

File No. SND/CT/20/000061
Government of India
Directorate General of Health Services
Central Drugs Standard Control Organisation
(Subsequent New Drugs Division)

FDA Bhawan, Kotla Road,
New Delhi-110002
Dated: 02-March-2021

To,
M/s Torrent Pharmaceuticals Limited,
Nr. Dinesh Hall, Ashram Road, Ahmedabad
Gujarat- 380009
India

Subject: Permission to conduct Phase IV Clinical trial of Tapentadol Hydrochloride Nasal Spray 225 mg/ml "Title- A Multicentric, Open-label, Single-arm, Non-comparative, Clinical Trial to Evaluate Safety and Efficacy of Tapentadol Nasal Spray in Patients with Moderate to Severe Pain" (Vide Protocol No. CT/TAPE/PAIN/20/04_01, Version No: 02, Dated 27/10/2020) - Reg.

CT NOC No.: SND/CT/20/000061

Sir,

With reference to your Application No. SND/CT04/FF/2020/22176 dated 10.10.2020 please find enclosed herewith the permission in Form CT-06, CT NOC No. **SND/CT/20/000061** to conduct the subject mentioned clinical trial under the provisions of New Drugs and Clinical Trial Rules, 2019.

This permission is subject to the conditions, as mentioned below.

Yours faithfully,

VENUGOPAL
GIRDHARILAL
SOMANI

Digitally signed by VENUGOPAL
GIRDHARILAL SOMANI
DN: c=IN, o=MINISTRY OF HOME AFFAIRS,
ou=CDSCO DGHS, postalCode=431401,
st=Maharashtra,
2.5.4.30=17580134548626480632379a1471b,
f04c9f9a2de55c83f0bc2134c39961af7,
cn=VENUGOPAL GIRDHARILAL SOMANI
Date: 2021.03.02 16:38:01 +05'30'

(Dr. V. G. Somani)
Central Licensing Authority

Conditions of Permission

- (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

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 of the New Drugs and Clinical Trials Rules, 2019;
- (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 the Chapter VI of the said Rules 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 the Chapter VI and details of compensation provided in such cases shall be

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) Informed Consent Documents (ICD) viz. Patient Information Sheet (PIS) and Informed Consent Form (ICF) complete in all respect & must be got approved from the respective Ethics committee and submitted to CDSCO before enrolling first subject at the respective site.

FORM CT-06

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

**PERMISSION TO CONDUCT CLINICAL TRIAL OF NEW DRUG OR
INVESTIGATIONAL NEW DRUG****CT NOC NO.: SND/CT/20/000061**

The Central Licensing Authority hereby permits **M/s Torrent Pharmaceuticals Limited Nr. Dinesh Hall, Ashram Road, Ahmedabad Gujarat- (India)- 380009** to conduct clinical trial of the new drug or investigational new drug as per **Protocol No: CT/TAPE/PAIN/20/04_01, Version No: 02, Dated 27/10/2020** in the below mentioned clinical trial sites.

2. Details of new drug or investigational new drug:

Names of the new drug:	Tapentadol Hydrochloride Nasal Spray 225 mg/ml
Therapeutic class:	Analgesic
Dosage form:	Nasal Spray
Composition:	Each Spray (0.1 ml) contains: Tapentadol Hydrochloride.....22.5 mg Benzalkonium chloride (50%).....0.02%w/v (As Preservative)
Indications:	For treatment of moderated to severe post-operative pain in hospital admitted patients.

