PPMI Recruitment Strategies: Genetic Cohort & Registry

Genetic Kick-off Meeting
September 17, 2013
Prodromal PPMI: Genetic At-risk Cohort

- Ultimate goal of PPMI is to follow individuals who are unaffected but will develop PD
Prodromal Cohort: Hyposmia and RBD
Genetic Cohort: LRRK2+/SNCA+

Objective #1: Fill the funnel

Objective #2: Convert qualified leads to enroll

Goal: Enroll subjects with potential to phenoconvert to PD
Recruitment and the GCC

- Screen to identify LRRK2+/SNCA+ individuals
- Review previous molecular testing to confirm LRRK2+/SNCA+

Pool of individuals eligible for PPMI

Pool of family members eligible for PPMI
Identifying individuals for the genetic cohort

- PD patients reporting a LRRK2/SNC A mutation
- Unaffected individuals reporting a LRRK2/SNC A mutation

Review testing results

- PD patients at ↑ risk for LRRK2/SNCA (fam hx of PD, AJ dissent)
- Unaffected individuals at ↑ risk for LRRK2/SNCA (fam hx of PD, AJ dissent)

Consent for testing
Local site outreach to LRRK2+/SNCA+ individuals

Clinical Sites
- Identify LRRK2+ and SNCA+ PD and unaffected individuals within clinic population
- Hold a seminar to discuss and invite LRRK2+/SNCA+ individuals and their family members to participate in the study
- Set an appointment with LRRK2+/SNCA+ individuals to review family history in detail
- Provide brochures for LRRK2+/SNCA+ to provide to family members

Medical Community Outreach
- In regions in which testing occurs broadly among physicians (eg France), host a meeting with local MDs to discuss the study and encourage referral
- For individuals that have provided samples, but do not know their gene status, offer them the opportunity to learn their status through participation in this study
- Provide feedback to referring MDs regarding ultimate enrollment of their patient and status in the study

Review genetic testing results and provide to Genetics core for approval
Local site outreach to genetic at-risk populations

Local Community Outreach

- Connect with key individuals in Jewish communities to:
  - Arrange for local seminar/educational sessions re LRRK2+
  - Place study flyers/brochures in synagogues and Jewish community centers
  - Newsletter stories/information about LRRK2+/SNCA+ PD connection and study participation

Clinic Outreach

- Reach out to other neurologists and movement disorder specialists in your area to engage them in identifying individuals in their practice at risk based on family hx of PD and family background
- Provide info about PD genetic risk factors and study in institution newsletter/websites

Complete genetic testing and counseling for those willing
Centralized Outreach via MJFF and FTF to Individuals with Genetic Risk

MJFF Outreach and Communications

- Key talking point in all constituents presentations and events
- Newsletter (e-newsletter and paper)
- Social Media
- Online Ads
- MJFF Website
- National and local media

Fox Trial Finder

- Messages to individuals with LRRK2+/SNCA+ individuals to share trial opportunity with family members

Complete genetic testing and counseling
Genetic Cohort Recruitment Materials

- Brochure for PD/At-risk LRRK2+ individuals to distribute to family members
- Brochure to provide to individuals of Ashkenazi Jewish dissent with PD relative regarding LRRK2 screening
- MD pocket cards to distribute to community physicians as a reminder regarding the study and basic criteria

- Language for newsletter articles/blurbs to be provided to
  - local synagogues and Jewish Community Centers
  - local center/institutional newsletters

- PPMI Study Update Newsletter Articles
  - Short and long newsletter stories that can be shared with local PD groups and included in your own center publications about the progress of the study
Genetic At-risk Recruitment
Additional Key Points

• Think broadly regarding screening procedures – this will likely be site specific

• Engage individuals in at-risk populations to serve as ambassadors

• There will be additional funding available for travel for LRRK2+ and SNCA+ subjects – if travel is the issue, please ask
  – Subjects in PPMI have been willing to travel longer distances to participate than we projected

• Once a LRRK2+/SNCA+ individual is identified, be diligent about getting detailed family history...even if they do not ultimately participate, other family members may
Breaking Apart the Process

Identifying Related Individuals
Recruiting Across the Age and Disease Spectrum

• PD patients at any stage of disease to identify those with a LRRK2/SNCA mutation
  – Some can complete a more extensive protocol (Genetic Cohort)
  – Some cannot due to disease progression (Genetic Registry)
• Recruit family members of the LRRK2+/SNCA+ proband
  – Some family members may carry the mutation
  – Some family members will not carry the mutation
• Mutation carrier
  – Some too young to be expected to develop disease soon
  – Some at a critical age and more extensive protocol desired
Identifying Related Individuals in PPMI

