



National Research Ethics Service

Cambridgeshire 2 Research Ethics Committee

Victoria House
Capital Park
Fulbourn
Cambridge
CB21 5XB

RECEIVED

Telephone: 01223 597693
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11 November 2009

12 NOV 2009

Professor Ieuan A. Hughes
Professor of Paediatrics
University of Cambridge
Department of Paediatrics
Box 116, Level 8
Addenbrookes Hospital
Cambridge
CB2 0QQ

Dear Professor Hughes

Study Title: Correlation of mutant androgen receptor (AR) function and pubertal outcome in partial androgen insensitivity syndrome (PAIS).
REC reference number: 09/H0308/158
Protocol number: 1

Thank you for your letter of 27 October 2009, responding to the Committee's request for further information on the above research and submitting revised documentation.

The further information has been considered on behalf of the Committee by the Chair.

Confirmation of ethical opinion

On behalf of the Committee, I am pleased to confirm a favourable ethical opinion for the above research on the basis described in the application form, protocol and supporting documentation as revised, subject to the conditions specified below.

Ethical review of research sites

The favourable opinion applies to all NHS sites taking part in the study, subject to management permission being obtained from the NHS/HSC R&D office prior to the start of the study (see "Conditions of the favourable opinion" below).

Conditions of the favourable opinion

The favourable opinion is subject to the following conditions being met prior to the start of the study.

Management permission or approval must be obtained from each host organisation prior to the start of the study at the site concerned.

For NHS research sites only, management permission for research ("R&D approval") should be obtained from the relevant care organisation(s) in accordance with NHS research

This Research Ethics Committee is an advisory committee to East of England Strategic Health Authority
The National Research Ethics Service (NRES) represents the NRES Directorate within the National Patient Safety Agency and Research Ethics Committees in England

governance arrangements. Guidance on applying for NHS permission for research is available in the Integrated Research Application System or at <http://www.rdforum.nhs.uk>. Where the only involvement of the NHS organisation is as a Participant Identification Centre, management permission for research is not required but the R&D office should be notified of the study. Guidance should be sought from the R&D office where necessary.

Sponsors are not required to notify the Committee of approvals from host organisations.

Other conditions specified by the REC

Where the reference to PALS has been inserted into the Information Sheets, the abbreviation needs to be explained in full (Patient Advice and Liaison Service) and the relevant contact details should be given ie.

Box 53, Cambridge University Hospitals NHS Foundation Trust, Hills Road, Cambridge CB2 0QQ, Tel: 01223 216 756, pals@addenbrookes.nhs.uk

In making the revision referred to above, the version number and date of the information sheets will change so the reference to them in paragraph 1 of the consent forms will also change. Accordingly, copies of the revised information sheets and consent forms should be provided to the committee for information. They will not receive ethical review.

It is the responsibility of the sponsor to ensure that all the conditions are complied with before the start of the study or its initiation at a particular site (as applicable).

Approved documents

The final list of documents reviewed and approved by the Committee is as follows:

<i>Document</i>	<i>Version</i>	<i>Date</i>
Covering Letter		14 August 2009
REC application	AB/145657/1	14 August 2009
Protocol	1	05 August 2009
Investigator CV		04 December 2008
Check list AB/145657/1		14 August 2009
Evidence of insurance or indemnity	Letter from UOC team	11 August 2009
Data collection sheet		20 July 2009
Workpackage description		
Signed CI declaration page	AB/145657/1	23 August 2009
Covering Letter	from Harriet Miles	24 August 2009
Participant Information Sheet: Parent/guardian	2	27 October 2009
Participant Information Sheet: 14 years and upwards	2	27 October 2009
Participant Information Sheet: 11-14 ys	2	27 October 2009
Response to Request for Further Information		27 October 2009
Participant Consent Form: parent/guardian	2	27 October 2009
Participant Consent Form: 14 years and upwards	2	27 October 2009
Participant Consent Form: Assent 11-14 ys	2	27 October 2009

Statement of compliance

The Committee is constituted in accordance with the Governance Arrangements for Research Ethics Committees (July 2001) and complies fully with the Standard Operating Procedures for Research Ethics Committees in the UK.

