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Advanced 2nd Edition Immune-Regulation Reagents

CD112R - NEW & UNIQUE Immune Checkpoint Receptor

CD112R [PVRIG; Poliovirus receptor-related immunoglobulin domain-containing protein] is a new high affinity receptor of CD112/Nectin-2. It inhibits TCR-mediated signals upon binding to CD112/Nectin-2. CD112R as a new regulator of the immune system may become a new attractive cancer immunotherapy target.

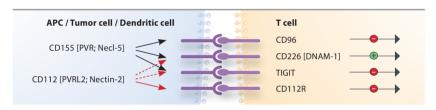


FIGURE: Regulation and activation of Tlymphocytes depend on signaling by the T cell receptor (TCR) and also by cosignaling receptors of the Poliovirus receptor (PVR)-like proteins that deliver negative (— →) or positive (— →) signals.

NEW

CD112R (mouse):Fc (human) (rec.)AG-40B-0170 10 μg | 3 x 10 μg

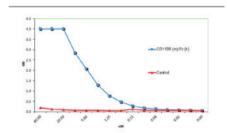


FIGURE: Binding of CD112R (mouse):Fc (human) (rec.) (AG-40B-0170) to Nectin-2/CD112 (mouse) (rec.).

LATEST INSIGHT

CD28/B7 Pathway – Importance in PD-1 Therapy

PD-1–targeted therapies have been a breakthrough for treating certain tumors and can rejuvenate T cells to unleash the anticancer immune response. Recently, it was shown that PD1 inhibits T cell activation by suppressing positive co-stimulation through CD28, instead of TCR itself. CD28 is required for PD-1 therapies to kill cancer cells efficiently and eliminate chronic viral infections in mice and is a critical target for PD-1 blockade.

LIT: Rescue of exhausted CD8 T cells by PD-1-targeted therapies is CD28-dependent: A.O. Kamphorst, et al.; Science 355, 1423 (2017) • T cell costimulatory receptor CD28 is a primary target for PD-1-mediated inhibition: E. Hui, et al.; Science 355, 1428 (2017)

| CD28 (human):Fc (human) (rec.) CHI-HF-210CD28 | 200 μg | CD86 (human):Fc (human) (rec.) CHI-HF-210CD86 | 100 μg |
|---|--------|--|----------------------------|
| CD28 (mouse):Fc (mouse) (rec.) CHI-MF-110CD28 | 200 μg | CD86 (mouse):Fc (mouse) (rec.) CHI-MF-110CD86 | 100 μg |
| CD80 (human):Fc (human) (rec.) CHI-HF-210CD80 | 100 µg | PD-L1 (human):Fc (human) (rec.) (no CHI-HF-220PDL1 | on-lytic) 100 μg |
| CD80 (mouse):Fc (mouse) (rec.) CHI-MF-110CD80 | 100 µg | PD-L1 (mouse):Fc (mouse) (rec.) (non-lytic) CHI-MF-120PDL1 100 μg | |

New Role of IL-33 – Expansion of T Regulatory Cells (Treg)

IL-33 has emerged as a multifunctional protein, with roles in driving potent Type 1 and Type 2 immunity, as well as facilitating profound Treg expansion *in vitro* and *in vivo*. IL-33-expanded Treg express the IL-33 receptor ST2 and classical markers associated with Treg phenotype and suppressor function. IL-33 may potentially be used to promote/maintain organ transplant tolerance or suppress autoimmunity and has been associated with Treg-mediated wound healing.

LIT: Poor repair of skeletal muscle in aging mice reflects a defect in local, interleukin-33-dependent accumulation of regulatory T cells: W. Kuswanto, et al.; Immunity 44, 355 (2016) • Expansion of regulatory T cells in vitro and in vivo by IL-33: B.M. Matta & H.R. Turnquist; Methods Mol. Biol. 1371, 29 (2016)

UNIQUE

NEW Highly Active Human IL-33 Proteins

IL-33 (oxidation resistant) (human) (rec.)

AG-40B-0160 Untagged 10 μg | 100 μg AG-40B-0167 His-Tag 10 μg | 100 μg

LIT: Oxidation of the alarmin IL-33 regulates ST2-dependent inflammation: E.S. Cohen, et al.; Nat. Commun. 6, ID8327 (2015)

UNIQUE

Antibody inhibiting the binding of mouse IL-33 to ST2/IL-1RAcP

IL-33 (mouse), mAb (rec.) (blocking) (Bondy-1-1)

AG-27B-0013 $100 \ \mu g \\ AG-27B-0013PF \qquad \qquad Preservative Free \qquad 100 \ \mu g \ | \ 500 \ \mu g \ | \ 1mg \\$

LIT: Regulation of de novo adipocyte differentiation through crosstalk between adipocytes and pre-adipocytes: T.D. Challa, et al.; Diabetes 64, 4075 (2015)

Visit www.adipogen.com for a Complete Panel of IL-33 Proteins, Antibodies & ELISA Kits!

