# Immune checkpoint inhibitor

Drug discovery toward immune checkpoint LILRB4 and its prognostic application for cancer patients as a biomarker

#### Overview

A myeloid checkpoint (CP), ILT3/LILRB4 (B4) is well-known and expected for cancer treatment, but its physiological ligand was unknown. This invention is about the physiological ligand of B4, fibronectin (FN), and its therapeutic use.

Following patterns can be considered for immunoregulation by blocking the binding between B4 and its ligand FN:

- (1) FN analog (competitively binds to B4-FN)
- (2) Anti-B4 antibody (acts on B4 and inhibits B4-FN binding) \*
- (3) Anti-FN antibody (acts on FN and inhibits B4-FN binding)

\*Note: Anti-B4 antibody has been exclusively licensed to Company X, but small molecules and peptides that interact with B4, as well as (1) and (3) are free to develop.

Further, B4 as a biomarker for lung cancer patients' prognosis prediction was verified by our original B4 monoclonal antibody that inhibits B4-FN binding (Figure C, Patent ②).

# Possible applications

- □ Drugs for treating autoimmune, cancer, inflammatory or allergic diseases associated with B4
- Diagnostic agent that predicts the effectiveness of cancer immunotherapy
- Diagnostic agents that predict cancer prognosis

#### **IP Data**

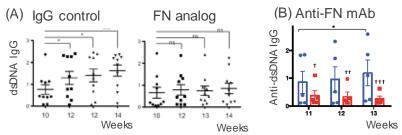
IP No. : ①WO2021/029318, ②WO2023/002943 Inventor : TAKAI Toshiyuki, SU Mei Tzu, *et al.* 

Admin No. : ①T18-289, ②T20-3069

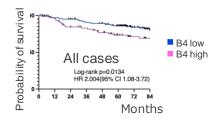
# Mechanism<sup>2</sup> Fibronectin FN30 B4/gp49B Cell membrane Cytoplasm Regulation Regulation Proinflammatory Attach, growth, locomotion

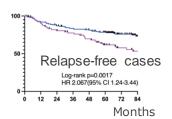
## Features · Outstandings

Treatment of FN analog (A) and anti-FN antibody (B) to SLE mice suppressed the generation of autoantibodies (dsDNA lgG)



(C) High B4 expression relates to a poor prognosis in lung cancer patients





#### **Related Works**

- [1] International Immunology, 33(8), pp. 447–458.
- [2] International Immunology, 34(8), pp. 435–444.

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