

Warsaw, 23/10/2018

REQUEST FOR PROPOSAL NR 66/2018- IBD

I. Ordering Party

<p>OncoArendi Therapeutics S. A. ul. Żwirki i Wigury 101 02-089 Warszawa VAT ID: 728 27 89 248</p>	<p>Contact person: Michał Młacki e-mail: m.mlacki@oncoarendi.com Tel. +48 798 352 548</p>
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II. OBJECT FOR THE REQUEST:

In vivo study of chitinase inhibitor efficacy in the rat model of inflammatory bowel disease (IBD) induced with 2,4,6-trinitrobenzenesulfonic acid (TNBS), in which there is observable granulomas formation

The order carried out as a part of the projects titled:

— **IBD:** „PRE-CLINICAL AND CLINICAL RESEARCH OF CANDIDATE FOR INNOVATIVE DRUG IN ASTHMA AND INFLAMMATORY BOWEL DISEASES THERAPY” (POIR.01.01.01 00-0168/15)

co-financed by the National and European Union Funds and because of the competitiveness principle

III. THE FORM OF THE ORDER:

- III.1 The request is not made under The Public Procurement Law (Journal of laws of 2013, item 907 as mentioned).
- III.2 This order is carried out in accordance with the principle of competitiveness, openness, transparency and equal access.
- III.3 The Ordering Party reserves the right to cancel this procedure without providing reasons and also to complete the procedure without choosing the winner tender.
- III.4 In the course of examination and evaluation of the offers the Ordering Party may require Contractors to present explanations concerning the content of submitted bids.
- III.5 The Ordering Party reserves the right to change the content of this request. If the changes can affect the content of tenders submitted in the procedure, the Ordering Party shall extend the tender submission deadline. Any changes made shall be provided promptly to all tenders to which the request was sent and shall be binding on them.
- III.6 This procedure does not set the obligation for OncoArendi Therapeutics SA to sign any formal contracts.
- III.7 It is not possible to make and offer for part of order.

IV. CONDITIONS FOR PARTICIPATION IN THE PROCEEDINGS:

- IV.1 The Request for offers relates to potential Contractors whose scope of business activity is in full compliance with the subject of this Request.
- IV.2 The offers may be issued by parties who:
 - A) Have the necessary qualifications to carry out the described activity and have the appropriate



technical potential and personnel capable of performing the contract.

- B) Are in a good economic and financial standing, which assures proper execution of the project in the declared time.
- C) Will pursue the contract in a way that is beneficial to the environment by minimizing the consumption of materials, raw materials energy, etc. (with environmental certificates such as ISO 14001:2015)

As a proof of the above, the Ordering Party requires that the Contractor submit, along with the tender, a statement about fulfilling conditions for participation in the proceedings. The model statement is attached as Appendix 2 to this request for proposal.

IV.3 Excluded from the proceedings shall be those contractors who are personally or equity related to the Ordering Party by in particular:

- A) participation in the company, in a civil or limited partnership;
- B) holding at least 10% shares or interests;
- C) serving a function of a member of the supervisory organ, a member of the management organ or proxy;
- D) having family ties, such as by marriage, by lineage at first or second degree, by adoption, guardianship or custody.

As a proof of the above the Ordering Party requires that the Contractor submit, along with the tender, a statement about not being related to the Ordering Party. The model statement is attached as Appendix 3 to this request for proposal.

IV.4 Issuing the offer represent the full acceptance of the rules set in this Request.

V. DETAILED DESCRIPTION OF THE OBJECT OF THE REQUEST:

CPV Code: 73111000-3 – Laboratory research services

The contract research service aims to test the therapeutic effects of chitinases inhibitor in a 21-day-long rat model of inflammatory bowel disease (IBD), in which 2,4,6-Trinitrobenzenesulfonic acid (TNBS) is applied per-rectum. Particularly, the service evaluates the inhibitor efficacy on development, number and severity of granulomas in colon wall (granulomatous inflammation).

