APPROVED
AS MODIFIED
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WIRB®

OPTIONAL GENE EXPRESSION ADDENDUM TO INFORMED CONSENT FORM

TITLE: PHASE III, RANDOMIZED, DOUBLE-BLIND, PLACEBO-CONTROLLED, PARALLEL-GROUP, STUDY OF AB103 AS COMPARED TO PLACEBO IN PATIENTS WITH NECROTIZING SOFT TISSUE INFECTIONS (NSTI)

ACCUTE (AB103 Clinical Composite Endpoint Study in Necrotizing Soft Tissue Infections)

PROTOCOL NO.: ATB-202
WIRB® Protocol #20151312

SPONSOR: Atox Bio Ltd.

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STUDY COORDINATOR(S): Debra Burris, RN
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An optional part of this study involves the collection of a blood sample from you for gene expression (RNA expression or transcriptome profile) research. Before you decide whether you would like participate in this part of the study we would like to give you some more information. Your rights to participation as set forth in this addendum are identical to the rights that have been presented to you in the main ICF. This substudy has been evaluated by the ethic committees of the involved sites, the specific study assessments of this substudy cannot be charged to you or your insurance, the trial is also covered by a no fault insurance according to the same conditions and guarantees of confidentiality (protection of your privacy).

Atox Bio Ltd. will be referred to as sponsor throughout this document.

WHAT IS THE BACKGROUND TO AND PURPOSE OF THE GENETIC RESEARCH?

Cells in your body contain a type of molecule called ribodeoxyribonucleic acid, or "RNA" for short. RNA is made from DNA and making RNA is called gene expression. RNA is a genetic material that has a major role in making proteins. Proteins are the building blocks of your body, cells, and organs. Proteins then help direct growth, control how the body functions or reacts to challenges like infections or injuries. Researchers can gain an understanding of the type of proteins the body can make in response to infections and how that response can be affected by treatment by studying the type and amount of RNA (gene expression) produced in certain cells like the white blood cells (which are involved in fighting infections). You are being asked to donate a blood sample for gene expression research that will help determine how your body responds to the infection you are experiencing and how AB103 may affect that response. To help with this process medical information collected about you in the main study will be compared with the results of the gene expression studies. You are being asked to take part in the genetic research because you are already taking part in the study with AB103 in patients with necrotizing soft tissue infections which we will call "the main study" in the rest of this document. We will use the words "this RNA expression research" for the RNA transcriptome research we are now asking you to take part in.

The blood sample you donate will not be used for other purposes or be used to store your DNA (your genes).

The study sponsor (Atox Bio) will not make any test results of this RNA expression research available to you, any insurance company, your employer, your family, the Study Doctor, or any other physician who treats you now or in the future. You should be aware that the study sponsor has no obligation to conduct this RNA expression research, or any additional research on your blood sample or RNA.

WHAT WILL HAPPEN TO ME IF I TAKE PART?

If you decide to donate a blood sample, we will collect about 1 teaspoon (5 mL) of blood from you at five different times during the study-before you get the blinded study drug, at 4-6 hours, 24 hours, 4 days and 7 days after receiving the blinded study drug. RNA will be extracted from your blood sample. In this process, most of the original blood sample will be used up but a small amount will be kept as a "backup" in case of problems in the testing of your RNA.

The remaining blood sample will be stored with similar samples from other people at a secure central laboratory contracted by the study sponsor. RNA and blood samples from this genetic research will be destroyed 5 years after the main study is completed. Your RNA may be studied at any time before this. The results from this gene expression research may be analysed along with results from other research.

WHAT ARE THE POSSIBLE RISKS/BENEFITS AND INCONVENIENCES OF TAKING PART?

The taking of a blood sample may cause some discomfort and bruising (and there is a potential for infection). Approximately 25 mL (five teaspoons) of blood will be used for this test. Where possible these blood samples will be taken from a vein in your hand or arm at the same time that other blood tests are being performed in the main study.

There is no direct benefit to you in taking part in this gene expression research. However, this research may contribute to our understanding of necrotizing soft tissue infections and its treatment, and may eventually lead to improvements in treatment.

DO I HAVE TO TAKE PART?

It is up to you whether to donate a sample for this gene expression research or not. You may refuse to donate a sample at any time without penalty or loss of benefits to which you are otherwise entitled. You will receive the same treatment and care in the main study whether or not you donate a blood sample for this research as described in this document. **If you decide not to donate a sample, you can still take part in the main study**.

WHAT RIGHTS DO I HAVE TO THE RESULTS OF THE GENETIC REESEARCH?

The purpose of this gene expression study is not to provide you with test results nor make any results available to you, any insurance company, your employer, your family, the Study Doctor, or any other physician who treats you now or in the future.

Any information derived directly or indirectly from this gene expression research, as well as any patents, diagnostic tests, drugs, or biological products (such as antibodies developed as a result of characterisation of a gene or protein) developed directly or indirectly as a result of this gene expression research, are the sole property of the study sponsor (and its successors, licensees, and assigns) and may be used for commercial purposes. You have no right to this property or to any share of the profits that may be earned directly or indirectly as a result of this genetic research. However, in signing this form and donating a blood sample for gene expression research, you do not give up any rights that were proposed to you in the main informed consent.