After ethical review

Now that you have completed the application process please visit the National Research Ethics Service website > After Review

You are invited to give your view of the service that you have received from the National Research Ethics Service and the application procedure. If you wish to make your views known please use the feedback form available on the website.

The attached document "*After ethical review – guidance for researchers*" gives detailed guidance on reporting requirements for studies with a favourable opinion, including:

- Notifying substantial amendments
- Adding new sites and investigators
- Progress and safety reports
- Notifying the end of the study

The NRES website also provides guidance on these topics, which is updated in the light of changes in reporting requirements or procedures.

We would also like to inform you that we consult regularly with stakeholders to improve our service. If you would like to join our Reference Group please email referencegroup@nres.npsa.nhs.uk.

09/H0308/158

Please quote this number on all correspondence

Yours sincerely



Dr Rowan Burnstein
Chair

Email: Ros.Cook@eoe.nhs.uk

Enclosures: "After ethical review – guidance for researchers"

Copy to: R&D Department at Cambridge University Hospitals NHS Foundation Trust

Dr Harriet Miles-by e-mail

Research and Development Department

R&D ref: A091713

23 November 2009

Professor Ieuan A Hughes
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www.addenbrookes.org.uk

Dear Professor Hughes

Re: 09/H0308/158 Correlation of mutant androgen receptor (AR) function and pubertal outcome in partial androgen insensitivity syndrome (PAIS).

In accordance with the Department of Health's Research Governance Framework for Health and Social Care, all research projects taking place within the Trust must receive a favourable opinion from an ethics committee and approval from the Department of Research and Development (R&D) prior to commencement.

R&D have reviewed the documentation submitted for this project, and has undertaken a **site specific assessment** based on the information provided in the SSI form, and I am pleased to inform you that we have no objection to the research proceeding within Cambridge University Hospitals NHS Foundation Trust.

Sponsor: Cambridge University Hospitals NHS Foundation Trust and University of Cambridge

Funder: European Consortium Grant

End date: 01/10/2012

Protocol: Version 1 dated 05/08/2009

The project must follow the agreed protocol and be conducted in accordance with all Trust Policies and Procedures especially those relating to research and data management.

You and your research team must ensure that you understand and comply with the requirements of the NHS Confidentiality Code of Practice and the Data Protection Act 1998 and are aware of your responsibilities in relation to the Human Tissue Act 2004, Good Clinical Practice, the NHS Research Governance Framework for Health and Social Care, Second Edition April 2005 and any further legislation released during the time of this study.

Members of the research team must have appropriate substantive or honorary contracts with the Trust prior to the study commencing. Any additional researchers who join the study at a later stage must also hold a suitable contract.

If the project is a clinical trial under the European Union Clinical Trials Directive the following must also be complied with:

the EU Directive on Clinical Trials (Directive 2001/20/EC) and UK's implementation of the Directive: The Medicines for Human Use (Clinical Trials) Regulations 2004;

- the EU Directive on Principles and Guidelines for Good Clinical Practice (EU Commission Directive 2005/28/EC); and UK's implementation of the Directive: The Medicines for Human Use (Clinical Trials) Amendment Regulations 2006;

Amendments

Please ensure that you submit a copy of any amendments made to this study to the R&D Department.

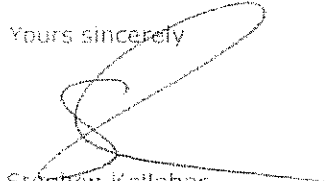
Annual Report

It is obligatory that an annual report is submitted by the Chief Investigator to the research ethics committee, and we ask that a copy is sent to the R&D Department. The yearly period commences from the date of receiving a favourable opinion from the ethics committee.

Please refer to our website www.cuh.org.uk/research for all information relating to R&D including honorary contract forms, policies and procedures and data protection.

Should you require any further information please do not hesitate to contact us.

Yours sincerely



Stephen Kelleher
R&D Manager

Cambridgeshire 2 Research Ethics Committee

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11 FEB 2009

10 February 2009

Professor Ieuan Hughes
Department of Paediatrics
Box 116, Level 8
Cambridge University Hospitals NHS Foundation Trust, Addenbrooke's Hospital
Hills Road
Cambridge
CB2 0QQ

Dear Professor Hughes

Study title: XY Sex Reversal Syndromes in Childhood (a diagnostic study)

REC reference: 96/129

Amendment numbers: 3 and 4

Amendment date: 21 January 2009

The above amendments were reviewed at the meeting of the Sub-Committee of the REC held on 09 February 2009.