The experiment should consist of 5 groups:

- non-TNBS group;
- TNBS group with administration of vehicle;
- TNBS group with reference compound (dosing once a day qd);
- TNBS group treated with chitinases inhibitor at lower dose (dosing once a day qd);
- TNBS group treated with chitinases inhibitor at higher dose (dosing once a day qd);

Every group should consist of minimum 10 rats, except control (non-TNBS) group which should consist of minimum 5 animal. The compounds administration should be conducted once a day (qd) in therapeutic scheme with 2-week-long treatment (from 7th-20th day after TNBS administration).

Chitinases inhibitor will be provided by the Ordering Party, reference compound will be provided by Contractor.

At 21st day after TNBS administration, animals should be euthanized and following material should be collected:

- Blood, 0.5 ml, collected to the tube with anticoagulation agent, plasma should be flash-frozen;

- After macroscopic evaluation, 3 fragments of transversely cut colon (1 cm in width, collected in such way, that the whole circumvent is present in samples) should be fixed in 10% neutral buffered formalin. These fragments should be collected from similar regions in every animal (irrespective of the pathological lesions), from areas separated by about 5 cm in colon. During preparations these fragments should be rolled into "swiss-roll", process to paraffin blocks and cut in such a way that the whole circumvent of the colon is visible;
- 3 fragments (about 1 cm²) collated from similar regions in every animal (irrespective of the pathological lesions), from areas separated by about 5 cm in colon tissue – flash-frozen;
- 3 fragments (about 1 cm²) collated from similar regions in every animal (irrespective of the pathological lesions), from areas separated by about 5 cm in colon tissue – preserved in RNA-later, incubated in it at +4°C for minimum 12h and then flash-frozen;
- Remaining colon fragments should be preserved in 10% neutral buffered formalin for possible future analyses by Ordering Party (OncoArendi).

The primary readouts should be as follows:

- Body weight changes through the experiment (measured minimum 5 times/week) and potential mortality;
- Colon length at the end of study during material collection;
- Macroscopic evaluation and scoring of pathological changes in colon;
- Histological analysis of pathological changes in 3 fragments of colon separated by about 5 cm (at least scores for general inflammation, granulomatous inflammation, erosion, ulceration - on hematoxylin-eosin stained samples of colon)

Plasma, frozen pieces of colon, paraffin-embedded tissue, formalin-fixed fragments of colon and histological slides will be transferred to Ordering Party laboratories after study final at Contract Research Organisation's own expense.

The detailed study plan and study timelines will be agreed with the Ordering Party before signing an agreement.

VI. EVALUATION OF THE OFFERS:

VI.1 Price – weight: 80% (80 pts.)

A) In this criterion points will be calculated according to the formula below:

$$Pc = \frac{C_{min}}{C_{evaluated}} \times 80$$

Pc – Points received

C_{min} – The smallest Net price

C_{evaluated} – Net price of the offer being evaluated

80 – weight of the criterion (80%)

VI.2 Payment deadline – weight: 20% (20 pts.)

In the payment deadline, points will be awarded according to the formula:

20 points – when the payment deadline is over than 30 days from the invoice delivery

15 points – when the payment deadline is between 21-29 days from the invoice delivery

10 points – when the payment deadline is between 14-20 days from the invoice delivery

5 points – when the payment deadline is between 7-13 days from the invoice delivery

0 points – when the payment deadline is less than 6 days from the invoice delivery

20 – weight of the criterion (20 %)

VI.3 In the case of two or more tenders with equal number of points awarded, to guarantee performance of the contract in a manner favorable to the environment, by providing minimize the consumption of

materials, raw materials, energy etc. any environmental certifications will be taken onto consideration (eg. ISO 14001:2015).

If the abovementioned does not allow to choose the best offer, the Ordering Party shall call Contractors who submitted equally evaluated offers to submit, within the period specified, additional offers. Contractors cannot offer higher prices than offered in the tenders.

