



Minutes of CRIC Scientific Steering Committee Meeting

Date: November 3rd, 2017

Meeting time: 12am-3pm **Lunch:** Sandwich is served at 11.30-12.00 am

Place: Forskningsenheden 7831, 3rd floor, Tagensvej 22, 2100 Copenhagen N (same building as usual)

Invited members: Bodil Steen Rasmussen (BSR), John A. Myburgh (JAM), Ville Pettilä (VP), Thorbjørn Grøfte (TG), Ingrid Egerod (IE), Jakob Kjellberg (JK), Jan Bonde (JB), Christian Gluud (CG), Morten H Bestle (MHB), Hans-Henrik Bülow (HHB), Robert Winding (RW), Theis Lange (TL), Helle L. Nibro (HN), Morten H. Møller (MHM), Jens Winther (WIN), Lone Musaeus Poulsen (LMP), Jørn Wetterslev (WET), Anders Perner (AP).

Invited non-members: Lars Winther, Innovationsfonden (LW), Birgit Agerholm Larsen, CRIC project manager (BAL)

Non-attendees: JAM, VP, TG, JB, HN, WIN

Minutes according to agenda

1. Welcome (AP) and presentation (all)
 - a. A welcome to all attendees.
2. Challenges 2017/2018 (AP)
 - a. GCP monitoring in UK – assumed fraud with timesheets
 - i. Comment: Danish sites experience inconsistency in the requests from GCP between sites and regions
 - b. ICU patients may experience to be considered and categorized unconscious in one trial and conscious in another trial.
 - c. Funding from Medicinpuljen to HOT-ICU and AID-ICU, while awaiting decisions from Novo Nordisk Foundation (in December) (three applications - from Køge, Aalborg and Rigshospitalet).
 - d. Financing – is still an issue in general, and AP may have succeeded to put clinical trials on the political agenda, but it remains to shown if total funding available for trials will be increased. An appearance for ‘Sundhedsudvalget’ in the Parliament



- planned for last week has been postponed. Post meeting note: RE-scheduled for Nov 14th, 2017
- e. Different attempts under consideration for financing of the CRIC office beyond 2019.
3. SUP-ICU (MHM – short resume)
 - a. SUP-ICU inclusion has closed as of October 22nd, 2017 – 90 days follow-up and 1-year follow-up may remain.
 - b. Short summary of the status from www.sup-icu.com front page.
 - c. A separate account sheet has been made for SUP-ICU expenses only – we estimate total to be around 11 mill DKK
 - d. Authorship/co-authorship for SUP-ICU – suggestion to follow the protocol for the first publication, and to offer an authorship for the 1-year follow-up publication for sites with 25 or more included patients in excess of the 50 patients cut-off, or just in total if the 50 patients cut-off was never reached. This was decided.
 - i. Authorship restrictions from publishers were discussed in general. Post meeting note. The NEJM has no max. on the no. of authors.
 - e. Comment: A lot of e-mails encouraging to step up for SUP-ICU have been experienced as overwhelming at some sites – expiration of medicine soon has definitely been the cause of the pushing e-mails. Suggestion to remember to highlight accomplishments from both small and large sites, i.e. milestones passed.
 4. HOT-ICU (BSR – short resume)
 - a. Status on active sites in DK (Aalborg, Køge, RH4131, Randers)
 - b. Dialog with foreign sites – Switzerland, Norway, Finland (negative), Sweden (inconclusive due to remaining legal issues with the consent procedure in general), UK – Cardiff sites involved, on hold for the remaining part of the UK, which may have a similar trial in UK.
 - c. Comment: What if? – just to consider a plan B should it be necessary
 - d. Acknowledgement of the post.doc. funding in HOT-ICU.
 5. AID-ICU (LMP – short resume)
 - a. Protocol - only a few minor questions from LMS to reply to before approval.
 - b. Expected start-up in Køge will be March 2018, and other Danish sites will follow in May 2018
 - c. Dialog with foreign sites – Spain, Finland and Italy
 - d. Overlap between trials may be an issue
 6. CTU (WET – short resume and post meeting notes)
 - a. Overview of reviews on pharmacological prevention and management of delirium in ICU patients
 - i. Protocol published December 2016 in *Systematic Reviews*
 - ii. Preliminary report made June 2016
 - iii. *To be done: Co-authors will be forwarded the 1st draft review within a couple of weeks. We aim to submit the final report (to BMJ or BMJ Open) before Christmas 2016.*
 - b. Systematic review on haloperidol

- i. Protocol finished, author group will be sent the final version for feedback next week.
 - ii. Protocol will be submitted to Acta Anaesthesiologica Scandinavica
 - iii. *To be done: update searches, data extraction, analysis, write report*
 - c. Systematic review on oxygen for ICU patients
 - i. Preliminary report made Jan 2017
 - ii. Cochrane protocol published April 2017
 - iii. *To be done: Do final (all) analysis, write report, we aim to submit the report before Christmas 2017.*
 - d. Systematic review on oxygen for critically ill patients
 - i. Protocol registered on Prospero
 - ii. Searches made and run, screening and data extraction done
 - iii. *To be done: compare data extraction, do analysis, write report (+ update searches)*
 - e. Systematic review on stress ulcer prophylaxis for ICU patients
 - i. A meeting will be held Nov 2017 where the content/deadline dates for the review will be discussed. We aim to submit the final paper ultimo March 2018 (as closest as possible to the publication of the SUP-ICU RCT).
 - ii. *To be done: write protocol, run searches, screen and data extract, do analysis, write report.*
 - f. A methodological study investigating clinical heterogeneity versus statistical heterogeneity for interventions in the intensive care patients.
 - i. *To be done: write protocol, arrange stay in Holland, data extraction and analysis, write report.*
- 7. Biostatistics (TL – short resume)
 - a. Axel Jensen has received a grant and will not be available for the projects after Summer 2018
 - b. We are working on a new method to increase statistical power when analyzing outcomes not normally distributed and of the type "Days alive outside ICU/hospital within 90 days" used in all our trials.
- 8. VIVE (KORA) (JK – short resume)
 - a. Challenges in the new organization VIVE but with no major impact on the project.
 - b. Christine has signed-in as a PhD student.
- 9. Opening new research programs (5-10 min each)
 - a. Classic II has applied for Novo Nordisk Foundation funding.
- 10. CRIC Conference (Q1) 2018
 - a. Postponed until we know if 'Landstingsalen' is a possibility
- 11. CRIC (AP/BAL)
 - a. Fundraising (BAL)
 - i. Awaiting decisions from Novo Nordisk Foundation re. HOT-ICU, AID-ICU and CLASSIC, which will be the 4th clinical program in CRIC, if granted
 - b. Budget/account (BAL)
 - i. SUP-ICU is on track – awaiting case-money and GCP expenses but the budget should have room for both.

- ii. CRIC account is on track – have excess of money on the account planned to be spent on AID and HOT, so we show compliance with the expenses in the budget, but the timeline is postponed.
- c. Conflict of interest
 - i. Post meeting note: Unfortunately, it was not mentioned at the meeting, but it is time to renew the Conflict of Interest Form – it will be forwarded to all members of the SSC.

12. Next meeting

- a. Next SSC meeting is to be held April 19th, 2018 at 11.30am – 2.30pm
(lunch/sandwich will be served at 11.00 am)
- b. Next All Stakeholder meeting is to be held April 19th, 2018 at 9.00am-11.00am.

Referent / BAL