TECHNOLOGY BRIEF

Clinical immunodiagnostic – immune cell specific monoclonal antibody

Broad Specificity HLA-DR Monoclonal Antibody

Lead Inventors:

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Background

HLA-DR plays a central role in antigen presentation and immune regulation. It is expressed on professional antigen presenting cells (APCs) and is upregulated during immune activation. The high degree of polymorphism in the HLA-DR gene has prevented the use of antibodies to this important receptor in manufacturing, histology, and flow-cytometry.

Our team aims to address these challenges with a novel antibody product exhibiting broad HLA-DR reactivity with a modular architecture (Fig. 1) allowing the fusion to nanoparticles, detection agents like fluorophores, or surfaces for diverse reagent-based applications.

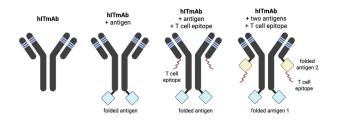


Fig. 1. Modular Architecture of HITmAb enabling fusion of diverse antigens and peptides.

Description of the Invention

Drs. Brian Barber and Jean-Philippe Julien have engineered a fully humanized immune targeting monoclonal antibody (HITmAb). Derived from murine antibody 44H10 and engineered using CDR grafting onto an IgG1 framework, HITmAb binds HLA-DR with nanomolar affinity and maintains high stability and specificity posthumanization. The antibody demonstrates broad allele reactivity (Fig. 2) essentially reacting with HLA-DR in all humans, thereby eliminating the need for MHC typing when used with individual humans. Additionally, HITmAb has cross-species compatibility (demonstrated in humans, rabbits, ferrets, macagues), and exhibits exceptional stability and favorable manufacturing characteristics (Fig. 3).

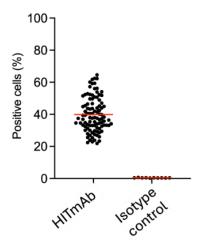


Fig. 2. Broad specificity of HITmAb. Data showing HITmAb binding to 100% of blood donor derived PBMCs of 100 random samples from a diverse population.

Keywords: monoclonal antibody, HLA-DR specific, flow cytometry, cell enrichment, ELISA

Commercial Applications

Potential applications for a reagent HITmAb include:

- Sorting for HLA-DR positive cells in blood flow cytometry
- As a reagent in immunobiological analysis of human tumour tissues to identify the presence of HLA-DR positive APCs
- As a manufacturing reagent to remove HLA-DR positive cells from a bulk mixture (e.g. stem cell transplantation applications)

Publications & Patents

Patents pending in Canada (CA 3250551) and the US (US 18/863143).

Kassardjian et al., Antibodies 13, 57 (2024). DOI: 10.3390/antib13030057

Kassardjian et al., Cell Reports 42, 112391 (2023). DOI: 10.1016/j.celrep.2023.112391

Seeking strategic partnership/licensing to develop reagent products specifically recognizing HLA-DR expressing immune cells

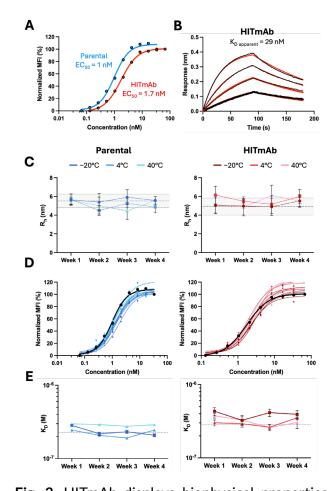


Fig. 3. HITmAb displays biophysical properties and thermostability similar to the originating mouse parental antibody. (A) Flow cytometry titration curves comparing parental and HITmAb binding to BJAB cells. (B) BLI binding of HITmAb to HLA-DR, where black lines represent measured binding and red curves correspond to the data fitted to a 1:1 binding model. (C-E) Biophysical characterization of parental and HITmAb kept at various storage temperatures for up to four weeks, including hydrodynamic radii and polydispersity measured by DLS (C), binding to BJAB cells measured by flow cytometry (D) and binding to recombinant HLA-DR measured by BLI (E).