New Parkinson's Genetic Risk Factor Discovered in African Populations

Editor's Note: In August 2023, GP2 scientists published this finding in The Lancet Neurology. Read the paper at bit.ly/gba1.

The GP2 study — that BLAAC PD contributes to — found a gene change in some people with Parkinson's disease. The change is in the *GBA1* gene.

But what is the *GBA1* gene? The *GBA1* gene tells the body how to make a protein called GCase. The GCase protein plays a role in the cell's "recycling center." This part of a cell removes damaged or faulty cell parts. Changes in the *GBA1* gene can reduce GCase activity, which means the cell's recycling center doesn't work correctly. Without that function, damaged or faulty cell parts can build up, harming the cell. Scientists believe this build-up may lead to the cell death that causes Parkinson's symptoms in people with a *GBA1* change, but more research is needed.



NOVEL RISK VARIANT rs3115534

When GBA1 is mutated:

- Misfolded alpha-synuclein ("the Parkinson's protein") clumps together
- Lysosomes
 (the cell's recycling centers) do not work properly
- Mitochondria (the cell's powerhouse) are damaged
- Stress occurs in part of the cell that manages how cells communicate

This change in the *GBA1* gene was found in people with African ancestry. The finding is from genetic information shared by BLAAC PD participants like you and from another study in Nigeria. While having this variant may increase risk of Parkinson's, it does not guarantee disease.

As a result of this finding, we are starting to understand causes of Parkinson's in people of African descent.

As scientists learn more about this gene change, there may be other findings. Some gene changes may be linked to faster or slower progression or to certain symptoms. That information could help doctors monitor disease more closely. It may help you make lifestyle choices as well. You may decide to exercise more or change your diet.

This finding also shows the importance of partnering with people of many backgrounds.

Most genetics studies have been in people of European descent. This finding shows scientists have more to learn by working with diverse groups.

There are treatments currently in testing to correct *GBA1* and GCase function in Parkinson's disease. The scientists who found the gene change in people of African descent are talking to the researchers testing these treatments. They may decide to open their trials to test for this new *GBA1* gene change.

Right now, your doctor cannot tell you if you have this variant. There are tests that share

information on other changes in the *GBA1* gene. (Not all genetic tests look for all variants.)

Scientists can now learn more about this newly discovered *GBA1* variant. Soon it may be added to those existing *GBA1* tests.

In the future, you may be able to help test new treatments.

Learn more about this finding and Parkinson's genetics at michaeljfox.org.

BLAAC PD on the Map

Eleven sites across the United States are collecting valuable data and samples from volunteers like you.

BLAAC PD Site Locations:

Alabama – Birmingham

Florida — Gainesville

Illinois - Chicago (2 sites)

Louisiana — Shreveport and New Orleans (2 sites)

Maryland – Baltimore

Missouri - St. Louis

Ohio - Cleveland

South Carolina - Charleston

Texas - Houston

BLAAC PD is part of GP2. GP2 is gathering genetic information from groups with different backgrounds. Learn more at gp2.org.

