AGENZIA ITALIANA DEL FARMACO

AIFA Call 2023 for Independent Research

SECONDARY PREVENTION IN CARDIO-CEREBROVASCULAR DISEASES IN GENDER MEDICINE

Principal Investigator (PI)

Principal Investigator PEC

Please indicate the certified email (PEC) address of Scientific Responsible. If not available, please report the PEC address of the PI Centre.

Date of birth (dd/mm/yy)

Grant in AIFA Call past editions

Please indicate if PI received AIFA grant in the past editions.

YES ○

NO ◉

If yes, please indicate the year(s) :

Proposal title

Running title (max 50 characters)

Acronym title (max 10 characters)

Key words (max 10 keywords)

Keyword 1	
Keyword 2	
Keyword 3	
Keyword 4	
Keyword 5	
Keyword 6	
Keyword 7	
Keyword 8	
Keyword 9	
Keyword 10	

Thematic area

Please indicate the most appropriate area for your study

- ○ Secondary prevention of myocardial infarction
- ○ Secondary prevention of stroke
- ○ Secondary prevention of hearth failure

Institutional Address (Public Institution or Non Profit Institution)

Please indicate the address of the Institution which applies.

Institutional Address PEC

Please indicate the certified email (PEC) address of proposing Institution

PI Centre Address (Public Institution or Non Profit Institution)

Please indicate the address of the centre where Principal Investigator is employed. The centre here indicated is the Coordinating Centre.

Operative Units Institutional Address

Please list all the Clinical Centres directly involved in patients' recruitment and any Centres involved in other tasks/activities as statistics, administrative, management, etc. The foreign Centres, if applicable, should be listed after the national centres. Please detail responsible investigator, name and address of the Collaborating Centres.

Pharmacological Treatment

Please list every drug used both in treatment and control groups. If a therapeutic class is used instead of a specific drug, please indicate the class.
The drugs/drug classes that can only be included are: antithrombotics (acetylsalicylic acid and other antiplatelets; anticoagulants), statins, beta-blockers, ACE-1 inhibitors, sartans and SGLT-2 inhibitors.

Drug			**Marketed**
Active ingredient/ Drug class (max. 50 chars)	Reimburse-ment class within the NHS (A,C,H)	Drug for treatment group **(T)** Drug for standard group **(C)** *(Can be clicked both)* Not Applicable **(NA)**	Patent / Off patent

Drug			**Marketed**
Active ingredient/ Drug class (max. 50 chars)	Reimburse -ment class within the NHS (A,C,H)	Drug for treatment group **(T)** Drug for standard group **(C)** *(Can be clicked both)* Not Applicable **(NA)**	Patent / Off patent

Total study duration

Please note that the maximum allowed study duration (from AC/CE approval to last patient follow-up visit) is 36 months.

Study duration in months:

Study timing and recruitment

Timing (in months)
Enrollment phase
Treatment and follow-up
Estimation of number of patients to be recruited

Synopsis (max 4.000 characters)

Please describe: Background, Objectives, Methods, Expected results.

Background (max 4.000 characters)

Please describe an updated review of relevant literature related to disease, available treatments and therapeutic regimen information.

Rationale and Innovativeness (max 4.000 characters)

Please describe the clinical question that will be investigated in the study and what research findings will add to existing evidence. If applicable, please describe innovativeness of the study.

Impact on the National Health Service (max 4.000 characters)

Please describe the impact on the Italian National Health Service (NHS) and transferability on general population.

Objectives of the study (max 4.000 characters)

Please report the purpose of the study, the primary and secondary objectives of the study according to the statistical hypothesis.

Study Design (max 4.000 characters)

Please describe the type of experimental design selected for clinical trial in order to answer the clinical question and to produce valid results.

Study Population (max 4.000 characters)

Please report study poulation characteristics and the clinical setting (hospital, general practice, etc.) where the study will be conducted; enrolment procedure and accrual time. Please specify withdrawal criteria and procedures.

Inclusion criteria (max 4.000 characters)

Exclusion criteria (max 4.000 characters)

Intervention (max 4.000 characters)

Please provide detailed information about treatments (or other type of intervention) for each group (treatment and control). The following items must be included:

- Dose (and dose escalation) and the dosage form, packaging and labelling of the IMP;
- Duration of treatment (including number and duration of the cycles, if applicable) and the follow-up period for each IMP/trial treatment group/arm of the trial;
- Route of administration;
- Medication/treatment (including rescue medication) permitted and not permitted before and/or during the trial;
- Procedure for monitoring subjects compliance;
- Description of the "stopping rules" or "discontinuation criteria" for individual subjects, part of trial, and entire trial;
- Accountability procedures for the IMP, including placebo and comparator if any.

Endpoints and Outcomes (max 4.000 characters)

Please indicate the primary and secondary endpoints and related outcome measures; the relation between subjective/objective evaluation of endpoints, justifications to support the validity of any surrogate or composite endpoints, if applicable.

Methods (max 6.000 characters)

A description of the measures taken to minimize/avoid bias, including (but not limited to):

Randomization. Please report methods used to generate the random allocation sequence. Central randomization should be preferred; other randomization procedures should be adequately motivated. Include a description of maintenance of trial treatment randomisation codes and procedures for breaking the code.

Blinding (masking). Please describe whether or not the personnrl involvedrin administering interventions and assessing outcomes is aware of group assignment and if not, how the success of masking is assessed.

Data Collection. Please report data that will be collected; the forms/tools used for the retrieval of information and their validity and reliability; the measures/indicators used; the potential sources of bias in the retrieval of information regarding study subjects and interventions/treatments; duration and frequency of follow up; missing data. When the use of an electronic clinical reporting form (e-CRF) is envisaged, only validated systems that address traceability are acceptable (for instance, excel spreadsheets do not represent an adequate system for recording data). Please include in this section the identification of source data definition.

Statistical considerations (max 4,000 characters)

Please describe in detail the statistical hypothesis (e.g. superiority, equivalence or non-inferiority for the primary endpoint(s).

Sample size estimates. Please report the estimate of sample size and how it is determined, indicating information on the following components of sample size estimation: the power,the significance levels,the underlying event rate in the population under investigation,the size of the treatment effect, the standard deviation, the expected proportion of drop-outs. Adjustment for other factors affecting the sample size calculation (e.g. expected compliance rates), should also be reported. For studies of equivalence - non inferioriority the maximum acceptable difference should be stated. For composite endpoints, please indicate how different parameters contribute to their definition and to sample size calculation.

Statistical analysis. Please report the main statistical analyses that will be carried out. Definition of the study population for main analysis, error probabilities, brief description of the statistical techniques, methods for additional analyses, subgroup analyses (if planned).

Explanation of each interim analysis (if planned) and predefined stopping rules should also be clearly stated.

Timing (max 4.000 characters)

Please report the duration of the study (patient enrolment; treatment duration, follow up, etc.); predefined check points for the evaluation of work in progress.

Feasibility (max 4.000 characters)

Organizational characteristics. Please describe the participating centres, the specialties and experience needed for conducting the study. In case of multicentre study specify:

- the institutions/units in charge of coordinating the study, assigning treatment, monitoring the procedures;
- the presence of steering committees and/or data monitoring committees (when applicable);
- the presence and the organization of centralized laboratories (when applicable).

Feasibility. Please describe previous experience of the principal investigator; previous experience of the institution that will coordinate the study; available technology that may be relevant for the study.

GCP and Ethical aspects (max 4.000 characters)

Good Clinical Practices. Clinical studies are required to be conducted in accordance with Good clinical practices (GCP). Please discuss the benefit/risk ratio, the specific hazards of the study (e.g. risks for the patients, complexity of the study design, validity of the information retrieval, etc.), the procedures of risk minimisation (e.g. training activities, review of eligibility prior to randomization, data verification, drug accountability, drug reconciliation, etc.), the characteristics and frequency of monitoring activities and the institution(s) that will be in charge of this task.
Ethical aspects. Please describe the potential risks for study subjects, either related to physical/psychological domains or to a possible excessive interference with the subject privacy, and the procedures that will be followed to prevent these potential risks. As above mentioned, it is not necessary to include the documentation required by the Italian ethics committees.
Insurance. Please include a comment about the application of the law about the study insurance.

References (max 4.000 characters or max 20/25 references)

Please report only the references that are strictly relevant to the study proposal. References should include authors (when there are more than 6 authors, report the first 3 authors), title, book or journal, year, volume number and page numbers. For books, the publisher should also be reported.

