File No. CT/118/20-DCG(I)

GOVERNMENT OF INDIA
CENTRAL DRUGS STANDARD CONTROL ORGANISATION (Headquarter)
(Directorate General of Health Services)
Ministry of Health & Family Welfare
FDA Bhavan
ITO, Kotla Road New
Delhi - 110002 (Delhi) Phone No.: 91-11-23216367
Fax No.: 91-11-23236973
E-Mail : dci@nic.in

सत्यमेव जयते

File No. CT/20/000119

To,

M/s Bayer Pharmaceuticals Pvt. Ltd.,
Bayer House, Central Avenue, Hiranandani Estate,
Thane , Maharashtra (India) - 400607.

Sir,

With reference to your application No. GCT/CT04/FF/2020/22499 (GCT/118/20) dated 14-11-2020, please find enclosed herewith the permission in Form CT-06 for conduct of clinical trial titled, "**A randomized, double-blind, placebo-controlled Phase 3 study of darolutamide in addition to androgen deprivation therapy (ADT) versus placebo plus ADT in men with metastatic hormone-sensitive prostate cancer (mHSPC)", Protocol number 21140 Original Clinical Study Protocol, dated 24/Sep/2020** under the provisions of New Drugs and Clinical Trial Rules, 2019

The permission granted by the Central Licensing Authority to conduct clinical trial shall be subject to following conditions, namely:-

(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 Licencing 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 Licencing 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 Licencing 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 Licencing 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 Licencing 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 Licencing 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;

(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 Chapter VI and details of compensation provided in such cases shall be intimated to the Central Licencing Authority within thirty working days of the receipt of order issued by Central Licencing 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 Chapter VI and details of compensation provided in such cases shall be intimated to the Central Licencing Authority within thirty working days of receipt of the order issued by the Central Licencing 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 Licencing Authority who may be accompanied by officers of the State Licencing Authority or outside experts as authorised by the Central Licencing 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 Licencing 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 Licencing 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) The permission to initiate clinical trial granted under rule 22 in form CT-06 or automatic approval under rule 23 in Form CT 4A shall remain valid for a period of **two years** from the date of its issue, unless extended by the Central Licencing Authority.

Yours faithfully,

VENUGOPAL
GIRDHARILA
L SOMANI

(Dr. V. G. Somani)
Drugs Controller General (India)
Central Licencing Authority
Stamp

File No. CT/118/20-DCG(I)

FORM CT-06
(See rules 22,25,26,29 and 30)

PERMISSION TO CONDUCT CLINICAL TRIAL OF NEW DRUG OR INVESTIGATIONAL NEW DRUG

1.The Central Licensing Authority hereby permits **M/s. Bayer Pharmaceuticals Pvt. Ltd., Bayer House, Central Avenue Hiranandani Estate, Thane (West) (India) - 400607** to conduct clinical trial of the new drug or investigational new drug as per **Protocol number 21140 Original Clinical Study Protocol, dated 24/Sep/2020** in the below mentioned clinical trial sites [As per Annexure].-

2. Details of new drug or investigational new drug and clinical trial site [As per Annexure].

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.

Place: New Delhi
Date _____

VENUGOPAL GIRDHARILAL SOMANI

Digitally signed by VENUGOPAL GIRDHARILAL SOMANI
DN: c=IN, o=MINISTRY OF HOME AFFAIRS, ou=CDSCO DGHS, postalCode=431401, st=Maharashtra, 2.5.4.20=173d03345df62d489632379a1471bfdae9f0b2bea56c83bfbbe2154e399b1af7, cn=VENUGOPAL GIRDHARILAL SOMANI
Date: 2021.02.04 17:31:07 +05'30'

(Dr. V. G. Somani)
Drugs Controller General (India)
Central Licencing Authority
Stamp

Note: The permission to initiate clinical trial granted under rule 22 in form CT-06 shall remain valid for a period of **two years** from the date of its issue, unless extended by the Central Licencing Authority.

