## IVMB0489

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IVMB0489	Clone:	IMC-1121B	Target:	VEGFR2
500 µg			<b>Isotype</b> :	Human lgG1ĸ
formation				
Human	Human		Host Species	<b>s</b> : Human
: Biosimilar Rec	Biosimilar Recombinant Human Monoclonal Antibody		Expression Host: HEK-293 Cells	
	IVMB0489 500 µg formation Human	IVMB0489 Clone: 500 µg formation Human	IVMB0489 Clone: IMC-1121B 500 μg formation Human	IVMB0489 Clone: IMC-1121B Target: 500 μg Isotype: formation Human Host Specie

## Immunogen Information

Background: Vascular endothelial growth factors (VEGF) and VEGF receptors (VEGFR) play an essential role in angiogenesis<sup>1</sup>. There are three VEGFRs: VEGFR-1, VEGFR-2, and VEGFR-3. VEGFR-1 and VEGFR-2 are responsible for angiogenesis, and VEGFR-3 affects lymphogenesis. In the pathogenesis of diseases including diabetes mellitus, rheumatoid arthritis, and cancer, new blood vessel formation is highjacked. Changes at the VEGF/VEGFR-2 axis are particularly potent at allowing VEGF-induced proliferation, migration, and vascular endothelial cell differentiation during tumor angiogenesis. Additionally, VEGFR-2 is upregulated in tumor vascular endothelial cells, and VEGF levels are associated with poor prognosis and resistance to chemotherapy. Consequently, the VEGF/VEGFR axis is a prime anti-cancer target. Blocking VEGF/VEGFR-2 with Ramucirumab inhibits tumor growth in animal models and cancer patients<sup>2</sup>, <sup>3</sup>, <sup>4</sup>, and Ramucirumab is approved by the US Food and Drug Administration for treatment of various cancers<sup>5</sup>. Ramucirumab blocks all known VEGFs from binding to VEGFR-2<sup>4</sup>, <sup>6</sup>. Ramucirumab specifically and potently inhibits VEGFR-2 by binding to the VEGF-binding domain at an epitope located within VEGFR-2 extracellular Ig domain 37. Ramucirumab inhibits VEGF/VEGF-2 interaction 1, VEGFR-2 phosphorylation<sup>7</sup>, VEGFinduced VEGFR-2 activation<sup>1</sup>, VEGF-stimulated cellular migration<sup>6</sup> and proliferation<sup>1</sup>, and prolongs the survival of leukemia-inoculated mice<sup>6</sup>. Ramucirumab (IMC-1121B) was fully humanized from chimeric antibody IMC-1121<sup>1</sup>, which was constructed from a Fab fragment (Hu-1121 Fab) isolated by immunopanning against VEGFR-2 under stringent conditions using a VL-shuffled library and the VH gene segment Hu- 2C6 Fab<sup>6</sup>, <sup>7</sup>. The original library



was constructed from spleen cells of mice immunized with a soluble form of VEGFR-2<sup>8</sup>. Ramucirumab was converted into a full length bivalent IgG1 antibody from the Fab fragment 1121<sup>7</sup>.

Ramucirumab, clone IMC-1121B, a non-therapeutic biosimilar antibody for research use only was developed recombinantly and has the same variable regions as the original therapeutic.

Endotoxin Level: < 1.0 EU/mg as determined by the LAL method

- Applications: ELISA
- Synonyms: Ramucirumab, VEGFR-2

Antigen Distribution:VEGFR-2 is widely expressed by vascular endothelial cells, some vascular tumors,<br/>carcinomas, malignant melanomas, and lymphomas. Certain leukemia cells express<br/>functional VEGFR on the cell surface.

Immunogen: Human VEGFR2

- Formulation:This biosimilar antibody is aseptically packaged and formulated in 0.01 M phosphate<br/>buffered saline (150 mM NaCl) PBS pH 7.2 7.4 with no carrier protein, potassium, calcium<br/>or preservatives added. Due to inherent biochemical properties of antibodies, certain<br/>products may be prone to precipitation over time. Precipitation may be removed by aseptic<br/>centrifugation and/or filtration.
- Specificity: This non-therapeutic biosimilar antibody uses the same variable region sequence as the therapeutic antibody Ramucirumab. Ramucirumab activity is directed against human vascular endothelial growth factor receptor-2 (VEGFR-2; also known as kinase insert domain-containing receptor, KDR).

 Product Preparation:
 Recombinant biosimilar antibodies are manufactured in an animal free facility using onlyin vitroprotein free cell culture techniques and are purified by a multi-step process including the use of protein A or G to assure extremely low levels of endotoxins, leachable protein A or aggregates.

Storage & Handling:