







#### **OUR VERSATILITY**

The reactivity, adaptability and flexibility of a human-sized compagny.



#### **OUR PERSONALIZED ADVICE**

A Scientific Advisory Board of clinical experts to understand your needs and projects.



#### SECONDARY RESISTANCE MODELS

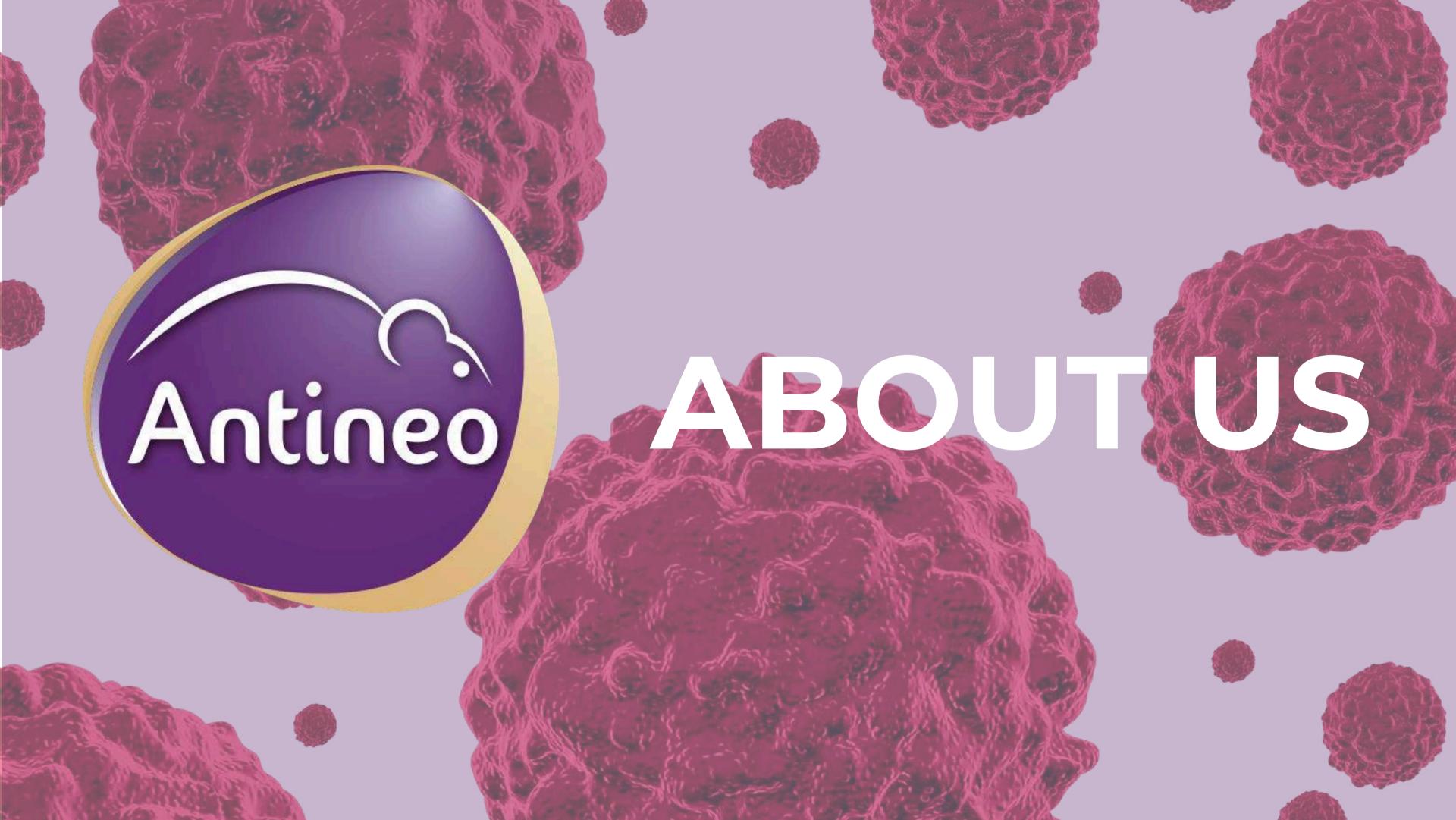
Secondary resistance models to standards of care established and characterized through RNAseq and immunophenotyping of the tumor microenvironment.



#### **OUR COMPETITIVE PRICES**

To provide our customers with excellent value for money compared to the competition.





## **Spin-off of the Team Anticancer Antibodies** Centre de Recherche en Cancérologie (CRCL)

## Company

CRO created in May 2015, specialized in proof-ofconcept studies in oncology



## Location

BioParc Rockefeller **Bâtiment BIOSERRA 2** Lyon

Regulation



## • CIR agreement 2020-2025

• Fully authorized animal house and personnel



## **Team**

Scientists and efficient team recognized for its expertise in onco-pharmacology the at international level



## **Customers**

French and international clients composed pharmaceutical, biotechnology companies and academic institutions



