47040/20/09/13

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@nb.nic.in, cdscog@gmail.com

File No: CT/11/13-DCG (I) Dated: 31/01/14

To

M/s. Boehringer Ingelheim India Pvt. Ltd.,
1102,11th Floor, Hallmark, Business Plaza,
Gurunanak Hospital, Near Gurunanak Hospital,
Bandra (East), Mumbai – 400 051.

Subject: Permission for conducting a phase III clinical study entitled, "A randomised, double-blind, placebo-controlled, phase III study to evaluate the efficacy and safety of afatinib (BIBW 2992) as adjuvant therapy after chemo-radiotherapy in primary unresected patients with stage III, IVa, or IVb locoregionally advanced head and neck squamous cell carcinoma." – regarding.

Clinical Trial NOC No. GCT/22/13

Reference: Your letter no. BI/DCGI/CT/132/2013 dated 17/09/13 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: 1200.131 versions 1.0 dated 22-01-2013** submitted to this Directorate.

1. Dr Murali Voona, Mahatma Gandhi Cancer research institute, Vishakapatnam-530017, Andhra Pradesh.
2. Dr Radheshyam Naik, HCG Bangalore Institute of Oncology, Kalinga Rao Road, Sampangi Rama Nagar, Bangalore
3. Dr Ravi Mohan, King George Hospital, Cancer Clinical Trial New Medicine Block, King George Hospital, Visakhapatnam - 530 002, AP
4. Dr. Rejnish Kumar, Regional Cancer Centre, Division of Head & Neck Radiation oncology, Medical college campus, Thiruvananthapurum, Kerala--695 011

5. Dr Sachin Almel, P. D. Hinduja National Hospital & Research Center, Mahim (West), Mumbai-400016
6. Dr. Neeti Sharma, Acharya Tulsi Regional Cancer Treatment & Research Institute, Sardar Patel. Medical college and Associated Group of Hospitals, Department of Medical Oncology, Bikaner -334001
7. Dr. A. K. Anand, Max Cancer Centre, Max Super Speciality Hospital Department of Radiation Oncology, New Delhi
8. Dr Minish Jain, Ruby Hall Clinic, 40 Sassoon Road, Pune-411001
9. Dr. Kirushna Kumar, Meenakshi Mission Hospital, Department of Oncology, Lake area, Melur Road, Madurai-625107
10. Dr. Chiramana Haritha, M.S.Patel Cancer Centre, ShriKrishna Hospital & Medical Research Centre, Gokal Nagar, Kramsad
11. Dr Amit Joshi, Tata Memorial Hospital, Dr. E Borges Marg, Parel, Mumbai - 400012, Maharashtra,
12. Dr Ghanshyam Biswas, Sparsh Hospital and Critical care, A-407, Sahed Nagar, Behind Metro House, Bhubaneshwar,
13. Dr Srinivasan V, Kamakshi Memorial Hospital, Pallikkaranai, Chennai
14. Dr. Hemant Malhotra, Birla Cancer Centre, SMS Medical College & Hospital, J L N Marg, Jaipur
15. Dr RK Grover, Delhi State Cancer research Institute, Dilshad Garden, Delhi
16. Dr BK Mohanti, Fortis Memorial research hospital, Gurgaon, Opposite to HUDA city centre metro station
17. Dr Meenu Walia, Dharamshila Hospital and research centre. Vasundra Enclave, Delhi 110095
18. Dr. Sachin Hingmire, Deenanath mangeshkar Hospital and Research Centre, Erandwane, Pune-411004
19. Dr Maheboob Basade, Jaslok Hospital and Research Centre Department of Medical Oncology, 15 - Dr. Deshmukh Marg, Pedder Road, Mumbai
20. Dr Apurva Ashokbhai Patel, The Gujarat Cancer & Research Institute, M. P. Shah Cancer Hospital, Asarwa, Ahmedabad

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)