

SIGNAL

A Phase 2, multi-center, randomized, double-blind, placebo controlled study in subjects with late prodromal and early manifest Huntington disease (HD) to assess the safety, tolerability, pharmacokinetics, and efficacy of VX15/2503

Clinical Trial Status:

This study is not currently enrolling

Clinical Trial Principal Investigator:

Dr. James Boyd

Clinical Trial Protocol Description:

In this study, researchers are looking at a monoclonal antibody, VX15/2503, as a potential treatment for HD. This monoclonal antibody is a class of drug that binds to a molecule, and may block it from causing inflammation in the brain of those with HD. It may specifically protect against the inflammation that has been shown to affect thinking, movement and behaviors in those with HD, as it has been shown to do so in animal models.

The main objective of the SIGNAL study is to evaluate the safety and tolerability of monthly intravenous (IV) administration of an investigational product (IP), VX15/2503, as compared to placebo. In this study there are two cohorts; Cohort A receives the IP for 12 months, in which the first 6 months are blinded (it is unknown if the subject is receiving placebo or the active drug) and the last 6 months they receive the active drug, VX15/2503. Cohort B receives blinded IP for 18 months, which means for these subjects they do not know (nor does the investigator) if they are receiving the active drug, VX15/2503 or the placebo for the 18 month treatment period.

Secondary objectives of this study are:

- How does the body's immune system respond to the study drug
 - Does the study drug affect brain tissue size, as seen on MRI (magnetic resonance imaging) brain scans
- Does the study drug affect brain activity as seen on FDG-PET (fluorodeoxyglucose-positron emission tomography) and TSPO-PET (Translocator Protein-positron emission tomography)
- Does the study drug affect thinking, movements, and/or behaviors
- How does the body take in, distribute, and remove the study drug from its system (pharmacokinetics)
- Does the study drug affect biomarkers found in the blood

Individuals who have tested positive for the HD gene and either 1) have early signs of HD but have not been given a clinical diagnosis of HD by a physician, or 2) have early signs of HD and a physician has determined that they meet the criteria for a clinical diagnosis of HD, are invited to participate. There will be an initial screening visit to determine if a patient is eligible to participate in the study.

Clinical Trial Eligibility Criteria:

In order to participate you must meet the following criteria:

- Have undergone genetic testing with a known CAG repeat greater than or equal to 36
- Fulfill one of the following criteria:

- Late prodromal HD (based on a CAP score)
- Early manifest HD (based on a functional assessment and clinical diagnosis of HD)
- Are at least 21 years of age
- Are taking stable doses of your medications
- If female, you must be of non-childbearing potential, or practice a highly effective method of contraception (e.g. oral contraceptives, barrier methods of birth control, intrauterine devices, partner with vasectomy or sexual abstinence)
- If male, you must agree to use a reliable method of birth control during the course of the study
- If female, you must be using a reliable form of contraception, or be surgically sterile, or be postmenopausal

You will not be eligible to participate if any of the following criteria apply to you:

- Have had previous neurosurgery for HD or another movement disorder
- Have participated in an investigational drug or device study within 30 days of baseline
- Have a history of suicidal ideation within 60 days of baseline, or are a suicide risk, as judge by the Investigator
- Have a clinically significant laboratory or ECG abnormalities
- Have a history of infection with human immunodeficiency virus (HIV), hepatitis C, and/or hepatitis B
- Have a history of substance abuse within the last 12 months
- Are pregnant or breastfeeding
- Have had a known allergy to any ingredient in the study drug
- Have a condition which would exclude MRI participation (e.g. pacemaker, severe claustrophobia, metal fragments in the eyes or body)

Note: this is a partial list of the inclusion and exclusion criteria

Clinical Research Coordinator:

Emily Houston

(802) 656-8974

Emily.houston@med.uvm.edu

Please contact the CRC with any questions