ANEWERA

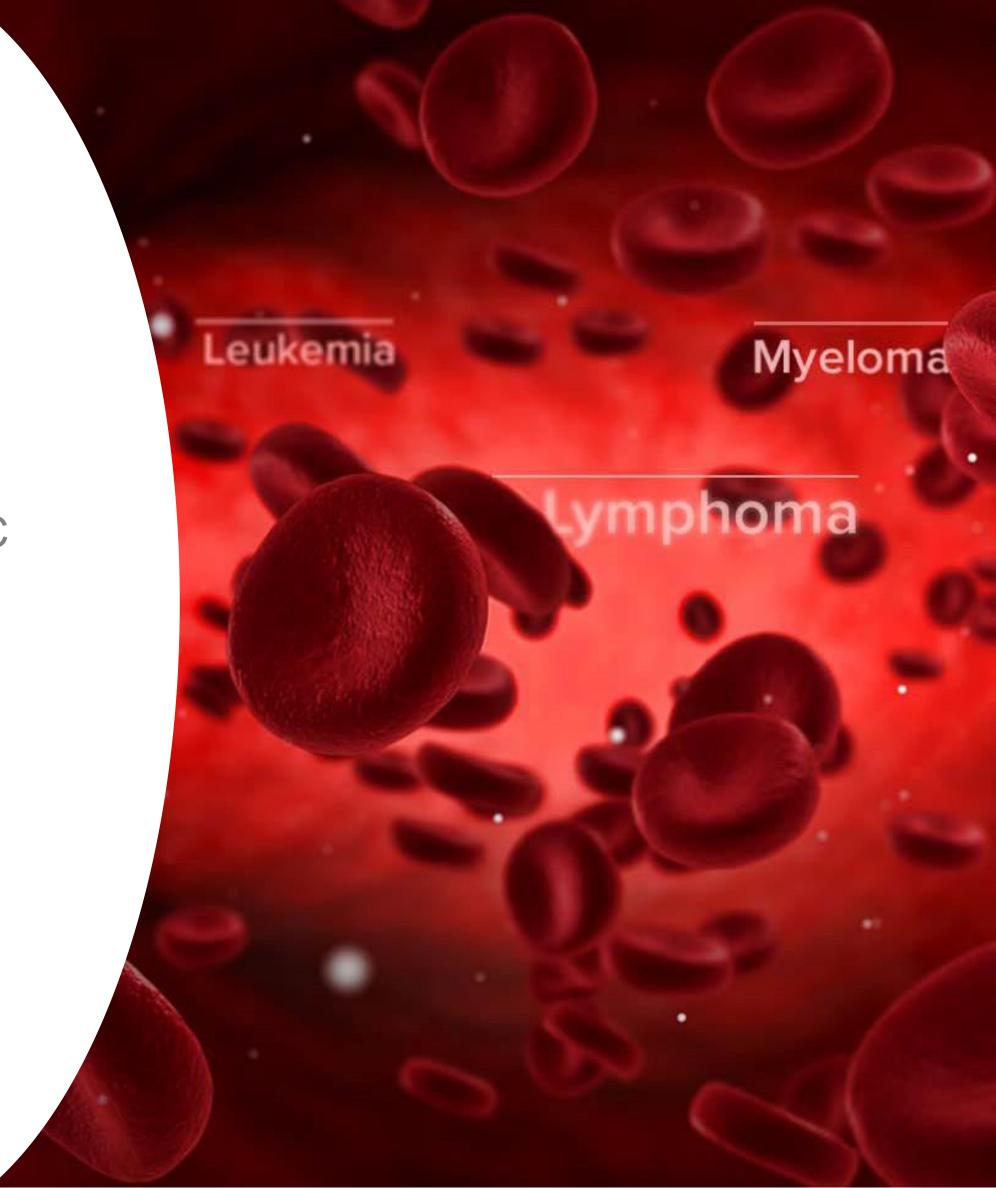
for the modeling of blood and autoimmune diseases and drug development

Advanced Hematopoietic PBMC Chimera – ApbHC

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lmmugenyx

A new era for the modelling of blood / autoimmune diseases and drug development

Immugenyx has developed a novel type of humanized mice that possess a functional human immune system:

Advanced Hematopoietic peripheral blood Chimeras ('ApbHC')

What are ApbHC?