- 200+ studies
- 40 sponsors
- 6 ongoing partnerships







Renaud Marin-Sidgwick CEO



Marie Tautou, PhD
Study Director
Head of Business Development



Charles Dumontet, MD-PhD
CSO - Consultant



Aurélie Cadiou PhD student



Jean-Philippe Druet Executive Assistant



**Doriane Mathé Study Manager** 



Marine Fellmann Study Manager

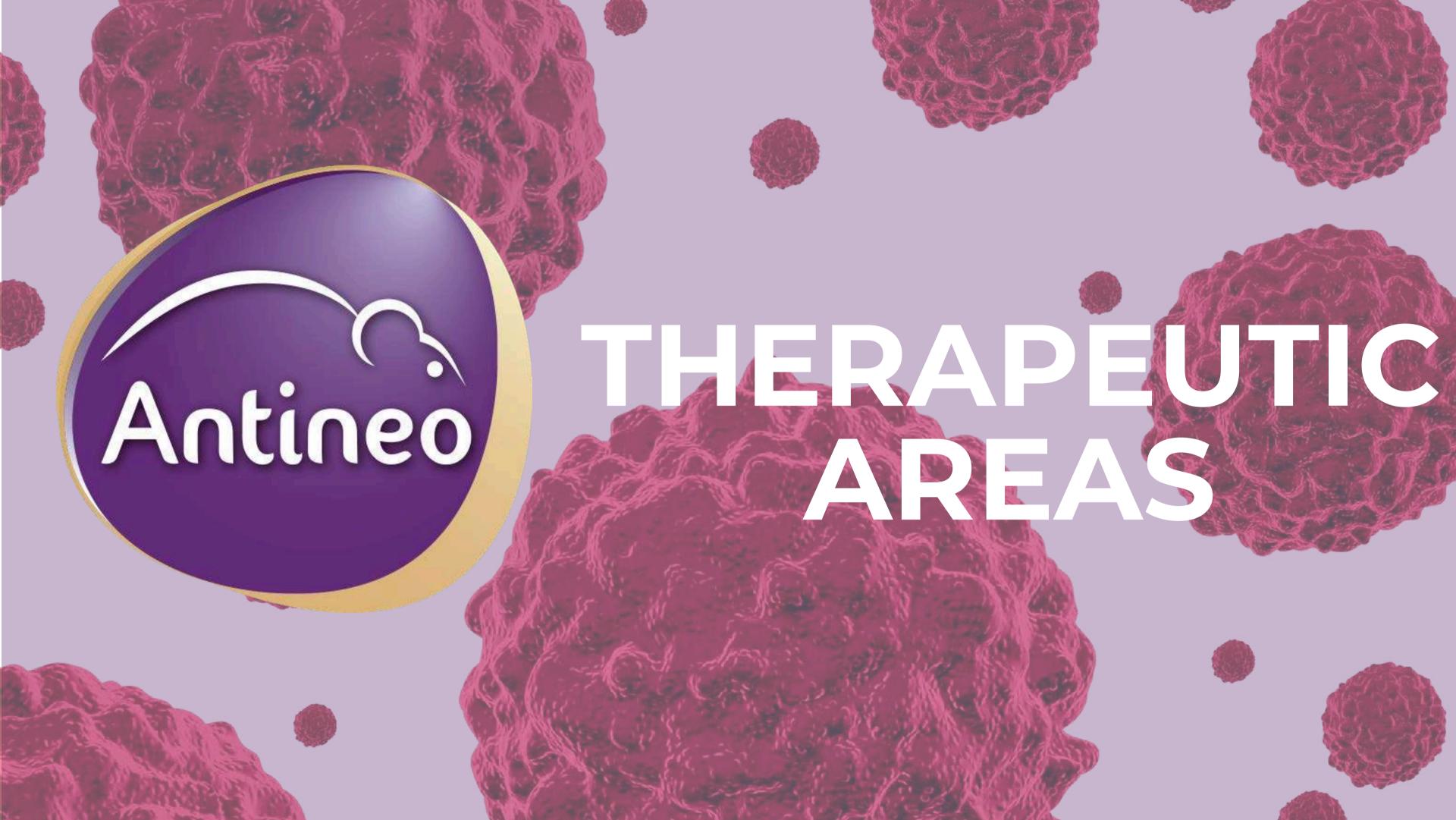


Pierre-Antoine Choffour Study Manager



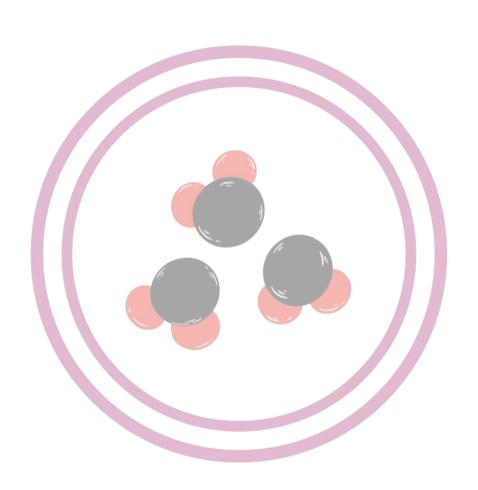
