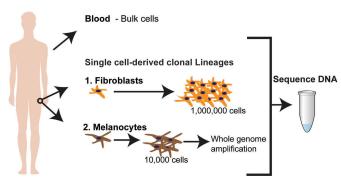


Even skin shielded from the sun accumulates genomic DNA changes from UV light

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Schematics of the study design. From each donor, we obtained blood for whole-genome sequencing. In addition, we obtained skin biopsies from the hips of the donors from which fibroblasts and melanocyte clonal lineages were obtained. Fibroblasts were grown up to a million cells and their DNA was directly used for whole-genome sequencing, while melanocytes grew up to 10,000 cells and the DNA was whole-genome amplified and thereafter sequenced. Credit: Saini N et al., 2021, *PLOS Genetics*

For the first time, scientists have measured the different types of genomic DNA changes that occur in skin cells, finding that mutations from ultraviolet (UV) light is especially common, but Black individuals have lower levels of UV damage compared to white people. Dmitry Gordenin and colleagues at the National Institute of Environmental Health Sciences, report these findings January 14 in *PLOS Genetics*.

The DNA in our skin cells suffer damage from sources inside and outside the body, leading to genomic changes such as mutations that may lead to cancer. UV light is the major source of these mutations, but byproducts of cellular metabolism, like <u>free radicals</u>, and DNA copying errors that occur during <u>cell division</u> also cause genomic

changes. These mutation-causing mechanisms are well known, but previously, no one had been able to accurately measure the relative contributions from each source.

In their new paper, Gordenin and his colleagues quantified the amounts of each type of genomic changes by sequencing the genomes of skin cells donated from 21 Black and white individuals, ranging in age from 25 to 79. The researchers discovered that the total amount of genomic changes from metabolic byproducts accumulates as a person gets older, while the amount of genomic changes caused by UV damage is unrelated to a person's age. Additionally, they showed that genomic changes from UV light is common, even in skin cells typically shielded from the sun, but it was less prevalent in Black donors compared to white donors.

The researchers suspect that Black individuals may be better protected from UV light due to having higher levels of the skin pigment melanin.

Supporting this idea, is the fact that Black people have much lower rates of skin cancer compared to white people. Overall, the new study provides an accurate estimate of the genomic changes that occur in skin cells due to different types of DNA damage, and establishes the normal range of somatic genomic changes across a wide range of ages and of different races, providing a baseline for future research.

The authors add, "The new study provides an accurate estimate of the genomic changes that occur in skin cells due to different types of DNA damage, and establishes the normal range of somatic genomic changes across a wide range of ages and of different races, providing a baseline for future research."



More information: Saini N, Giacobone CK, Klimczak LJ, Papas BN, Burkholder AB, Li J-L, et al. (2021) UV-exposure, endogenous DNA damage, and DNA replication errors shape the spectra of genome changes in human skin. *PLoS Genet* 17(1): e1009302. journals.plos.org/plosgenetics ...

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