



File No. BIO/CT/22/000005

Dated 25-Mar-2022

To,
M/s Enzene Biosciences Ltd.,
Plot no A 22, A/1/2 Chakan Industrial Area,
Phase 2, Khalumbre Chakan,
Pune – 410501,India.

Subject: Application for grant of permission to conduct Phase IV clinical trial entitled “A phase IV, open label, single-arm, multi-centre clinical study to evaluate the safety and efficacy of Denosumab in patients with postmenopausal osteoporosis” as per Protocol Number: ALK23/DEN3 Version: 2.1 Date: 10-Mar-2022 -regarding

Ref.:Your Application No. BIO/CT04/FF/2022/30050 dated 14-Jan-2022

Sir,

With reference to your Application No.: BIO/CT04/FF/2022/30050 dated 14-Jan-2022, please find enclosed herewith the permission in Form CT-06 for conduct of subject Phase IV clinical trial under the provisions of New Drugs and Clinical Trial Rules, 2019.

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

- (I) The firm should submit the insurance certificate for the subject trial before initiation of the trial to this office.
- (II) 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 Licensing Authority under rule 8;
- (III) 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;

- (IV) 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;
- (V) The Central Licensing 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;
- (VI) 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;

- (VII) 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;
- (VIII) Status of enrolment of the trial subjects shall be submitted to the Central Licensing Authority on quarterly basis or as appropriate as per the duration of treatment in accordance with the approved clinical trial protocol, whichever is earlier;
- (IX) Six monthly status report of each clinical trial, as to whether it is ongoing, completed or terminated, shall be submitted to the Central Licensing Authority electronically in the SUGAM portal;
- (X) In case of termination of any clinical trial the detailed reasons for such termination shall be communicated to the Central Licensing Authority within thirty working days of such termination;
- (XI) 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 Licensing 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;
- (XII) 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 Licensing Authority within thirty working days of the receipt of order issued by Central Licensing Authority in accordance with the provisions of the said Chapter;
- (XIII) 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 Licensing Authority within thirty working days of receipt of the order issued by the Central Licensing Authority in accordance with the provisions of the said Chapter;
- (XIV) The premises of the sponsor including his representatives and clinical trial sites, shall be open for inspection by officers of the Central Licensing Authority who may be accompanied by officers of the State Licensing Authority or outside experts as authorised by the Central Licensing 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;
- (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 Licensing 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.

Yours faithfully,

**VENUGOPAL
G SOMANI**

Digitally signed by VENUGOPAL G SOMANI
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(Dr. V.G. Somani)

Drugs Controller General (India)

FORM CT-06

(See rules 22, 25, 26, 29 and 30)

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

The Central Licencing Authority hereby permits M/s Enzene Biosciences Ltd., Plot no A 22, A/1/2 Chakan Industrial Area, Phase 2, Khalumbre Chakan, Pune – 410501, India to conduct clinical trial of the new drug or investigational new drug study titled “A phase IV, open label, single-arm, multi-centre clinical study to evaluate the safety and efficacy of Denosumab in patients with postmenopausal osteoporosis” as per Protocol Number: ALK23/DEN3 Version: 2.1 Date: 10-Mar-2022 in the below mentioned clinical trial sites.

2. Details of 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: 25-Mar-2022

**VENUGOPAL G
SOMANI**

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(Dr. V.G. Somani)

Drugs Controller General (India)
Central Licencing Authority

CDSCO

CDSCO

MINISTRY OF HEALTH, GOVERNMENT OF INDIA
सत्यमेव जयते

Annexure:**Details of new drug or investigational new drug:**

Names of the new drug or investigational new drug:	Denosumab Injection 60 mg in 1mL	
Therapeutic class:	RANK ligand inhibitors	
Dosage form:	single use pre-filled syringe for subcutaneous injection	
Composition:	Name of Ingredients	Quantity per ml
	Denosumab IH	60mg
	Sorbitol JP, Ph.Eur, BP, NF, IP	47.0mg (4.7%)
	Glacial acetic acid Ph.Eur, BP, JP, USP, IP	1.0mg(17 mM)
	Polysorbate 20 B.P, EP, JP, NF,IP	0.1 mg(0.01%)
	Sodium hydroxide IP,USP,BP,EP,JP,NF	q.s. to adjust pH to 5.2
	Water for Injection IP, USP	q.s to 1ml
Indications:	Postmenopausal Osteoporosis	

