


| | | | |
|---|-------------|-----------------------------|------------------------------------|
|  | | Financial Requirements 2023 | Clinical trial Center |
| AAHRPP-DSQ-070 | Version 7.0 | | Date d'application : 20/06/2023 |

Dear Sponsors / CRO,

The main budget only includes the procedures performed by the Principal Investigator and the Study Coordinator.

Separate exhibits to the main contract must be drawn up with different technical departments involved in a study. The fees in this document constitute a basis for work. Each exhibit must still be validated by the responsible of the department.

All the fees are inclusive of the institutional overhead.

The fees are not exhaustive and will be adapted according to the specificities of your study. The protocol and the imaging manual procedure will be sent to the departments.

Any other department not mentioned in this document may be involved in clinical research. Their specific fees will be provided on a case-by-case basis.

1. NEGOTIATION OF GENERAL BUDGET

Our budget is established as follows:

- Principal Investigator working time x hourly rate (180€/hour)
- Study Coordinator working time x hourly rate (80€/hour)
- If not standard of care: Use of the facility if applicable
 - Outpatient clinic : a cost per consultation in function of the specialization of the Principal Investigator
 - One-day clinic: 325€ for the monotherapy or 375€ for the combitherapy
 - Hospitalization: 1.048€ (Official INAMI price for the Cliniques Universitaires Saint-Luc)
- If not standard of care: technical acts which will be mentioned in a separate budget.

| Institutional additional costs (PI budget) – not negotiable | | |
|--|--------------|--|
| Start-up fee : | | |
| Interventional trials | 4.000€ | This fee is independent of the patient recruitment. It covers the time spent by the investigator and by the study coordinator before the study starts and the access badges to the patients' files. The payment must be made upon signature of the agreement. |
| Observational, retrospective or phase 4 studies | 1.500€ | |
| Maintenance fee : | | |
| Interventional trials | 1.200€ /year | This fee is applicable upon the signature of the contract, and each year thereafter, and covers the administrative study-related fees. |
| Observational, retrospective or phase 4 studies | 600€/year | |

| Institutional additional costs (PI budget) – not negotiable | | |
|--|--------------------------------|--|
| Archiving fee (if not organized by the sponsor) | 150€/box | For 25 years (size : 340mm x 390mm x 360mm) |
| Audit fees : For the PI For the study coordinator | 180€/h 80€/h | In case of CRO or sponsor driven audit |
| Administrative amendment fee Protocol amendment involving budget | 250€ 500€ | For each amendment to the investigator agreement sent for signature and requested by the sponsor. |
| Patient travel fees | / | What is foreseen? In respect of the GDPR, in case of the use of a third-party vendor, the patient must have the choice with a system of payment through the Institution. |
| Screen failures | / | Best option : Same fee than the screening visit |
| Particular fees | / | Such as long lasting trials, transfer of material from other sites, etc... > To be discussed per protocol. |
| Transfer of archival tissue from another hospital to Saint-Luc | 180€ | |
| SAE or Adverse Event of Special Interest | 200€/event | |
| Unscheduled visit | 250€ + procedures performed | |
| Phase 1 trial specific fees (in addition of the institutional additional costs) | | |
| Protocol start-up fee | 3.000€ | Payable upon signature of the contract |
| Premature closure fee | 5.000€ | Compensation for premature closure due to sponsor's decision without possibility of inclusion |

Explanation of the difference between the Start-up fee and the Maintenance fee

- ✓ The PI start-up fee covers :
 - feasibility questionnaire
 - pre study visit - qualification
 - protocol and IB evaluation

- EC submission process
 - contract-budget evaluation
 - contacts with different hospital services involved in study
 - initiation visit
 - patient roadmaps preparation according to flow chart
 - investigator meeting
 - trainings required by sponsor
 - preparation of documents required by sponsor : CV and other qualifications, lab normal values, lab certificates
 - lab samples preparation-shipment procedures training – IATA certification
 - administration : lab kits request, dry ice
 - control of calibration and maintenance of study equipment such as : centrifuges, fridges, freezers, ECG, tensiometer
- ✓ The Annual Maintenance fee : This fee covers all the administrative tasks which are study-related and not patient-related.
- Administrative charge for EC submission
 - Pharmacovigilance
 - Activation vendors
 - Technical issue (related to e-tablet/electronic system)
 - Lab kits re-ordering
 - Re-training (not included in Start-up fee)
 - Conference call (Investigator)

