**UTOPIAN Primary Care Trials Group – Session 4 Minutes**
Wednesday, May 27th, 2020 from 4:00 p.m. to 5:00 p.m., Zoom teleconference

**Attendance:**
- Andrew Pinto (AP) – Chair
- Aashka Bhatt (AB)
- Noah Crampton (NC)
- Giles Pereira (GP)
- Marjan Moeinedin (MM)
- Rahim Moineddin (RM)
- Braden Gregory O’Neill (BGO)
- Michelle Greiver (MG)
- Eva Grunfeld (EG)
- Ross Upshur (RU)
- Donatus Mutasingwa (DM)
- Carolyn Steele Gray (CSG)
- Sumette Kalia (SK)
- Ann Burchell (AB)
- Sheila Dunn (SD)
- Rosemarie Lall (RL)
- Sumeet Kalia (SK)
- Tony D’Urzo (DU)
- Peter Selby (PS)
- Chris Meaney (CM)
- Joanne King (JK)
- Jennifer Rayner (JR)

**Regrets:**
- Payal Agarwal (PA)
- Aisha Lofters (AL)
- Abhimanyu Sud (AS)

<table>
<thead>
<tr>
<th>Item</th>
<th>Topic</th>
<th>Minutes</th>
<th>Action</th>
<th>Responsible</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>Introductions (Andrew Pinto)</td>
<td>• Andrew Pinto introduced those present on the phone.</td>
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<td>2</td>
<td>Review and approval of April 30, 2020 draft meeting minutes (All)</td>
<td>• Minutes of the previous meeting were approved by those present.</td>
<td>Approved</td>
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| 3    | Learning topic: Adaptive Trial Design (Dr. Ross Upshur) | • Clinical trials, their design and analysis are constantly evolving field  
• Structure of RCT: Standard architecture:  
  o Inception Cohort (sampling, inclusion and exclusion criteria)  
  o Method of Randomization (allocation to Treatment and Control groups)  
  o Observer groups over time and measure the outcomes (relevant issue is the difference between the Treatment and Control Groups) | | |
• Theory of RCT:
  o Randomization balances known and unknown co-variates, such that we have an un-biased estimator (outcome measure) that tells us the difference between the Treatment and Control
  o We are trying to set up a form of inference on the basis of the data

• Adaptive Designs for Clinical Studies:
  1. Model-based/Continual Assessment Designs
  2. Group Sequential/Sample-Size Re-Estimation Designs
  3. Group Sequential/Response Adaptive Designs
  4. Adaptive Randomization Designs

• Characteristics of Adaptive Designs:
  o Streamlined
  o Flexible
  o Optimized
  o Data-driven
  o Systematic
  o Decision-oriented
  o Validity
  o Integrity
  o Bayesian
  o Simulation
  o Real-time
  o Robust
  o Cost-efficient
  o Sequential learning
  o Dynamic

• 8 Common Types of Adaptations
  o Stopping early (or late, i.e. extending accrual) with a conclusion of superiority or futility
  o Adaptively assigning doses to more efficiently assess the dose-outcome relations
  o Adding or dropping arms or doses (perpetual motion machines)
  o Seamless phases of drug development within a single trial
  o Changing the proportion of patients randomized to each arm
  o Adaptively identifying in on an indication or responder population
  o Changing accrual rate
  o *Many of these 'types' can be set up in advance (i.e. planned adaptations)

• Adaptive Trials Components:
  o Interim Analysis: Frequent
  o Randomization: Variable
  o Number of Arms: Few to many
Discussion of trial proposals and ongoing work (All)

- Aashka is maintaining a list of all the COVID-related studies happening in the DFCM and across UTOPIAN sites. In addition, we are also maintaining a list of investigators who are connected to different sites and the different trials they are connected to.
  - The advantage of keeping track of this information, is that when trial ideas emerge from our work, we can quickly link these ideas with sites and investigators.
- 13 COVID-19 trials ongoing at DFCM and UTOPIAN sites (2 funded, 11 pending funding)
  - We will continue to update this list, and share it with this group in a frequent communication
- We have secured an email: covid.trials@utoronto.ca
- Clinical Trials Bootcamp:
  - A series of sessions that will run at lunchtime over a two-week span during the summer (similar to a summer institute model)
  - Will cover the basics of trials
  - We will be seeking people to present

Meeting adjourned at 5:00 p.m.
Next meeting: Thursday, June 18, 2020; 4:00 p.m.-5:00 p.m. (virtual)