



**A user guide to the clinical datasets and biosamples  
available from Enroll-HD**

**Version 2015-01-R1**

**Enroll-HD**

**A worldwide observational study for Huntington's disease families**

**A CHDI Foundation Project**

**Access to Enroll-HD Data and Biosamples – Summary Guide**

Enroll-HD is a worldwide longitudinal observational study of Huntington's disease (HD) - this summary guide provides an overview of the policies and the accompanying documentation relevant to gaining access to and using the clinical data and biosamples from the study.

The overarching goal of Enroll HD is to accelerate progress towards the development of effective therapeutics for HD. To do this the study has three main aims:

### **1. To provide a platform to support the design and conduct of clinical trials**

#### Objectives:

- Providing a resource to identify, develop and qualify novel assessment tools, clinical endpoints and biomarkers
- Collecting longitudinal data to inform disease modeling studies, and
- Facilitating the identification of potential trial participants informing the selection of potential trial participants using data to estimate and quantify slopes/rates of disease progression (providing “run-in” data).

### **2. To improve the understanding of the dynamic phenotypic spectrum and the disease mechanisms of HD**

#### Objectives:

- Collecting natural history data covering the cognitive, behavioral and motor domains permitting estimates of rates of progression in HD and allowing insights into the neurobiology of HD,
- Collecting data and biologic samples to identify genetic and environmental factors influencing and/or modifying the HD
- Phenotype and disease progression
- Promoting interrogatory studies that may provide clues to the pathogenesis of HD.

### **3. To promote the development of evidence-based guidelines to inform clinical decision making and improve health outcomes for the participant/family unit**

#### Objectives:

- Assisting in the identification of beneficial interventions (clinical, pharmaco-therapeutic, non-pharmacologic)
- Facilitating the dissemination and implementation of currently proposed best clinical practices
- Providing a platform for conducting outcome research
- Promoting exploratory data analysis projects that may identify processes to further improve the healthcare of affected individuals and their families.

To achieve these goals, Enroll-HD was designed to ensure that as many researchers as possible worldwide could access the clinical data and renewable biosamples in a rapid, transparent, straightforward manner, lowering barriers to entry into HD research while safeguarding the identity of research participants. Data included in the released datasets are de-identified and coded, which means the data has gone through a process of removing identifying information that could connect a participant's identity to the information.

The Enroll-HD website hosts periodic datasets that will be made available to any researcher that is employed at a recognized research organization—from academia, government institutions, nonprofit organizations, or public and private companies; a researcher gains access to the periodic datasets by undergoing a straightforward process to verify that they are employed by a recognized research organization, setting up a user account, and executing the requisite data use agreement.

All researchers can download the periodic dataset(s). To encourage collaboration among researchers, researchers are asked to provide a brief description (50-200 words maximum) of their proposed project—this is automatically posted on the Enroll HD website along with the researchers name and affiliation.

## **DATA RELEASE – PERIODIC and SPECIFIED DATASETS**

Enroll-HD clinical data consists of longitudinal data from participants' annual visits from countries around the world. Each participant has completed a baseline visit and a set of general forms. At least one on-site monitoring visit has been completed for each participant in the dataset.

**Periodic datasets** are time-stamped cuts of the total Enroll-HD clinical dataset. These periodic datasets are available for download, along with accompanying documentation to aid in your use

of the data. The first public data release from Enroll-HD provides all data that met the approved criteria for release as of January 1, 2015 (11am GMT). Additional datasets will be made available; the current expectation is at approximately 6-monthly intervals.

Researchers may also request **specified datasets** that include Enroll-HD data from participants not included or contain non-standard data fields, or data in different file formats. Such requests require completion of the Specific Dataset Request Form.—Requests for specific data will be referred to the Scientific Publication Review Committee for prioritization, feasibility, and approval and will take longer to process.

## **MONITORING OF DATA**

Enroll-HD utilizes risk-based monitoring (RBM) employing a number of approaches: Electronic Data Capture (EDC) system edit checks, remote and onsite data review, medical monitoring and centralized statistical monitoring. Defined triggers are activated during the monitoring process, signaling the need for increased monitoring to mitigate increased risk. Using this strategy and combining several linked risk-based approaches in sequence improves monitoring efficiency and effectiveness. Edit checks help to identify potentially erroneous data or data entry mistakes before data are reviewed by the project team. Remote and onsite monitoring occurs sequentially (at the participant level) while medical and statistical monitoring of aggregated data and listings occur on a rolling basis.

## **DATA INCLUSION CRITERIA**

Not all the data in the Enroll-HD database are included in the periodic datasets; all of the following criteria must be met for inclusion:

- 1) Participants must have properly signed informed consent forms (ICF) for their data to be included in the released datasets. This means that each participant's ICF has been verified at the research site as part of the monitoring process.
- 2) The dataset only includes participants that are enrolled in "Enroll-HD".
- 3) The dataset only includes data from the Enroll-HD study.
- 4) Participants that do not have a research CAG determination are not included.

