26497/20.06.14

GOVERNMENT OF INDIA
Directorate General of Health Services
Central Drugs Standard Control Organization
Office of Drugs Controller General (India)
(Global Clinical Trial Division)
FDA Bhawan, Kotla Road, New Delhi-110002
Te No: 01123236965, Fax: 01123236971
E-mail: dci@nic.in

File No: CT/14/14 – DCG (I) Dated: 27/08/14

To,

M/s.Boehringer Ingelheim India Pvt. Ltd.
1102, 11th Floor, Hallmark Business Plaza,
Guru Nanak Hospital Road, Near Guru Nanak Hospital
Bandra (East), Mumbai-400051

Subject: Permission for conducting a Phase III clinical trial entitled "A Phase III, case series clinical study of the reversal of the anticoagulant effects of dabigatran by intravenous administration of 5.0 g idarucizumab (BI 655075) in patients treated with dabigatran etexilate who have uncontrolled bleeding or require emergency surgery or procedures"– regarding.

Clinical Trial NOC No.:- GCT/59/14

Reference: -Your letter no. BI/DCGI/CT/075/2014 dated 18 Jun 2014 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: 1321.3 Version 1.0 dated 23rd Dec 2013** submitted to this Directorate.

1. Dr. Upendra Kaul, Fortis Escorts Heart Institute, Okhla Road, New Delhi-110025
2. Dr. Iyengar Shamanna Seshadri, Dept. of Cardiology, Manipal Hospital, 98, HAL Airport Road, Bangalore-560017, Karnataka.
3. Dr. Ram Kumar Raja Ram, Dept, of Cardiology, 4-A, Dr. J.J. nagar, Mogappair, Chennai-600037
4. Dr. Gurappa Shetty Gojanur, Narayana Health, Narayana Institute of Cardiac Sciences 258/A, Bommasandra Industrial Area, Anekal Taluk, Bangalore-560099, Karnataka.
5. Dr. Keshava R, Fortis Hospital, 14- Cunningham Road, Sheriff's Chamber, Bangalore-560052, Karnataka.
6. Dr. Bhuwnesh Kandpal, Dept. of Cardiology, Sir Ganga Ram Hospital, Sir Ganga Ram Hospital Marg, Rajinder Nagar, New Delhi-110060
7. Dr. Naresh Kumar Goyal, Max Super Speciality Hospital (A Unit of Max Healthcare Institute Limited) FC 50, C & D Block, Shalimar Bagh, New Delhi-110088
8. Dr. Anupama V Hegde, Dept. of Cardiology, M.S. Ramaiah Clinical Research Centre, 1st Floor, M.S. Ramaiah Advanced Learning Centre, Gate-4,

Gnanagangotri campus, M.S. Ramaiah Medical College & Hospitals, New BEL Road, MSRIT Post, Bangalore-560054, Karnataka.

9. Dr. Varunkumar Bhargava, CARE Hospital, 3-Farmland, Panchsheel Square, Nagpur-440012, Maharashtra.
10. Dr. Brian Pinto, Holy Family Hospital and Medical Research Centre, St. Andrew's Road, Bandra (W), Mumbai-400050
11. Dr. Dhiman Kahali, B.M. Birla Heart Research Centre, 1/1 National Library Avenue Kolkata-700027, West Bengal.
12. Dr. Nakul Sinha, Sahara India Medical Institute, Viraj Khand Gomti Nagar Lucknow-226010
13. Dr. Pravin K. Goel, Dept. of Cardiology, Sanjay Gandhi Post Graduate Institute of Medical Sciences, Raebareli Road, Lucknow-226014, Uttar Pradesh.
14. Dr. Viveka Kumar, Max Super Speciality Hospital, (A Unit of Devki Devi Foundation) 2, Press Enclave Road, New Delhi-110017
15. Dr. Arun Garg, Medanta- The Medicity, Sector-38, Gurgaon Haryana-122001
16. Dr. M.V. Padma Srivasatava, All India Institute of Medical Sciences, Room No. 708, 7th Floor, Dept. of Neurology, C.N. Centre, Ansari Nagar, New Delhi-110029
17. Dr. santosh Kumar Dora, Asian Heart Institute, G/N Block, Bandra-Kurla Complex, Bandra (E), Mumbai-400051
18. Dr. Ravindra Buaditta Gulati, Ruby Hall Clinic, 40, Sassoon Road, Pune-411001
19. Dr. Sharad Chandra, Dept. of Cardiology, King George's Medical University, Lucknow-226003.

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)