- Mice with high levels of human hematopoietic chimerism made via transplantation of proprietary processed human peripheral blood mononuclear cells
- Possess a variety of T Cells in the peripheral blood and a variety of human immune cells in the spleen and bone marrow.
- Chimeric animals generate human IgM and IgG and immunogen-specific human IgM and IgG easily detectable in peripheral blood.
- Chimeric animals continue generating antibodies that were developed by a human donor of blood that was used to make the ApbHC.
- Fast and inexpensive to make

Validated:

- Collaboration with a major biopharmaceutical company on the development of AHC as a tool for assessment of immunogenicity of biologics
- Collaboration with J&J (Janssen) to develop an in vivo tool for modeling and development of treatments of Lupus (LSE)
- Used to test efficacy of CDX antibody
- AML engraftment into AHC is demonstrated and used to test CDX bispecific antibody

Possible applications:

- Assessment of immunogenicity of biologics
- An in vivo tool for the modelling and development of treatments of autoimmune diseases
- A tool for the rapid generation of human antibodies in response to human-specific pathogens (Biodefense)
- A tool for modelling blood diseases such as AML and testing novel treatments that involve reprogramming of the immune system (multi-specific antibodies for immune cell redirection, CAR-T etc.)
- An in vivo tool to study the physiology of human plasma cells and plasma cellassociated diseases such as multiple myeloma



