SMART Trial

ETHICAL APPROVAL AND PROTOCOL AMENDMENTS

Ethical Approval: Granted by NRES Committee South Central – Southampton B on 28th September 2011.

1. Substantial Amendment. Date: 13/10/2011.
   Accepted by REC 18/10/2011

1. Changing the definition of a procedure tract metastasis to a 7cm margin from the procedure site (protocol sections 4.3.2, 6.1)
2. Changes to the Inclusion/Exclusion Criteria
   a. Changing exclusion criteria 7 to exclude patients with chest wall nodules within 7cm of the procedure site. (IRAS form sections A6-2, A17-2; protocol sections 2.2; 3.3)
   b. Clarification that drains inserted by either a seldinger technique or blunt dissection can be enrolled (IRAS form sections A6-2, A17-1; protocol sections 2.2, 3.2)
3. Changing the minimisation criteria to include surgical procedure or other and removing plan for chemotherapy. (IRAS form section A61; protocol sections 2.2, 4.2.2)
4. Changes to the secondary endpoints
   a. Adding health economic analysis to the research questions and secondary endpoints. (IRAS form sections A11, A58; protocol sections 2.1; 6.1)
   b. Secondary endpoint number 3 should read ‘analgesia requirements’ (IRAS form section A11; protocol section 6.1)
5. Clarification in the main protocol that chemotherapy should not be started until 2 weeks after the radiotherapy is completed (protocol section 4.3.2).
6. Changes to the Radiotherapy Trial Specific Procedure
   a. Adding a line regarding taking photos of the radiotherapy field
   b. Clarifying when a trial radiotherapy CRF needs to be completed
   c. Clarifying that the volume to be treated should include the PTV and that it should be acceptable to the treating oncologist.
   d. Standardising the wording in the main protocol relating to the radiotherapy protocol to ‘trial specific procedure’ (rather than SOP) (protocol sections 4.2.3, 4.3.2)
7. Only reporting of SAEs after each trial clinic visit (protocol section 7.1).
8. Changes to the consent form:
   a. Only 3 copies of the consent form to be taken
   b. Clarification that information will be forwarded to Oxford and Bristol
c. Asking for consent to obtain information from NHS registers
9. Changes to the Patient information sheets:
   a. Addition regarding obtaining information from NHS registers.
10. The extra boxes for patient details have been removed from the GP letters.
11. Database management to be provided by the trial team at the Oxford Respiratory Clinical Trials Unit. (IRAS sections A6-2, A36, A38; protocol section 8)
12. Increasing the number of recruiting centres to 17 and changes to PIs and recruiting centres:
   a. At Plymouth Hospitals NHS Foundation Trust, the PI will be Mr Adrian Marchbank
   b. At Royal Devon and Exeter NHS Foundation Trust, the PI will be Dr Liz Toy
   c. At Portsmouth Hospitals NHS Trust, the PI will be Dr Lesley Bishop
   d. At Great Westen Hospitals NHS Foundation Trust, the PI will be Dr Anthony Kerry
   e. Llandough Hospital has been removed as a recruiting centre
   f. Addition of Blackpool, Dorchester and Torbay as recruiting centres (IRAS form Part C; Protocol section 3.1)

2. Non substantial Amendment. Date 11.01.2012
   Accepted by REC 11.01.2012
   Alteration of wording of standard consent form and semistructured interview consent form.

3. Non substantial Amendment. Date 12.01.2012
   Accepted by REC 12.01.2012
   Addition of additional recruiting centres.

4. Substantial Amendment. Date: 01.02.2012
   Accepted by REC 07.03.2012
   1. Removal of stratifying by centre (protocol section 4.2.2; IRAS section 61-1)
   2. Change need to inform the trial team if a different RT beam is used to documenting it on CRF, clarification of wording regarding field, CTV and PTV and adding contact details of trial Clinical oncologists for advice (RT TSP)
   3. Updating TSC membership (Protocol section 10.2)
   4. Removal of site specific logos from VAS score sheets and questionnaires
   5. Changes to VAS score format for administration purposes
   6. Clarification of requirement for centres to have recent audit data for radiotherapy quality assurance. (protocol section 3.1)

5. Non substantial Amendment. Date 05.03.2012
   Accepted by REC 16.03.2012
   Clarification of wording in VAS score

6. Non substantial Amendment. Date 30.03.2012
   Accepted by REC 11.04.2012
   Addition of new sites
7. Substantial Amendment. Date 16/07/2012  
Accepted by REC 01.08.2012

1. Addition of 2 extra questions to the semi-structured interview  
2. Change to inclusion criteria to include large bore pleural procedures within 35 days of the pleural intervention (Protocol section 3.2, IRAS sections A6-2, A17-1)  
3. Change to timeframe for radiotherapy to within 35 days (but a maximum of 42 days) (Protocol section 4.2.3, IRAS section A13)  
4. clarification to wording about SAEs (Protocol section 7.1)  
5. Change of name in research team (Protocol section “research team contact details”, IRAS sections A3, A63)  
6. Removal of sites from the R&D form (Blackpool, Southampton, Isle of Wight, Salisbury, UCLH, The Royal Brompton, South Devon) (protocol section 10.3, IRAS section C)

8. Non-substantial Amendment. Date 25/07/2012  
Accepted by REC 06.08.2012

1. Change of PI at UHB from Dr. Paula Wilson to Dr. Charles Comins (maternity cover)

9. Non-substantial Amendment Date 16/10/2012  
Accepted by REC 31.10.2012

1. Clarification regarding patient withdrawals (protocol section 5)  
2. Extension of recruitment period from 15 to 24 months

10. Substantial Amendment Date 30/10/2012  
Accepted by REC 07.12.2012

Change to Semi-structured interview TSP  
Adding option of an extra semi-structured interview for patients who develop a PTM after their 6 month follow up (protocol 4.3.3 & semi-structured interview PIS)  
Clarifying wording of semi-structured interview PIS.

11. Non-substantial Amendment Date 27/02/2013  
Acknowledged by REC 06.03.203

Addition of new site – Glangwili General Hospital, Carmarthenshire

12. Non-substantial Amendment Date 13/03/2013  
Acknowledged by REC 02.05.2013

Addition of new PIC site – Royal Brompton and Harefield NHS Trust

13. Non-substantial Amendment Date 12/04/2013  
Acknowledged by REC 02.05.2013

Addition of new PIC site – Worcester Royal Hospital
14. Non-substantial Amendment Date 26.04.2013
Acknowledged by REC

Addition of new sites – Royal Berkshire NHS Foundation Trust and The Royal Marsden NHS Foundation Trust

15. Non-substantial Amendment Date 21.08.2013
Acknowledged by REC

1. Extension of recruitment period from 24 to 36 months
2. Addition of PIC site – Heatherwood and Wexham Park Hospitals NHS Foundation Trust (King Edward VII Hospital)