File No. 12-42/2018-DC (Pt-Hetero-SND)
Government of India
Directorate General of Health Services
Central Drugs Standard Control Organisation
(Subsequent New Drugs Division)

Dated 17 JUN 2019

To,

M/s. Hetero Labs Ltd.,
Hetero Corporate, Industrial Estates,
H. No: 7-2-A2, Sanathnagar, Hyderabad – 500018,
Telangana State, India.

Subject: Permission for conducting Phase-III clinical trial titled "A Phase III, Multi-Center, Double blind, Comparative, active-control, parallel group, randomized, study to evaluate the efficacy and safety of Acotiamide ER tablet 300 mg in Indian Adult Patients with Functional Dyspecpsia-post prandial distress syndrome"- regarding.

CT NOC No. CT/SND/53/2019

Sir,

This Directorate has no objection to your conducting the subject mentioned clinical trial as per the provisions of Drugs & Cosmetics Rules under supervision of the following investigator and as per the **Protocol No: HCR/III/ACOFD-PDS/01/2015, Phase III, and Version 1.0, Dated 18-01-2018** submitted to this Directorate.

S. No	Investigator and Trial site	Ethics Committee Name and Registration Number
1	Dr. Malladi Umadevi MBBS, MD (General Medicine), In Patient Block, 5th Floor, Department of Gastroenterology, Gandhi Hospital, Musheerabad, Secunderabad, Telangana – 500003.	Institutional Ethics Committee, Gandhi Medical College/Hospital, Musheerabad, Secunderabad, Telangana – 500003. ECR/108/Inst/AP/2013/RR-16
2	Dr. Mukesh Agarwala MBBS, MD (Medicine), Marwari Hospital and Research Centre, S.J Road, Athgaon, Guwahati-781008.	ECR-HEC, Marwari Hospital and Research Centre, S.J Road, Athgaon, Guwahati-781008. ECR/487/Inst/AS/2013/RR-16

3	Dr. Mukesh Kumar Jain., MBBD, MD (internal Medicine), Department of Gastroenterology, Room No. 20, First Floor, Dhanvantri OPD Block, SMS Hospital, Jaipur - 302004	Office of the Ethics Committee, SMS Medical college & Attached Hospitals, Jaipur, F-1, Division of Allergy & Pulmonary Medicine, Dhanvantri OPD Block SMS Hospital, Jaipur, Rajasthan – 302004. ECR/26/Inst/RJ/2013/RR-16
4	Dr. Deo Nithi Mishra, MBBS, MD (Medicine), Nirmal Hospital, Opp. MLB Medical College, Gate No. 3, Jhansi (U.P) – 284128.	Institutional Ethics Committee, Nirmal Hospital Intuitional Ethics Committee, Nirmal Hospital Campus, Opp. MLB Medical College, Gate No. 3, Kanpur Road, Jhansi (U.P) – 284128. ECR/325/Inst/UP/2013/RR-16
5	Dr. Maitreyee Bandopadhyay, MBBS, DTM & H, MD (Medicine), NRS Medical College & Hospital, 138 AJC Bose Road, Kolkata – 700014, West Bengal, India.	Ethics Committee NRS Medical College, 138 AJC Bose Road, Kolkata – 700014. ECR/609/Inst/WB/2014
6	Dr. S.K. Sinha, MBBS, MD (Medicine), Nehru Hosptial, PGIMER Sector 12 Chandigarh, 160012.	Intuitional Ethics Committee, PGIMER, Sector 12 Chandigarh, 160012. ECR/25/Inst/CH/2013/RR-16
7	Dr. B. S.S.V. Ratna Giri MBBS, MD (General Medicine), DM (Gasteroloenterology), New Govt. General Hospital, (Associated by Govt. Siddhartha Medical college), Govt. Siddhartha Medical College Campus, Gundala, Vijaya wada – 520008. Andhra Pradesh.	Institutional Ethics Committee, Siddhartha Medical College & Govt. General Hospital (IEC SMV & GGH), Govt. Siddhartha Medical College Campus, Gundala, Vijaya-wada – 520008. Andhra Pradesh. ECR/633/Inst/AP/2014/RR-17
8	Dr. Rohit Sureka, MBBS, DNB. (Gen. Medicine), Jindal Superspeciality Hospital, S.P. Mukharjee Nagar, Bharatpur – 321001, Rajasthan.	Institutional Ethics Committee Jindal Superspeciality Hospital, S.P. Mukharjee Nagar, Bharatpur – 321001, Rajasthan. ECR/992/Inst/RJ/2017
9	Dr. Wagh Amol Nanasaheb, MBBS, (General Surgery), OPD. No.5, Department of General Surgery, Grant Government Medical College and Sir JJ Group of Hospitals, Byculla Mumbai-400008.	Institutional Ethics Committee, Grant Government Medical College and Sir JJ Group of Hospitals, Byculla Mumbai-400008. Maharashtra, India. ECR/14/Inst/UP/2013-RR-16

