Numer zapytania	Z18/5914/1
Tytuł zapytania	Bioequivalence studies - Colistimethate sodium - powder for suspension preparation
Kupiec prowadzący:	Topij, Ewa
Osoba kontaktowa w sprawach merytorycznych:	Łojszczyk, Ilona
Data złożenia:	2024-03-28 15:24:45
Waluta:	PLN

TERMINY W ZAPYTANIU

Data i godzina rozpoczęcia przyjmowania ofert:	2024-03-28 16:00:00
Data i godzina zakończenia przyjmowania ofert:	2024-04-10 23:00:00
Termin zadawania pytań (do kiedy?):	2024-04-04 15:00:00

Załączniki	tak

Treść zapytania

FIRST STAGE - Request for proposal - RFP

Subject: Bioequivalence studies for *Colistimethate sodium*

Background

Tarchomińskie Zakłady Farmaceutyczne “Polfa” Spółka Akcyjna (Polfa Tarchomin S.A.) is one of the oldest and most experienced pharmaceutical companies in the Central Europe. Our history goes back to 1823. Nowadays there are four pillars of our generic business: Anti-infectives (oral and injectable forms), Alimentary Tract & Metabolism products, Central Nervous System drugs and Dermatologicals.

Currently, Polfa Tarchomin S. A. develops *Colistimethate sodium* generic product and we would like to invite you to prepare a proposal for: 1) conducting comparative bioavailability study (full service) 2) conductingcomparative bioavailability study including pk parameters for bioequivalence assessment of medicinal products described below.

The project is co-finansed from the state budget under a grant awarded by the Medical Research Agency.

A. Project and product information

Tested product - *Colistimethate sodium TZF 1 mln IU***,** lyophilized powder for preparing a solution for injection, infusion and inhalation

Comparator- *Colistin TZF 1 mln IU,* lyophilized powder for preparing a solution for injection, infusion and inhalation

Reference- *Colomycin 1 mln IU,* lyophilized powder for preparing a solution for injection, infusion and inhalation

Registration

Assumed study design - Bioequivalence

IMPD availability - 01.2025

Currently, Polfa is developing three strengths (1 mln IU, 2 mln IU, 3 mln IU) of *Colistimethate sodium*, powder for

infusion, injection and inhalation. Polfa plans to conduct the study only for 1 mln IU.

We plan single dose 3-arms comparative bioavailability study under fasting condition with 1 mln IU dose of *Colistimethate sodium* (tested product), *Colistin TZF 1 mln IU* (comparator product) and *Colomycin 1 mln IU* (reference product). The primary objective of the study will not be to assess the bioequivalence, but we plan to make the assessment of the BEQ pk parameters as the subject of additional analysis / secondary objective. In the planned study two analytes will be evaluated: colistin A and colistin B (free colistin). This aim of this study will be assessment of the potential differences in the composition of individual polymyxins and the degree of sulphomethylation translate into the bioavailability of active colistin, and thus may be relevant to its efficacy and safety. The exact parameters which are planned to be evaluated will be defined in the study protocol and will include: C_{max}, T_{max}, AUC, $T_{1/2}$, MRT, CL, V_d and the ratio of these parameters.

SmPC of Colomycin 1 mln IU is available on website:

https://www.medicines.org.uk/emc/product/1094/smpc/print

This is just for information purposes.

We are interested in full service for one single dose comparative bioavailability study including the clinical conduct, bioanalytical part, data management, medical writing, project management, RA/EC submission etc.

For sample size assumptions we have no useful data we could share, therefore, we would appreciate your suggestions. Please provide at least two sample size options, i.e. assuming T/R 5 and 7.5%.

Proposal for comparative bioavailability / bioequivalence studies

The offer should contain at least:

1. information about the proposed studies (1- BA and 2 - BA with BEQ assessment): study design, number of subjects, number of sample per subject per period, total PK samples, analytical method
2. proposal of timelines assuming the above IMPD availability, duration of the study from start of documentation preparation until draft study report
3. costs calculations for:
    1. clinical part (preparation of the study design according to EU requirements and essential documentation for RA/EC submission, submission to RA/EC including fees, clinical conduct including all medical and safety laboratory assessments and PK sampling and remuneration to subjects, data management and integrated study report (ICH-E3), QA during the study);
    2. analytical part (method development and validation, if applicable, PK samples analysis including ISR, reports);
    3. PK and statistical part (pharmacokinetic and statistical analysis and report);
    4. other costs needed (for example monitoring, PK and documents transport, analytical reference standards, overheads etc.). Please list these costs if needed.

Additional information expected from the bidder

We kindly ask you to provide the information listed in the attached Excel file named ‘BIO offering TOC’. Please provide the information either in the offering document or in the Excel document as it is most convenient for you.

B. Conditions for participation

1. Present registry documents (National Court Register EDG or equivalent, Certificate of Fiscal Residence).
2. Confirmation of an appropriate experience in realization of similar services. Supplier should have experience, knowledge and abilities, should be demonstrated by realization of at least ten (10) studies over the past 3 years. Documents should include information about contractor, study specification and delivery date.
3. Clinical site capacity of admission of all volunteers in one round / one period.
4. Preparation the full scope of services specified in the RFI/TOC:
    1. Study planning;
    2. Preparation of study documentation, submission of applications to the Bioethics Committee/relevant authorities, obtaining appropriate opinions and consents;
    3. Clinical part: staff training, volunteer recruitment, administration of products, collection of samples, subjects examination;
    4. Bioanalytical part: method development and validation, test sample analysis;
    5. Pharmacokinetic and statistical part - preparation of study results;
    6. Preparation of the final study report;
    7. Study insurance (in cooperation with the Sponsor);
    8. Preparation and archiving of study documentation.
5. Compliance with European formal requirements by the center (appropriate certificates: GCP, GLP, Certificate of Laboratory Accreditation, ISO certification, the results of own audits / inscpetions) including full compliance with the Regulation 536/14, CTIS submission service (for studies conducted in Europe).