Details of clinical trial site:

S. No.	Name and Address of Clinical Trial Site	Ethics Committee Details	Name of Principal Investigator
1	Medical College and hospital ,MCH building, 4thFloor, 88 College street,Kolkata-700073 West Bengal	Institutional Ethics committee for Human Research ,Medical College and hospital, 88 College street, Kolkata -700073 EC Reg no: ECR/287Inst/WB/2013/RR-19	Dr. Kaushik Basu
2	Sancheti Institute for orthopaedics and Rehabilitation, 16, Shivaji Nagar, Pune-411005	Institutional Ethics committee Sancheti Institute for Orthopaedics and Rehabilitation 16, Shivajinagar, Pune 411005 EC Reg no: ECR/90/Inst/MH/2013 /RR-16	Dr. Parag Sanchet
3	Medilink Hospital and research centre, 1st floor, 132 ft ring road, Satelite, Ahmedabad- 380015	Medilink Ethics Committee, 1 st floor, 132 ft ring road, Satelite, Ahmedabad-380015. EC Reg. No.: ECR/344/Inst/GJ/2013 /RR-19	Dr. Sunil Maheshwari
4	Department of orthopaedics, All India Institute of Medical sciences, Bhubneshwar, Sijua, Patrapada, Bhubneshwar- 751019	Institutional Ethics committee, All India Institute of Medical Sciences, Sijua, PO-dumduma, Bhubneshwar, Odisha-751019 EC Reg no: ECR/534/Inst/OD/201 4/RR-17	Dr. Sujit Kumar Tripathy
5	Department of Orthopedics, St. John's Medical College and Hospital, Banglore-560034	Institutional Ethics committee, St. John's medical college, sarjapur road, Bangalore,560034 EC Reg no: ECR/238/Inst/KA/201 3/RR-19	Dr. Joe Joseph Cherian
6	Getwell Hospital and Research Institute, 20/1, Khare Marg, Dhantoli, Nagpur- 440012	Getwell Institutional Ethics Committee, Getwell Hospital and Research Institute, 20/1, Khare Marg, Dhantoli, Nagpur-440012 EC Reg no: ECR/109/Maha/2013/ RR-19	Dr. Jayashri Shembalkar

7	Prime Care Hospital, 4th floor, Rudra Arcade, B-wing, Helmet Char rasta, Near Mayflower Hospital, Drive in Road, Ahmedabad- 380052	Highland Super Speciality Hospital, Kolshet Road, above D-Mart, Dhokali, Thane West, Thane, Maharashtra 400607 EC Reg no: ECR/344/Inst/GJ/2013 /RR-19	Dr. Pankaj Agarwal
8	Department of endocrinology, Super Speciality Block, King George Hospital, Maharani-peta, Visakhapatnam- 530002, Andhra Pradesh	Institutional ethics committee, King George Hospital, Maharani-peta, Collector office junction, Visakhapatnam,530002, Andhra Pradesh EC Reg no:ECR/197/Inst/KGH/20 13/RR-20	Dr. Bongi Vivekanand
9	Highland Super Speciality Hospital, Kolshet Road, above D-Mart, Dhokali, Thane West, Thane, Maharashtra 400607	Vedant Hospital Institutional Ethics Committee, Vedant Hospital, Ghodbandar Road, Kasarvadavalli, Thane west-400615 EC Reg no: ECR/1154/INST/MH/ 2018	Dr Babasaheb Dada Chavan

