CLINICAL STUDY PROTOCOL AMENDMENT

A Randomized, Double-Blind, Vehicle-Controlled, Parallel Group Phase 2a Study to Determine Safety, Tolerability, Pharmacokinetics, and Pharmacodynamic Activity of NFX-179 Gel in Subjects with Cutaneous Neurofibromas

Protocol No.	NFX-179-NF1-201
Protocol/Amendment Date:	01-APR-2021
Amendment No.:	3
Supersedes:	N/A
Sponsor:	NFlection Therapeutics 714 Woodcrest Road, Wayne, PA 19087 United States
Medical Monitor:	Guy Webster, M.D. Ph.D. 24-hour telephone: (302) 559-8684 Email: GWebster@Nflectionrx.com

PROTOCOL AMENDMENT INVESTIGATOR SIGNATURE PAGE

Protocol Number: NFX-179-NF1-201
Amendment Number: 3

INVESTIGATOR COMMITMENT:

I will provide copies of the protocol, any subsequent protocol amendments and access to all information provided by NFlection to the investigational center staff under my supervision. I will discuss this material with them to ensure that they are fully informed about the Investigational Medicinal Product and the study protocol.

I agree to conduct this clinical study according to the attached protocol, except when mutually agreed to with NFlection in writing. I also agree to conduct this study in compliance with all local regulatory requirements, Good Clinical Practices, as well as with the requirements of the appropriate Institutional Review Board(s) /Ethics Committee(s) and any other Institutional requirements.

_____

Printed Name of Investigator

_____

Signature of Investigator

_____

Date

PROTOCOL AMENDMENT APPROVAL PAGE

Protocol Number: NFX-179-NF1-201
Amendment Number: 3

A Randomized, Double-Blind, Vehicle-Controlled, Parallel Group Phase 2a Study to Determine Safety, Tolerability, Pharmacokinetics, and Pharmacodynamic Activity of NFX-179 Gel in Subjects with Cutaneous Neurofibromas

Protocol No.	NFX-179-NF1-201
Protocol/Amendment Date:	01-APR-2021
Amendment No.:	3
Sponsor:	NFlection Therapeutics 714 Woodcrest Road, Wayne, PA 19087 United States

C. Powala 04-02-2021

Christopher Powala
Chief Executive Officer
NFlection Therapeutics

Date

4/2/21

Guy Webster, M.D., Ph.D.
Chief Medical Officer
NFlection Therapeutics

Date

TABLE OF CONTENTS

1 AMENDMENT HISTORY

Revised protocol 1 footer date: 24-JUL-2020

Previous amendments: Amendment 2, footer date 25-MAR-2021

2 AMENDMENT SUMMARY

The following section of the NFX-179-NF1-201 revised protocol (footer date 24-JUL-2020) as modified by amendment 2 (footer date 25-MAR-2021) and with a final NFlection Therapeutics approval signature date of 27-JUL-2020 is amended:

- Section 8.3.3 Efficacy analyses.

3 AMENDMENT RATIONALE

This protocol change is made to allow for Target cNF Tumors that are found not to be qualifying neurofibromas to be excluded from sensitivity analyses of efficacy results.

4 PROTOCOL CHANGES

Section 8.3.3 Efficacy analyses:

Previous paragraph 1:

"…Tumor volume is the primary measurement of interest; however, the ruler measurements of tumor length and height will also be analyzed as described below."

Changed paragraph 1:

"…Tumor volume is the primary measurement of interest; however, the ruler measurements of tumor length and height will also be analyzed as described below. Any Target cNF Tumors confirmed not to be qualifying neurofibromas based on histological evaluation of the excised tissue may be excluded from sensitivity analyses of efficacy results."