Government of South Australia
SA Health

Central Adelaide Local Health Network
Royal Adelaide Hospital Human Research Ethics Committee
Level 3, Hanson Institute, IMVS Building
Royal Adelaide Hospital
North Terrace
Adelaide, South Australia, 5000
Telephone: +61 8 8222 4139
Fax: +61 8 8222 3035
Email: rah.ethics@health.sa.gov.au

Approval Date: 7 March 2014

Prof G L Ludbrook
Pain & Anaesthesia Research Clinic
ROYAL ADELAIDE HOSPITAL

Dear Prof Ludbrook,

HREC reference number: **HREC/14/RAH/45**

Project Title: **"A Phase I, placebo and positive-controlled, dose-escalation study to determine the safety, pharmacodynamics and pharmacokinetics of a single intravenous injection of HSK3486 in healthy subjects." Protocol No: HSK3486 SAD.**

RAH Protocol No: 140225.

Thank you for submitting the above project for ethical and scientific review. This project was first considered by the Royal Adelaide Hospital Human Research Ethics Committee HREC at its meeting held on 27 Febraury 2014. I am pleased to advise that your protocol has been granted full ethics approval and meets the requirements of the *National Statement on Ethical Conduct in Human Research.* The documents reviewed and approved include:

- **NEAF Submission: AU/1/C1A6112**
- **Protocol, Version 2, 7 March 2014**
- **Participant Information Sheet & Informed Consent Form, Version 4, 4 March 2014**
- **Investigator's Brochure, Edition 1.2, 4 January 2014**
- **Chemistry Manufacturing and Control, updated Version, 16 January 2014**
- **Study Advertisement, Version 2, 4 February 2014**

Please quote the RAH Protocol Number allocated to your study on all future correspondence.

GENERAL TERMS AND CONDITIONS OF ETHICAL APPROVAL:

- For all clinical trials, the study must be registered in a publicly accessible trials registry prior to enrolment of the first participant.
- Adequate record-keeping is important. If the project involves signed consent, you should retain the completed consent forms which relate to this project and a list of all those participating in the project, to enable contact with them in the future if necessary. The duration of record retention for all clinical research data is 15 years.
- You must notify the Research Ethics Committee of any events which might warrant review of the approval or which warrant new information being presented to research participants, including:
  (a) serious or unexpected adverse events which warrant protocol change or notification to research participants,
  (b) changes to the protocol,
  (c) premature termination of the study
- The Committee must be notified within 72 hours of any serious adverse event occurringat this site.
- Approval is valid for **5 years** from the date of this letter, after which an extension must be applied for. Investigators are responsible for providing an annual review to the RAH REC Executive Officer each anniversary of he above approval date, within 10 workings days, using the Annual Review Form available at: http://www.rah.sa.gov.au/rec/index.php
- The REC must be advised with a report or in writing within 30 days of completion.

Should you have any queries about the HREC's consideration of your project, please contact Mrs Heather O'Dea on 08 8222 4139, or rah.ethics@health.sa.gov.au .

You are reminded that this letter constitutes ethical approval only. You must not commence this research project at a SA Health site until separate authorisation from the Chief Executive or delegate of that site has been obtained.

This Committee is constituted in accordance with the NHMRC's *National Statement on the Ethical Conduct of Human Research* (2007). The HREC wishes you every success in your research.

Yours sincerely,

A/Prof A Thornton
CHAIRMAN
RESEARCH ETHICS COMMITTEE

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ROYAL ADELAIDE HOSPITAL
North Terrace
Adelaide SA 5000
Tel: +61 8 8222 4000
Fax: +61 8 8222 5939
ABN 80 230 154 545
www.rah.sa.gov.au

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Level 3, Hanson Institute
Tel: (08) 8222 4139
Fax: (08) 8222 3035
Email: rah.ethics@health.sa.gov.au

Date: 7 March 2014

Research Ethics Committee :-
Meeting held on: 27 February 2014

Protocol: HSK3486 SAD
"A Phase I, placebo and positive-controlled, dose-escalation study to determine the safety, pharmacodynamics and pharmacokinetics of a single intravenous injection of HSK3486 in healthy subjects."

TO WHOM IT MAY CONCERN

Please be advised of the above Protocol approval by the Royal Adelaide Hospital Research Ethics Committee. It is the policy of both the RAH Research Ethics Committee & Investigational Drug Subcommittee that the names of the regular members of these Committees are not released.

The general makeup of the Committees is as listed below (effective at the above Meeting review date):

Research Ethics Committee
Male – Chairperson, Hospital Scientist employee Royal Adelaide Hospital
Female – One Qualified Lawyer
Male – One Layman
Female – One Laywoman
Female – One Laywoman, Philosophy, University of Adelaide
Male – One Minister of Religion
Male – One Counselling Professional.
Female – One Nurse, employee Royal Adelaide Hospital
Male – Clinician, Graduate of Medicine, employee SA Pathology
Male – Clinician, Graduate of Medicine, employee SA Pathology
Male – Clinician, Graduate of Medicine, employee SA Pathology
Male – Clinician, Graduate of Medicine, retired employee SA Pathology
Female – Clinician, Graduate of Medicine, employee Royal Adelaide Hospital
Female – Clinician, Graduate of Medicine, employee Royal Adelaide Hospital
Female – Clinician, Graduate of Medicine, employee Royal Adelaide Hospital
Male – Clinician, Graduate of Medicine, employee Royal Adelaide Hospital
Male – Clinician, Graduate of Medicine, employee Royal Adelaide Hospital
Male – Clinician, Graduate of Medicine, employee Royal Adelaide Hospital

Investigational Drug Subcommittee
Chairperson - Senior Staff Specialist in Clinical Pharmacology, Royal Adelaide Hospital.
Three senior members of the University of Adelaide, Department of Clinical Pharmacology.
Two members of the Royal Adelaide Hospital, Pharmacy Department.
Three members of the Royal Adelaide Hospital, Clinical Pharmacology.
Chairman, Research Ethics Committee.

The REC composition is in accord with the guidelines for Institutional Ethics Committee composition as set out in the NHMRC National Statement on Ethical Conduct in Human Research (2007). .

The committee is serviced by me (undersigned) in my role as Executive Officer.
Proxy members have been appointed to assure that there is full representation at all meetings.

Please be advised that no Investigators of the abovementioned study were present at either the Investigational Drugs Subcommittee Meeting or the Research Ethics Committee Meeting, at the time of discussion or recommendations.

The Hospital's Research Ethics Committee is governed by the NHMRC National Statement on Ethical Conduct in Human Research (2007).

Yours sincerely,

Mrs Heather O'Dea
Executive Officer, RESEARCH ETHICS COMMITTEE