

<b>Name Manufacturer</b>	<b>MEDITES PHARMA, spol. s r.o.</b>
<b>Department</b>	Quality Control
<b>Certificate</b>	Certificate of Analysis

We MEDITES PHARMA, spol. s r.o. hereby declare that:

**1. Product identification**


<b>Product name:</b>	CITRASOL 4%
<b>REF:</b>	601031
<b>Lot Number</b>	A0043020
<b>Manufacturing Date</b>	2020 09 26
<b>Expiry Date</b>	2022 08
<b>Packaging type</b>	18 x 250 ml in a cardboard box
<b>Packaging material</b>	non – PVC bag; connectors: Tulip red cap, injection connector secondary packaging
<b>Unit volume</b>	250 ml

**2. Finished product Analysis Compliance**


Each production batch was sampled, tested and released to MEDITES PHARMA in accordance with specification NR - C115 CITRASOL.

<i>Parameter</i>	<i>Specification</i>	<i>Result</i>	<i>Reference</i>
<b><i>Physical and Chemical parameters</i></b>			
Identification of Active Compound	Identity test for presence of Sodium ( Na <sup>+</sup> ) and Citrates has to be satisfactory (passed).	Comply	PRG452
Clarity	The product must be clear.	Comply	PRG452
Actual acidity – pH	6.7 – 7.4.	Comply	PRG452
Aluminium	< 30 µg/L	Comply	PRG452
Available volume	The volume of the solution in the bag must not be lower than the volume printed on the PP bag.	Comply	PRG452
Assay of sodium	9.08 - 9.54 g/L	Comply	PRG452
Assay of citrates	24.3 - 26.8 g/L	Comply	PRG452
Contamination by particles under the visibility level	Particles of size ≥ 10 µm max. 25/ml. Particles of size ≥ 25 µm max. 3/ml.	Comply	P/407/20
<b><i>Microbiological parameters</i></b>			
Limit test for bacterial endotoxins	< 0.25 IU/ml	Comply	2192/20
Sterility	The product has to be sterile.	Comply	20/0583

### 3. QC Final Approval

Designated Authority	Name	Job title	Signature	Date
Prepared By	Petra Šrámková	QA / QC Specialist		
Reviewed By	Radmila Doleželová	QA / QC manager		23.11.2020

### 4. Final Release Approval

Designated Authority	Name	Job title	Signature	Date
Reviewed By	Radmila Doleželová	QA / QC manager		19.11.2020

**MEDITES PHARMA**  
spol. s r. o.  
756 61 Rožnov pod Radhoštěm  
ORJ



## Certificate of Analysis

Project No:	PRG452	Issue Date:	09/10/2020
Client:	MEDITES PHARMA, spol. s r.o., Rožnov pod Radhoštěm	Laboratory:	ALS Czech Republic, s.r.o.   Pharmaceutical
Address:	1. máje 2625 Rožnov pod Radhoštěm 756 61 Czech Republic		Na Harfe 336/9 - Prague 9 - Vysocany Czech Republic 190 00
Contact:	Radmila Dolezelova	Contact:	F&P Client Service
Email:	radmila_dolezelova@meditespharma.cz	Email:	czsupport.pharma@alsglobal.com
Telephone:	571115905	Telephone:	+420 226 226 998
Order No:		Date of test:	05/10/2020 - 08/10/2020
Sampled By		Certificate No.:	TPRG000437-1 Final

### General Comments:

This report shall not be reproduced except in full, without prior written approval from the laboratory.  
The laboratory declares that the test results relate only to the listed samples.  
The information about sample representativity is not mentioned.  
The sample expiry date is not mentioned.

### Responsible person:

Responsible for accuracy of reported results are persons listed below. The results were approved and authorized for reporting according to the procedures specified in the document CZ\_SOP\_D06\_GMP\_02 Processing of GMP Project which is in accordance with the procedures specified in FDA document CFR Title 21, Part 11.

