檔 號: 保存年限:

衛生福利部食品藥物管理署 函

地址:11561 臺北市南港區昆陽街161-2號

聯絡人:蘇子婷

聯絡電話: 0227877148 傳真: 0227877178

電子信箱:daisyhaha@fda.gov.tw

受文者:中華民國西藥代理商業同業公會

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速別:普通件

密等及解密條件或保密期限:

附件:原料藥廠違反GMP警訊乙份(A21020000I105110391600-1.pdf)

主旨:歐洲理事會European Directorate for the Quality of Medicines & HealthCare (EDQM) 通報印度原料藥廠「Wo ckhardt Limited」(廠址:Plot No. L-1, M. I. D. C. Ar ea Chikalthana, India 431 210 Aurangabad, Maharash tra) 嚴重違反GMP乙案,詳如說明段,請轉知所屬會員知照。

說明:

一、英國衛生主管機關Medicines and Healthcare Products Regulatory Agency (MHRA) 查核旨揭原料藥廠,判定嚴重違反GMP,EDQM正式發布「DECISION TO SUSPEND A CERT IFICATE OF SUITABILITY」,凍結(SUSPEND)「OXYBUTY NIN HYDROCHLORIDE」之CERTIFICATE OF SUITABILITY(CEP)品質證明文件2年,待再次通過查廠後始得恢復。因前述處置將屆滿2年,EDQM另於105年6月28日發布旨揭工廠「WITHDRAWAL OF CERTIFICATES OF SUITABILITY」,吊銷(WITHDRAW)上述品項之CEP證明文件。

二、英國MHRA亦於104年1月16日發布「STATEMENT OF NON-COM





PLIANCE WITH GMP」(詳如附件),所有該廠製造之原料 藥品項皆受影響。

- 三、承上,且英國MHRA已啟動相關後續處置,包括:
 - (一)使用旨揭藥廠之原料藥者,應考慮變更原料來源(Variation of the marketing authorisations)。
 - (二)英國MHRA未啟動回收作業,仍建議使用受影響原料藥之 產品應評估是否回收。
- 四、鑒於旨揭原料藥廠之製造品質無法符合GMP之要求,可能 對藥品製造品質帶來影響與危害,請轉知所屬會員釐清相 關輸台製劑產品是否使用旨揭原料藥廠所生產原料藥,並 應依說明段三所述辦理。

正本:中華民國西藥商業同業公會全國聯合會、中華民國西藥代理商業同業公會、台北市西藥代理商業同業公會、中華民國開發性製藥研究協會、中華民國藥品行銷暨

副本: 〒2015-02-19文 交 13:196:37章







Medicines and Healthcare Products Regulatory Agency

Report No: UK API 8913 Insp GMP 89/3/18322-0006 - NCR

STATEMENT OF NON-COMPLIANCE WITH GMP

Exchange of information between National Competent Authorities (NCAs) of the EEA following the discovery of serious GMP non-compliance at a manufacturer

Part 1

Issued following an inspection in accordance with:

Art. 111(7) of Directive 2001/83/EC as amended

The competent authority of United Kingdom confirms the following

The manufacturer: WOCKHARDT LIMITED

Site address: L-1, MIDC, JALGAON ROAD, CHIKALTHANA, AURANGABAD, IN-431 210, India

From the knowledge gained during inspection of this manufacturer, the latest of which was conducted on **2013-11-15**, it is considered that it does not comply with the Good Manufacturing Practice requirements referred to in

- The principles and guidelines of Good Manufacturing Practice laid down in Directive 2003/94/EC
- The principles of GMP for active substances referred to in Article 47 of Directive 2001/83/EC.



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Signatory: Confidential

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The statement of non-compliance referred to invaragraph 111(7) of Directive 2001/83/EC and 80(7) of Directive 2001/82/EC, as amended, shall also be required for imports coming from third countries into a Member State.

Part 2

Human Medicinal Products

1 NON-COMPLIANT MANUFACTURING OPERATIONS

Include total and partial manufacturing (including various processes of dividing up, packaging or presentation), batch release and certification, storage and distribution of specified dosage forms unless informed to the contrary;

1.4	Other products or manufacturing activity				
	1.4.1 Manufacture of				
	1.4.1.4 Other: Active starting materials (See part 3 for more details)(en)				

4. Non-Compliant Other Activities - Active Substances:

All active substances manufactured at this facility

Part 3

1. Nature of non-compliance:

1. A critical deficiency was cited with regards to data integrity of GMP records, entries were seen to be made when personnel were not present on site, documentation was seen that was not completed contemporaneously despite appearing to be completed in this manner. 2. A second critical deficiency was cited regarding potential product contamination, this included the use of inappropriate materials close to product e.g. asbestos coated PTFE seals for centrifuge manways. 3. A major deficiency was cited with regards to equipment and facility, maintenance, design and qualification. Examples included, inappropriate pressure differentials that were not in line with the original design but had not been changed using change control, cleaning validation that was not sufficiently robust to confirm cleaning practices and maintenance issues, such as the failure to spark test glass lined reactor vessels for integrity especially following maintenance.

Action taken/proposed by the NCA

Requested Variation of the marketing authorisation(s)

Marketing Authorisation Holders should be requested to remove the site from applicable authorisations by variation

Recall of batches already released

No recall action is proposed.

Additional comments

The initial statement of serious non-compliance permitted continued supply of critical APIs. Following a company decision to cease supply of API to the EU market, this statement of non-compliance was updated to cover all active substances in January 2015. Re-inspection by an EU national; competent authority will be required prior to gaining approval to supply the EU market in future.

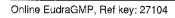
Products manufactured at site, if known	Products	Dosage Form	Reference Member State, National or EMA
Human Medicinal	Captopril		CEP 1998-052
Products	Oxybutynin HCI		CEP 2003-020
	Pramipexole Dihydrochloride monohydrate		CEP 2012-149

Name and signature of the authorised person of the Competent Authority of United Kingdom

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Medicines and Healthcare Products Regulatory Agency

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Issuance Date: 2015-01-16

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