

# Workgroup on International Clinical Trials Update

## Translation and adaptation of Clinician Rated Outcomes

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# Background

- Large International clinical trials are the current norm for CNS drug development
- A scientific session focused on this occurred at a previous ISCTM meeting
- Based on this , and the issues raised, a workgroup was initiated to examine issues associated with this phenomenon
- Chaired by Amir Kalali and Richard Keefe
- Other core members Elizabeth Pappadopulos and Monika Vance
- As with all ISCTM workgroups, the goal is to focus on one specific issue at a time
- Many topics were considered to be examined and the workgroup members were polled to prioritize one initial topic
- E.g. See tomorrow's talk on regional differences in placebo response

# Initial Topic for Focus

- Increased access to international subjects may threaten study integrity because of the complexities of cross-cultural research and assessment
- While there are numerous guidelines across therapeutic areas for the translation and adaptation of patient reported outcomes (PROs; eg., QOL), there is a dearth of recommendations for clinician reported outcome measures (CROs)
- Based on discussion, consensus and survey findings the ISCTM workgroup on International Clinical Trials has launched an initiative to develop practical guidelines for the translation and adaptation of Clinician Rated Outcome measures to improve signal detection in CNS global trials

# Method and Progress Update

- ✓ Four workgroups reaching consensus on our objective and scope of work
- ✓ Completion of a survey to identify current practices, knowledge gaps, and needs
- ✓ Review of existing PRO guidelines
- ✓ Workgroup endorsement of key objectives
- ✓ Commitment by individuals to contribute areas of expertise in manuscript development
- Guideline development and workgroup endorsement
- Publication

# Key Guideline Objectives

- Develop practical guidelines for Clinician Rated Outcome measures
- Present a range of recommendations that weigh benefits against cost/time
- Provide basic primer for individuals without expertise linguistic translation/adaptation to support knowledgeable decision-making

# Survey

- Sent to ISCTM membership with several reminders
- 78/442 responses (18%); 70 useable survey entries
- 44.7% had no formal coursework in Psychometrics
- >50% of respondent reported deep or moderate expertise in neuroscience, clinical trial design, statistics and tests and measurement

# Key Insights from Survey Results

- Greatest threats to Clinician Rated Outcome measure integrity:
  - scales that focus on symptoms behaviors that vary across cultures
  - item misunderstanding
  - item content that is not discussed openly in that culture
- Three most important reasons for selecting a measure: Use in other trials for indication, psychometric properties, validation in other languages/cultures
- CRO translation difficulties recognized only after the trial is complete, and contributes to the recruitment of inappropriate subjects

# Next steps

- Complete first draft of guidelines and gain workgroup endorsement for content
- Finalize of complete guidelines and gain endorsement
- Submit for publication