Annexure:

Details of new drug or investigational new drug:

Names of the new drug or investigational new drug	Darolutamide
Therapeutic class:	androgen receptor inhibitor
Dosage form:	Tablets
Composition:	Darolutamide = 300.0000 milligram (mg) In House Specification Active
Indications:	metastatic hormone-sensitive prostate cancer

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Details of clinical trial site:

Sr. No.	Names and address of clinical trial site	Ethics committee details	Name of investigator
1.	HCG Cancer Center, Sola Science City Road, Nr. Sola Bridge, S.G. Highway, Ahmedabad-380060, Gujarat, India	HCG Multispecialty Ethics Committee, HCG Hospitals-Ahmedabad, 1, Maharashtra Society, Mithakhali, Ellisbridge, Ahmedabad-380006 ECR/92/Inst/GJ/2013/RR-19	Dr. Ashish Kaushal
2.	Christian Medical College and Hospital, Department of Medical Oncology, Ludhiana-141008, Punjab, India	Institutional Ethics Committee, Christian Medical College and Hospital, Brown Road, Ludhiana, Punjab-141008 ECR/120/Inst/PB/2013/RR-19	Dr. Ashwin Philips
3.	Regional Cancer Centre, Medical College Campus, Trivandrum-695011, Kerala, India	Human Ethics Committee, First Floor, Regional Cancer Centre, Medical College Road, Thiruvananthapuram, Kerala-695011 ECR/21/Inst/Ker/2013/RR-19	Dr. Francis V. James
4.	Sparsh Hospital & Critical Care (P) Ltd., Plot no-A/407, Sahid Nagar, Bhubaneshwar-751007, Odisha, India	Institutional Ethics Committee, Sparsh Hospital & Critical Care (P) Ltd, A/407, Saheed Nagar, Bhubaneswar, Odisha, India, 751007 ECR/68/Inst/OR/2013/RR-19	Dr. Ghanashyam Biswas
5.	All India Institute of Medical Sciences, Room # 133,Dr. Bhimrao Ambedkar Institute of Rotary Cancer Hospital (Dr BRA IRCH), Ansari Nagar, New Delhi-110029, Delhi	Institute Ethics Committee, Room # 102, All India Institute of Medical Science, Ansari Nagar, New Delhi-110029 ECR/538/Inst/DL/2014/RR-20	Dr. Haresh KP
6.	Kailash Cancer Hospital and Research Centre, Muniseva Ashram, Goraj, Waghodia, Vadodara-391760, Gujarat, India	Institutional Review Board, Kailash Cancer Hospital and Research Centre, Muniseva Ashram, Goraj, Waghodia, Vadodara-391760, Gujarat, India ECR/49/Inst/GJ/2013/RR-19	Dr. Niraj Bhatt
7.	Bhagwan Mahaveer Cancer Hospital and Research Centre, Jawahar Lal Nehru Marg, Jaipur-302017, Rajasthan, India	Institutional Ethics Committee, Bhagwan Mahaveer Cancer Hospital and Research Centre, Jawahar Lal Nehru Marg, Jaipur (Raj) – 302017,India ECR/19/Inst/RJ/2013/RR/20	Dr. Pawan Agarwal
8.	HealthCare Global Enterprises Ltd, No.8, HCG Towers, P. Kalinga Rao Road, Sampangi	HCG-Central Ethics Committee, #8, P. Kalinga Rao Road, Sampangi Ram Nagar,	Dr. Raghunath S K

	Ram Nagar, Bengaluru-560027, Karnataka, India	Bangalore-560027 ECR/386/Inst/KA/2013/RR-19	
9.	HCG Manavata Cancer Centre, Behind Shivang Auto, Mumbai Naka, Nashik-422002, Maharashtra, India	Manavata Clinical Research Institute, Ethics Committee, HCG Manavata Cancer Centre, Behind Shivang Auto, Mumbai Naka, Nashik-422002, Maharashtra, India ECR/500/Inst/MH/2013/RR-20	Dr. Rajnish Nagarkar
10.	Rajiv Gandhi Cancer Institute and Research Centre, Sector-5, Rohini, New Delhi-110085	Institutional Review Board, Rajiv Gandhi Cancer Institute and Research Centre, Sector-5, Rohini, New Delhi-110085 ECR/10/Inst/DL/2013/RR-19	Dr. Sudhir Kumar Rawal
11.	Tata Medical Center, 14, MAR(E-W), DH Block (Newtown), Action Area I, Newtown, Kolkata, West Bengal-700160	Tata Medical Center – Institutional Review Board, 14 MAR(E-W), Rajarhat, Newtown, Kolkata, 700160, West Bengal, India ECR/269/Inst/WB/2013/RR-19	Dr. Sujoy Gupta