# Engineered Anti-CTLA-4 Antibody for Tumor Delivery: Toward Human Clinical Validation Within Two Years

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## Vision

- ➤ We are creating a next-generation anti-CTLA-4 therapy engineered with a collagen-binding domain (CBD), with recurrent glioblastoma (rGBM) as the initial indication. By achieving tumor-selective delivery, we address the safety and efficacy limitations of conventional checkpoint inhibitors (CPI).
- > This innovation not only provides a much-needed option for patients with rGBM but also establishes a versatile platform. The CBD technology can be extended to other cancer types and modalities, unlocking multiple therapeutic pipelines and broader market opportunities.

# Marketability

- > The initial indication is rGBM, where treatment options remain extremely limited and prognosis is poor. Achieving clinical PoC would unlock significant value.
- > We plan to expand into CPI-approved tumors such as NSCLC, a rapidly growing market with best-in-class potential, supported by a roadmap from rGBM human PoC to Phase I basket trial across multiple tumor types enabling partnerships and licensing.

## Innovation

- ➤ A novel approach that engineers CTLA-4 to act specifically within the tumor microenvironment.
- ➤ CBD-mediated collagen targeting allows selective accumulation in the tumor, balancing efficacy with safety.
- The technology can be extended to other immunotherapies and modalities, providing broad potential as a versatile platform.

# Partnering

#### [ Expected partners ]

Pharmaceuticals • Biotech/Drug Discovery Service • CMO/CDMO/CRO/SMO • Venture capitals

## [ Expectation ]

Out-licensing and co-development, Talent acquisition and leadership build-up, Series A financing

# Research Outline

Key Words: #Engineered antibody, #Immuno-oncology, #Drug Delivery

#### <u>Concept</u>

➤ A von Willebrand factor (vWF)-derived CBD is fused to an anti-CTLA-4 antibody, enabling selective binding to tumor collagen. This design reduces systemic exposure and strengthens local immune responses.

#### Nonclinical Data

- ➤ In a rGBM model, treatment extended survival and achieved complete remission in some cases.
- The tumor immune microenvironment was favorably remodeled, supporting durable antitumor responses.
- Compared with conventional anti-CTLA-4 antibodies, systemic toxicity was markedly reduced, indicating an improved safety profile.

### Research Plan

- Our initial focus is rGBM, with a microdose clinical study planned to establish human PoC.
- After establishing PoC, we plan to move into additional tumor types responsive to checkpoint inhibitors, expanding the scope of our therapy.

#### Goal: Creating Novel therapy with a Collagen-Targeted Drug Delivery Platform **Business Model:** Collagen-binding domain (CBD)-based cancer drug delivery Continuous pipeline creation using CBD • By conjugating CBD to therapeutics, selective delivery to tumors Stage 1: CBD-Ipilimumab for rGBM-tumor expansion with high collagen expression is achieved. CBD-lpilimumab/rGBM • Adding CBD to CTLA-4 antibodies significantly prolongs survival CBD-lpilimumab/expansion Stage 2: CBD-based engineering of other drugs 2nd pipeline: CBD-Protein 3rd pipeline: CBD-Ab Stage3: Expansion into other indications Liver cirrhosis, pulmonary fibrosis Plan: Clinical trial of CBD-CTLA-4 antibody in rGBM Stage 4: Theranostics Preparation for GMP Fastest path: initiate **CBD PET Probe** microdosing trial in the manufacturing U.S. within 2 years Accelerated preclinical studies

#### **Reference**

➤ Ishihara J, et al. Targeted antibody and cytokine cancer immunotherapies through collagen affinity. *Sci Transl Med*. 2019;11(487):eaau3259.