Medical Director's signature for validation of institutional and specific costs

Jean-Louis Vanovershelde

Signé numériquement par
CONNECTIVE NV - Connective
eSignatures de la part de Jean-Louis
Vanovershelde
Date : 19/06/2023 13:40:20

2. NEGOTIATION OF ADDITIONAL SERVICES

Negotiation is carried out with the contact person of each service: CoFi contact should be informed of the status.

| Department | Contact | Responsible signature |
|---|--|--|
| Laboratory (local analysis) | damien.gruson@saintluc.uclouvain.be soumaya.wasmine@saintluc.uclouvain.be | Prof. D. Gruson, Head of Clinical Laboratory (<i>and referent for clinical research</i>) |
| Pathology (for archived or fresh human material) | biobanque@saintluc.uclouvain.be selda.aydin@saintluc.uclouvain.be | Prof S. Aydin – Head of Biobank |
| Radiology | muriel.decooman@saintluc.uclouvain.be (commercial studies in oncology except hematology) valerie.rosseels@saintluc.uclouvain.be (other commercial studies) | Mrs Muriel De Cooman or Mrs Valérie Rosseels – Clinical research coordinator of radiology department |
| Nuclear Medicine | francois.jamar@saintluc.uclouvain.be | Prof F. Jamar |
| Pharmacy | pharmaetudes@saintluc.uclouvain.be | Mrs D. Wouters – Head of Pharmacy |
| Cardiology | virginie.donnay@saintluc.uclouvain.be thomas.paulus@saintluc.uclouvain.be christophe.beuloye@saintluc.uclouvain.be | Prof C. Beuloye |
| Rheumatology | dominique.dienst@saintluc.uclouvain.be marie.mzoughui@saintluc.uclouvain.be | Prof F. Houssiau |

3. BUDGET OF TECHNICAL DEPARTMENTS

3.1. Pharmacy

| * = mandatory fields | Frequency | Transfer study | Non-transfer study | Non-transfer study with preparation | Device |
|---|---|----------------|----------------------------|-------------------------------------|------------------------------|
| | | (€ HTVA) | (€ HTVA) | (€ HTVA) | (€ HTVA) |
| FIXED COSTS* | | | | | |
| Administrative cost incl SIV and COV | per study :1 | 550 € | 800 € | 1100 € | 600 € |
| Administrative cost (study not started, signed agreement, without SIV) | if applicable | 250 € | 400 € | 400 € | 300 € |
| Archiving 25 years | per study :1 | 120 € | 120 € 220 € > 2 binders | 120 € 220€ > 2 binders | 120 € 220€ > 2 binders |
| Audit | / h of pharmacist | 130 € | 130 € | 130 € | 130 € |
| ANNUAL FEES* | | | | | |
| Maintenance (work/follow-up of the Pharmacist in the cause of the Study: CRA, monitoring visits, etc...) | from first reception / month until COV pharmacy | 35 € | 40 € | 50 € | 40 € |
| Storage fee room temperature (volume <=2x40x60x30cm) | / month | - | 35 € | 40 € | - |
| Storage fee (fridge or freezer) (volume <=2x40x60x30cm) | / month | - | 40 € | 40 € | - |
| Storage freezer (volume <= 2x 40x60x30) | / month | - | 50 € | 50 € | - |
| Additional fee if volume > 2x (40x60x30) | / month | - | 10 € | 10 € | - |
| VARIABLE COSTS* | | | | | |
| Reception fee : IMP at room temperature | / shipment | 55 € | 55 € | 55 € | 100 € (first reception) /55€ |
| Reception fee : fridge T° or congelation | / shipment | 65 € | 65 € | 65 € | |

