

## **Research Project Proposal**

# Academic year 2021-2022

# Máster en Investigación Biomédica

Project Nº 29

Title: Role of interleukin- $1\beta$  in the infiltration and polarization of macrophages in adipose tissue

**Department/Laboratory** Functional Metabolomics Laboratory, Department of Endocrinology & Nutrition. Clínica Universidad de Navarra.

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### **Summary**

**Background:** The obesity-associated low-grade chronic inflammation results from the interaction between adipocytes and cells from the immune system, mainly macrophages. Obesity induces a phenotypic switch from an anti-inflammatory M2-polarized state to a pro-inflammatory M1 state mediated through different cytokines.

**Hypothesis:** This project addresses the hypothesis that the adipose tissue excess and the glycemic state underlay the changes in the gene expression of the proinflammatory cytokine interleukin- $1\beta$ . In this way, interleukin- $1\beta$  may play a role in the macrophage polarization, aggravating the inflammatory state of obese patients. In addition, the blockade of interleukin- $1\beta$  using siRNA may contribute to improve the inflammation of adipose tissue associated to obesity.

Objectives and Methods: The involvement of interleukin- $1\beta$  in M1 polarization will be determined in human adipocytes and macrophages cell cultures as well as the potential use of blockade of interleukin- $1\beta$  in the improvement of the obesity-associated inflammatory state. In addition, the effect of conditioned medium secreted by adipocytes, with normal expression of interleukin- $1\beta$  or silenced, on gene expression profile of macrophages will be studied. Moreover, the relationship with other inflammatory markers as well as extracellular matrix components will be also studied.

#### The following **techniques** will be used:

### Sample processing:

- Serum, plasma and buffy coat extraction
- Cellular isolation from adipose tissue
- RNA isolation from adipose tissue and peripheral blood mononuclear cells
- Protein extraction from adipose tissue

## Biology molecular techniques:

- Nucleic acid and protein quantification and quality assessment
- Analysis of gene expression by Real-time PCR
- Analysis of protein expression by Western-blot

### Analytic techniques:

- ELISAs
- Large-scale cytokine analyses Multiplex (Luminex<sup>™</sup> 200)
- Immunohystochemical analysis of proteins

## Human macrophage and adipocyte cell cultures

| ı | yes |   | Does the project include the possibility of supervised animal manipulation to |
|---|-----|---|---|
|   |     |   | complete the training for animal manipulator?                                 |
|   | no  | X |   |
|   |     |   |   |