Certificate of Analysis

Bestätigung der ordentlichen Prüfung

Product: Produkt	Norspan 5 CL VK2		
		Article-No.: Artikel-Nr.	3319713
Country: Importland	Chile	**Order No.:** Auftrags-Nr.	1060689
LTS Lot-No.: LTS Ch.-B.	7210227AA	**Customer-Lot No.:** Kunden Ch.-B.	70967A114
Date of manufacture: Herstelldatum	05.10.2017	**Expiry Date:** Verfalldatum	09.2019

Test parameters Prüfpunkte	**Specified** Soll	**Result** Ergebnis
Appearance	Corresponds to requirement	corresponds
Identity by Marquis	Violet spot after application of one drop of Marquis' reagent	corresponds, violet spot
Identity by HPLC	The ratio of the retention times of the peaks of active principle and standard should range between 0.95 - 1.05.	Retention time corresponds
Dimension reservoir 25x25 +- 0.5mm	Corresponds to requirement	corresponds
Dimension coloured web 45x45 +- 2.0mm	Corresponds to requirement	corresponds
Content / mean	4.9 - 5.4 mg/TTS	5.2 mg/TTS
Content uniformity EP 2.9.6 test C	Average content of 10 units between 90-110% of content stated on label claim; individual content of each unit is between 75-125% of average content	corresponds to EP 2.9.6 test C
Content uniformity EP 2.9.40 acc. value	nmt. 15.0	4.2
Content uniformity EP 2.9.40 level	Content uniformity EP 2.9.40	corresponds EP 2.9.40 level 1 (n=10)
In vitro release 0,5h / mean	15 - 35 %	27 %
In vitro release 0,5h / min	nlt. 15 %	27 %
In vitro release 0,5h / max	nmt. 35 %	28 %
In vitro release 2h / mean	30 - 70 %	57 %
In vitro release 2h / min	nlt. 30 %	56 %
In vitro release 2h / max	nmt. 70 %	58 %
In vitro release 24h / mean	nlt. 80 %	102 %
In vitro release 24h / min	nlt. 80 %	101 %
In vitro release 24h / max	nlt. 80 %	103 %
In vitro release 0,5h / mean	120 - 280 µg/cm2	217 µg/cm2
In vitro release 0,5h / min	nlt. 120 µg/cm2	213 µg/cm2
In vitro release 0,5h / max	nmt. 280 µg/cm2	221 µg/cm2
In vitro release 2h / mean	240 - 560 µg/cm2	458 µg/cm2
In vitro release 2h / min	nlt. 240 µg/cm2	452 µg/cm2

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Test parameters Prüfpunkte	**Specified** Soll	**Result** Ergebnis
In vitro release 2h / max	nmt. 560 µg/cm2	462 µg/cm2
In vitro release 24h / mean	nlt. 640 µg/cm2	818 µg/cm2
In vitro release 24h / min	nlt. 640 µg/cm2	808 µg/cm2
In vitro release 24h / max	nlt. 640 µg/cm2	825 µg/cm2
In vitro release 0,5h / mean	0.75 - 1.75 mg/TTS	1.35 mg/TTS
In vitro release 0,5h / min	nlt. 0.75 mg/TTS	1.33 mg/TTS
In vitro release 0,5h / max	nmt. 1.75 mg/TTS	1.38 mg/TTS
In vitro release 2h / mean	1.5 - 3.5 mg/TTS	2.9 mg/TTS
In vitro release 2h / min	nlt. 1.5 mg/TTS	2.8 mg/TTS
In vitro release 2h / max	nmt. 3.5 mg/TTS	2.9 mg/TTS
In vitro release 24h / mean	nlt. 4.0 mg/TTS	5.1 mg/TTS
In vitro release 24h / min	nlt. 4.0 mg/TTS	5.0 mg/TTS
In vitro release 24h / max	nlt. 4.0 mg/TTS	5.2 mg/TTS
Rel.subst.: Norbuprenorphine	nmt. 1.0 %	< 0.1 %
Rel.subst.: Buprenorphine-N-oxide	nmt. 0.5 %	< 0.1 %
Rel.subst.: Individual unknown	nmt. 0.5 %	< 0.1 %
Rel.subst.: Total	nmt. 1.5 %	< 0.1 %
Pouch integrity	Tightness of pouches corresponds	corresponds = Pouches tight
Imprint TTS / Pouch	Imprint consistent and legible	Imprint corresponds
Peel adhesion / mean	3.0 - 50.0 Newton/TTS	14.5 Newton/TTS
Release strength / mean	10 - 200 cN/TTS	40 cN/TTS
Residual solvents: Toluene	nmt. 0.002 %	< 0.001 %
Residual solvents: Isopropanol	nmt. 0.002 %	< 0.001 %
Residual solvents: Ethylacetate	nmt. 0.070 %	0.006 %
Residual solvents: n-Heptane	nmt. 0.050 %	0.003 %
Residual solvents: Acetyl acetone	nmt. 0.020 %	< 0.001 %
Residual solvents: Ethanol	nmt. 0.050 %	0.002 %
Microbiology - periodical monitoring	MIBI - periodical monitoring - performed once a year	This batch was not tested

Used API	6506481, Buprenorphine Geel 2, A17AB0438, Janssen Pharmaceutica
Used API	6506481, Buprenorphine Geel 2, A17AB0456, Janssen Pharmaceutica

This report was approved and released by the responsible Head of QC electronically, and is valid without manual signature.

Certificate of Compliance

Bestätigung der ordentlichen Herstellung

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		Article-No.: Artikel-Nr.	3319713
Country: Importland	Chile	**Order No.:** Auftrags-Nr.	1060689
LTS Lot-No.: LTS Ch.-B.	7210227AA	**Customer-Lot No.:** Kunden Ch.-B.	70967A114
Date of manufacture: Herstelldatum	05.10.2017	**Expiry Date:** Verfalldatum	09.2019

Manufacturing site: Herstellungsstätte/Ort	LTS Lohmann Therapie-Systeme AG, Lohmannstr.2, 56626 Andernach, Germany
Manufacturing license No.: Nr. Herstellungserlaubnis	DE_RP_01_MIA_2011_0023
GMP Certificate No.: GMP Zertifikatsnummer	DE_RP_01_GMP_2016_0024

OOS / Deviations, OOS / Abweichungen	**Number, Nummer**
No confirmed OOS-Results / Keine bestätigten OOS-Ergebnisse	-
No reportable Deviations / Keine zu berichtenden Abweichungen	-

Customer specific Info / Kundenspez. Infor	**Number, Nummer**
Quantity released:	15980 ST

Registration No:	F-22124/15
Strength/Potency:	5 mg/TTS 5 µg/1 h Buprenorphin
Dosage Form:	TTS
Package size and type:	2 Pouches per box
Customer Material No:	201053/95005088

This document was signed electronically and is valid without manual signature.

Release for shipment

I hereby certify that the above information is authentic and accurate.
This batch of product has been manufactured, including packaging/labelling and quality control at the above mentioned site in full compliance with the GMP requirements of the local Regulatory Authority and with the specifications in the Marketing Authorization of the importing country.

The batch processing, packaging and analysis records were reviewed and found to be in compliance with GMP.

Date	**Time**	**Signatur Qualified Person**
16.01.2018	14:38:37	Dr. Anita Noeske

LTS Lohmann Therapie-Systeme AG Lohmannstr. 2 D-56626 Andernach
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