Name/Position	Authorization Date
Eva Hudečková, Pharma Section Supervisor	09/10/20 07:43
Miroslava Juričová, Metals Section Coordinator	08/10/20 17:22



Issue Date: 09/10/2020

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Project: PRG452

Client: MEDITES PHARMA, spol. s r.o., Rožnov pod Radhoštěm



Client Sample ID	Prod: CITRASOL 4% Citrasol 4% 250 ml Rcvd Dt: 01/10/2020 Sched Dt: 01/10/2020 Sample Date: 01/10/2020 Description: Citrasol 4% (250 ml); č.š. A0043020					
Laboratory Sample ID	PRG680					
Client Sampling Date/Time	01/10/2020 00:00					
Test	Method	Result	Units	Specification	Eval	Flag
Al	METL-ALMSFXVJ0-PRG	<10	µg/L	<30 µg/L	Pass	\$2
Na	METL-NAMSFVC2-PRG	9.40	g/L	9.08 - 9.54 g/L	Pass	\$2
Na - ID	METL-IDENOES-NA-PRG	Pass	-	Pass	Pass	\$2
Citrates	PHAR-CITRA2#M1-PRG	25.6	g/L	24.3 - 26.8 g/L	Pass	\$1
Actual acidity (pH)	PHAR-PHP2-ELE#M1-PRG	7.0	-	6.7 - 7.4	Pass	\$1
Citrates	PHAR-CITR2-ID#MC-PRG	Pass	-	Complies to test	Pass	\$1
Clarity	PHAR-SENZL1#MC-PRG	The sample is clear.	-	The sample is clear.	Pass	\$1
Available volume	PHAR-AVOLUM1#M0-PRG	250	mL	min. 250 ml	Pass	\$1

S2 Analysis was performed by ALS testing site Na Harčé 336/9, Prague 9, 190 00  
S1 Analysis was performed by ALS testing site Poděbradská 540/26, Prague 9, 190 00

If the client does not specify the date and time of sample collection, the laboratory will specify the date on sample delivery in parentheses, instead. If the time of sample collection is specified as 0:00 it means that the client did specify the date but not the time.

**The end of result part of the certificate of analysis**

**Brief Method Summaries**

Analytical Methods	Method Descriptions
METL-ALMSFXVJ0-PRG	Metals by ICP-MS - fixation
METL-IDENOES-NA-PRG	Metals by ICP-OES-A - fixation
METL-NAMSFVC2-PRG	Metals by ICP-MS - fixation
PHAR-AVOLUM1#M0-PRG	Volume; G184
PHAR-CITR2-ID#MC-PRG	Identification; G174
PHAR-CITRA2#M1-PRG	Assay of citric acid / citrate; Medites
PHAR-PHP2-ELE#M1-PRG	pH; G183 - Medites
PHAR-SENZL1#MC-PRG	Clarity/Coloration; G182

**LAL TEST**  
**Limit test for bacterial endotoxins (BET)**  
**in CITRASOL 4%...**

The test was carried out using work equipments and procedure in accordance with SOP: 10.05.06.02 LAL test.

Controlled product: CITRASOL 4% 250 ml Lot No.: A0043020

Endotoxin limit: 0,25 EU/ml Analysis No.: 2192/20

LAL Reagent: Pyrotell (ACC) Declared sensitivity  $\lambda$  0,03 EU/ml

Lysate: --- Lot No.: 518-11-902

- dissolved in reconstitution buffer

Endotoxin: --- Lot No.: 170

Validation of the kit (confirmation of sensitivity) carried out on: 14.5.2020

**Preparation of the product:**

Dilution:

Quantity of undiluted product in ml: 1,0 Initial pH: 6,97

Quantity of buffer (Tris ...1...) in ml: 7,0

Quantity of water for BET in ml: ---

**Test result CITRASOL 4%:**

Determination	Sample	Positive control	Negative control
		1. 50 $\mu$ l 0,125 IU endotoxin/ml + 50 $\mu$ l buffer TRIS 1 2. 50 $\mu$ l 0,125 IU endotoxin/ml + 50 $\mu$ l sample 1:3 buffer TRIS 1	100 $\mu$ l buffer TRIS 1
1.	-	+	-
2.	-	+	-

