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SZABO-SCANDIC HandelsgmbH

Quellenstraße 110, A-1100 Wien

T. +43(0)1 489 3961-0

F. +43(0)1 489 3961-7

mail@szabo-scandic.com

www.szabo-scandic.com

[linkedin.com/company/szaboscandic](https://www.linkedin.com/company/szaboscandic) 

Benralizumab

Cat. No.:	HY-P9923
CAS No.:	1044511-01-4
Target:	Interleukin Related; Apoptosis
Pathway:	Immunology/Inflammation; Apoptosis
Storage:	Please store the product under the recommended conditions in the Certificate of Analysis.

BIOLOGICAL ACTIVITY

Description	Benralizumab (MEDI-563) is an interleukin-5 receptor α (IL-5R α)-directed cytolytic monoclonal antibody that induces direct, rapid and nearly complete depletion of eosinophils via enhanced antibody-dependent cell-mediated cytotoxicity. Benralizumab can be used for severe eosinophilic asthma ^[1] .									
IC₅₀ & Target	IL-5R α									
In Vitro	<p>Benralizumab (MEDI-563) binds to recombinant human and cynomolgus monkey IL-5Rα extracellular domains with a dissociation constant of 11 and 42 pM, respectively^[2].</p> <p>Benralizumab (1 pM-100 nM, 48 h) inhibits IL-5-induced proliferation of CTLL-2 cells transfected with recombinant human IL-5R$\alpha\beta$ with an IC₅₀ of 0.3 nM^[2].</p> <p>Benralizumab binds only to the constructs containing human IL-5Rα D1^[2].</p> <p>Benralizumab mediates ADCC (antibody-dependent cell-mediated cytotoxicity) of human eosinophils and basophils with EC₅₀s of 0.9 pM and 0.5 pM, respectively, and mediates eosinophil apoptosis^[2].</p> <p>MCE has not independently confirmed the accuracy of these methods. They are for reference only.</p> <p>Cell Proliferation Assay^[2]</p> <table border="1"> <tr> <td>Cell Line:</td> <td>CTLL-2 human IL-5R$\alpha\beta$ cells</td> </tr> <tr> <td>Concentration:</td> <td>1 pM-100 nM</td> </tr> <tr> <td>Incubation Time:</td> <td>48 h</td> </tr> <tr> <td>Result:</td> <td>Inhibited cell proliferation with an IC₅₀ of 0.3 nM.</td> </tr> </table>		Cell Line:	CTLL-2 human IL-5R $\alpha\beta$ cells	Concentration:	1 pM-100 nM	Incubation Time:	48 h	Result:	Inhibited cell proliferation with an IC ₅₀ of 0.3 nM.
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Result:	Inhibited cell proliferation with an IC ₅₀ of 0.3 nM.									
In Vivo	<p>Benralizumab (MEDI-563) (0-30 mg/kg; i.v.; once every 3 weeks for 12 weeks) reduces peripheral blood and bone marrow eosinophil^[2].</p> <p>MCE has not independently confirmed the accuracy of these methods. They are for reference only.</p> <table border="1"> <tr> <td>Animal Model:</td> <td>Female and male cynomolgus monkeys^[2]</td> </tr> <tr> <td>Dosage:</td> <td>0.1, 1, 10, and 30 mg/kg</td> </tr> <tr> <td>Administration:</td> <td>Intravenous injection, once every 3 weeks for 12 weeks</td> </tr> </table>		Animal Model:	Female and male cynomolgus monkeys ^[2]	Dosage:	0.1, 1, 10, and 30 mg/kg	Administration:	Intravenous injection, once every 3 weeks for 12 weeks		
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Result:

Blood eosinophil counts decreased close to the limit of detection after the first administration at all dose levels and remained undetectable. Eosinophil precursors in the bone marrow were reduced 80% or greater in all doses 3 days after the last administration (terminal necropsy) and remained undetectable until 18 days after the last dose in the 30 mg/kg group. The profound effect observed in the bone marrow was specific for the eosinophil lineage.

REFERENCES

- [1]. Kolbeck R, et al. MEDI-563, a humanized anti-IL-5 receptor alpha mAb with enhanced antibody-dependent cell-mediated cytotoxicity function. *J Allergy Clin Immunol.* 2010 Jun;125(6):1344-1353.e2.
- [2]. Menzies-Gow A, et al. Corticosteroid tapering with benralizumab treatment for eosinophilic asthma: PONENTE Trial. *ERJ Open Res.* 2019 Sep 25;5(3). pii: 00009-2019.

Caution: Product has not been fully validated for medical applications. For research use only.

Tel: 609-228-6898

Fax: 609-228-5909

E-mail: tech@MedChemExpress.com

Address: 1 Deer Park Dr, Suite Q, Monmouth Junction, NJ 08852, USA