



ENAMINE, Confidential

Quotation Date: May 16, 2023

Attn.: [REDACTED]
Scientist [REDACTED] group
Institute of Organic Chemistry and Biochemistry of the CAS
Flemingovo nám. 2
166 10 Praha 6
Czech Republic
[REDACTED]

Quotation #BEIO20230516YK-2

This Quote is pursuant to the Framework Service Agreement ("Agreement") between Enamine Ltd ("Enamine") and Institute of Organic Chemistry and Biochemistry of the CAS dated July 10, 2019. The parties agree that Enamine shall perform the work in accordance with this Quote and subject to all the terms and conditions of the Agreement.

#	Study title	Report delivery time*	Rate, EUR	Qty	Price, EUR
1.	Pharmacokinetic Study in CD-1 mice (cassette dosing): 3 compounds , IV delivery route, one dose level (1 mg/kg each cpd); 7 time points; 3 mice per time point, 22 mice in total; plasma samples analysis.	4 weeks	3,215.00	1	3,215.00
TOTAL amount, EUR:					3,215.00

*After receipt of the test articles at Enamine facility

Notes: At least 3 mg of each dry compound is required to run the PK study as described. We do not need to know structures of the molecules for testing. However, we ask our customers to provide brutto formulas for all studies involving MS detection.

Pharmacokinetic Study Design

Background and Service Details: This pharmacokinetic (PK) study will be run in **male CD-1 mice**. The goal is to determine levels of the test articles in blood plasma over time after a single dose. This PK study will involve cassette dosing of **3 compounds**, **intravenous delivery route** (IV), one dose level (1 mg/kg for each compound) and **7 time points** (e.g. 5 min, 15 min, 30 min, 60 min, 120 min, 240 min, and 480 min). **3 animals per each time point** group will be used in this study, plus one control animal ("0") – 22 mice in total. We will use one specific dosing formulation selected by the customer (DMSO – PBS (7%:93%, v/v)). We will prepare plasma samples by terminal sampling at each time point. The compound concentrations in plasma will be measured by LC-MS/MS. The service includes development and validation of the analytical procedures. We use Applied Biosystems/PE Sciex API3000 mass spectrometer (if needed API 4000QTrap, API5000) and Shimadzu HPLC.



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Deliverable: A detailed study report in electronic format including description of the study design, analytical method, measured test article concentrations, calculations of relevant PK parameters and graphs using Phoenix WinNonlin software.

Sample Submission: For a PK study with above design, **3-4 mg** of each dry compound is required. We do not need to know structures of the molecules for testing. However, we ask our customers to provide brutto formulas for all studies involving MS detection.