

## WHAT IS THE NATURAL HISTORY STUDY OF CTNNB1?

This natural history study being led by Principal Investigators Dr. Wendy Chung and her team of clinicians from Columbia University Irving Medical Center is an opportunity for patients and families from the CTNNB1 community to advance clinical understanding of CTNNB1 disorder and bring us closer to treatment. Through participation in clinical assessments, CTNNB1 patients will provide a greater insight into how symptoms develop and change over time across various ages. Participation also ensures that the spectrum of phenotypes is represented across the entire range of pathogenic mutations in the CTNNB1 gene.

The natural history study also prepares the CTNNB1 community for clinical trials. Clinical trials require validated endpoints and outcome measures to reliably assess outcomes when evaluating the effectiveness of potential therapeutics. This study ensures that meaningful endpoints are identified for measurement in future clinical trials and that the most appropriate tools for assessing these outcomes are established.

Additionally, we hope to improve the standard of care for families facing CTNNB1 Disorder and treating medical communities.

## WHAT DOES THE NATURAL HISTORY STUDY INVOLVE?

The study involves clinical assessments to investigate multiple domains of CTNNB1 disorder, including motor function, cognition, nervous system issues, and the brain's electrical activity. Assessment will be conducted by a specialist. The visit includes:

Motor Function Assessment (Physical Therapy)

Neurological Exam

Research EEG

Neurocognitive Assessment

Blood Sample Collection

Photos and videos to capture physical changes over time

Eligibility to participate in each assessment will be determined by the study team based on what the participant can comfortably engage in. Each eligible assessment will be optional and can be withdrawn from at any time.

Participation in the natural history study of CTNNB1 does not constitute a clinical appointment for medical advice. You are welcome to discuss medical history and issues with the clinicians, but all components of a full clinical appointment are not assessed during the clinical research evaluations and therefore it may not be possible for the clinicians to answer specific medical questions.