WHAT STEPS ARE TAKEN TO ENSURE THE GENETIC RESULTS ARE KEPT CONFIDENTIAL?

Special precautions are taken to ensure that the research in this gene expression study will be carried out with a very high degree of confidentiality. When your blood sample is drawn, it will be labelled with the same code that is given to you in the main study. The link between this code and your personal identifiers, such as your name, will be kept by your study doctor only for 15 years after the end of the study. Then it will be destroyed.

Staff from the company, Criterium, whose job it is to make sure the research has been done properly by checking the records at the Study Doctor's site, will also be able to identify you from your medical files but will not have access to the results of the gene expression research in this study. Apart from these persons, other research staff at Criterium will not know your identity. Regulatory authorities, including the FDA, which also may wish to check that this gene expression research has been done properly, will also have access to your files and know your identity.

The data and results of this gene expression research may be reviewed with collaborators and published. Neither your name nor any other information that identifies you personally will appear in any publications or reports.

Sometimes patients have been required to furnish information from genetic testing for health insurance, life insurance, and/or a job. A Federal law, the Genetic Information Nondiscrimination Act of 2008 (GINA), generally makes it illegal for health insurance companies, group health plans, and employers with 15 or more employees to discriminate against you based on your genetic information.

CAN I WITHDRAW MY CONSENT?

You may withdraw your consent to the use of your sample in genetic research at any time. If you withdraw your consent **before** your blood sample is sent for genetic research, the Study Doctor will arrange to have it destroyed. If you withdraw your consent **after** your blood sample has been sent for gene expression research the study sponsor and the Study Doctor will ensure that your blood sample and any RNA that has been extracted from it are destroyed. However, if genetic research has already been performed the study sponsor is not obliged to destroy results of this research. In this case only the blood sample and any RNA extracted will be destroyed.

INFORMED CONSENT STATEMENT

Consent and Assent Instructions:

Consent: Subjects able to provide consent must sign on the subject line below
Consent is provided by the Legally Authorized Representative for adult subjects unable to consent
For subjects under 18, consent is provided by the parent or guardian

Assent: Verbal assent is required for all subjects who are capable of giving assent using the Assent section below.

I,_____(name of subject), have read and I understand all the information in this informed consent addendum. I have been given the chance to discuss it and ask questions. All my questions have been answered to my satisfaction. I voluntarily consent to take part in this genetic study and I am free to withdraw from the genetic research at any time without my medical care or legal rights being affected. I understand I will receive a copy of this informed consent addendum.

By signing this informed consent, I have not given up any of the legal rights, which I otherwise would have as a subject in a research study. I authorise the collection, use and disclosure of my medical information in accordance with this form.

_____ _____

Signature of subject
(Please date your own signature at the time of signing)

Date of signature

_____

Printed name of subject (BLOCK CAPITALS)

_____ _____

Signature of person conducting the informed consent discussion
(Please date your own signature at the time of signing)

Date of signature

_____

Printed name of person conducting the informed consent discussion (BLOCK CAPITALS)

_____ _____

Signature of legally accepted representative, parent or guardian
(Please date your own signature at the time of signing)

Date of signature

_____

Printed name of legally accepted representative, parent or guardian (BLOCK CAPITALS)

_____

Relationship of legally accepted representative, parent or guardian to subject (BLOCK CAPITALS)

_____ _____

Signature of impartial witness
(in the event the subject is unable to read or write) (Please date your own signature at the time of signing)

Date of signature

_____

Printed name of impartial witness (BLOCK CAPITALS)

CONSENT OF THE SUBJECT TO CONTINUE TO BE IN THE STUDY

Your legal representative gave his/her consent for you to give an additional blood sample for gene expression research. This is because you were not able to make your own decision due to your illness. Your condition has now improved. You are being asked to decide whether you agree with the consent given by your legal representative. Your decision is voluntary. This means your decision is up to you.

You have read this information in this form and someone has explained to you what the analysis of your blood sample for genetic research means. Your questions have been answered to your satisfaction.

_____ _____

Signature of subject
(Please date your own signature at the time of signing)

Date of signature

_____

Printed name of subject

_____ _____

Signature of person administering this consent
(Please date your own signature at the time of signing)

Date of signature

_____

Printed name of person conducting the informed consent discussion (BLOCK CAPITALS)

ASSENT SECTION:

Statement of person conducting assent discussion:

1. I have explained all aspects of the research to the subject to the best of his or her ability to understand.
2. I have answered all the questions of the subject relating to this research.
3. The subject agrees to be in the research.
4. I believe the subject's decision to enroll is voluntary.
5. The study doctor and study staff agree to respect the subject's physical or emotional dissent at any time during this research when that dissent pertains to anything being done solely for the purpose of this research.

_____ _____

Signature of Person Conducting Assent Discussion — Date

_____

Printed name of Person Conducting Assent Discussion

Statement of Parent or Guardian: