The first 10-year population-based coronary disease genomewide association study (GWAS) in more than 100,000 participants to personalize cardiovascular prevention in Spain. A project of the CORDELIA Study (Collaborative cOhorts Reassembled Data to study mEchanisms and Longterm Incidence of cArdiovascular diseases)

Isaac Subirana (PhD), Anna Camps (MSc) & Jaume Marrugat (PhD, PI) on behalf of CORDELIA investigators contact: isubirana@imim.es

## PURPOSE



Southern Europe is lacking sufficiently large cohorts and DNA biobanks to organize genome-wide association studies (GWAS) with cardiovascular disease (CVD), which come mainly from case-control studies.

1) To identify the genetic characteristics associated with the 10-year incidence of CVD in the Spanish population using a GWAS on ~102,000 participants form 24 Spanish cohorts already created, followed and duly combined.
 2) To test new 10-year genetic risk score (GRS) for CVD adapted to the characteristics of the Spanish population, and to validate previously proposed GRS.



## METHODS

# Study design and population

- A collaborative and multicenter prospective cohort of 31
  population-based pooled cohorts recruited in Spain in the
  last 30 years, with more than 167.000 participants, of whom
  > 102.000 have DNA samples still available.
- Spanish natives or residents 35 and 84 years old, 50% women, free of acute myocardial infarction, stroke or peripheral arterial disease at the time of recruitment.



#### Statistical analyses

- Data management: data collection, data merging, quality control reports, ...
- Genetic analyses: Genome Wide
  Association (GWAS) of observed and imputed SNPs, multiple testing, population stratification adjustment, Hardy-Weinberg Disequilibrium,
   Genetic and Polygenetic Risk Scores (GRS, PGR), ...
- Predictive models: assessment of
  prediction capacity and accuracy of
  multivariate linear, logistic regression
  and Cox models.
- Elaborate reports of intermediate and final results.

## Work plan



### CONCLUSIONS

#### Student will:

Clean and prepare complex data from big multicentric cohort study.
 Perform Genome-Wide-Association Study on more than 100.000 individuals.
 Evaluate prediction capacity of genetic scores using multivariate regression models and survival techniques.



Ia Unión Europea NextGenerationEU







d'Investigacions Mèdiques



**CORDELIA** CVD GWAS is:

Feasible at 3 years

- Cost-effective
- Promising