6. Confirmation of compliance with the above mentioned requirements.
7. Conducting the study according to the determined by the Sponsor timelines: 30.11.2025 – Study Report.
8. An unblemished CRO reputation– verification i.e. via official EMA inspection documents including concerns about the reliability of the company (if available), publicly available sources (i.e. web sites).

C. Terms of the offer submission

1. **Deadline for submitting Offer is 04-04-2024 at 23:00 CET**. The deadline for Answers submission may change, all Suppliers will be informed by e-mail from the Procurement system.
2. Any Supplier may submit only one offer.
3. Partial offers are not allowed.
4. The offer with the required Attachments should be submitted in the Logintrade Procurement system (https://polfa-tarchomin.logintrade.net/). In order to submit an Offer, the electronic form must be completed and saved. The completed Attachments should be uploaded to the system.
5. In addition, the Supplier should present terms of maintenance services and may also prepare an offer in his own preferable format and upload it into the system.
6. Supplier's Offer should be comprehensive in Polish or English, in electronic form.
7. The offer should include at least the following information: supplier name and address, bid inquiry number, price, delivery date (time required for delivery), date of payment.
8. The tenderer, before the deadline for submitting bids, has the right to:
    1. withdraw the offer by written notification by e-mail indicated for the submission of tenders,
    2. change the offer – notification of the changes must be submitted by the same rules as the submitted offer, appropriately marked with the note "CHANGING THE OFFER".
9. Offer validity: 90 days from the deadline for submitting offers.
10. The tenderer shall bear all possible costs related to the preparation and submission of the offer. The Contracting Party does not provide reimbursement of the costs of participation in the procedure.

D. Additional information about participation in the tender

1. The procurement procedure will be conducted in two stages.

- The first stage RFP - Request for proposal.
- The second Stage RFQ - Request for quotation - after verification of the received documentation in the RFP.

1. Before proceeding with the procedure, it is recommended to read the operating manual of the purchasing system available at https://polfa-tarchomin.logintrade.net/.
2. The dates of commencement and completion as well as the configuration of each stage of the procedure will be presented in the Procurement System and may be changed about which all interested Contractors will be immediately notified by e-mail via the system.
3. Technical support during the procedure. During the procedure, the contractor may use the technical assistance of the Logintrade Procurement platform (https://polfa-tarchomin.logintrade.net/).

E. General provisions

1. The submission of an application for registration in the procedure or an offer is tantamount to accepting all terms of the procedure without reservations.
2. The Contracting party reserves the right to cancel the procedure without giving any reason.
3. The Suppliers are not entitled to submit any claims to the Contracting Party in connection with the inquiry and the proceedings conducted under the project, including costs and damages, in particular in the event of cancellation of the procedure by the Contracting Party or selection of another Supplier.
4. Withdrawal by the Contracting Party from concluding the contract in the event of notifying the Supplier about the selection of its offer, it may not be the basis for claims for the costs of participation in the procedure.
5. In the course of evaluating the submitted applications, the Contracting Party may demand explanations from the bidders regarding the content of the submitted documents.
6. If the application does not contain all the required elements, the Contracting Authority may, in justified cases, call the Supplier to supplement it.
7. The Contracting Party reserves the right to record sound during technical meetings with Bidders during the tender procedure. Sending the offer will be treated as giving consent to record the meetings.

F. Cooperation and communication

1. All questions should be submitted via the Procurement Platform, the Message tab of the proceeding.
2. All responses will be provided to Suppliers via the Procurement Platform, the Message tab of the procedure in question.
3. The persons authorized to contact the bidders on Monday-Friday between 07.00-15.00

Please attach a signed or annotated - confidentiality agreement (draft attached)

LISTA ZAŁĄCZNIKÓW

Lp.	Dokumenty
1.	RFP-BioequivalencestudiesKolistynaABM.pdf
2.	TOC_2024Colistimethate.xlsx
3.	Confidentiality Agreement.doc

PRODUKTY

Lp.	Produkt	Indeks/Nr produktu	Ilość	Jednostka miary	Kategoria zakupowa
1.	Clinical operations		1	szt.	Badania biorównoważności
2.	Bioanalytics		1	szt.	Badania biorównoważności
3.	Project management, protocols and CTA submission		1	szt.	Badania biorównoważności

KRYTERIA FORMALNE (WARUNKI UDZIAŁU W POSTĘPOWANIU):

Lp.	Kryterium
1.	Termin płatności: 30 dni
2.	Miejsce dostawy: siedziba
3.	Koszt transportu: po stronie dostawcy
4.	Dodatkowe warunki formalne: **Signed or annotated - confidentiality agreement (draft attached)**

DODATKOWE PYTANIA DO OFERTY

Lp.	Pytanie
Brak pozycji	

SKŁADANIE OFERT

Zezwól na składanie ofert częściowych	nie
Zezwól na składanie ofert na zamienniki	nie
Zezwól na dodatkowe uwagi do produktów	nie
Zezwól na korygowanie ofert do momentu zakończenia przyjmowania ofert	tak
Zezwól na składanie ofert w przypadku braku spełniania kryteriów formalnych	nie
Zezwól na składanie ofert w innych walutach	nie
Zezwól na składanie ofert na inne ilości	nie
Zezwól na składanie ofert wariantowych	nie