What will this study involve?

THIS RESEARCH STUDY INVOLVES UP TO A MAXIMUM OF TWO VISITS TO THE STUDY SITE.

At the first visit, before any study procedures start, the study doctor's staff will describe this study in detail. If you wish for your child to participate, you must then sign a document called an Informed Consent Form. Your child may be asked to sign an Assent Form (depending on their age/ability) to indicate that they understand what will happen, including any potential risks and benefits of participating.

You will speak with the study doctor about your child's medical history and then, the doctor's staff will take a small blood sample.

This blood sample will be sent to a laboratory for testing to see whether there is a particular sign of AADC deficiency in the blood (abnormally high levels of 3-O-methyldopa (3-OMD)). If no sign is found, the study doctor will inform you and your child of this, and their participation in the study will be complete.

If a sign of AADC deficiency is found in the first blood sample, the study doctor will invite you and your child to return to the study site where a second larger blood sample will then be taken. Genetic testing will also be performed to see if AADC deficiency can be confirmed.

You and your child will be informed about the results of all the tests done using your child's blood samples by your child's study doctor.

What are the next steps?

If you think participating in this research study might be right for your child, please talk with your child's doctor. Then, please contact us as directed below. Contacting us does not obligate your family in any way to participate in this study.

FOR MORE INFORMATION ABOUT THIS RESEARCH STUDY, PLEASE

 MED-UK-AADC-2000007
EE/AADC/UK/20/0035d | Date of preparation: August 2020 UK Version 08-28-20-v2

WHEN YOUR CHILD HAS CLINICAL FEATURES SUGGESTIVE OF CEREBRAL PALSY THAT GO UNEXPLAINED, A RARE GENETIC DISORDER MAY BE THE CAUSE.

If your child has symptoms of weak muscle tone (hypotonia), painful muscle stiffness, uncontrollable movement of the hands and legs, low blood pressure (hypotension), fainting, low blood sugar (hypoglycaemia), impaired sweating, abnormal weight gain, inability to regulate body temperature, vision problems (droopy eyelids and pupil constriction), developmental delay without any clinical or neuroradiological (MRI brain) evidence of cerebral palsy, please consider our research study which may help identify a cause.

REVEAL-CP™

WELCOME and thank you for your interest in the REVEAL-CP™ Study.

This research study is for people of any age who have symptoms of weak muscle tone (hypotonia), painful muscle stiffness, uncontrollable movement of the hands and legs, low blood pressure (hypotension), fainting, low blood sugar (hypoglycaemia), impaired sweating, abnormal weight gain, inability to regulate body temperature, vision problems (droopy eyelids and pupil constriction), developmental delay without any clinical or neuroradiological (MRI brain) evidence of cerebral palsy. This includes but is not limited to ~~patients~~ with a current or previous diagnosis of cerebral palsy.

The purpose of this study is to learn how many ~~patients~~ with symptoms that look like cerebral palsy, but with no known reason for these symptoms, may actually have a rare genetic condition called Aromatic L-amino Acid Decarboxylase (AADC) deficiency.

Participants will receive diagnostic testing to determine whether their symptoms are caused by AADC deficiency. This diagnostic testing, which may include genetic analysis, is conducted through blood testing. This study does not involve any medicines or treatments.

What is AADC deficiency?

AADC deficiency is a rare genetic disorder that affects the brain, causes abnormal muscle tone, and affects how a person develops. AADC deficiency may be the reason for a child's symptoms.

AADC deficiency is very rare. Because it is so rare, and because the symptoms are similar to symptoms of other ~~diseases~~, including cerebral palsy, diagnosis can be difficult.

AADC deficiency is a disorder that interferes with the way the cells in the nervous system talk to each other through substances called neurotransmitters. In AADC deficiency, a genetic mutation (change within a gene) leads to a decrease in the amount of neurotransmitters made.

There is no cure for AADC deficiency, but there are some medicines that doctors use to help treat the symptoms of AADC deficiency.

What are the symptoms of AADC deficiency?

A child with AADC deficiency may have some or all of the following symptoms. Many of the symptoms of AADC deficiency are similar to cerebral palsy symptoms, and every child is different, so your child's symptoms may vary.

Some of the most common symptoms of AADC deficiency are:

- Low or abnormal muscle tone
- Abnormal movements, especially involuntary eye movements*
- Delays in development. A child with AADC deficiency may be unable to:
  - Lift and control his or her head
  - Crawl, sit, or stand without support
  - Walk
  - Babble or say words

*Children with AADC deficiency often have movement problems, including involuntary eye movements (called oculogyric crises). These are moments when a child's eyes suddenly roll upward involuntarily.

Who can take part in the REVEAL-CP™ Study?

Your child may be eligible to take part in this research study if they:

- Have previous or current clinical features suggestive of cerebral palsy with no underlying cause found; this includes but is not limited to ~~patients~~ with a current or previous diagnosis of cerebral palsy
- Are not taking carbidopa-levodopa

There are also additional requirements to participate. The staff at the study site will explain the other requirements in more detail.

WHAT ARE THE POTENTIAL BENEFITS OF PARTICIPATING?

If your child participates, you and your child will learn whether or not your child has AADC deficiency. We expect that only a small percentage of ~~patients~~ taking part in this study will have AADC deficiency.

PTC Therapeutics, Inc., the company sponsoring this study, hopes to learn more about both AADC deficiency symptoms and conditions that present like AADC deficiency.

DOES IT COST ANYTHING TO PARTICIPATE?

You or your child will not receive any payment for taking part in this study and you will not have to pay for the study visits or for the procedures needed for your child to take part.

All reasonable study-related travel expenses will be reimbursed to and from your home address and to your local research site, including but not limited to receipts for personal car fuel, train/bus tickets, taxi fares, and parking.

ARE THERE ANY RISKS TO PARTICIPATING IN THIS RESEARCH STUDY?

If your child participates in this study, it is possible that they may experience some discomfort during the blood samples. The study doctor's staff can discuss any concerns you may have in more detail.