| * = mandatory fields | Frequency | Transfer study | Non-transfer study | Non-transfer study with preparation | Device |
|--|-----------------------|---|--------------------|--|----------|
| | | (€ HTVA) | (€ HTVA) | (€ HTVA) | (€ HTVA) |
| Return of recycling boxes (per study) | per study :1 | 150 € | 150 € | 150 € | 150 € |
| Global Dispensation | / dispensation | 25 € | - | - | 25 € |
| IMP Nominative Dispensation | /dispensation and log | - | 35 € | 35 € (medication without preparation) | - |
| Special handling (special procedure for dispensation, blinding, repackaging...) | /dispensation | 10 € | 10 € | 10 € | 10 € |
| Standard arm per os: Nominative Dispensation | /dispensation and log | - | 10 € | 10 € | 10 € |
| Co-medication Nominative dispensation | /dispensation and log | - | 10 € | 10 € | 10 € |
| FEEES IF PREPARATION AT THE PHARMACY | | | | | |
| IMP preparation (incl diluent, sterile preparation in the clean room, necessary equipments for the operating staff and accountability log) | / preparation | - | - | 120 € | - |
| Preparation of ATMP / radiopharmacy | - | - | - | 220 € | - |
| Standard arm preparation (INAMI) (incl diluent and accountability) | / preparation | - | - | 10 € | - |
| Special material (filters, needleless system, administration material, ...) | / preparation | - | - | delivered or invoiced | - |
| FEE IF APPLICABLE | | | | | |
| Randomization | per study :1 | 60 € | 60 € | 60 € | 60 € |
| Accountability, return in IWRS /eCRF | Per patient | 50 € | 50 € | 50 € | 50 € |
| Destruction in IWRS /eCRF | Per destruction | 20 € | 20 € | 20 € | 20 € |
| Order: IMP /co-medication/material ordered by pharmacy (not delivered by sponsor) | per unit | price (ambulatory) + 10% (max 5 €/unit) | | | |

| * = mandatory fields | Frequency | Transfer study | Non-transfer study | Non-transfer study with preparation | Device |
|---|-----------------|----------------|--------------------|-------------------------------------|----------|
| | | (€ HTVA) | (€ HTVA) | (€ HTVA) | (€ HTVA) |
| NB: please provide the list of co-medications requested (please note that the oncology team is asking for the anti-emetic reimbursement) | | | | | |
| Emla (in case of pediatric studies) | 5,52 € per unit | - | - | - | - |
| Urine pregnancy test (if not provided by the sponsor) | 10 € per unit | - | - | - | - |
| Non automatic resupply: Order: IMP/co-medication/material ordered by pharmacy and delivered by sponsor (request) | per study :1 | 150 € | 150 € | 150 € | 150 € |
| Re-labelling | / relabeling | 80 € | 80 € | 80 € | 80 € |
| Duty call (return of the Pharmacist outside of the working hours: Monday to Friday 8H-17H) | / duty call | 330 € | 330 € | 330 € | 330 € |
| Storage of used/expired or returned medication and material (if more than 6 month at the pharmacy before return to sponsor or destruction) | per case | 110 € | 110 € | 110 € | NA |
| Destruction fee * | per year | 330 € | 330 € | 330 € | 330 € |
| Return to sponsor (exl cost for transport): used and unused IMP/non-IMP (per study) * | per study :1 | 150 € | 150 € | 150 € | 150 € |
| Remote Monitoring (Excl COV) | / 30 minutes | 140 € | 140 € | 140 € | 140 € |

Mention: 901E study number (will be provided at each study)

The exhibit must be signed by Mrs D. Wouters, Head of Pharmacy department.
In case of absence, the exhibit can be signed by Mrs Stéphanie Quennery.