Budget

B1. Operative Units

Units	Name	Institution	Responsible
CC			
UO1			
UO2			
UO3			
UO4			
UO5			
UO6			
UO7			
UO8			
UO9			
UO10			
UO11			
UO12			
UO13			
UO14			
UO15			
UO16			
UO17			
UO18			
UO19			
UO20			

Units	Name	Institution	Responsible
UO21			
UO22			
UO23			
UO24			
UO25			
UO26			
UO27			
UO28			
UO29			
UO30			
UO31			
UO32			
UO33			
UO34			
UO35			
UO36			
UO37			
UO38			
UO39			
UO40			
UO41			

Units	Name	Institution	Responsible
UOпн			
UOпо			
UOпп			
UOпр			
UOпс			
UOпт			
UOпу			
UOпф			
UOрл			
UOрм			
UOрн			
UOро			
UOрп			
UOрр			
UOрс			
UOрт			
UOру			
UOрф			
UOсл			
UOсм			
UOсн			

B2. Personnel

UO	Degree	Tasks	Contract	Duration (in months)	Salary per month (€)		% TES[5]	Salary × % time employed on study[3]		Total (€) (Salary × % time employed on study) × duration
					Gross[1]	Charges[2]		Gross	Charges	
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00

1 *Gross salary for employees*
2 *Related charges to be paid by institutions*
3 *When the % of time employed is 100%, report on this column the same values of the previous one*
4 *Total personnel cost for all units*
5 *% time employed on study*

UO	Degree	Tasks	Contract	Duration (in months)	Salary per month (€)		% TES[5]	Salary × % time employed on study[3]		Total (€) (Salary × % time employed on study) × duration
					Gross[1]	Charges[2]		Gross	Charges	
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00

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					Gross[1]	Charges[2]		Gross	Charges	
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00

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3 *When the % of time employed is 100%, report on this column the same values of the previous one*
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5 *% time employed on study*

UO	Degree	Tasks	Contract	Duration (in months)	Salary per month (€)		% TES[5]	Salary × % time employed on study[3]		Total (€) (Salary × % time employed on study) × duration
					Gross[1]	Charges[2]		Gross	Charges	
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00

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5 *% time employed on study*

UO	Degree	Tasks	Contract	Duration (in months)	Salary per month (€)		% TES[5]	Salary × % time employed on study[3]		Total (€) (Salary × % time employed on study) × duration
					Gross[1]	Charges[2]		Gross	Charges	
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00

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4 *Total personnel cost for all units*
5 *% time employed on study*

UO	Degree	Tasks	Contract	Duration (in months)	Salary per month (€)		% TES[5]	Salary × % time employed on study[3]		Total (€) (Salary × % time employed on study) × duration
					Gross[1]	Charges[2]		Gross	Charges	
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00

1 *Gross salary for employees*
2 *Related charges to be paid by institutions*
3 *When the % of time employed is 100%, report on this column the same values of the previous one*
4 *Total personnel cost for all units*
5 *% time employed on study*

UO	Degree	Tasks	Contract	Duration (in months)	Salary per month (€)		% TES[5]	Salary × % time employed on study[3]		Total (€) (Salary × % time employed on study) × duration
					Gross[1]	Charges[2]		Gross	Charges	
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
							0,00%	€ 0,00	€ 0,00	€ 0,00
									Total personnel cost[4]	€ 0,00

1 *Gross salary for employees*
2 *Related charges to be paid by institutions*
3 *When the % of time employed is 100%, report on this column the same values of the previous one*
4 *Total personnel cost for all units*
5 *% time employed on study*

B3. Supplies

Please specify the cost of the main categories of supplies for the entire study.

Categories of Supplies	Brief description, if needed (max 200 chars.)	Total budget for the entire project and for all participating centres (€)
1. Hardware		
2. Software		
3. Consumable materials/Stationery		
4. Device		
5. Labs material		
6. Other costs for supplies		
Total (€)		€ 0,00

B4. Services

Please specify the cost of the main categories of services for the entire study.