Flore Sarraute
Apprentice





# Oncology and immuno-oncology

## **Small molecules**



## **Biological molecules**

Antibodies, peptides



# Cellular and gene therapies





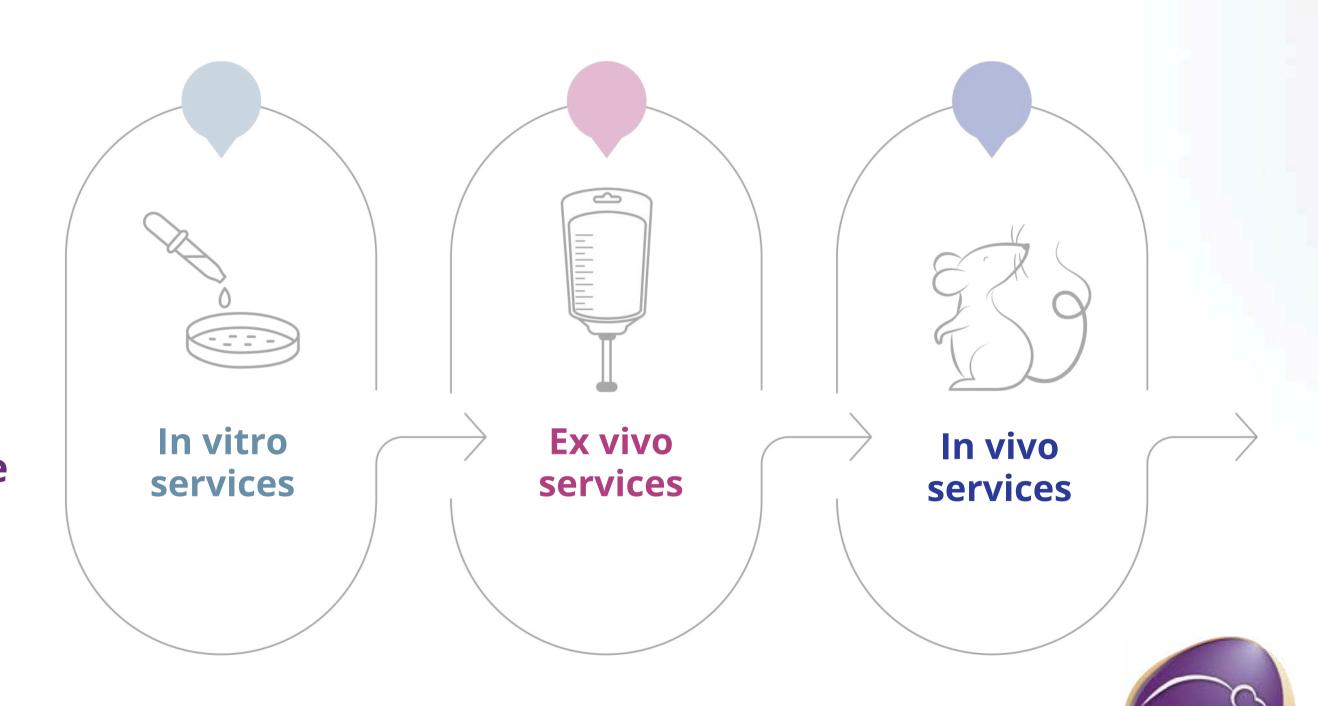


## Antineo's services



Optimize and accelerate the development of our customers' compounds

Provide advice, expertise and services





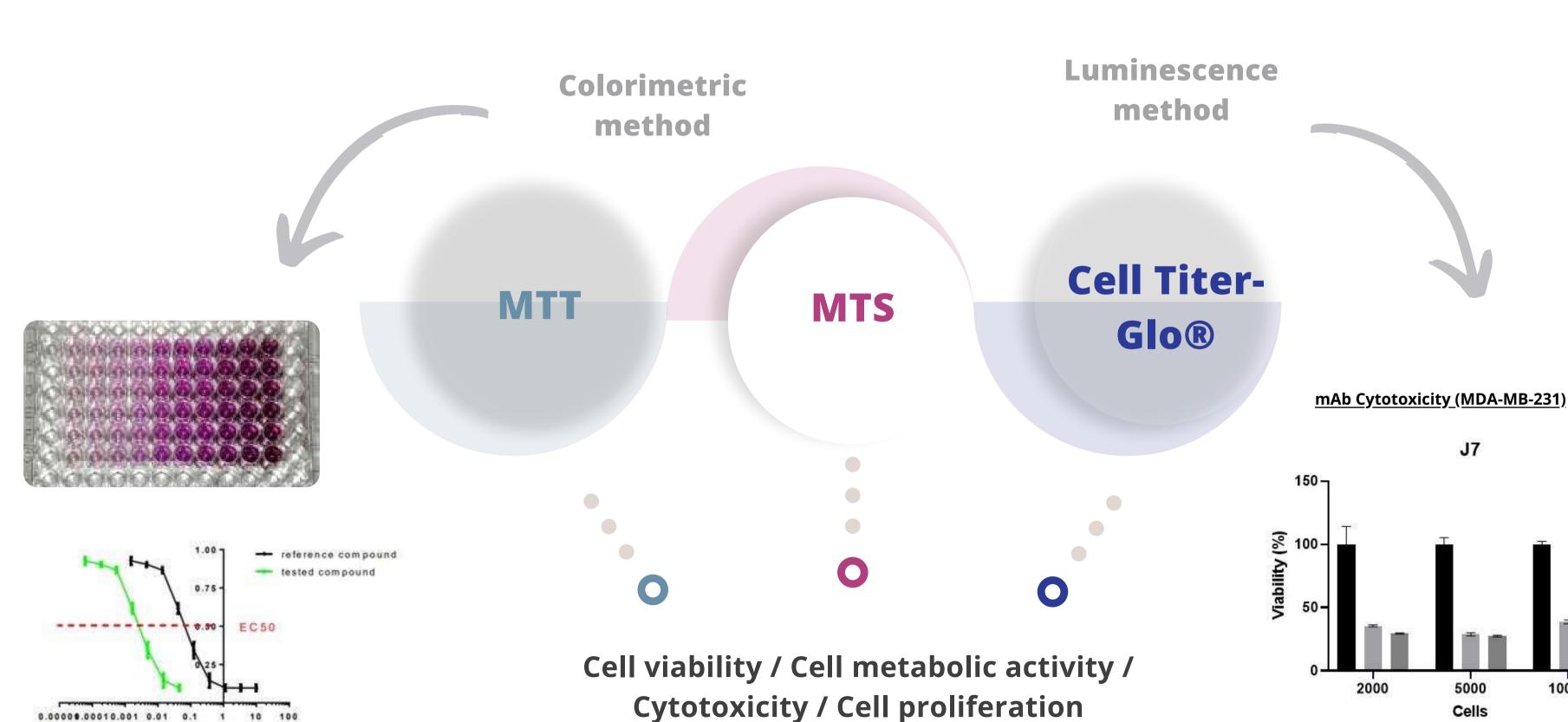
# Cytotoxicity assays

concentration (µg/m L)



