Guideline for obtaining 180-day follow-up

There are four outcomes to be obtained at follow-up 180 days after randomisation:

- Mortality
- Cognitive function
  - MoCA 5 minute/Telephone (”Mini MoCA”)
- Health-Related Quality of Life (HRQoL)
  - EQ-5D-5L index values
  - EQ VAS

Retention strategies

- The 180-day follow-up is conducted per domain. Follow-up data for cognitive function and HRQoL should be collected as close to day 180 as possible but may be collected within a 14-day period (in both directions) and the obtained values may be reused across domains if the follow-ups occur within a 14-day period of day 180 for other domains, as specified in the core protocol. However, mortality is registered according to the status on exactly day 180 after randomisation in each domain, with values reused if randomisation occurred on the same date.
- When obtaining participant consent, please assure that the participants are informed about the 180-day follow up and that we will ***contact them by telephone.***
- Similarly, inform the relatives about the half-year follow-up when obtaining their consent.
- If the participants cannot be reached at 180-day follow-up, aim to contact the participant daily for 2 weeks. It is also okay to use days beyond the 2 weeks, if it is realistic obtaining the follow-up – the most important thing is to get the answers.
- If the participant is assumed to be at work, then try to make contact during late afternoon.
- If the participant chooses to respond electronically, it should be via secure electronic post.
- If the participant chooses physical mail, the time for obtaining answers is extended by another 4 weeks, with a reminder 2 weeks after the letter has been posted. The date to entered in the EasyRF is the date the questionnaire is fully completed. ***Remark: the cognitive function can only be assessed by telephone.***

We cannot contact any participants who rejected to be contacted or who withdrew with a wish for no further data registration.

Procedure at follow-up:

- The trial follow-up will be conducted by the site's trial staff using a procedure aiming at reducing risk of bias. This means that trial staff should not review the allocations before doing the follow-up.
- Staff should **not access participants' electronic health records** at 180-day follow-up, except to assess whether the patient is alive or not.
- Access EasyRF at 180 days follow-up using either of these log-in options:
    - If the **participant is dead** – use your standard access for EasyRF and register death as an event including the date of death. Then the follow-up forms will disappear and cannot be completed.
    - If the **participant is alive** – use your 180 days follow-up log-in for EasyRF
- Allocation should not be discussed with the participant (or their proxy) before the Mini MoCA and EQ-5D-5L + EQ VAS has been completed.
- **Please read the instructions** for both questionnaires beforehand, to obtain as objective answers as possible. To ensure a consistent approach to follow-up, cognitive function should be assessed first to reduce fatigue from the HRQoL questionnaire.
- It is important to register whether HRQoL is obtained by proxy.
- If by proxy it is important that the answer is given on behalf of what proxy believes the patient would have answered.

1. Ensure **participant is alive and has consented** before making contact
2. Try to contact the participant for a 4 week period (+/-2 weeks from day 180)
3. Ask the participant to sit in an undisturbed room **without a clock or calendar**.
4. 1st obtain Mini MoCA
5. 2nd obtain EQ-5D-5L + EQ VAS
6. Not obtainable -> register reason in eCRF

Follow-up data are entered directly into the EasyRF, which is the source data.

Cognitive function is assessed with a 4-item cognitive test. The MoCA 5 minute/Telephone is a shorter version than the full MoCA. The Mini MoCA takes 5 minutes, and you will need the instructions (short guide) together with the test. Staff using **Mini *MoCA needs to be certified*** in the full MoCA. Mini MoCA can only be assessed by telephone or face to face and not by mail and not by a proxy (e.g. a relative).

HRQoL is assessed with the EQ-5D-5L. It is a 5-item questionnaire together with 5 response levels for each item and a VAS scale score (0-100); it takes 5 minutes to complete. We recommend conducting the interview by phone, but it is also possible to answer the questionnaire by physical mail. If the participant is too frail or cognitively impaired to consistently complete EQ-5D-5L, try to obtain the participant's HRQoL by proxy (a relative) *asking the proxy to respond from the participant's point of view*. There is specific questionnaire for proxies assessing the HRQoL from the participant's point of view.

Mini MoCA certification: This is done here: https://www.mocatest.org/training-certification/ and takes 1 hour. The full MoCA instructions are very helpful. The certificate is valid for two years and is charged USD 125, which will be refunded. E-mail an invoice to contact@incept.dk.

Resources for each 180-day follow-up you conduct. The payment for conducting follow-up is outlined in the document 'Case money for participating in INCEPT' placed in the Site File #16.1.1.

Contact details: contact@incept.dk or phone +45 3545 7450.