



### **TECHNOLOGY OFFER**

#### SUPPORTING INNOVATION AND TECHNOLOGY TRANSFER IN ONCOLOGY

#### **NUCANTHERM**



NEW NUCLEOSIDE ANALOGUES FOR THE TREATMENT OF AZACITIDINE-RESISTANT MDS AND AML PATIENTS

# CONTEXT & BACKGROUND

This program concerns the development of new nucleotide analogues for the treatment of Azacitidine-resistant MDS and AML patients. We developed a series of new compounds with anti-leukemic activities in MDS and AML. The mechanism of action of these compounds has been partly elucidated and involves both apoptosis and autophagic cell death. The compounds are effective in vitro on Azacitidinesensitive and resistant MDS/AML cell lines as well and on MDS and AML bone marrow patient's samples ex vivo with a high efficiency on bone marrow samples of Azacitidine-resistant patients. The compounds are also efficient in grafted models of MDS or AML in mice.



New nucleoside analogues for cancer therapy



To treat patients suffering MDS/AML

#### SCOPE

ONCOHEMATOLOGY / MYELODYSPLATIC SYNDROM /ACUTE MYELOID LEUKEMIA

#### **KEYWORDS**

KEYWORDS Azacitidine / Resistance / Nucleoside Analogues / Acadesine/ Apoptosis /Autophagy



#### **DEVELOPMENT & MATURATION STAGE**

A licence may be established which concerns, firstly the use of these new nucleoside analogues for the treatment of MDS / AML and secondly a compagnon test for the evaluation of azacytidine resistant in newly diagnosed MDS and AML patients. Two products have been tested in animal studies. Our results indicated a beneficial effect of these compounds in mice xeno-transplanted with azacytidineresistant SKM1 cells. However, both molecules seem to act as pro-drugs. Taking into account the difficulties encountered to identify the active compounds, they were not further developed. We are currently validating new analogues for in vivo assays in mice, before a possible clinical phase I trial in humans



Azacitidine-resistant MDS/AML Patients

**TARGET PROFILE** 

Apoptosis/Autophagic cell death



- Patent on «triazole nucleoside analogs for cancer therapy» : WPT/2012/143624

- license on the companion diagnostic test (patent : FR12/00584/PCT10669) can be considered



These compounds are effective on Aza-resistant cell lines and patient's samples

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#### **INDUSTRIAL APPLICATIONS & OPPORTUNITIES**

Phase I study to be scheduled after validation of new analogues for in vivo assays



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