**Determination of IC50 / EC50** 

Synergy / Antagonism assay

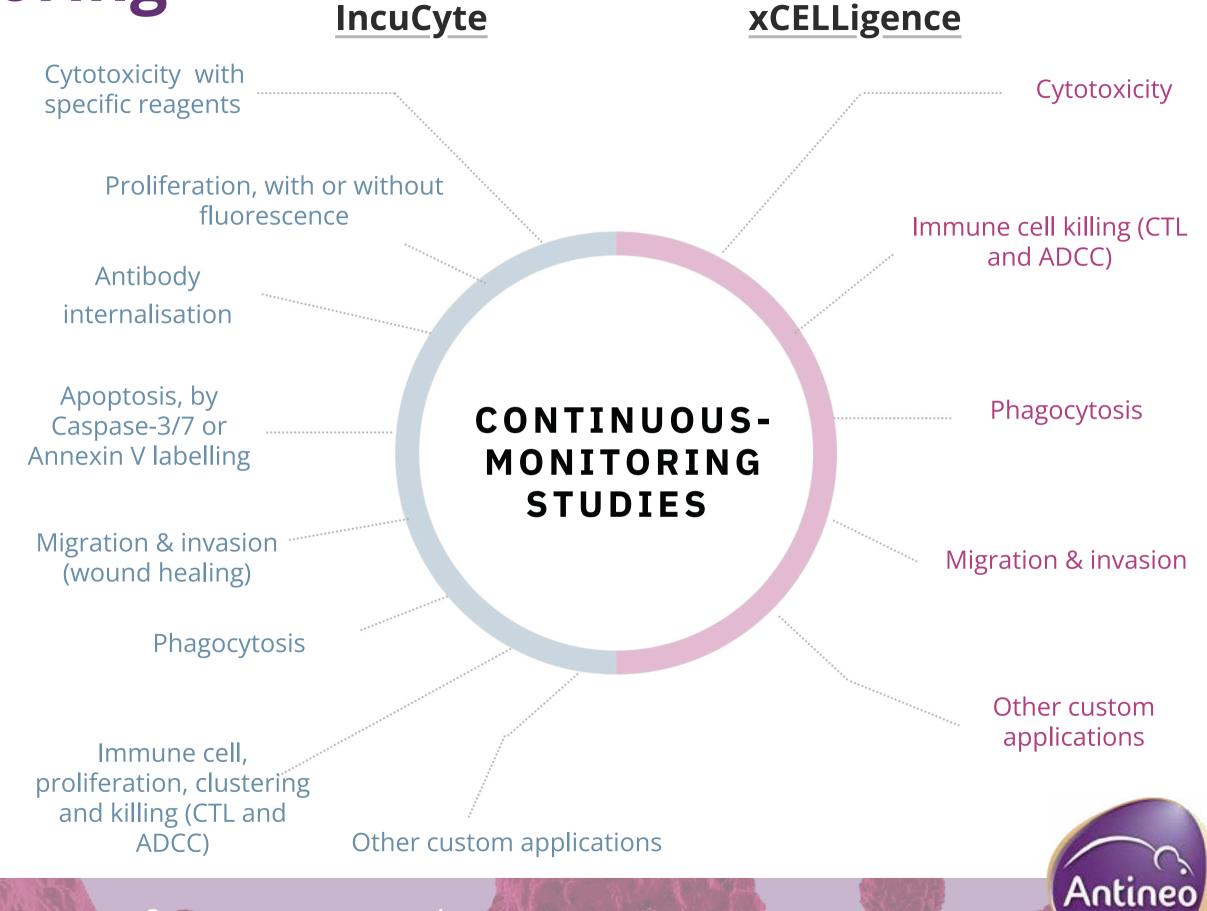


# Continuous-monitoring

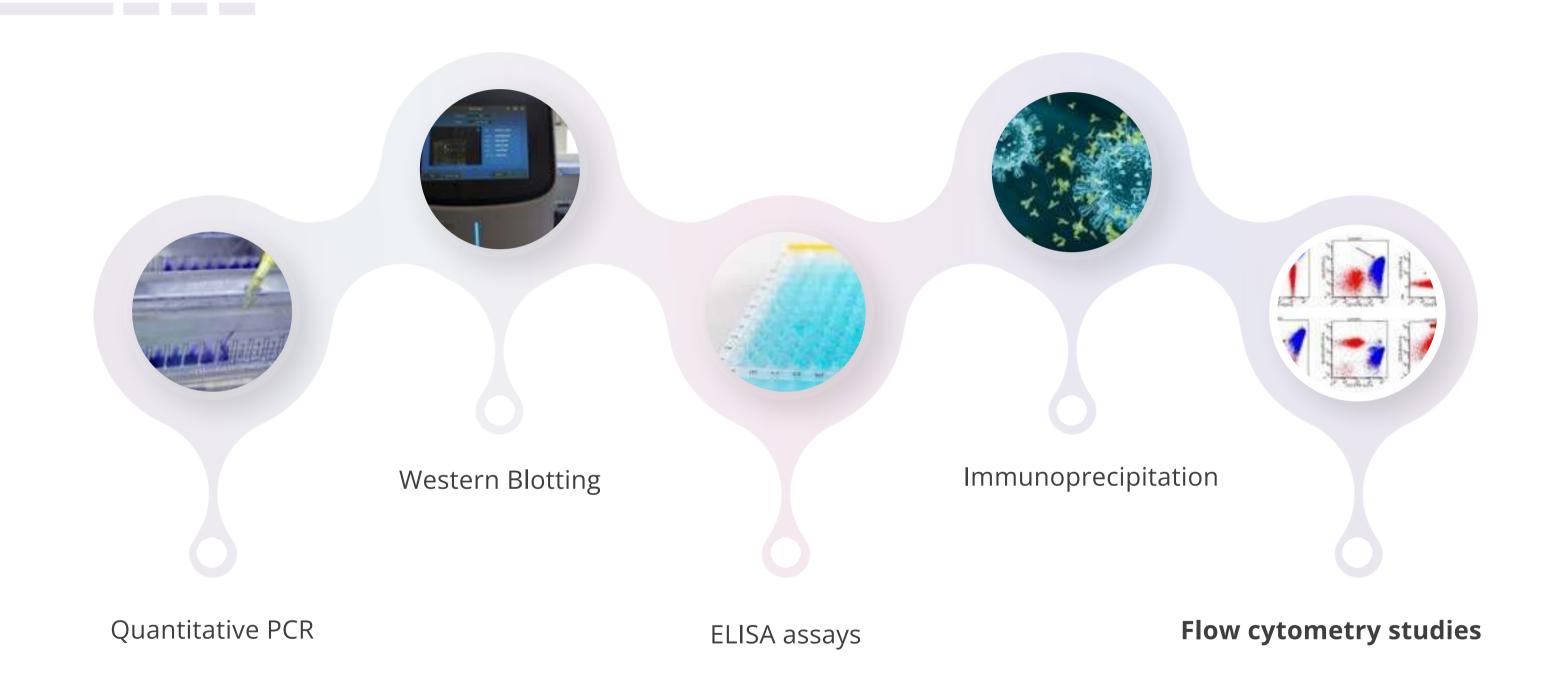
studies



Realize a real-time analysis of a variety of cellular and immunological processes



