

**Internship Proposition**  
**(one page max)**  
**Master 2 GP Immunology & ImmunIntervention (I<sup>3</sup>)**  
**2024-2025**



**Lab: CR2TI UMR 1064**

**Team: 1 supervised by E Chiffoleau/R Josien**

**Name and position of the supervisor: E Chiffoleau, CR Inserm**

**Email of the supervisor: [Elise.Chiffoleau@univ-nantes.fr](mailto:Elise.Chiffoleau@univ-nantes.fr)**

**Candidate (if internship filled):**

**Title of the internship: Investigate the function of the immune checkpoint CLEC-1 in the cross-presentation of dead cell-associated antigens by conventional type 1 dendritic cells.**

**Summary of the internship proposal:**

We have previously shown that the CLEC-1 receptor expressed by dendritic cells corresponds to a sensor of cell death and acts as an immune checkpoint to prevent acute immune response and collateral tissue damage following injury. Thus, we showed that CLEC-1 signaling blockade by genetic ablation or antagonistic antibodies can restore an effective anti-tumor immune response and prolongs survival in pre-clinical tumor mouse models. Mechanistically, we observed that blocking CLEC-1 increases the cross-presentation of dead cell-associated antigens by conventional type 1 dendritic cells (cDC1). However, the underlying mechanisms and the precise step in which CLEC-1 limits antigen cross-presentation in cDC1 remains to be elucidated.

The aim of this internship is to investigate the localization and function of CLEC-1 in cDC1 in key processes of antigen cross-presentation such as:

- 1) localization in endo/phagosomes following internalization of necrotic material,
- 2) phagosome maturation,
- 3) antigen priming,
- 4) secretion of cytokines and chemokines
- 5) CD8<sup>+</sup> T activation.

This will be achieved in mouse cDC1 using *Clec1a* gene-deficient mice in *in vitro* and *in vivo* models using antigen-specific systems (OVA and OT-1 transgenic mice).

In the long term, a better understanding of the role of CLEC-1 in antigen presentation by cDC1 should help to consider its blockade as a new therapeutic tool for restoring an effective anti-tumor immune response in cancer patients.

Option(s) linked to the project:

- Clinical Research Profile (Recherche Clinique)
- Data Analyst Profile (Recherche et Analyse de Données Biologiques)
- Experimental Biology Profile (Recherche Expérimentale)

Form to be sent by email to : [gpi3@univ-nantes.fr](mailto:gpi3@univ-nantes.fr)