

File No CT/20/2013-DCG(I)

Clinical Trial NOC No:- GCT/26/13

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Government of India

Directorate General of Health Services
Central Drugs Standard Control Organization
Office of Drugs Controller General (India)
(Global Clinical Trial Division)

36925/31/07/13

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File No: CT/20/13-DCG (I)

Dated: 02/01/14

To

M/s. Klinera Corporation India,
401, Hill view idle. Estate, L.B.S Marg,
Ghatkopar(w), Mumbai - 400 050.

Subject: Permission for conducting a phase III clinical study entitled, "A phase 3, randomized, double-blind study to evaluate the safety and efficacy of Tenofovir Alafenamide (TAF) 25 mg QD versus Tenofovir Disoproxil Fumarate (TDF) 300 mg QD the treatment of HBeAg-Negative, Chronic Hepatitis B." - regarding.

Clinical Trial NOC No. GCT/26/13

Reference: Your letter no. nil 26/07/13 on the subject mentioned above.

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 investigators and as per the **Protocol No: GS-US-320-0108 versions 1.0 dated 12-07-2013** submitted to this Directorate.

1. Dr. Akash Shukla, Seth GS Medical College and KEM Hospital, Mumbai, Maharashtra, 400012
2. Dr. Shrikant Mukewar, Midas Multispeciality Hospital, Nagpur, Maharashtra,
3. Dr. Pravin Rathi, BYL Nair Hospital & T N Medical College, Mumbai, Maharashtra,
4. Dr. S. K. Acharya, All India Institute of Medical Sciences, New Delhi, Delhi, 110029
5. Dr. V. G. Mohan Prasad, VGM Hospital, Coimbatore, Tamil Nadu, 641005
6. Dr. S. K. Sarin, Institute of Liver and Biliary Sciences, New Delhi, Delhi, 110070
7. Dr. Abhijit Chowdhury, Institute of Post Graduate Medical Education And Research, Kolkata, West Bengal, 700020
8. Dr. P. N. Rao, Asian Institute of Gastroenterology, Hyderabad, Andhra Pradesh,
9. Dr. Aejez Habeeb, Center for Liver Research & Diagnostics, Deccan College of Medical Sciences and Allied Hospitals, Hyderabad Andhra Pradesh, 500058
10. Dr. Ajit Sood, Dayanand Medical College & Hospital, Ludhiana, Punjab, 141001
11. Dr. Kaushal Madan, Medanta-The Medicity, Gurgaon, Haryana, 122001

12. Dr. Vivek Saraswat, Sanjay Gandhi Postgraduate Institute of Medical Sciences, Lucknow, Uttarpradesh, 226014
13. Dr. Rajiv Mehta, Nirmal Hospital Pvt. Ltd, Surat, Gujarat, 395002
14. Dr. Ajay Duseja, Postgraduate Institute of Medical Education & Research, Chandigarh, Chandigarh, 160012
15. Dr. Anil Arora, Sir Ganga Ram Hospital, New Delhi
16. Dr. Babadev Goswami, Institute of digestive and liver Disease, Dispur Hospital, Guwahati

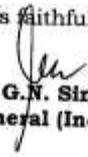
Kindly note that the clinical trial permission is subject to the following conditions:

- a. Clinical trial shall be conducted in compliance with the approved protocols, requirements of Schedule Y, Good Clinical Practice Guidelines issued by this Directorate and other applicable regulations;
- b. Approval of the Ethics Committee shall be obtained before initiation of the study;
- c. Clinical trials shall be registered at Clinical trials Registry of India 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 ten days of its occurrence as per Appendix XI and in compliance with the procedures prescribed in Schedule Y;
- 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 rule 122 DAB and the procedures prescribed under Schedule Y, 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 Schedule Y, 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. However, under no circumstances the number of trials should be more than three at a time.
- 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. You are also informed that audio visual recording of the informed consent process of each trial subject, including the procedure of providing information to the subject and his/her understanding on such consent is required to be done duly adhering to the principles of confidentiality. Such audio-visual recording and related documentation shall be preserved. All the sponsors /investigators /institutes/Organizations and other stake holders involved in conduct of clinical trials in the country are hereby directed to adhere to the above requirement of audio-visual recording of informed consent process of trial subjects with immediate effect.
- m. 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.

Yours faithfully,


(Dr. G.N. Singh)
Drugs Controller General (India)