Categories	Brief description (max 200 chars.)	Total budget for the entire project and for all participating centres (€)
1. Data collection *(e.g.: e-CRF)*		
2. Study Monitoring		
A. Number of Expected site visit per centre		
B. Number of Clinical centres		
C. Number of Total site visits (*automatic* A x B)		0,00
D. Average cost for site visit (*) (*)*Excluding personnel and/or travels already budgeted in table B2 and/or B6*		
E. Total cost (C x D)		€ 0,00
3. C.R.O. Activities (different from a.m. categories)		
4. Insurance		
5. Publication cost		
6. Collaboration contracts/ consulting (*)		
7. Other		
Total (€)		€ 0,00

(*) "Partita IVA" Included

B5. Drug cost

Please specify the cost of the drugs needed to conduct the entire study (if applicable)

	Brief description (max 200 chars.)	Total budget for the entire study (€)
1. Costs for drug(s) not used according to approved indications, if not provided free of charge by a Company or paid by others (*)		
2. Costs related to drug(s) management (e.g. relabelling)		
3. Costs for placebo (*)		
4. In case the drug(s) will be provided free of charge(s), please specify the provider (*):		
Total (€)		€ 0,00

(*) Costs described in row 1, 3 and 4 are not applicable for this Call

B6. Meetings, conferences, workshops, travels

Please describe the cost of events and specify if the same cost is related to partecipation or to organization

	Brief description, if needed (max 200 chars.)	**Total budget for the entire project and for all participating centres (€)**
1. Coordination meetings		
2. Participation in scientific conferences (in Italy or abroad)		
3. Organization of scientific conferences related to the study project		
4. Travel, food and accomodation		
5. Other (specify)		
Total (€)		€ 0,00

B7. Other

Please describe the costs arising from categories not mentioned above.

Categories	Brief description (max 200 chars.)	Total budget for the entire study and for all participating centres (€)
Total (€)		€ 0,00

B8. General Costs/Overhead

Please specify the impact on financing of each categories. The total of these costs and overhead cannot affect more than 10% of the sum of the expense items previously represented, with the exception of the drug cost.

Categories	**% Incidence (tot max 10%)**	**Total budget for the entire study and for all participating centres (€)**
General costs (brief description)	0,00%	
Overhead	0,00%	
Total (€)		€ 0,00

B9. Overall expected costs for each item of the study as indicated below

Please specify the overall expected costs for each item of the study.

Items	**Total (€)**
Personnel	€ 0,00
Supplies	€ 0,00
Services	€ 0,00
Meetings, conferences, workshops, travels	€ 0,00
Other	€ 0,00
Sub-total (€)	€ 0,00
General costs/Overhead (max 10% of sub-total)	€ 0,00
Drugs cost	€ 0,00
Total (€) (Sub-total + Overhead + Drugs cost)	€ 0,00

Overhead check

If the reported value is a number smaller than zero, the overhead exceed 10% of subtotal in contrast with the Call AIFA requirements.

€ 0,00

B10. Distribution of costs between coordinating and participating centres

Please specify the percentage of budget allocated to each centre. Compile for all UOs inserted into section B.1.

Operative Unit	**% Budget**	**Total Costs (€)**
Coordinating centre	%	
UO1	%	
UO2	%	
UO3	%	
UO4	%	
UO5	%	
UO6	%	
UO7	%	
UO8	%	
UO9	%	
UO10	%	
UO11	%	
UO12	%	
UO13	%	
UO14	%	
UO15	%	
UO16	%	
UO17	%	
UO18	%	
UO19	%	
UO20	%	

Operative Unit	% Budget	Total Costs (€)
¦ hнм	%	
UOнн	%	
UO2o	%	
UOнп	%	
UOнр	%	
UOнс	%	
UOнт	%	
UOну	%	
UOнф	%	
UOoл	%	
UOo1	%	
UOoн	%	
UOoo	%	
UOoп	%	
UOoр	%	
UOoс	%	
UOoт	%	
UOoу	%	
UOoф	%	
UOпл	%	
UOпм	%	

Operative Unit	% Budget	Total Costs (€)
¦ hпн	%	
UOпо	%	
UOпп	%	
UOпр	%	
UO4c	%	
UOпт	%	
UOпу	%	
UOпф	%	
UOрл	%	
UOрм	%	
UOрн	%	
UOро	%	
UOрп	%	
UOрр	%	
UOрс	%	
UOрт	%	
UOру	%	
UOрф	%	
UOсл	%	
UOсм	%	
UOc2	%	
Total (€)	**0,00 %**	€ 0,00

B.11 Budget Annotation (not mandatory)

List of the investigators in charge of the units dedicated to data analysis and to GCP monitoring of the study (max 4.000 characters)

Please report the investigators responsible for the units dedicated to data analysis and GCP monitoring of the study (when applicable). Please be sure to follow applicable monitoring requirements.

Additional documents may be attached in order to explain the study protocol (e.g. reporting form, rating scale, Gantt, etc)