Highly Active IL-36 Family Proteins

IL-36 α β , γ are members of the IL-1 family with proinflammatory effects and involvement in first-line defence against microorganisms. IL-36 γ is associated with skin, lung and bowel inflammation and was shown to be a specific biomarker for psoriasis.

IL-36lpha (aa 8-160) (mouse) (rec.) (untagged)

AG-40B-0098 10 μg

IL-36 β (aa 31-183) (mouse) (rec.) (untagged) AG-40B-0099 10 μ g

IL-36γ (aa 13-164) (mouse) (rec.) (untagged)

AG-40B-0100 IL-36Ra (aa 3-156) (mouse) (rec.) (untagged)

AG-40B-0096 $10 \mu g \mid 3 \times 10 \mu g$

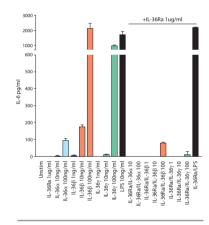
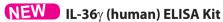


FIGURE: IL-36Ra (aa 3-156) (mouse) (rec.) (AG-40B-0096) inhibits IL-36 α , IL-36 β , IL-36 γ -dependent expression of IL-6 in mouse BMDCs.

UNIQUE IL-36γ & IL-37 ELISA Kits



AG-45B-0008 96 wells

IL-36 γ is a specific marker for Psoriasis. This assay is highly specific for human IL-36 γ . Does not cross-react with human IL-36 α and IL-36 β .

Sensitivity: 3 pg/ml **Range:** 3.9 to 250 pg/ml

Sample: Serum, Cell Culture Supernatant

IL-37 (human) ELISA Kit

10 µg

AG-45A-0041Y 96 wells | TwinPlex

IL-37 is a new anti-inflammatory cytokine. This assay detects natural and recombinant monomeric and dimeric human IL-37.

Sensitivity: 10 pg/ml **Range:** 16 to 1000 pg/ml **Sample:** Serum, Plasma

LIT: Homodimerization attenuates the anti-inflammatory activity of interleukin-37: A.M. Ellisdon, et al.; Sci. Immunol. 2, eaaj1548 (2017)



Plasma Cell Survival & Proliferation Blocking Antibodies

anti-BAFF (mouse), mAb (blocking) (Sandy-2)

AG-20B-0063 100 μg AG-20B-0063PF Preservative Free 100 μg | 500 μg

Functional Application: Depletion of B cells in vivo. Inhibition of mouse BAFF binding to BAFF-R and TACI (BCMA not tested); blocks BAFF activity in mice.

LIT: The B cell-stimulatory cytokines BLyS and APRIL are elevated in human periodontitis and are required for B cell-dependent bone loss in experimental murine periodontitis: T. Abe, et al.; J. Immunol. 195, 1427 (2015)

anti-APRIL (mouse), mAb (rec.) (blocking) (Apry-1-1)

AG-27B-0001 100 μg AG-27B-0001PF Preservative Free 100 μg AG-27B-0001B Biotin 100 μg

Functional Application: Depletion of plasma cells. Inhibits binding of mouse APRIL to mouse BCMA and TACI.

LIT: Production of the plasma-cell survival factor APRIL peaks in myeloid precursor cells from human bone marrow: T. Matthes, et al.; Blood 118, 1838 (2011)

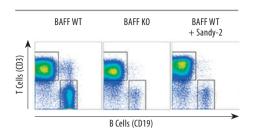


FIGURE: anti-BAFF (mouse), mAb (Sandy-2) (Prod. No. AG-20B-0063) blocks the action of endogenous BAFF *in vivo*.

METHOD: Wild type C57BL/6 mice were treated at day 0 (single administration) with monoclonal antibody anti-BAFF (mouse), mAb (Sandy-2) (at 2mg/kg). Lymph nodes were prepared at week 2 and analyzed by FACS for the presence of T (CD3) and B (CD19) cells. Untreated BAFF WT and KO mice were analyzed in parallel.

Multimeric & Mutant Cytokines with Increased Activity

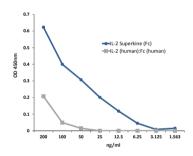
IL-2 Superkine (Fc)

AG-40B-0111 $10 \mu g \mid 3 \times 10 \mu g$

Compared to IL-2, IL-2 Superkine does not require CD25 to be active and induces superior expansion of cytotoxic CD8+ T and NK cells, leading to improved antitumor responses *in vivo*. IL-2 Superkine activates only poorly T regulatory cells (Tregs), meaning less toxic effects *in vivo*.