# Characterisation of samples





# Flow Cytometry - (FACS)

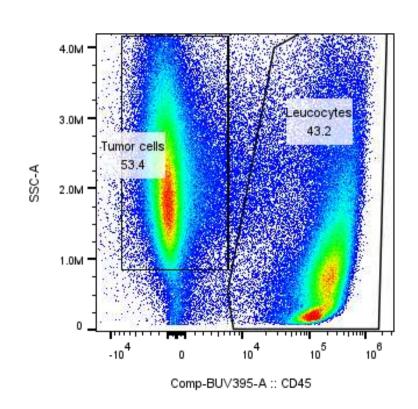


Qualitative and quantitative multiparametric analyses

**BD FACS CANTO™** 

Classical

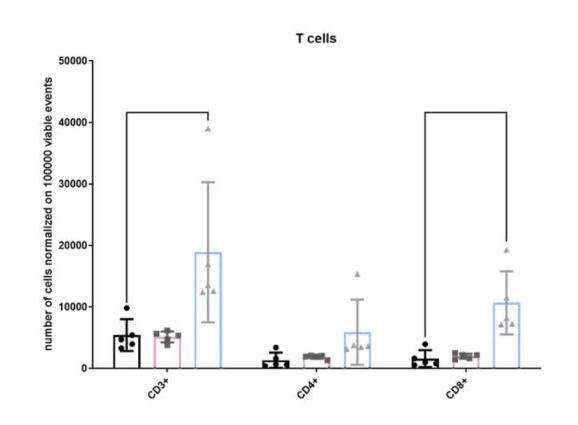
3 lasers: 8 markers\* maximum



## LSR FORTESSA<sup>TM</sup>

Classical

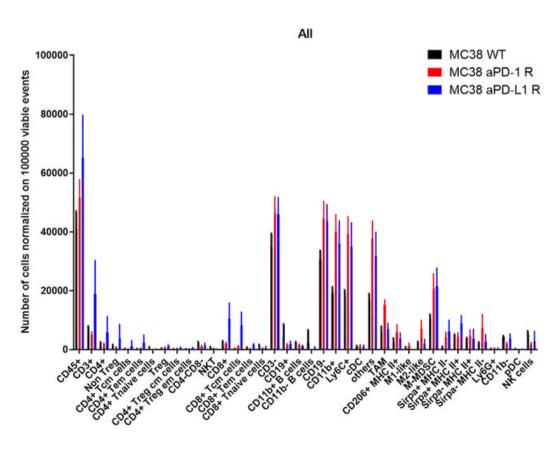
4 lasers: 18 markers\* maximum



#### **CYTEK**

Spectral

5 lasers : up to 29 markers\*



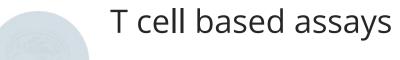


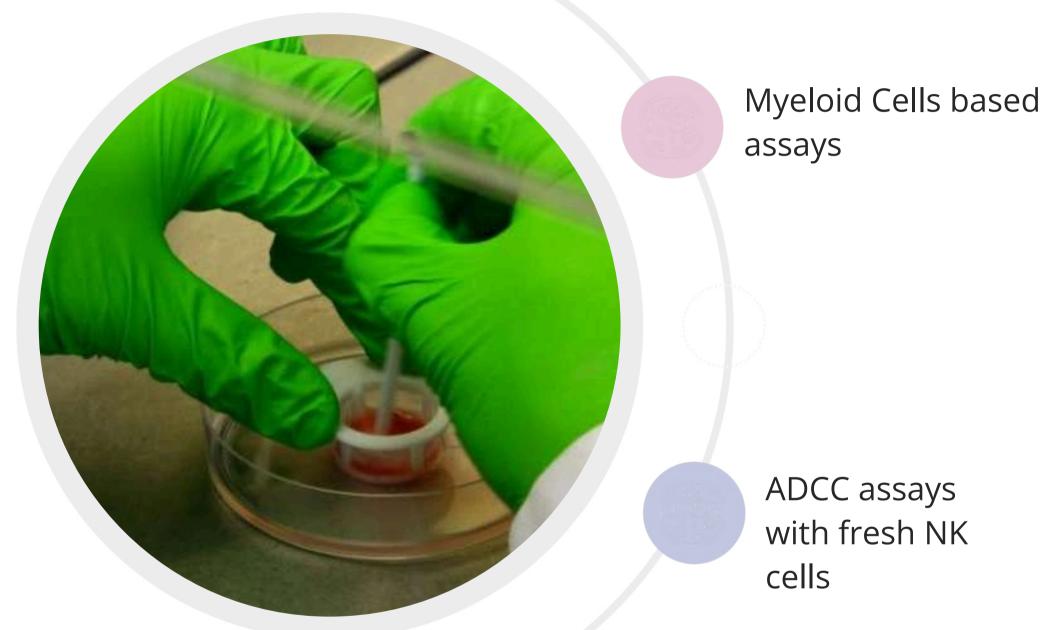
# Immunology services

Isolate the cells of interest and characterise your target molecule by Flow Cytometry



- Analysis on fresh samples : on blood products (blood bags) and by products (buffy coat)
- Most assays can be performed as end point and continuous-monitoring studies





ADCC assays with fresh NK cells

ADCP assays with fresh myeloid cells



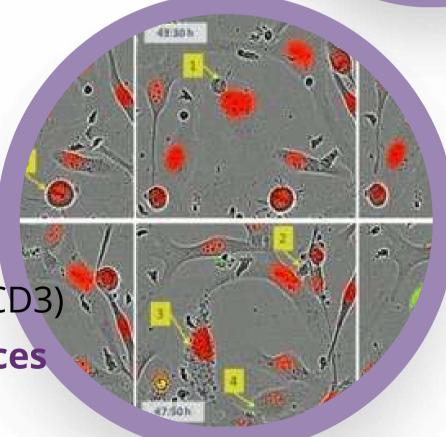
# Development of immunotherapies



Highlight the target and decipher the mechanisms of action of therapeutic antibodies

Immunology ex vivo assays (T cell activation by IFNy measurements,
 CTL assays, Macrophage polarisation etc.)

- In vitro or ex vivo ADCC, ADCP and CDC assays (calcein release)
- Original methods for in vivo assessment of ADCC and CDC activities
- Titration / Internalization / By-stander effect (ADC) ...
- Original in vitro and in vivo assays for bispecific antibodies (anti-CD3)
- A unique panel of tumour models presenting secondary resistances to immunotherapies





## **Animal facilities**

- High standards facility with strictly controlled environment
- Multiple types of experiments: study cancer progression, assess the effectiveness and innocuity of novel therapies,...
- Wide range of animal models from immune-competent (C57Bl/6, CD1...) to immune-deficient (SCID-CB17, NSG, NOG, NOD-SCID...) and humanized models (BRGSF-HIS, huNOG-EXL)

- Capacity over 3,000 mice
- Ethical compliance and expertise in animal experimentation







## **Standard of care therapies**

- As reference for the tested compound
- For comparison studies
- For combinaison / synergy studies

## **Choice of tumor models**

- 100+ cell-derived xenograft models
- 40 murine syngeneic models for immuno-oncology
- Subcutaneous or orthotopic implantation





# Protocol adapted to our clients' compounds

- Route (IV / IP / PO / IT)
- Galenic formulations (liposome encapsulation)
- Schedule of injection
- Schedule and duration of follow-up
- Weekly updates
- Choice of end-point (with control or individual ethical end-points)

# In vivo analysis



Recommendations on the choice of the best indication and model



Systemic and haematological toxicity of your compounds in rodents (VetScan / MS9)



Pharmacokinetics properties in mouse and rat



Antitumor efficacy in human or mice tumour models



Combination / comparison with gold standards



Demonstrate the antitumor activity of a novel agent in animal models, as well as defining the dosage and schedule that is both efficient and non-toxic



Orthotopic models\*



Immunophenotyping of the tumour micro environment



An original offer of secondary resistances to reference therapies (CDX and syngeneic)



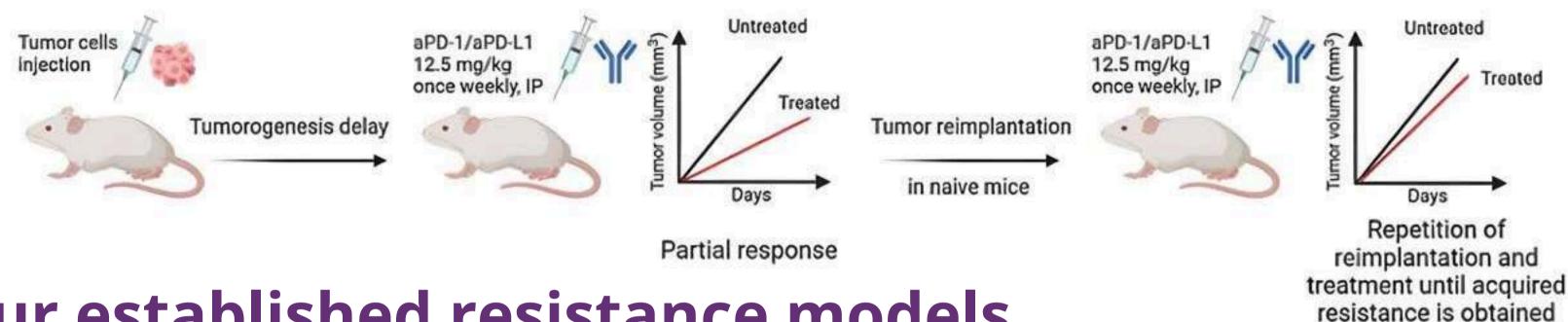
The development of models of resistance

**Antineo** 

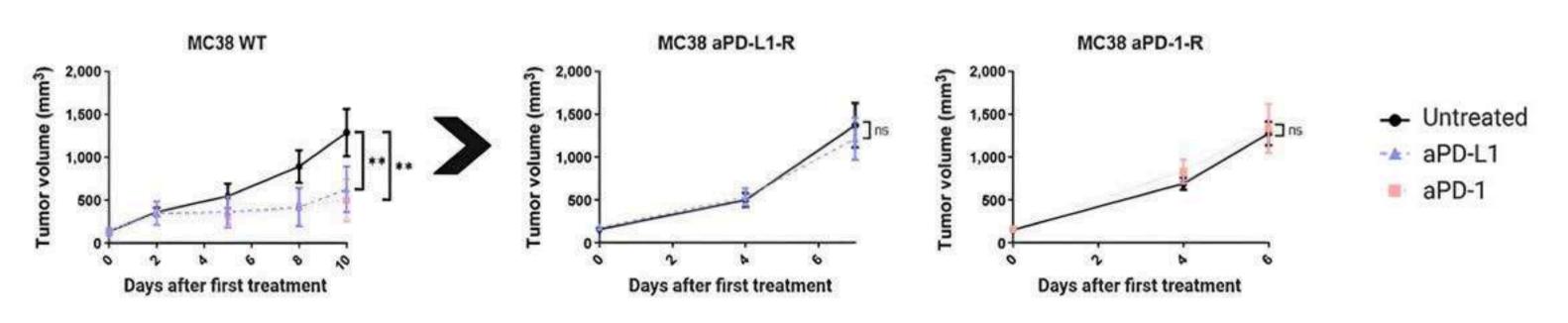
\*In Vivo Syngeneic Tumor Models with Acquired Resistance to Anti–PD-1/PD-L1 Therapies



## Acquired resistance to anti-PD(L)1



## Our established resistance models



\*In Vivo Syngeneic Tumor Models with Acquired Resistance to Anti-PD-1/PD-L1 Therapies



**CDX Models** Syngeneic Models Lymphoma Colon Follicular Lymphoma - (RL model):
• Rituximab / GA101 / R-CH0P / R-DHAP) MC38 model anti-PD1 / anti-PDL-1 Mantle Cells Lymphoma - (Granta model) : Rituximab Diffuse large B cells lymphoma - (Toledo model): Bladder Rituximab MBT-2 model Burkitt's lymphoma anti-PD1 Raji model : RituximabDaudi model : CAL-101 MB49 model anti-PD1 / anti-PDL-1 Myeloma Plasma cells myeloma RPMI8226 model **Kidney**  Daratumumab **RENCA** model **SECONDARY** anti-PD1 / anti-PDL-1 Multiple myeloma MM.1S Daratumumab **RESISTANCES TO** REFERENCES Melanoma **THERAPIES Breast** B-raf anti-PD1 / anti-PDL-1 Tubular Adenocarcinoma BT474 T-DM1 N-Ras anti-PD1 / anti-PDL-1 MDA-MB-361 model • Trastuzumab T-DM1 Tyr N-Ras models anti-PD1 / anti-PDL-1 **Colorectal / Gastric BRAND NEW:** Pancreas Kic8 model Gastric carcinoma NCI-N87 PD1/PDL1/Gemcitabine Trastuzumab Lymphoma Bespoke services: On demand

development of resistant models

(Syngeneic or CDX models)

P388 model

anti-PD1 / anti-PDL-1

## Syngeneic Models

Follicular Lymphoma - (RL model):

Rituximab / GA101 / R-CHOP / R-DHAP)

Mantle Cells Lymphoma - (Granta model):

Rituximab

Diffuse large B cells lymphoma - (Toledo model) :

RPMI8226

Rituximab

Burkitt's lymphoma

Raji model : RituximabDaudi model : CAL-101

#### Myeloma

Plasma cells myeloma model

Daratumumab

Multiple myeloma MM.1S

Daratumumab

#### **Breast**

Tubular Adenocarcinoma BT474

T-DM1

MDA-MB-361 model

Trastuzumab T-DM1

# SECONDARY RESISTANCES TO REFERENCES THERAPIES

## Bladder

MBT-2 model

anti-PD1

MB49 model

• anti-PD1 / anti-PDL-1

#### **Kidney**

RENCA model

anti-PD1 / anti-PDL-1

#### Melanoma

B-raf

anti-PD1 / anti-PDL-1

N-Ras

anti-PD1 / anti-PDL-1

Tyr N-Ras models

anti-PD1 / anti-PDL-1

#### **Colorectal / Gastric**

Gastric carcinoma NCI-N87

Trastuzumab

Bespoke services: On demand development of resistant models (Syngeneic or CDX models)

#### **BRAND NEW:** Pancreas

Kic8 model

• PD1 / PDL1 / Gemcitabine

#### **Lymphoma**

P388 model

Colon

MC38 model

• anti-PD1 / anti-PDL-1

anti-PD1 / anti-PDL-1

## Partner platforms













CIQLE

Microscopy platform for Immunohistochemistry (IHC)



High throughput sequencing, microdissection and single cell technologies

### **HAWKCELL**

Platform for Magnetic Resonance Imaging (MRI)

### **IMTHERNAT**

PET-Scan (Radiolabelling)

### **ANAQUANT**

Detection and quantification of proteins by mass spectrometry



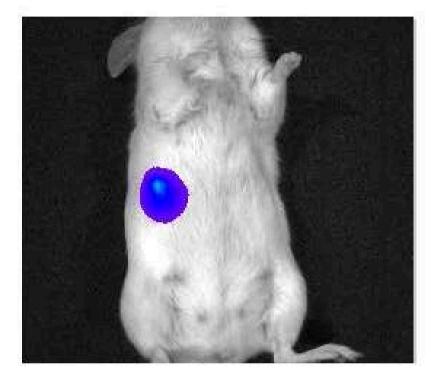
Immune omics analysis (Single cell)



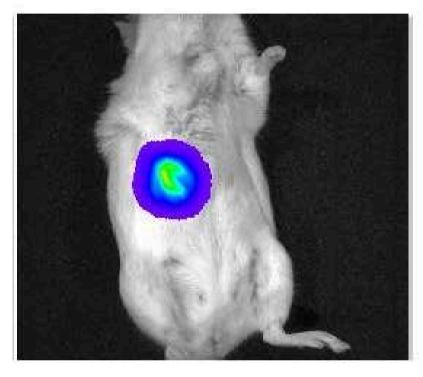


## **IVIS® Lumina Series III Imager**

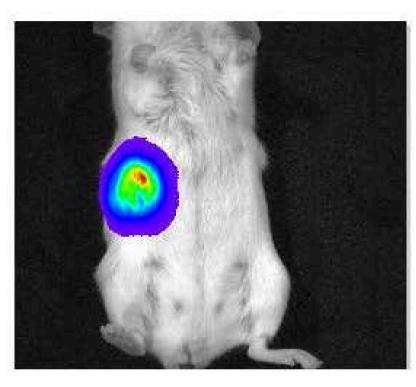
## MDA-MB-231 cell line



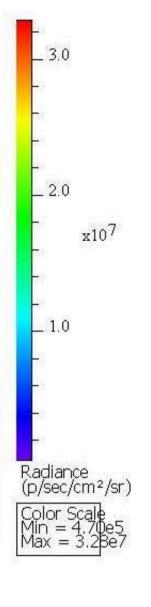




2023-04-12



2023-04-26



- (D) =

Hepatic metastases of an MDA-MB-231 Luciferase (+) model by intrasplenic implantation