3.2. Radiology

| | Frequency | COSTS (€ HTVA) |
|--|--------------------------|-------------------|
| RADIOLOGY START-UP FEES AND ADDITIONAL COSTS FOR COMMERCIAL STUDIES | | |
| Start-up fees (study without OR with central reader). After contract signature, the Radiology Department will draw up an invoice for start-up fees to the attention of the Sponsor. | - | 350 € or 600 € |
| Additional cost if Phantom/Test to send for qualification | - | 400 € |
| Additional cost if Phantom/Test to resend during study for parameters modification OR modification of central reader during the study OR implementation of a central reader during the study WITH parameters modification | - | 400 € |
| Additional cost if Phantom/Test to resend during study for verification of parameters and calibration OR modification of central reader during the study OR implementation of a central reader during the study WITHOUT parameters modification | - | 200 € |
| Additional cost if sponsor ask to upload image of other CT/MRI/X ray exam (example: pre-study exam, routine exam, not performed for the study ...) | per upload | 50 € |
| Additional cost if set up by the referent radiologist before start of study | Radiologist fee per hour | 150 € |
| RADIOLOGY FEES FOR EXAMINATIONS FOR COMMERCIAL ONCOLOGIC STUDIES | | |
| CT brain/neck/chest/abdomen/pelvis/bone/other site with IV contrast NON SOC | - | 610 € |
| CT brain/neck/chest/abdomen/pelvis/bone/other site without IV contrast NON SOC | - | 510 € |
| CT brain/neck/chest/abdomen/pelvis/bone/other site with or without IV contrast SOC (Fee per examination considered as « standard of care » : covers the Data management and radiologist's workload) | - | 160 € |
| MRI brain/neck/abdomen/pelvis/bone/soft tissue/bone narrow/whole body/other | - | 610 € |
| Fee for tumor measurement done by radiologist | per exam | 150 € |
| Chest X ray | - | 86,59 € |
| Localized bone X-ray if visible lesion at Bonescan (scintigraphy in Nuclear Department) | - | 159,06 € |
| Fee included examination, institutional overhead and radiology data management. Valid with our standard acquisition parameters. | | |

| | Frequency | COSTS |
|--|-----------|----------|
| | | (€ HTVA) |
| RADIOLOGY FEES FOR TUMOR BIOPSY PERFORMED IN RADIOLOGY DPT FOR COMMERCIAL ONCOLOGIC STUDIES | | |
| Liver/deep node biopsy (with US) | - | 891,89 € |
| Liver/deep node biopsy (with CT scan) | - | 942,27 € |
| Renal biopsy (with US) | | 876,01 € |
| Renal biopsy (with CT scan) | | 926,40 € |
| Breast biopsy (Microbiopsy with US) | - | 503,12 € |
| Breast biopsy (Macrobiopsy with stereotaxy) | - | 950,43 € |
| Breast biopsy (punch) | - | 421,55 € |
| Pulmonary biopsy (with CT scan) | - | 782,13 € |
| Bone biopsy (with CT scan) | - | 890,40 € |
| Soft tissue biopsy (with CT scan) | - | 698,86 € |
| Node/Cutaneous biopsy with US (if not possible in dermatology department) | - | 528,69 € |
| Fee included examination (US or CT scan + biopsy), institutional overhead, material (needle ...), medication, radiology data management and interventional radiologist fees. | | |

These fees are given for information and are not final.

For each new study, a budget will be established by the radiology coordinators according to the protocol and vendor requests. This budget will be submitted for approval before being finalized.

The exhibit must be signed by Mrs Muriel De Cooman or Mrs Valérie Rosseels.

3.3. Laboratory

| | Frequency | COSTS |
|--------------|-----------|----------|
| | | (€ HTVA) |
| | | |
| Start-up fee | once | 500 € |

Laboratory fees: please contact Mrs Wasmine Soumaya → soumaya.wasmine@saintluc.uclouvain.be

Mention: Invoice number – 865E

The exhibit must be signed by Prof. Damien Gruson, Head of Laboratory Department.
In case of absence, the exhibit can be signed by Prof. Vincent Haufroid.

3.4. Nuclear Medicine

| | Frequency | COSTS |
|--|-----------|----------|
| | | (€ HTVA) |
| Start-up fee (Excluding technical procedures; e.g.phantom) | - | 250 € |
| MUGA | - | 486 € |
| Bone scan | - | 486 € |
| PET-SCAN | - | 1.782 € |
| CD/upload | - | 65 € |
| Can be adapted in function of the specificities or additional requirements of each protocol. | | |

Mention: Invoice number – 531E

The exhibit must be signed by Prof. François Jamar, clinical research responsible for Nuclear Medicine.
In case of absence, the exhibit can be signed by Prof. Renaud Lhommel, Head of the Department.

