Roadmap to **PATIENT-FOCUSED OUTCOME MEASUREMENT** in Clinical Trials

1. **Understanding the Disease or Condition**
   - **A. Natural history of the disease or condition**
     - Onset/Duration/Resolution
     - Diagnosis
     - Pathophysiology
     - Range of manifestations
   - **B. Patient subpopulations**
     - By severity
     - By onset
     - By comorbidities
     - By phenotype
   - **C. Health care environment**
     - Treatment alternatives
     - Clinical care standards
     - Health care system perspective
   - **D. Patient/caregiver perspectives**
     - Definition of treatment benefit
     - Benefit-risk tradeoffs
     - Impact of disease

2. **Conceptualizing Treatment Benefit**
   - **A. Identify concept(s) of interest (COI) for meaningful treatment benefit, i.e., How a patient:**
     - Survives
     - Feels (e.g., symptoms)
     - Functions
   - **B. Define context of use (COU) for clinical trial:**
     - Disease/Condition entry criteria
     - Clinical trial design
     - Endpoint positioning
   - **C. Select clinical outcome assessment (COA) type:**
     - Patient-Reported Outcome (PRO)
     - Observer-Reported Outcome (ObsRO)
     - Clinician-Reported Outcome (ClinRO)
     - Performance Outcome (motor, sensory, cognition)

3. **Selecting/Developing the Outcome Measure**
   - **A. Search for existing COA measuring COI in COU:**
     - Measure exists
     - Measure exists but needs to be modified
     - No measure exists
     - Measure under development
   - **B. Begin COA development**
     - Document content validity (qualitative or mixed methods research)
     - Evaluate cross-sectional measurement properties (reliability and construct validity)
     - Create user manual
     - Consider submitting to FDA for COA qualification for use in exploratory studies
   - **C. Complete COA development:**
     - Document longitudinal measurement properties (construct validity, ability to detect change)
     - Document guidelines for interpretation of treatment benefit and relationship to claim
     - Update user manual
     - Submit to FDA for COA qualification as effectiveness endpoint to support claims