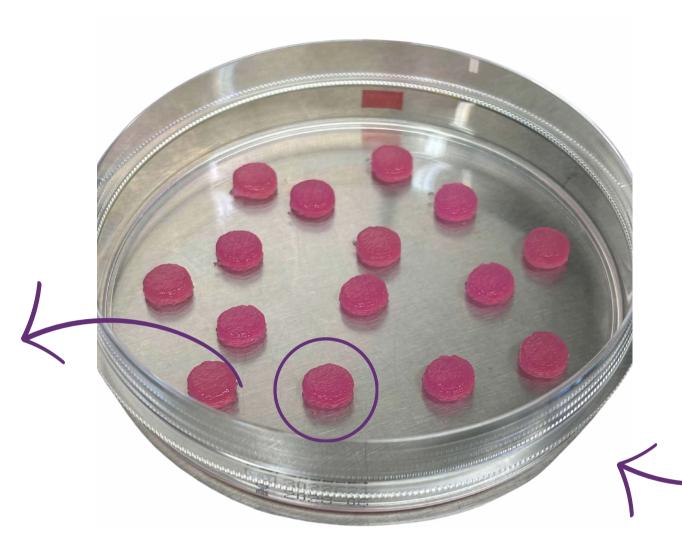


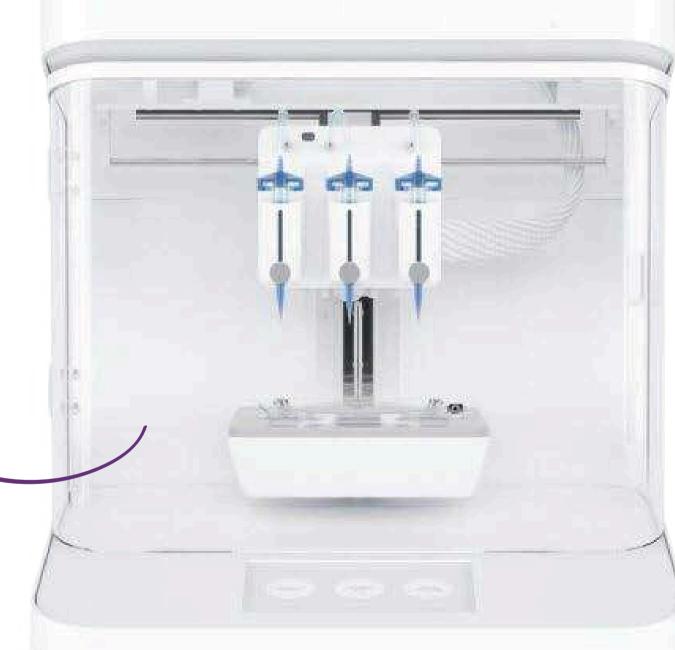
- Precise tumor monitoring and follow-up
- Animal saving
- Biodistribution and efficacy studies

# 3D Bioprinting - Ongoing development

## BT 474-GFP cell line









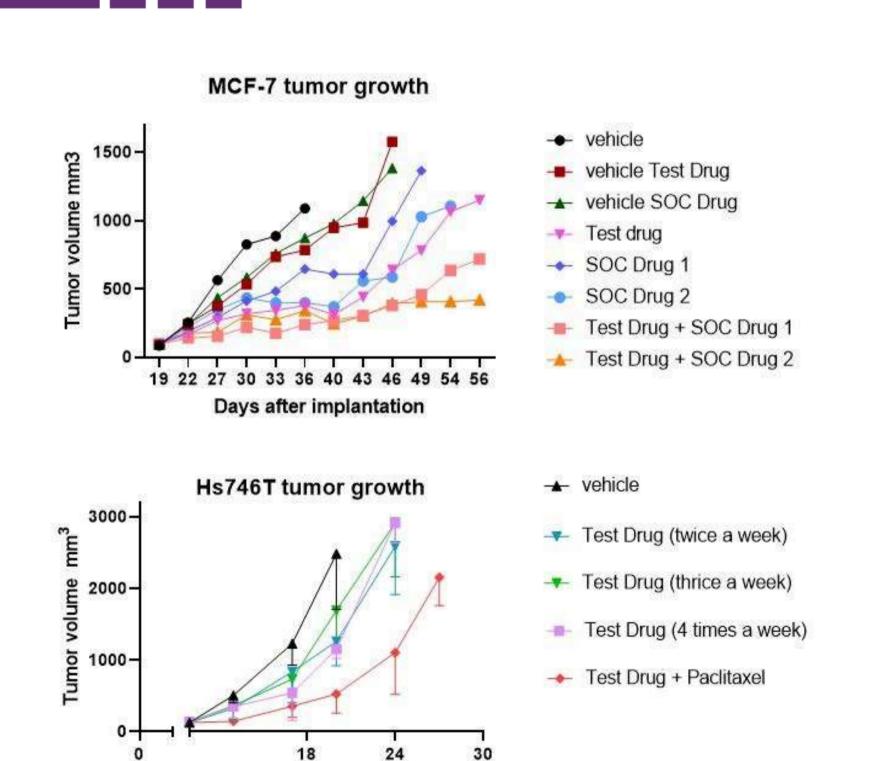
- Recapitulate the tumor microenvironment (TME)
- High-throughput screening
- Minimising animals used and costs



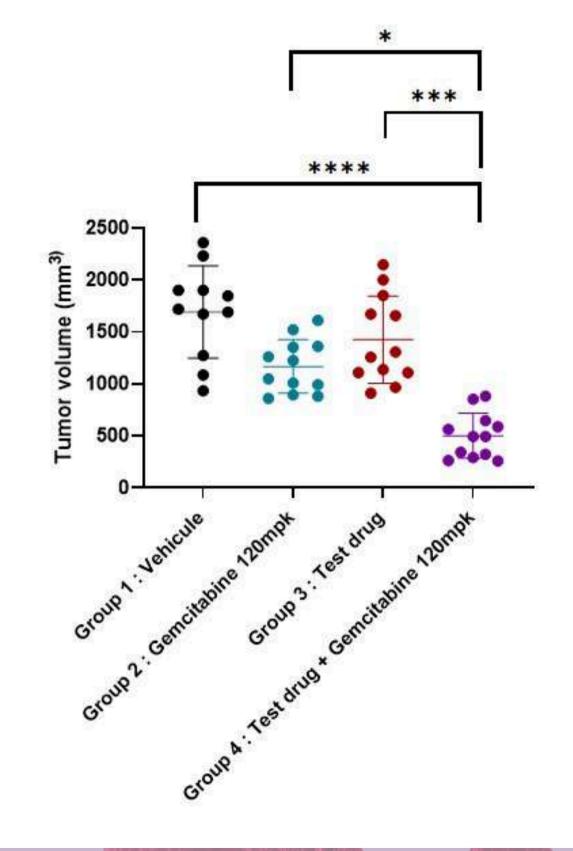


# **Efficacy study**

#### **KIC8 Gemcitabine-resistant tumor growth**

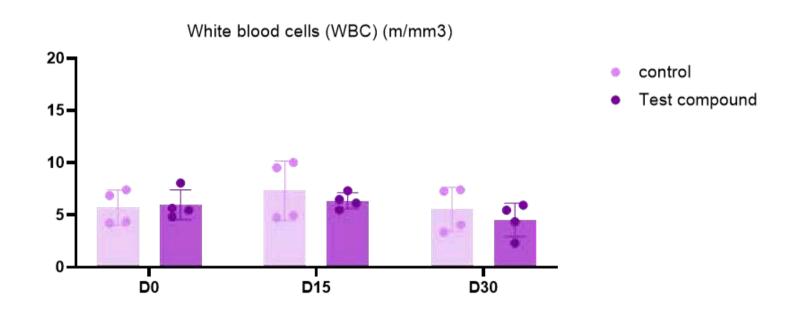


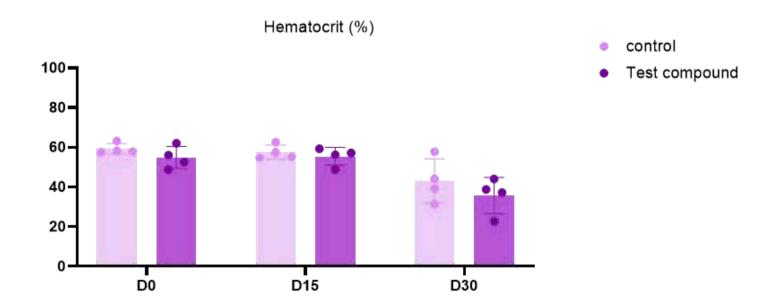
Days after implantation

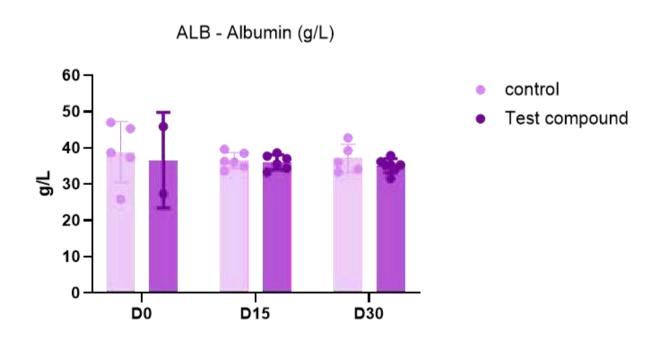


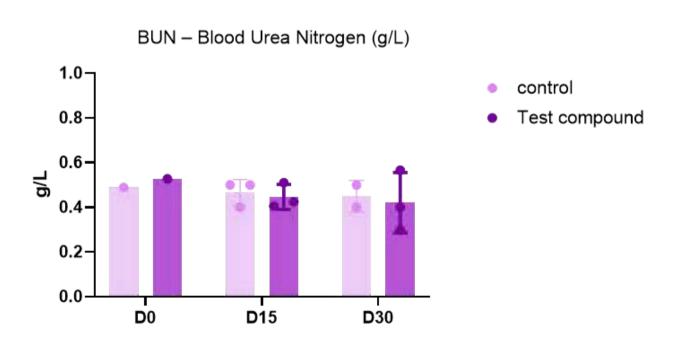


# Toxicity study - Haematological analysis





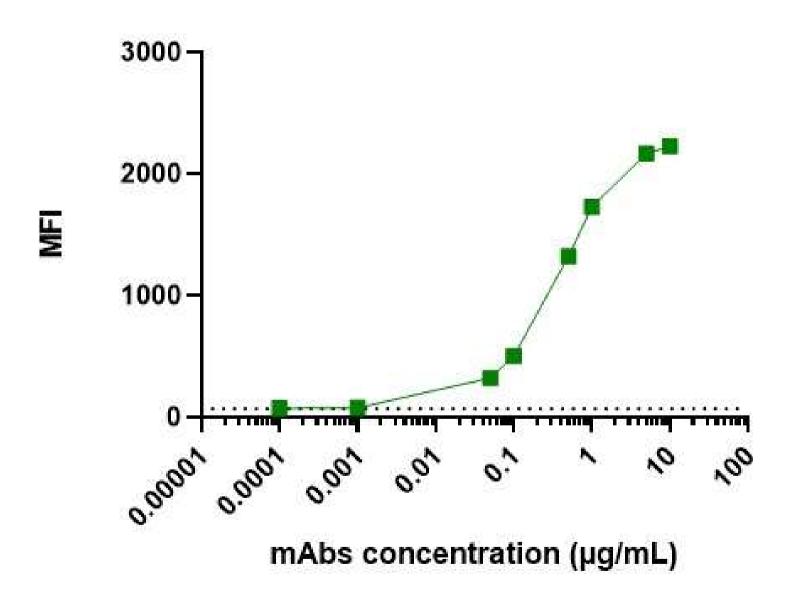


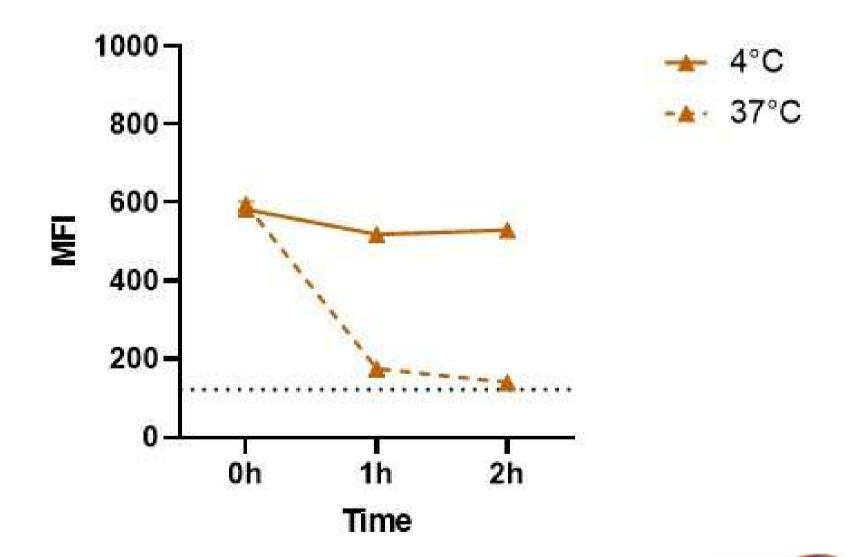




## mAb Titration assay (SK-BR3)

## mAb Internalization assay (SK-BR3)









# EVEXTA BIO-



