10	Dr. Gyan Prakash Rai, MBBS, MD, DM (Gastrology), Surya Super Speciality Hospital, a Unit of G.V. Meditech (P) Ltd, B 38/46 H Raman Niwas, Mahmoorganj, Varanasi – 221010, UP, India.	G.V. Metitech Ethics Committee, G.V. Meditech (P) Ltd, B 38/46 H Raman Niwas, Mahmoorganj, Varanasi – 221010, UP, India. ECR/14/Inst/UP/2013-RR-16

Kindly note that the clinical trial permission is subject to the following conditions:-

a) Clinical trial shall be conducted in compliance with the approved protocols, requirements of New Drugs and Clinical Trial Rules, 2019, Good Clinical Practice Guidelines issued by this Directorate and other applicable regulations.

b) Approval of Institutional Ethics Committee duly registered with CDSCO (under Rule 122DD of Drugs & Cosmetics Rules/ New Drugs and Clinical Trial Rules, 2019) should be obtained and submitted to this Directorate before initiation of the study.

c) Clinical trials shall be registered at Clinical Trials Registry of India (CTRI) before enrolling the first patient for the study.

d) Annual status report of each clinical trial, as to whether it is ongoing, completed or terminated, shall be submitted to the Licensing Authority, and in case of termination of any clinical trial the detailed reasons for the same shall be communicated to the said Licensing Authority.

e) Any report of serious adverse event occurring during clinical trial study to the subject, after due analysis, shall be forwarded within fourteen days of its occurrence as per New Drugs and Clinical Trial Rules, 2019.

f) In case of an injury or death during the study to the subjects, the applicant shall provide complete medical management and compensation in the case of trial related injury or death in accordance with New Drugs and Clinical Trial Rules, 2019 and the details of compensation provided in such cases shall be intimated to the Licensing Authority within thirty days of the receipt of the order of the said authority.

g) The premises of Sponsor including their employees, subsidiaries and branches, their agents, contractors and sub-contractors and clinical trial study sites shall be open to inspection by the officers authorized by the Central Drugs Standard Control Organization, who may be accompanied by an officer of the State Drug Control Authority concerned, to verify compliance to the requirements of New Drugs and Clinical Trial Rules, 2019, Good Clinical Practices guidelines for conduct of clinical trial in India and other applicable regulations.

h) The Sponsor including their employees, subsidiaries and branches, their agents, contractors and sub-contractors and clinical trial study sites and the Investigator shall allow officers authorized by the Central Drugs Standard Control Organization, who may be accompanied by an officer of the State Drug Control Authority concerned, to enter with or without prior notice, any premises of Sponsor including their employees, subsidiaries and branches, their agents, contractors and sub-contractors and clinical trial sites to inspect,

search and seize any record, data, document, books, investigational drugs, etc. related to clinical trial and provide adequate replies to any queries raised by the inspecting authority in relation to the conduct of clinical trial.

i) Clinical trial shall be conducted only at those sites which are institutes/hospitals having adequate emergency facilities and duly registered Institutional Ethics committees.

j) The sponsor shall ensure that the number of clinical trials an investigator can undertake should be commensurate with the nature of the trial, facility available with the Investigator etc.

k) The details of payment/honorarium/financial support/fees paid by the Sponsor to the Investigator(s) for conducting the study shall be made available to this directorate before initiation of each of the trial sites.

l) In addition to the requirement of obtaining written informed consent, an audio-video recording of the informed consent process in case of vulnerable subjects in clinical trial of New Chemical Entity or New Molecular Entity including procedure of providing information to the subject and his understanding on such consent, shall be maintained by the investigator for record; provided that in case of clinical trial of anti-HIV and anti-leprosy drugs, only audio recording of the informed consent process of individual subject including the procedure of providing information to the subject and his understanding on such consent shall be maintained by the investigator for record, as per provisions of New Drugs and Clinical Trial Rules, 2019.

m) **The bulk drug to be used in manufacturing of finished formulation intended to be used in the clinical trial and clinical trial batches of finished formulation shall be manufactured under GMP conditions using validated procedures and shall have ongoing stability programme.**

n) If the clinical trial batches are different from that of the primary batches for which data have been submitted, stability reports for clinical trial batches are to be submitted as per New Drugs and Clinical Trial Rules, 2019 for Drug substances and formulation along with Clinical Study Report.

o) It may kindly be noted that merely granting permission to conduct clinical trials with the drug does not convey or imply that based on the clinical trial data generated with the drug, permission to market this drug in the country will automatically be granted to you.

p) Informed consent documents (ICD) viz. Patient Information Sheet (PIS) and Informed Consent Form (ICF) complete in all respect as per the requirements specified in Appendix V of Schedule Y of the Drugs and Cosmetics Rules, 1945 must be got approved from respective Ethics Committee and submitted to CDSCO before enrolling first subject at the respective site.

Yours faithfully,

(Dr. S. Eswara Reddy)
Drugs Controller General (India)
& Licensing Authority