

F. No. ND/CT/23/000061
Government of India
Directorate General of Health Services
Central Drugs Standard Control Organization
New Drugs Division

FDA Bhawan, Kotla Road,
New Delhi-110002

To
M/s Hetero Labs Limited,
Hetero Corporate, 7-2-A2 Industrial Estate,
Sanath Nagar Hyderabad (India) – 500055.

Subject: Grant of permission to conduct Phase IV Clinical Trial title “A Phase IV, Post-marketing, Prospective, Multi-centric, Single Arm, Non-comparative Study to Evaluate the Safety and Efficacy of Polmacoxib 2mg in Adult Patients with Osteoarthritis of Hip/Knee”. (Protocol no. HCR/IV/POLMOA/07/2023, Version. 1, dated 10-07-2023) -regarding.

Reference: Letter Ref. No. ND/CT04/FF/2023/38962 dated 31-JUL-2023.

Sir,

With reference to your application no. ND/CT04/FF/2023/38962 dated 31-JUL-2023, please find enclosed herewith the permission in Form CT-06, No. **CT/ND/11/2024** to conduct the subject mentioned clinical trial under the provisions of New Drugs and Clinical Trial Rules, 2019.

This permission is subject to the conditions, as mentioned below.

RAJEEV SINGH
RAGHUVANSHI

Digitally signed by RAJEEV SINGH RAGHUVANSHI
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Yours faithfully,

(Dr. Rajeev Singh Raghuvanshi)
Central Licensing Authority

Conditions of Permission

- (i) Clinical trial at each site shall be initiated after approval of the clinical trial protocol and other related documents by the Ethics Committee of that site, registered with the Central Licensing Authority under rule 8;
- (ii) Where a clinical trial site does not have its own Ethics Committee, clinical trial at that site may be initiated after obtaining approval of the protocol from the Ethics Committee of another trial site; or an independent Ethics Committee for clinical trial constituted in accordance with the provisions of rule 7:

Provided that the approving Ethics Committee for clinical trial shall in such case be responsible for the study at the trial site or the centre, as the case may be:

Provided further that the approving Ethics Committee and the clinical trial site or the bioavailability and bioequivalence centre, as the case may be, shall be located within the same city or within a radius of 50 kms of the clinical trial site;

- (iii) In case an ethics committee of a clinical trial site rejects the approval of the protocol, the details of the same shall be submitted to the Central Licensing Authority prior to seeking approval of another Ethics Committee for the protocol for conduct of the clinical trial at the same site;
- (iv) The Central Licensing Authority shall be informed about the approval granted by the Ethics Committee within a period of fifteen working days of the grant of such approval;
- (v) Clinical trial shall be registered with the Clinical Trial Registry of India maintained by the Indian Council of Medical Research before enrolling the first subject for the trial;
- (vi) Clinical trial shall be conducted in accordance with the approved clinical trial protocol and other related documents and as per requirements of Good Clinical Practices Guidelines and the provisions of these rules;
- (vii) Status of enrolment of the trial subjects shall be submitted to the Central Licensing Authority on quarterly basis or as appropriate as per the duration of treatment in accordance with the approved clinical trial protocol, whichever is earlier;
- (viii) Six monthly status report of each clinical trial, as to whether it is ongoing, completed or terminated, shall be submitted to the Central Licensing Authority electronically in the SUGAM portal;
- (ix) In case of termination of any clinical trial the detailed reasons for such termination shall be communicated to the Central Licensing Authority within thirty working days of such termination;
- (x) Any report of serious adverse event occurring during clinical trial to a subject of clinical trial, shall, after due analysis, be forwarded to the Central Licensing Authority, the chairperson of the Ethics Committee and the institute where the trial has been conducted within fourteen days of its occurrence as per Table 5 of the Third Schedule and in compliance with the procedures as specified in Chapter VI of the New Drugs and Clinical Trials Rules, 2019;
- (xi) In case of injury during clinical trial to the subject of such trial, complete medical management and compensation shall be provided in accordance with the Chapter VI of the said Rules and details of compensation provided in such cases shall be intimated to the Central Licensing Authority within thirty working days of the receipt of order issued by Central Licensing Authority in accordance with the provisions of the said Chapter.
- (xii) In case of clinical trial related death or permanent disability of any subject of such trial during the trial, compensation shall be provided in accordance with the Chapter VI and details of compensation provided in such cases shall be intimated to the Central Licensing Authority within thirty working days of receipt of

the order issued by the Central Licensing Authority in accordance with the provisions of the said Chapter;

- (xiii) The premises of the sponsor including his representatives and clinical trial sites, shall be open for inspection by officers of the Central Licensing Authority who may be accompanied by officers of the State Licensing Authority or outside experts as authorized by the Central Licensing Authority, to verify compliance of the requirements of these rules and Good Clinical Practices Guidelines, to inspect, search and seize any record, result, document, investigational product, related to clinical trial and furnish reply to query raised by the said officer in relation to clinical trial;
- (xiv) Where the New Drug or Investigational New Drug is found to be useful in clinical development, the sponsor shall submit an application to the Central Licensing Authority for permission to import or manufacture for sale or for distribution of new drug in India, in accordance with Chapter X of these rules, unless otherwise justified;
- (xv) The Laboratory owned by any person or a company or any other legal entity and utilised by that person to whom permission for clinical trial has been granted used for research and development, shall be deemed to be registered with the Central Licensing Authority and may be used for test or analysis of any drug for and on behalf of Central Licensing Authority;
- (xvi) The Central Licensing Authority may, if considered necessary, impose any other condition in writing with justification, in respect of specific clinical trials, regarding the objective, design, subject population, subject eligibility, assessment, conduct and treatment of such specific clinical trial;
- (xvii) The sponsor and the investigator shall maintain the data integrity of the data generated during clinical trial.
- (xviii) Informed Consent Documents (ICD) viz. Patient Information Sheet (PIS) and Informed Consent Form (ICF) complete in all respect & must be got approved from the respective Ethics committee and submitted to CDSCO before enrolling first subject at the respective site.
- (xix) The Informed Consent Document including ICF and Patient Information Sheet should clearly mention in understandable language about the details of the drug therapy that the patient may or may not receive.
- (xx) **Indian version of WOMAC- OA index to be used for valid assessment of efficacy.**
- (xxi) **Serum NT Pro BNP test at screening and 6 weeks for Cardiovascular safety assessment.**

FORM CT-06*(See rules 22, 25, 26, 29 and 30)***PERMISSION TO CONDUCT CLINICAL TRIAL OF NEW DRUG OR INVESTIGATIONAL
NEW DRUG**

The Central Licensing Authority hereby permits M/s Hetero Labs Limited, Hetero Corporate, 7-2-A2 Industrial Estate, Sanath Nagar Hyderabad (India) - 500055 Telephone No.: 040-23704923 FAX: 040-23714250 E-Mail :LAKSHMI.G@HETERODRUGS.COM to Phase IV conduct clinical trial of the new drug Polmacoxib Capsules 2 mg as per protocol No. HCR/IV/POLMOA/07/2023, Version. 1, dated 10-07-2023 in the below mentioned clinical trial sites.

2. Details of new drug or investigational new drug:

Names of the new drug or investigational new drug:	Polmacoxib Capsules 2 mg
Therapeutic class:	Dual COX2 carbonic anhydrase I and II inhibitor
Dosage form:	Capsule
Composition:	Each Capsule contains Polmacoxib.....2.0 mg
Indications:	Indicated for the treatment of Idiopathic Osteoarthritis of Hip/Knee.

Details of clinical trial sites-

Sr. No.	Name of Principal Investigator & Trial sites	Ethics Committee Name/Registration Number
1.	Dr. Tadikonda Bhavani Prasad, HOD & Professor of Orthopaedics, Visakha Institute of Medical Sciences Department of Orthopaedics Visakhapatnam-530040, Andhra Pradesh, India	Visakha Institute of Medical Sciences S.No.97/2 Hanumathwaka, Chinnagadili Village, Visakhapatnam 530040, Andhra Pradesh, India ECR/1421/Inst/AP/2020
2.	Dr. Kiran Kumar Mukhopadhyay, NRS Medical College and Hospital Department of Orthopaedics, 138, AJC Bose Road Kolkata-700014, West Bengal, India	NRS Medical College and Hospital, 138 AJC Bose Road, Kolkata-700014 West Bengal, India ECR/609/Inst/WB/2014/RR-20
3.	Dr. Bhavesh Jesalpura, GMERS Medical College, Civil Hospital, Sola, Department of Orthopaedic Surgery, Solagram, Rd. Beside High Court, Ahmedabad-380060, Gujarat, India	GMERS Medical College, Sola, GMERS Medical College, Civil Hospital, Sola, High Way Near Gujarat High Court Ahmedabad, Gujarat -380060, India ECR/404/Inst/GJ/2013/RR-20
4.	Dr. Rakesh Verma, Jawahar Lal Nehru Medical College, Kala Bagh, Ajmer-305001, Rajasthan	Jawahar Lal Nehru Medical College, Kala Bagh, Ajmer-305001, Rajasthan ECR/1156/Inst/RJ/2018/RR-22

5.	Dr. Brijesh Patel, Hi-Tech Multispeciality Hospital Sector3-D, Plot No.1180, Gh Road, Nr. Gh-11/2 Bus stand Gandhinagar, Gujarat-382003.	Hi-Tech Multispeciality Hospital Sector 3-D,Plot No.1180,Gh Road, Nr.Gh-11/2 Bus stand Gandhinagar -382003 Gujarat ECR/1057/Inst/GJ/2018/RR-21
6.	Dr. Kishore Babu Sattaru, Government Medical College & Govt General Hospital (Old RIMSGGH) Department of Orthopaedics Srikakulam-532001, Andhra Pradesh	Government Medical College & Govt General Hospital Srikakulam -532001, Andhra Pradesh ECR/492/Inst/AP/2013/RR-20

3. This permission is subject to the conditions prescribed in part A of Chapter V of the New Drugs and Clinical Trials Rules, 2019 under the Drugs and Cosmetics Act, 1940.

RAJEEV SINGH
RAGHUVANSHI

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(Dr. Rajeev Singh Raghuvanshi)
Central Licensing Authority

New Delhi