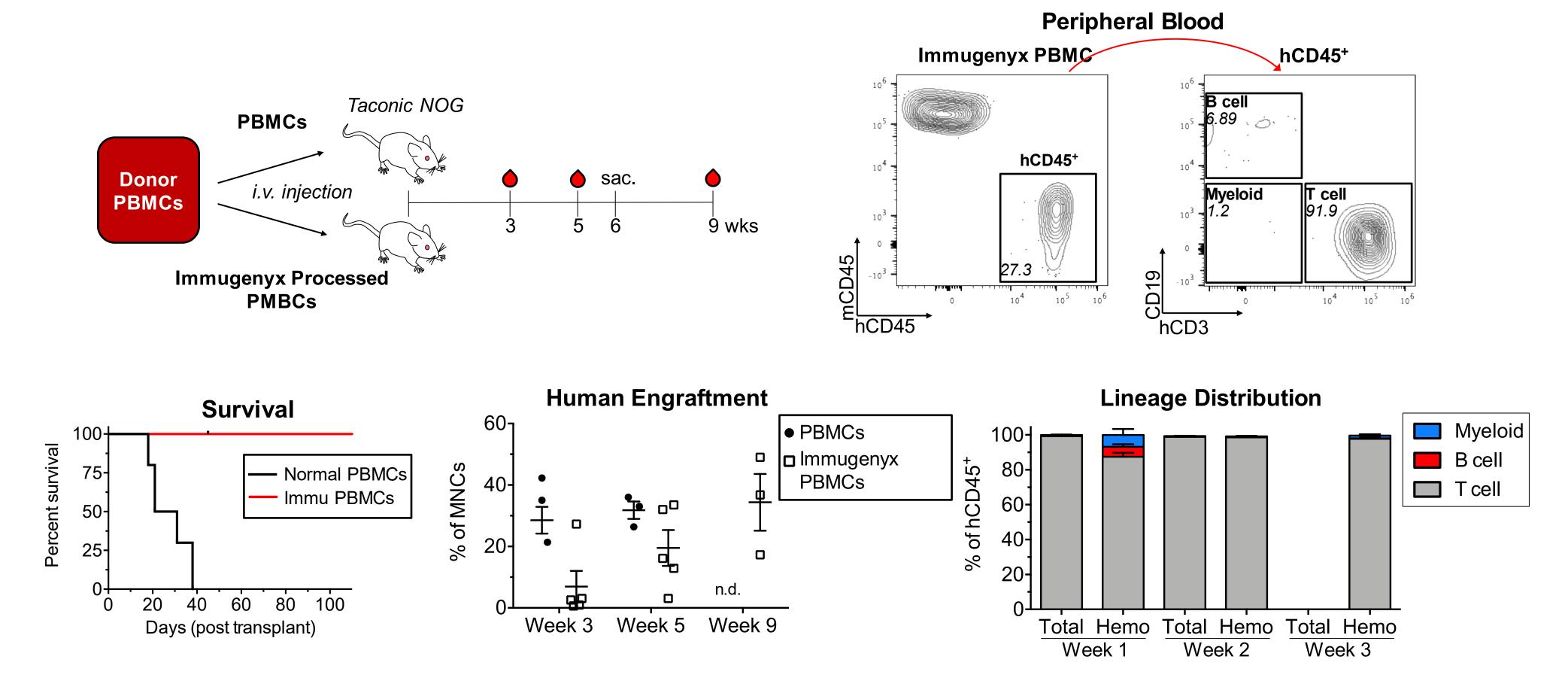
ApbHC

Advanced peripheral blood Hematopoietic Chimera

- 1. Properties
- 2. Application Examples



Immugenyx Processing of PBMCs Prior to Transplantation Prevents Development of GVHD and Retains Normal Engraftment Frequencies





Lymphocyte Engraftment of Hematopoietic Organs with Presence of Human IgG/IgM in **Plasma** Serum antibodies 30007 Taconic NOG Control **PBMCs** ■ PBMCs sac. 2 2000-(ng/mL) Hemo mice □ Immugenyx PBMCs Donor i.v. injection **PBMCs** 9 wks 1000 **Immugenyx Processed PMBCs** IgG IgM **Immugenyx PMBCs** Immugenyx PMB T cell frequency **Human engraftment** 100 T cell B cell **BONE MARROW** 91.4 80 hCD45⁺ 87.8 8.05 80 % of MNCs % of hCD45⁺ plasma 60 60 cell 40 20 BM SP PB **₽** Plasma cell **B** cell frequency SPLEEN 48.5 % of hCD45 $^{\scriptscriptstyle +}$ of hCD45⁺ 30 15 20 hCD45 CD38 68.4 CD19 10 hCD138 hCD45 hCD45 **S**B