VII. HOW TO PREPARE AND SUBMIT THE OFFER:

- VII.1 The offer should be signed by the person authorized to represent the institution.
- VII.2 Each contractor may submit only one offer.
- VII.3 Costs of the offer preparation shall be incurred by the offering party.
- VII.4 Offers must be submitted no later than: **30/10/2018 23:59** CET and must be written on the form as in Appendix 1 to the request for proposals.
- VII.5 Offers shall be issued only via email to: **k.czyzewska@oncoarendi.com**
- VII.6 The date of receiving the email shall be considered as a date of issuing the offer.
- VII.7 Offers that do not meet the deadline, are incomplete or sent to the wrong email address will not be taken into consideration.
- VII.8 Any questions concerning the Object of the tender should be addressed to m.mlacki@oncoarendi.com (+48 798 352 548) no later than 26/10/2018 23:59 (CET). Contact person is: Michał Młacki.
- VII.9 Any questions concerning the formal issues of the tender should be addressed to m.skrzek@oncoarendi.com (+48 22 552 67 24) no later than 26/10/2018 23:59 (CET).
- VII.10 The offer should include the validity date (at least 30 days from the submission deadline).
- VII.11 The price should be set in both Net and Gross.
- VII.12 The values in the offer (Net and Gross) should be rounded to two decimals with the mathematical rule of rounding the numbers (according to § 5 section 6 of the regulation of Ministry of Finance of 28 November 2008 (Journal of Laws of 2008, No. 212, item 1337, as mentioned).
- VII.13 The offer price should include VAT. The correct determination of VAT is responsibility of the contractor – in accordance with the provisions of the Act of 11 March 2004 on Goods and Services Tax (Journal of Laws of 2004 No. 54 item. 535 as mentioned).
- VII.14 The offer shall not be prepared in price variants.
- VII.15 The financial settlements between the Ordering Party and the contractor may be made in PLN, EUR, USD or GBP.

VIII. TENDER RESULTS:

Bidder will be informed about choosing his offer via email. Formal results will be also published on the Ordering Party's website (www.oncoarendi.com) and concurrency database.

IX. MOST IMPORTANT PROVISIONS OF THE AGREEMENT:

- IX.1 Supplier will be obligated to enter into the agreement including all conditions presented in the Request for the Offer in the place and time specified by the Ordering Party.

IX.2 It is not possible to introduce significant changes to the content of the agreement in relation to the content of the offer, which was the base for the Service Provider selection, unless:

- A) The amendments concern performing additional services by the Service Provider, not covered by the basic contract, provided they are necessary and the following conditions are met:
 - i. The change of the Service provider cannot be made due to to the economic or technical reasons, in particular concerning the interchangeability and interoperability of equipment, services or installations, ordered as part of basic contract.
 - ii. The change of the Service Provider would cause significant inconvenience or substantial cost increase to the Ordering Party.
 - iii. The value of any subsequent changes do not exceed 50% of the basic contract value.
- B) The amendment does not lead to change in the nature of the contact and the following conditions are met:
 - i. The need for the contract change is brought about by circumstances which the Ordering Party, acting with due diligence, could not foresee.
 - ii. The value of a change does not exceed 50% of the basic contract value.
- C) The amendment does not lead to change in the nature of the contract and the total value changes is less than 209 000 EUR, and at the same time is less than 10% of the basic value.

Any contract amendment must be done in writing, otherwise will not be valid.

IX.3 Information regarding contractual penalties:

- A) If the offered project execution time (as defined in the offer) extends for at least 15 days, the Service Provider shall pay the Ordering Party a contractual penalty of 1,5% of the net offer price for exceeding the time limit, and then another 2% of net offer price for each additional 15 days of delay. Moreover, the Ordering Party will gain a right to withdraw from the contract of the project execution time is exceeded by at least 60days. The Ordering Party may withdraw from the agreement until 31st Dec 2018.
- B) Due to the termination or withdrawal from the Agreement by either Party for reasons caused by the Service Provider, the Ordering Party will change a contractual penalty of 25% of net offer price.
- C) The formal basis for changing contractual penalties will be a debit note the Ordering Party delivers to the Service Provider. The Ordering Party shall be entitled to deduct contractual penalties from payments due to the Service Provider.
- D) The Ordering Party has the right to claim damages in the amount exceeding contractual penalties based on general principles.
- E) The Ordering Party has the right to claim damages in the amount exceeding contractual penalties based on general principles.
- F) Contractual penalties sum up.

X. APPENDENCIES:

- A) The offer form.
- B) Statement concerning fulfilment of all the requirements set out in part IV of the Request for offers.



- C) Statement concerning persona or/and capital connections between the Service Provider and the Ordering Party.
- D) Declaration of compliance with the information obligations provided for in Article 13 or Article 14 of the GDPR (Appendix No. 4).