said Catarina Pinho, researcher from the University of Porto and co-first author of the work. "There are previous studies showing a relationship between BCO2 and yellow coloration in other organisms, namely birds, and it is interesting that the same gene is being recruited for the same role in different branches of the animal tree of life."

The gene SPR is of particular interest. The metabolic byproducts of this gene's activity are closely linked to the production of neurotransmitters like serotonin and dopamine. These are chemicals that, also in humans, are responsible for the transmission of information between neurons.

"Our results implicate that these pigmentation <u>genes</u> lead to other sideeffects, such as potential differences in the functioning of the brain that could explain why lizards of different color behave differently or have slightly different habitat choices," said Professor Leif Andersson from Uppsala University, another senior author of the paper.

The team also found that the DNA sequences associated with different colors were extremely divergent. The divergence in DNA sequence was big enough to go beyond what one would expect to find between species. The differences were up to six times larger than what is normally found when comparing human and chimpanzee DNA. Further DNA analyzes showed that this genetic variation is shared with multiple lizard species that occur in the Mediterranean. The likely explanation for this pattern is that the DNA sequences associated with coloration are millions of years old and that some species have been exchanging them through natural hybridization. This highlights how hybridization is an important mechanism for the fast acquisition of new characteristics – even humans are an example of this, as shown by multiple evidence of hybridization



between modern and archaic humans like Neanderthals.

This international team of researchers has an ongoing research program focused on how color variation and associated characteristics arise and are maintained. Ongoing work leverages techniques and expertise in the fields of genome sciences, animal behavior, and neurobiology and aims to help decipher other interesting aspects of the biology of the color variation and what exactly the different colors are signaling to <u>lizards</u>, which is something that remains a mystery.

More information: Pedro Andrade et al. Regulatory changes in pterin and carotenoid genes underlie balanced color polymorphisms in the wall lizard, *Proceedings of the National Academy of Sciences* (2019). DOI: 10.1073/pnas.1820320116

Provided by Uppsala University

Citation: Two genes explain variation in color and behavior in the wall lizard (2019, March 1) retrieved 25 April 2024 from https://phys.org/news/2019-03-genes-variation-behavior-wall-lizard.html

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