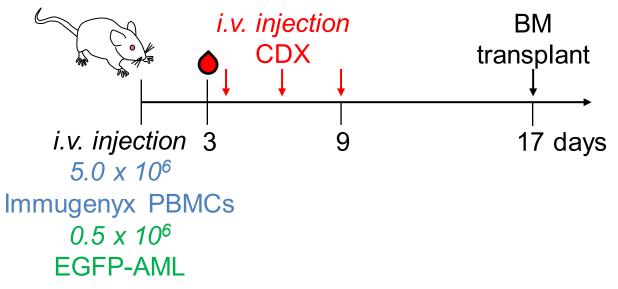


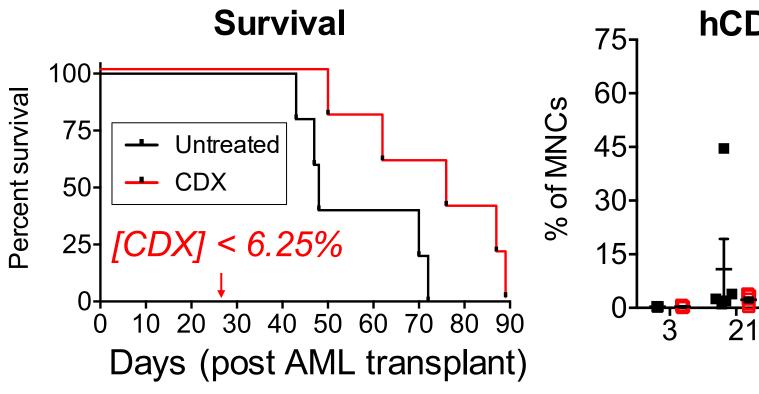


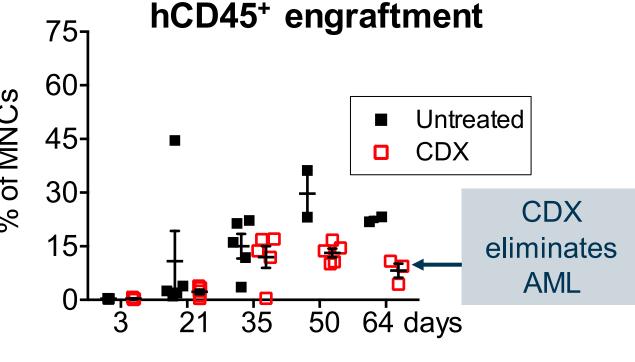
Bi-specific Antibody Test (Example)

Targeting of AML-derived cells in vivo

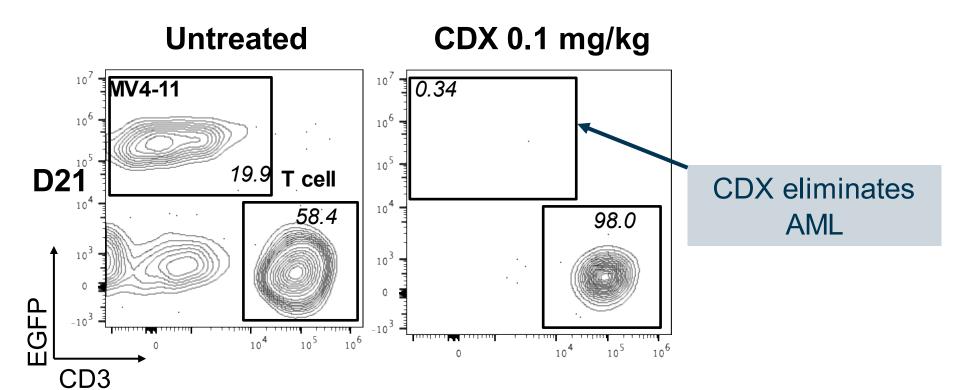
Immugenyx (Hemogenyx subsidiary) proprietary novel Advanced peripheral blood Hematopoietic Chimera humanized mouse model



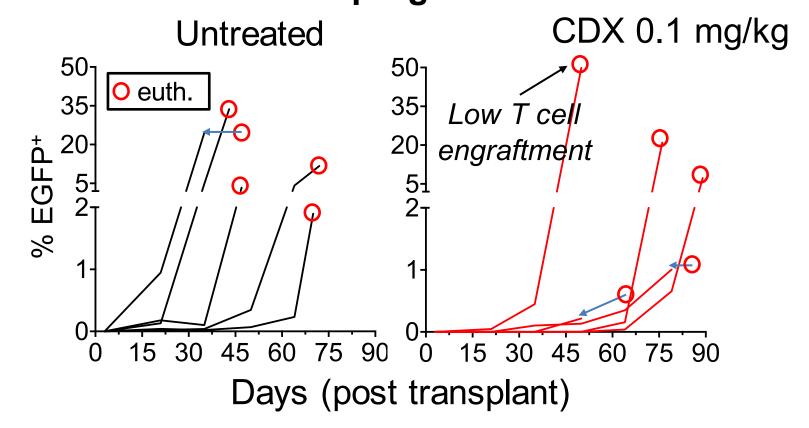




Peripheral blood hCD45⁺



AML progression





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