



28^E CONGRES DU CHO
12-15 OCTOBRE 2022
PRESQU'ILE DE GIEN



POSTER 7: Elucidating the role of N-Acetyl-Aspartate in normal hematopoiesis

Claire Lauvinerie 1, Arnaud Villacreces 1, Vanessa Desplat 1, Pierre-Yves Dumas 1, Isabelle Vigon 1, Jean-Max Pasquet 1, Amélie Guitart 1, *

1 : Cellules Souches Hématopoïétiques Normales et Leucémiques, INSERM UI312 BRIC, Université de Bordeaux, Bordeaux, France

Université de Bordeaux (Bordeaux, France), Institut National de la Santé et de la Recherche Médicale -INSERM

* Auteur correspondant

Mature blood cells have a limited lifespan and must be continually renewed. At the top of this hierarchical process are haematopoietic stem cells (HSC). It is well established that adult bone marrow (ABM) HSC are mainly quiescent. Conversely, HSC in the foetal liver (FL) proliferate extensively, calling for distinct bioenergetic requirements. The link between metabolism and functional capacities of HSC have mostly been studied in adult mouse models. Hence, knowledge on proliferating HSC remains elusive. To fill this gap, we conducted a metabolomic comparative analysis of quiescent ABM-HSC and proliferative FL-HSC. Our interest rapidly focused on a metabolite with a 10-fold higher abundance in FL-HSC than in ABM-HSC: the N-Acetyl-Aspartate (NAA). Although NAA is the second most prevalent metabolite in the brain, its function is still unclear. To investigate the role of NAA in adult and foetal haematopoiesis, we have used a transgenic mouse model knockout for *Nat81*, the gene coding for N-Acetyl-Aspartate Transferase, the enzyme catalysing the production of NAA in cells; in this mouse model NAA is not detectable in any tissue. Using flow cytometry, we have characterised the bone marrow of *Nat81*^{-/-} and *Nat81*^{+/+} adult mice, as well as the FL of E14.5 embryos. We have also conducted competitive haematopoietic reconstitution assays with both genotype to assess the role of NAA in HSC function. This project should address the question regarding the involvement of NAA in foetal and adult haematopoiesis regulation.

Keywords: stem cells, normal haematopoiesis, metabolism, N-Acetyl-Aspartate