**Test result CITRASOL CPD50 / ACD-A:**

Determination	Sample	Positive control	Negative control
		1. 50 $\mu$ l 0,125 IU endotoxin/ml + 50 $\mu$ l water for BET 2. 50 $\mu$ l 0,125 IU endotoxin/ml + 50 $\mu$ l sample 1:3 buffer TRIS 2	100 $\mu$ l water for BET
1.		---	---
2.		---	---

**Test result CITRASOL 0,5%:**

Determination	Sample	Positive control	Negative control
		1. 50 $\mu$ l 0,5 IU endotoxin/ml + 50 $\mu$ l water for BET 2. 50 $\mu$ l 0,5 IU endotoxin/ml + 50 $\mu$ l sample	100 $\mu$ l water for BET
1.		---	---
2.		---	---

**Evaluation:**

The sample complies test for BET according to Ph.Eur. 9: yes  no

QC: *Kralinková* Date of execution: 1.10.2020

QP: *Kralinková* Date of control: 1.10.2020

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Particle Sizing Systems, Inc.  
Santa Barbara, Calif., USA

10.06.02.10/16.9.2019

Particle Analysis Report

CITRASOL 4% 250ml A0043020 P/407/20  
File Name = A0043020.1CB Time Date = 9:39:56 10/ 1/2020  
Sensor Model: LE400-0.5 EXT S/N: 9811901 Cal. File: 9811901S.sns  
Date of Calibration: 06-11-19  
Control File: C:\PSS\SWS788\sws788.rcp

Elapsed Time of Data Collection = 40 Sec.  
Background File = NONE

Number of Pulls = 4 Volume of Pull = 5.0 ml

Large Volume Injectables

Cumulative Number of Particles Per Pull Volume

PULL	CH1	CH2
	10.0	25.0 (um)
1 *	20	0
2	15	0
3	23	1
4	21	0
MEAN #	19	0
#/ml	3	0
LIMIT	25	3
TEST	Passed	Passed

\* First pull was not included.

Operator's Name: sramkova

Operator's Signature *Sramkova* Date 1.10.2020

Supervisor's Approval *Wen* Date 1.10.2020

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IMUNA PHARM a.s.,  
Jarková 269/17  
082 22 Šarišské Michaľany


## Protokol o skúške / Protocol of analysis

Prípravok: Name of the product:	CITRASOL 4 %	Balenie / Packaging	250 ml
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Číslo šarže / Batch No.:	A0043020	Č. protokolu OKK / QCD protocol No.:	20/0583
Zadávateľ / Customer:	MEDITES PHARMA	Č. protokolu zadávateľa / Customer protocol No.:	Zmluva o dielo z 27.3.2017 /
Skúšku vykonal / Analysed by:	ITEST plus s.r.o.		The works contract from 27.3.2017

Skúška / Test	Akceptačné kritériá / Acceptance criteria	Výsledok / Result
Sterilita / Sterility (2.6.1.)	sterilný / sterile	sterilný / sterile
<b>Záver: Výsledky skúšok</b>	<b>VYHOVUJÚ</b>	<b>NR-C 115 10.08.13 časť 1</b>
<b>Conclusion: Results of tests</b>	<b>COMPLY</b>	<b>with specification NR-C 115 10.08.13 supplement 1</b>

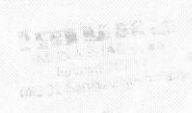
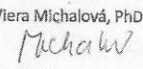
Protokol o skúške vypracoval pracovník OKK / Certificat of Analysis issued by worker of QC:

  
Mgr. Klaudia Vargová

Dátum / Date: 16.10.2020

Meno a podpis / Name and signature

Dátum / Date: 16.10.2020

  
Mgr. Daniel Žatko, PhD.  Ing. Viera Michalová, PhD.

Meno a podpis vedúceho OKK /

Name and signature of the Head of QC Department