3.5. Pathology

| | Unit | COSTS per unit |
|--|-----------|--|
| | | (€ HTVA) |
| Start-up fee | /study | 500 |
| Exit from the archives (FFPE & frozen blocks) | / case | 20 € |
| Case selection and evaluation | / case | 50 € |
| Histological analysis (with the report including the evaluation of the tumour cells) | /analysis | 300€ |
| Preparation of unstained slides on positively charged slides (white SuperFrost Plus slides) | / slide | 20 € |
| Preparation of unstained slides on standard slides (colored SuperFrost slides) | / slide | 20 € |
| Sections preparation: | | |
| - in RNase free tubes (FFPE blocks) | / block | 50 € |
| - in RNase free tubes (Frozen blocks) | / block | 60 € |
| - on slides (FFPE blocks) | / slide | 30 € |
| - on slides (Frozen blocks) | / slide | 40 € |
| Preparation of FFPE blocks | / block | 200 € |
| Preparation of frozen OCT embedded blocks | / block | 200 € |
| In Situ Hybridization (ISH) : HER2 | / slide | 450 € (including the pathologist case revision) |
| In Situ Hybridization (FISH): ALK | / slide | 250 € (including the pathologist case revision) |
| In Situ Hybridization (FISH): ROS1 | / slide | 250 € (including the pathologist case revision) |
| In Situ Hybridization (ISH): EBER | / slide | 300 € (including the pathologist case revision) |
| In Situ Hybridization (ISH): HPV high & low risk | / slide | 300 € (including the pathologist case revision) |

| | Unit | COSTS per unit |
|--|--------------|--|
| | | (€ HTVA) |
| PCR: HPV | / test | 300 € (including the pathologist case revision) |
| HE staining | / slide | 10 € |
| Special staining | / slide | 25 € |
| Pathology report | / hour | 300 € |
| Estimation of % tumor cells | / 15 minutes | 50 € |
| Immunohistochemistry (IHC) INAMI accredited | / slide | 250 € (including the pathologist case revision) |
| Immunohistochemistry (IHC) standard | / slide | 80 € |
| Immunohistochemistry (IHC) p16 | / slide | 110 € |
| Renal biopsy: | | |
| - 1 HE staining | / slide | 10 € |
| - 3 special stains | / staining | 25 € (total 75 €) |
| - 1 immunofluorescence markers | / slide | 40 € |
| - Pathology report | / hour | 300 € |
| Transmission electron microscopy (T.E.M.) | / sample | 900 € |
| Preparation of Material for NGS analysis: | | |
| - Biopsy | | Total cost: 260 € |
| ▪ 1 HE | / slide | 10 € |
| ▪ 10 unstained slides | / 10 slides | 200 € |
| ▪ Revision of the slide | / slide | 50 € |
| - Macroscopic specimen | | Total cost: 160 € |
| ▪ 1 HE | / slide | 10 € |
| ▪ 5 unstained slides | / 5 slides | 100 € |
| ▪ Revision of the slide | / slide | 50 € |

Mention: Invoice number – 868E

This exhibit will be signed by Prof Selda Aydin, head of the biobank.

3.6. Rheumatology

| | Frequency | COSTS |
|--|-----------|----------|
| | | (€ HTVA) |
| Start-up fee | - | 950 € |
| Lumbar vertebrae DXA (L1-L4 together) & proximal femur (total hip and neck) | - | 300 € |
| Wrist | - | 300 € |
| W body Composition | - | 345 € |
| QUALITY CONTROL IF CENTRAL READER IF REQUESTED | | |
| Total Baseline IQC | - | 297 € |
| Total Cross Calibration | - | 297 € |
| Monthly IQC Data Management and Submission | - | 90 € |
| Training if requested | - | 175 € |
| IF PATIENT YOUNGER THAN 5 YEARS OR PATIENT WITH COMPLICATION | | |
| Under narcosis | - | 150 € |
| DXA Lumbar & Hip | - | 360 € |
| DXA WRIST | - | 360 € |
| W body | - | 518 € |
| ADD COSTS FOR PAEDIATRICS STUDIES LESS THAN 18 Y | | |
| Growth curve for children less than 18 y Add for calculation | - | 35 € |

Per patient per DXA VISIT & scan INDEPENDENTLY FROM CRF completion. Screen failure will be paid without rate as independent of the department.