LIT: Exploiting a natural conformational switch to engineer an interleukin-2 'superkine': AM. Levin, et al.; Nature **484**, 529 (2012)

FIGURE: Binding of IL-2 Superkine (Fc) (Prod. No. AG-40B-0111) to IL-2Rß (human) is increased 10 fold compared to IL-2 (human):Fc (human).



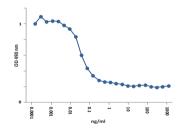
BAFF, Soluble (human) (60-mer) (rec.) (highly active)

AG-40B-0112 10 μg | 3 x 10 μg

Biological Activity: Increases B cell survival/proliferation. Increases CD21/CD23 expression on B cells *in vivo*. **Activates BAFF-R, TACI and BCMA receptors.** Works at concentrations <20ng/ml.

LIT: Mutation of the BAFF furin cleavage site impairs B-cell homeostasis and antibody responses: C. Bossen, et al.; Eur. J. Immunol. 41, 787 (2011)

FIGURE: BAFF, Soluble (human) (60-mer) (Prod. No. AG-40B-0112) binds and activates BCMA receptor.



TNF Ligands Multimeric Proteins

APRIL (human) (multimeric) (rec.)

AG-40B-0017 $10 \mu g \mid 3 \times 10 \mu g$

Binds to human and mouse BCMA and TACI. Binds to proteoglycans. Stimulates B cell proliferation.

APRIL (human) (H98) (multimeric) (rec.)

AG-40B-0088 $10 \mu g \mid 3 \times 10 \mu g$

Binds to human and mouse BCMA and TACI. Does not bind to proteoglycans. Stimulates B cell proliferation.

For a Complete Overview see our TNF Ligands Multimeric Proteins Product Flyer!

Fc (human):OX40L, Soluble (human) (rec.)

AG-40B-0172 $10 \mu g \mid 3 \times 10 \mu g$

Binds to human OX40 [CD134] and activates T cell proliferation.

CD40L (human) (multimeric) (rec.)

AG-40B-0010 $10 \mu g \mid 3 \times 10 \mu g$

Potent B cell activator.

FasL (human) (multimeric) (rec.) [MegaFasL™]

AG-40B-0130 $10 \mu g \mid 3 \times 10 \mu g$

Induces apoptosis of human Jurkat T cells at <0.2ng/ml.

Highly Active hIDO1 For Enzymatic Assays & Inhibitor Screenings

IDO1 (human) (rec.) (His) (highly active)

AG-40B-0161

50 µg

Specific Activity: >100'000U/mg protein with L-tryptophan as substrate (activity assay with catalase). One unit is defined as the amount of enzyme that produces 1nmol of N-formylkynurenine (NFK) per hour.

Potent Triazole IDO1 Inhibitor: MMG-0358

AG-CR1-3630

1 mg | 5 mg

Shows IC₅₀ values of 2nM in a cellular assay on mIDO1, 80nM in a cellular assay on hIDO1, 330nM in an enzymatic assay on hIDO1 at pH 6.5, and 71nM in an enzymatic assay on hIDO1 at pH 7.4. Active in vivo.

LIT: Rational design of 4-aryl-1,2,3-triazoles for indoleamine 2,3-dioxygenase 1 inhibition: U.F. Roehrig, et al.; J. Med. Chem. 55, 5270 (2012)

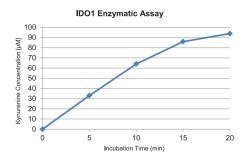


FIGURE: Enzymatic activity assay of IDO (human) (rec.) (His) (highly active) (Prod. No. AG-40B-0161).

NEW IDO1 (human) Enzyme + Inhibitor Set

1 Set

Includes 1x50 µg of Prod. No. AG-40B-0161 and 1x100 µg of Prod. No. AG-CR1-3630.

Highly Active DLL4 Mutant Protein



NEW DLL4 (human):Fc (human) (rec.) (highly active mutant)

AG-40B-0176

10 μg | 100 μg

Interacts with human Notch1 with >20 fold increase in affinity relative to WT DLL4 (as confirmed by ELISA).

> Visit www.adipogen.com for a Complete Panel of Notch-related Reagents!



In vivo Inflammasome Inhibitor

MCC950 . sodium salt (>98%)

AG-CR1-3615

1 mg | 5 mg | 10 mg

From the Manufacturer of **iNKT Stimulators**

α-Galactosylceramide (>96%) [KRN 7000]

AG-CN2-0013

250 μg | 1 mg

Potent Immunosuppressants

>> Cyclosporin A, FTY720, Mycophenolic acid, Rapamycin, and more!



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