• We want family members to participate in PPMI
  – Need a way to identify people as being from the same family
  – Family members may go to different sites
  – Family members may not mention that they have another family member in the study
Importance of Relatedness

• Many kinds of analyses will be performed using data from PPMI
  – Some analyses require (assume) subjects are independent (unrelated)
  – GCC can provide information to indicate how PPMI subjects are related
  – DNA studies later will confirm the reported relationships (and identify more)
Family History

- IU will have IRB approval to obtain family history information (with names) from study subjects
  - Subjects will be given a family history packet at their first PPMI study visit
  - Each packet is labeled with a packet ID
  - Subjects will complete the questionnaire and return all documents, including IU IC, directly to GCC
Family History Packet

Site gives subject Family History Packet

Site completes Family History Packet Distribution Fax and faxes to GCC

Subject takes home:
- IU Family History Substudy IC
- IU Family History Substudy HIPAA
- IU Family History Questionnaire
Family History Packet

Subject takes home and completes, Mails to GCC

Site completes and faxes to GCC

**PPMI Family History Substudy**

**FAMILY HISTORY QUESTIONNAIRE (FHQ)**

**Indiana University Informed Consent Statement for Parkinson’s Progression Marker Initiative (PPMI) – Family History Substudy**

**INTRODUCTION**

You are being asked to participate in the Family History substudy of the “Parkinson’s Progression Markers Initiative” (PPMI). Please read this form carefully. Ask the person presenting this form any questions that you have before making a decision about whether or not you will participate.
Linking Subject to Packet

• Site records on the Family Packet Distribution fax the PPMI Subject ID of the subject receiving this packet

• Site faxes the Family Packet Distribution fax to GCC

• **Now, PPMI Subject ID is linked to the Family History Packet ID**
Linking Family Members

- Subject sends the completed Family History Questionnaire, IU IC and HIPAA directly to GCC
- GCC will recognize related individuals through common family members
- All related individuals will receive the same family number, providing a link between them
5. Does the subject have a first degree relative (father, mother, sibling, child) who is also participating in the study? (0 = No, 1 = Yes)
Family Recruitment

Proband
Family Recruitment

Proband

Potentially recruit everyone in orange into PPMI
How to Discuss Testing Results with Family members

• MJFF has led the development of a new brochure
• Given to a PPMI subject with a genetic mutation (LRRK2 or SNCA)
• They can use this brochure as a way to discuss PPMI and their test results with other family members
Dear Friend,

In the years since my diagnosis, the love and support of my family have been vital to living well with Parkinson's disease. Today, the roles of all families supporting Parkinson's research is more crucial than ever.

Studying the genetics of Parkinson's disease could revolutionize the development of new treatments for patients worldwide. Families connected to people who carry genetic mutations of Parkinson's play a unique role in the pursuit of a cure.

A decision to get genetic testing in PPMI is an opportunity to join forces with thousands of families worldwide committed to speeding scientific progress toward cures for diseases that touch countless lives.

We're all in this together. I hope you'll consider joining our movement.

All my best,

Michael J. Fox

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Who is Eligible to Participate in PPMI?

Parents, children, brothers and sisters of an individual who carry a LRRK2 mutation are invited to receive genetic testing free of charge through the PPMI study. PPMI seeks volunteers both with and without Parkinson's disease. People who do not have Parkinson's disease must be 50 years or older to participate.

PPMI Sites Worldwide

Clinical sites in the following locations offer genetic counseling and testing for the LRRK2 gene free of charge.

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Play a Part in Parkinson's Research. Participate in PPMI.
888.PPMI or www.michaeljfox.org/PPMI/genetics.

BE PART OF THE GENETICS REVOLUTION: THE ROLE OF FAMILIES IN PARKINSON'S DISEASE RESEARCH

PARKINSON'S PROGRESSION MARKERS INITIATIVE
Play a Part in Parkinson's Research
Family Members and PPMI

• PPMI site is contacted by an individual who identifies themselves as a relative of a PPMI subject
• PPMI site schedules an appointment for the individual to come to the site and learn about PPMI
Family Member of a LRRK2/SNCA+ (No previous testing)

1. Identify individuals at ↑ risk for LRRK2/SNCA mutation
2. Individual signs Consent for PPMI Molecular Testing (PD or Unaffected)
3. Blood drawn and sent for testing with Laboratory Requisition Form
4. Enter Genetic Mutation Testing form into eClinical /FAX copy to GCC
Family Members

• Recruiting family members is critical
  – They provide the greatest opportunity to enroll individuals who carry a mutation but are not affected

• We want to be sure we maximize all the potential for family members for each and every PPMI study subject