Anti-Dxd Antibody [PSH0-36]

HA601133



Product Type: Mouse monoclonal IgG1, primary antibodies

Species reactivity: Species independent

Applications: ELISA
Clone number: PSH0-36

Description: Dxd (Exatecan derivative for ADC) is a potent DNA topoisomerase I inhibitor, with an IC50

of 0.31 μ M, used as a conjugated drug of HER2-targeting ADC (DS-8201a). Dxd is cytotoxic to human cancer cell lines of KPL-4, NCI-N87, SK-BR-3, and MDA-MB-468 with IC50s of 1.43 nM-4.07 nM, but the control IgG-ADC (Dxd is the payload) shows no inhibition on the four cell lines (with HER2 expression). DS-8201a (Dxd is the payload) displays significant suppression on the HER2-positive KPL-4, NCI-N87, and SK-BR-3 cell lines, with IC50 values of 26.8, 25.4, and 6.7 ng/mL, respectively, but with no such inhibition on MDA-MB-468. DXd is a part structure of DS-1062a which is a trophoblast cell surface protein 2 (TROP2)-targeting antibody-drug conjugate (ADC) comprised of a humanized anti-TROP2 monoclonal antibody, enzymatically cleavable peptide-linker, and DXd. DXd is an important part structure of DS-7300a, a novel B7-H3-targeting ADC, using DXd-ADC technology, which is composed of a humanized anti-B7-H3 mAb, an enzymatically cleavable tetra-

peptide-based linker, and a potent TOP1 inhibitor, DXd.

Immunogen: Dxd-OVA

Positive control: Dxd

Recommended Dilutions:

ELISA 1:1,000

Storage Buffer: PBS (pH7.4), 0.1% BSA, 40% Glycerol. Preservative: 0.05% Sodium Azide.

Storage Instruction: Store at $+4^{\circ}$ C after thawing. Aliquot store at -20° C. Avoid repeated freeze / thaw cycles.

Purity: Protein A affinity purified.

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Images

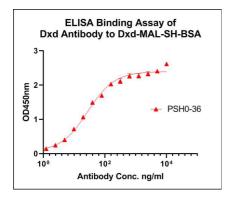


Fig1: Indirect ELISA analysis of Dxd was performed by coating wells of a 96-well plate with 50 μl per well of Dxd-MAL-SH-BSA diluted in carbonate/bicarbonate buffer, at a concentration of 1 μg/mL 2h at $37\,^{\circ}$ C. Wells of the plate were washed, blocked with 1% BSA blocking buffer, and incubated with 50 μL per well of Dxd monoclonal antibody (HA601133) serial diluted starting from a concentration of $10\,\mu\text{g/mL}$ for 1 hour at room temperature. The plate was washed and incubated with 50 μl per well of an HRP-conjugated goat anti-mouse IgG secondary antibody (HA1006) at a dilution of 1:5,000 for one hour at room temperature. Detection was performed using an Ultra TMB Substrate for 10 minutes at room temperature in the dark. The reaction was stopped with sulfuric acid and absorbances were read on a spectrophotometer at 450 nm.

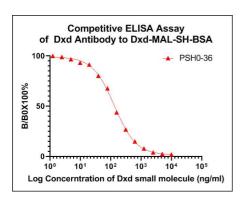


Fig2: Competitive ELISA analysis of Dxd was performed by coating wells of a 96-well plate with 50 μl per well of Dxd-MAL-SH-BSA diluted in carbonate/bicarbonate buffer, at a concentration of 1 μg/mL overnight at 4° C. Wells of the plate were washed, blocked with 1% BSA blocking buffer, and incubated with 50 μL per well of Dxd monoclonal antibody (HA601133) at concentration of 1 μg/mL with serial diluted Dxd starting from a concentration of 10μg/mL for 1 hour at room temperature. The plate was washed and incubated with 50 μl per well of an HRP-conjugated goat antimouse IgG secondary antibody (HA1006) at a dilution of 1:5,000 for one hour at room temperature. Detection was performed using an Ultra TMB Substrate for 10 minutes at room temperature in the dark. The reaction was stopped with sulfuric acid and absorbances were read on a spectrophotometer at 450 nm.

Note: All products are "FOR RESEARCH USE ONLY AND ARE NOT INTENDED FOR DIAGNOSTIC OR THERAPEUTIC USE".

Background References

- 1. Ogitani Y, Aida T, Hagihara K, et al. DS-8201a, A Novel HER2-Targeting ADC with a Novel DNA Topoisomerase I Inhibitor, Demonstrates a Promising Antitumor Efficacy with Differentiation from T-DM1Preclinical Efficacy of DS-8201a, a Novel HER2-Targeting ADC[J]. Clinical Cancer Research, 2016, 22(20): 5097-5108.
- 2. Okajima D, Yasuda S, Yokouchi Y, et al. Preclinical efficacy studies of DS-1062a, a novel TROP2-targeting antibody-drug conjugate with a novel DNA topoisomerase I inhibitor DXd[J]. 2018.

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