- 5) Only properly coded terms entered for pharmacologic therapies, indications for pharmacologic therapies, comorbid conditions, or reported events are included.
- 6) A participant is selected for the dataset if a “Baseline” visit has been performed with onsite review completed, a “General” visit with remote review completed and at least one annual additional visit (“Follow-up”, “Unscheduled Visit”, or “Phone Contact”) with remote review completed within the last 15 months prior to the planned date of the data cut. The additional visit criteria only applies for participants who are enrolled more than 15 months since date of the dataset. Examples for the selection of participants for a potential data cut 1/2015:
  - A participant with a Baseline & General visit done in 9/2014 is included.
  - A participant with a Baseline & General visit done in 10/2013 and no additional visits is not included.
  - A participant with a Baseline & General visit done in 9/2013 and a Follow Up in 8/2014 is included.
  - A participant with a Baseline & General visit done in 6/2012 and a Follow Up in 9/2013 and no additional visits is not included.
  - A participant with a Baseline & General visit done in 6/2012 and a Phone Contact in 5/2014 is included.
- 7) The dataset shall only include visits that are remotely reviewed at least once.
- 8) Participants who have discontinued from the study prematurely must have an End of Study form.

If any of the above criteria are not met then a participant’s data has not been included in this periodic dataset release. The Enroll-HD Operations Team continually monitors these parameters to maximize the amount of data that can be published with the dataset release.

#### **EXCLUDED DATA**

Certain data were aggregated or withheld from the periodic dataset because the data were deemed to increase the risk of potentially identifying individual participants or families, especially when combined with the broader dataset. If there are excluded data that you are interested in, please

complete a Specific Dataset Request form. This form is located on the Enroll-HD website ([www.Enroll-HD.org](http://www.Enroll-HD.org)).

### **BIOMATERIALS ACCESS POLICY**

Researchers can also request Enroll-HD biosamples by completing a request form and signing a Biosamples Use Agreement that are available on the Enroll-HD website. Currently there is no online catalog – to enquire about the type of biosamples available contact [biosamples@enroll-hd.org](mailto:biosamples@enroll-hd.org).

Biosamples require completion of the Biosamples Request Form and will take additional time to process; non-renewable biosample requests will also require Scientific Publication Review Committee approval.

### **PUBLICATION POLICY**

Many people contribute to creating and making these valuable resources available to researchers. These include CHDI Foundation, clinicians, site coordination staff, raters, a world-wide operations team, EDC programmers, biorepository personnel and scientists, assessment developers, statisticians, researchers and, most importantly, the Enroll-HD participants and their families and caregivers. In appreciation of all their efforts we request that each researcher adheres to the Enroll-HD publication policy when publishing work using this information (See Enroll-HD Publication Policy).

### **ACCOMPANYING DOCUMENTS FOR DATASETS**

There are several documents that are provided to help you with understanding and using the Enroll-HD resources. These documents are grouped as follows: 1) General Documents, 2) Study Detail Documents, and 3) Analysis Aides. While we hope these documents are comprehensive, there may be additional detail you need. For all questions and inquires please email us at [AccountSetup@Enroll-HD.org](mailto:AccountSetup@Enroll-HD.org).

#### General Documents

- 1.0 User Guide: Provides a summary of the study aims and objectives, how to access the data and biosamples, the quality control process used for the data and reference to the Publication Policy that should be adhered to by Researchers.
- 2.0 Data Handling Manual: Detailed description about how to handle the periodic dataset (for example, exporting, transforming and converting data files, handling missing and/or aggregated data values).
- 3.0 Publication Policy: Describes the full Publication Policy that should be adhered to by Researchers.
- 4.0 Enroll-HD ISCED: Supporting documents providing reference tables for each country's ISCED levels (education codes) mapping onto the respective national education system

#### Request Forms

- 5.0 Biosample Request Form: A form completed by Researchers who would like to access Enroll-HD biosamples. Details about the quantity and type of biosamples required should be given, along with a detailed project description.
- 6.0 Specific Dataset Request Form: A form completed by Researchers who would like to access specific datasets that are not included in the Periodic Dataset. Details about specific variables and the quantity of data should be given, along with a detailed project description.

#### Agreements

- 7.0 Data Use Agreement: Template of the Data Use Agreement that must be executed ahead of access to the dataset(s).
- 8.0 Biosamples Use Agreement: Template of the Material Transfer Agreement that must be executed before biosamples can be shipped to researchers.

#### Current Periodic Dataset Documents

- 9.0 Enroll-HD Protocol: Full description of the study aims, population, design, assessment protocol and procedures.
- 10.0 Data Dictionary: Describes all the data files and its variables used in the periodic dataset.

- 11.0 Enroll-HD Unusual Findings: Provides a summary of cases reported in the dataset that have unusual findings.
- 12.0 Quality Control Process: Detailed description of the quality control and de-identification procedures that have been applied to the periodic dataset.
- 13.0 Annotated CRF: Brochure of the Enroll-HD case report forms to illustrate the data capture for each data field captured in the periodic dataset. Includes the data variable names which can be cross-referenced to the Data Dictionary.
- 14.0 Coverage Chart: A description of the dataset available and number and type of biosamples.

**Revision History**

Document Name	Summary of Changes
Version 2015-01-R1	Initial version for first Enroll-HD Periodic Dataset