Ethical opinion

The members of the Committee present gave a favourable ethical opinion of the amendments on the basis described in the notice of amendment form and supporting documentation.

Approved documents

The documents reviewed and approved at the meeting were:

Document	Version	Date
Cover letter		30 January 2009
Notice of Substantial Amendment 3		21 January 2009
Clinical Information Data Collection Sheet	2	18 January 2009
New patient information sheet parents/guardians	1	18 January 2009

New patient information sheet over 14 ys	1	18 January 2009
New patient information sheet children <14ys	1	18 January 2009
New patient consent form parents/guardians	1	18 January 2009
New patient consent form over 14 ys	1	18 January 2009
New patient consent form children <14 ys	1	18 January 2009
Existing patient information sheet parents/guardians	1	18 January 2009
Existing patient information sheet over 14ys	1	18 January 2009
Existing patient information sheet children <14ys	1	18 January 2009
Existing patient consent form parents/guardians	1	18 January 2009
Existing patient consent form over 14ys	1	18 January 2009
Existing patient consent form <14 ys	1	18 January 2009
GP letter	1	18 January 2009

Amendment 4:

Document	Version	Date
Cover letter		30 January 2009
Notice of Substantial Amendment 4		21 January 2009

Membership of the Committee

The members of the Committee who were present at the meeting are listed on the attached sheet.

R&D approval

All investigators and research collaborators in the NHS should notify the R&D office for the relevant NHS care organisation of this amendment and check whether it affects R&D approval of the research.

Statement of compliance

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96/129:	Please quote this number on all correspondence
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Yours sincerely



Anna Bradnam
REC Assistant Co-ordinator

E-mail: Anna.Bradnam@eoe.nhs.uk

Enclosures List of names and professions of members who were present at the meeting and those who submitted written comments

Copy to: R&D Unit, Box 277, Cambridge University Hospitals NHS Foundation Trust,
 Hills Road, Cambridge CB2 0QQ

Cambridgeshire 2 Research Ethics Committee

Attendance at Sub-Committee of the REC meeting on 09 February 2009

<i>Name</i>	<i>Profession</i>	<i>Capacity</i>
Dr Rowan Burnstein (Chair)	Consultant in Intensive Care and Anaesthesia, SDU Director Neurocritical Care	Expert
Dr Nigel Hymas	Consultant Psychiatrist	Expert
Mrs Rebekah Ley	Lay Member	Lay Member

Also in attendance:

<i>Name</i>	<i>Position (or reason for attending)</i>
Anna Bradnam	REC Assistant Coordinator

Ethical Approval REC Ref 96/129

Cambridge Research Ethics Committee (LREC Cambridge2 Chaired by Dr Berrios)

- May 1991 DEC91/2 **Androgen Insensitivity Syndromes in Childhood.**
- March 1993 Three types of sample : blood,genital skin biopsy, gonadal tissue
- May 1996 Ethics committee reconstituted, Ethical approval granted for a further 3 years: **Androgen insensitivity syndromes in childhood**
REC ref 96/129.
- Feb 2001 Approval extended for a further 3 years retrospectively from May 1999
- August 2002 Chairmans extension 3 years, Dr Berrios
- August 2003 Histology of gonads clarified as part of diagnostic work.Dr Nick Coleman and Dr Ian Scott at the Cambridge MRC Cambridge Cancer Research Institute named as collaborators.
- May 2005 Progress report accepted (next progress report due 28/05/06 pending the committees decision on the substantial amendment) Dr Berrios
- July 2005 Substantial Amendment accepted 10/06/2005 (Amendment 2):
XY sex reversal syndromes in childhood (a diagnostic study).
- AIS syndromes and other sex reversal disorders eg. 17 β -hydroxysteroid dehydrogenase, 5 α -reductase and P450-oxido reductase enzyme deficiencies.
- Expected completion date – “an on going study” Annual progress report June 2005.

No end approval date on latest correspondence 13/7/05 Elaina Friend Committee Manager