DEXA will be paid for scans required by the protocol and as requested by the Principal Investigator. In the event it is not possible to obtain adequate scans.

DEXA will be paid, provided all reasonable measures are used in obtaining the scans. If adequate scans can't be obtained, it will be clearly documented in the medical report"

Mention: Invoice number – 517E

The exhibit must be signed by Prof. Frédéric Houssiau, Head of Rheumatology department. In case of absence, the exhibit can be signed by Prof. A. Nzeuseu.

3.7. Cardiology

| | Frequency | COSTS |
|---|-----------------|---------------|
| | | (€ HTVA) |
| Simple ECG (A separate exhibit with the cardiology department is not necessary if simple ECG is the only cardiology exam requested by the Protocol -->To be included in the PI budget) | / ECG | 60 € |
| Cardiac echography (A separate exhibit with the cardiology department is necessary) | / echo | 350 € |
| Simple ECG + Cardiac echography (A separate exhibit with the cardiology department is necessary) | / ECG / echo | 60 € 350 € |
| ECG with specific requirements (measures, interpretation...) (A separate exhibit with the cardiology department is necessary) | - | TBD |
| In case you provide the ECG machine, this procedure will be included in the data management fees of the main budget. | | |

Mention: Invoice number – 752E

The exhibit must be signed by Prof. Christophe Beauloye, Head of Cardiology department.

3.8. Pneumology

| | Frequency | COSTS |
|--|-----------|----------|
| | | (€ HTVA) |
| PROF. LIISTRO G. | | |
| Spirometry (FEV1 + FVC, FEV1/FVC) | - | 15,15 € |
| Spirometry + bronchoconstriction (Histamine) | - | 57,74 € |
| Residual volume (and TLC measurement) | - | 60,58 € |
| Spirography + bronchodilator test | - | 30,29 € |
| Spirography + bronchodilator test (under 7 years old) | - | 33 € |
| Diffusion capacity (= DLCO) | - | 60,58 € |
| Expired NO Measurement (Cost not reimbursed) | - | 12 € |
| Intellectual fee | / visit | 100 € |
| These tests can be combined. | | |

Signature by Prof. Pilette after validation of Prof. Liistro's budget

Mention: Invoice number – 545E

| | Frequency | COSTS |
|--|-----------|----------|
| | | (€ HTVA) |
| PROF. PILETTE Ch. | | |
| Bronchoscopy with broncho-alveolar lavage | - | 197,09 € |
| Intellectual fee | / visit | 75 € |

Signature by Prof. Pilette

Mention: Invoice number – 542E

The exhibit must be signed by Prof. Charles Pilette, Head of Pneumology department.

3.9. Ophthalmology

| | Frequency | COSTS per procedure |
|---|-----------|---------------------|
| | | (€ HTVA) |
| Ophthalmologist consultation (Including PI fee) | - | 110 € |
| Visual Field | - | 35 € |
| Funduscopy - dilated funduscopy | - | 10 € |
| Slit lamp | - | 10 € |
| Tonometry (Goldmann) - IOP | - | 10 € |
| Fluoroangiography with protocol | - | 275 € |
| Optical coherence tomography (OCT) (Mesure épaisseur de la rétine) | - | 90 € |
| Endothelial cells count by specular microscopy | - | 70 € |
| Visual acuity | - | 10 € |
| Eyes Ultrasound (bimonthly) (1/2 yx) | - | 25 € |
| Electroretinography | - | 75 € |
| Counting endothelial cells of the cornea (tachymetric corner) | - | 70 € |
| Schirmer Test | - | 10 € |
| Color fundus photography | - | 50 € |
| Ocular motility | | 50 € |

Mention: Invoice number –

The exhibit must be signed by the ophthalmologist who will be in charge of the study.