Details of clinical trial sites

Sr. No.	Name of Principal Investigator & Trial sites	Ethics Committee Name/Registration Number
1	Dr. Yash Sanghavi Anand Surgical Hospital Private Limited, Memco Cross RD, Naroda Road, Naroda, Ahmedabad - 382345, Gujarat, India.	Kanoria Ethics Committee Kanoria Hospital & Research Centre, Airport - Gandhinagar Highway, Village: Bhat, Dist. Gandhinagar-3 82428, Gujarat, India ECR/620/Inst/GJ/2014/RR-20
2	Dr. Sushil Mankar NKP Salve Institute of Medical Sciences and Research Centre and Lata Mangeshkar Hospital, Digdoh Hills, Hingna Road, Nagpur-440019.	Institutional Ethics Committee NKP Salve Institute of Medical Sciences and Research Centre and Lata Mangeshkar Hospital NKP Salve Institute of Medical Sciences and Research Centre and Lata Mangeshkar Hospital, Digdoh Hills, Hingna Road, Nagpur-440019 ECR/88/Inst/MH/2013/RR-I 9
3	Dr. Jaimin Vashi Shubham Multispecialty Hospital, ABC Complex, Near Rabari Colony Cross Road. Amraiwadi, Ahmedabad-380026.	Shubham Institutional Ethics Committee Shubham Multispecialty Hospital, 4h Floor, ABC Complex, Rabari Colony Char Rasta, N. H. No. 8, Amraiwadi, Ahmedabad-380026 ECR/853/Inst/GJ/2016/RR 20

4	Dr. Valsa Abraham Christian Medical &Hospital. Department of Anaesthesia, Christian Medical &Hospital, Brown Road, Ludhiana, Punjab, 141008.	Institutional Ethics Committee Christian Medical &Hospital, Brown Road, Ludhiana, Punjab, 141008. ECR/120/Inst/PB/2013/RR-19
5	Dr. Suraj Kumar Pattnayak Department of Surgery, Government Medical College & Govt. General Hospital (Old RIMSGGH), Srikakulam-532001, Andhra Pradesh	Institutional Ethics Committee Government Medical College & Govt. General Hospital, Srikakulam-532001, Andhra Pradesh. ECR/429/Inst/AP/2013/RR-20
6	Dr. Chandan Kumar Department of Orthopedics, GSVM Medical College, Swaroop Nagar, Kanpur-208002, UP	Ethics Committee, GSVM Medical College Room no 125 1 st Floor GSVM Medical college, Swaroop Nagar, Kanpur-208002, UP ECR/680/Inst/UP/2014/RR-17
7	Dr. Jadhav Jagdish Shankarrao Lifepoint Multispecialty Hospital, 145/1, Mumbai Banglore Highway, Near Hotel Sayaji, Wakad, Pune-411057, Maharashtra.	LPR Ethics Committee Lifepoint Multispecialty Hospital, 145/1, Mumbai Banglore Highway, Near Hotel Sayaji, Wakad, Pune- 411057, Maharashtra. ECR/751/INST/MH/2015/RR-18
8	Dr. Praveen Talawar Department of Anaesthesiology, AIIMS, Rishikesh	Institute Ethics Committee AIIMS, Rishikesh ECR/736/Inst/UK/2015/RR-18
9	Dr. Prashant Chandrakant Kamble Seth G.S. Medical College & KEM Hospital, Parel, Mumbai-400012	Institutional Ethics Committee UG PG Hostel Ground Floor KEM Hospital, Parel-Mumbai-400012 ECR/417/Inst./MH/2013
10	Dr. Mahakalkar Chandrashekhar Chintamanrao Acharya Vinoba Bhave Rural Hospital, Jawaharlal Nehru Medical College, Datta Meghe Institute of Medical Sciences (DU), Sawangi (M), Wardha- 442004, Maharashtra	Institutional Ethics Committee Datta Meghe Institute of Medical Sciences (DU), Research House, Near Food Court, , Datta Meghe Institute of Medical Sciences (DU), Sawangi (M), Wardha-442004, Maharashtra. ECR/440/Inst/MH/2013/RR-19

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.

New Delhi

Date: 02-March-2021

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=173d03345d62d489632379a14710
f04e90b28e056c33b0e2154e33961a7f,
cn=VENUGOPAL GIRDHARILAL SOMANI
Date: 2021.03.02 16:38:18 +05'30'

(Dr. V. G. Somani)
Central Licensing Authority