African iron overload

African iron overload is a condition that involves absorption of too much iron from the diet. The excess iron is stored in the body's tissues and organs, particularly the liver, bone marrow, and spleen. Humans cannot increase the excretion of iron, although some iron is lost through bleeding or when cells of the intestine (enterocytes) are shed at the end of the cells' lifespan. Iron levels in the body are primarily regulated through control of how much iron is absorbed from the diet.

African iron overload results from a diet high in iron. It is particularly associated with consumption of a traditional African beer that contains dissolved iron from the metal drums in which it is brewed. Some evidence suggests that a genetic predisposition to absorbing too much iron may also be involved.

In African iron overload, excess iron typically accumulates primarily in certain immune cells called reticuloendothelial cells. Reticuloendothelial cells include macrophages in the bone marrow and spleen and Kupffer cells, which are specialized macrophages found in the liver that help protect the body against foreign invaders such as viruses and bacteria. Later in the course of the condition, iron also accumulates in liver cells (hepatocytes). This pattern differs from that seen in a similar iron overload disorder called hereditary hemochromatosis, in which the excess iron accumulates primarily in the hepatocytes.

When too much iron is absorbed, the resulting iron overload can eventually damage tissues and organs. Iron overload in the liver can lead to chronic liver disease (cirrhosis). Cirrhosis increases the risk of developing a type of liver cancer called hepatocellular carcinoma. Iron overload in immune cells may affect their ability to fight infections. African iron overload is associated with an increased risk of developing infections such as tuberculosis. The excess iron also leads to a faster-than-normal breakdown of vitamin C in the body, so affected individuals are at increased risk of vitamin C deficiency problems such as scurvy.

People with African iron overload may have a slightly low number of red blood cells (mild anemia), possibly because the iron that accumulates in the liver, bone marrow, and spleen is less available for production of red blood cells. Affected individuals also have high levels of a protein called ferritin in their blood, which can be detected with a blood test. Ferritin stores and releases iron in cells, and cells produce more ferritin in response to excess amounts of iron.

Frequency

African iron overload is common in rural areas of central and southern Africa; up to 10 percent of the population in these regions may be affected. Men seem to be affected more often than women, possibly due to some combination of differences in dietary iron
consumption and women's shedding of excess iron through blood loss in menstruation and childbirth.

The prevalence of increased iron stores in people of African descent in other parts of the world is unknown; however, these individuals may be at higher risk of developing mildly increased iron stores than are people of European background.

Causes

African iron overload was first noted in rural central and southern African populations among people who drink a traditional beer brewed in uncoated steel drums that allow iron (a component of steel) to leach into the beer. However, not all individuals who drink the beer develop African iron overload, and not all individuals of African descent with iron overload drink the beer. Therefore, researchers are seeking genetic differences that affect the risk of developing this condition.

Some studies have indicated that a variation in the \textit{SLC40A1} gene increases the risk of developing increased iron stores in people of African descent. This variation is found in 5 to 20 percent of people of African descent but is not generally found in other populations.

The \textit{SLC40A1} gene provides instructions for making a protein called ferroportin. This protein is involved in the process of iron absorption in the body. Iron from the diet is absorbed through the walls of the small intestine. Ferroportin then transports iron from the small intestine into the bloodstream, and the iron is carried by the blood to the tissues and organs of the body. Ferroportin also transports iron out of reticuloendothelial cells in the liver, spleen, and bone marrow. The amount of iron absorbed by the body depends on the amount of iron stored and released from intestinal cells and macrophages.

The \textit{SLC40A1} gene variation that some studies have associated with increased iron stores in people of African descent may affect the way ferroportin helps to regulate iron absorption in the body. However, researchers suggest that this variation is not associated with most cases of African iron overload.

Inheritance Pattern

African iron overload seems to run in families, and high iron in a family's diet seems to be the major contributor to development of the condition. There also may be a genetic contribution, but the inheritance pattern is unknown. People with a specific variation in the \textit{SLC40A1} gene may inherit an increased risk of this condition, but not the condition itself. Not all people with this condition have the variation in the gene, and not all people with the variation will develop the disorder.
Other Names for This Condition

- African hemochromatosis
- African nutritional hemochromatosis
- African siderosis

Diagnosis & Management

Genetic Testing Information

- What is genetic testing?
  /primer/testing/genetictesting
- Genetic Testing Registry: African nutritional hemochromatosis

Research Studies from ClinicalTrials.gov

- ClinicalTrials.gov
  https://clinicaltrials.gov/ct2/results?cond=%22African+siderosis%22+OR+%22Siderosis%22

Additional Information & Resources

Health Information from MedlinePlus

- Encyclopedia: Hepatocellular Carcinoma
  https://medlineplus.gov/ency/article/000280.htm
- Health Topic: Cirrhosis
  https://medlineplus.gov/cirrhosis.html
- Health Topic: Iron
  https://medlineplus.gov/iron.html

Additional NIH Resources

- Dietary Supplement Fact Sheet: Iron

Patient Support and Advocacy Resources

- American Liver Foundation
  https://liverfoundation.org/

Scientific Articles on PubMed

- PubMed
  https://www.ncbi.nlm.nih.gov/pubmed?term=%28%28african+siderosis%29+OR+%28african+iron+overload%29+AND+english%5Bla%5D+AND+human%20%5Bmh%5D+AND+%22last+3600+days%22+AND+5Bdp%5D
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http://omim.org/entry/601195

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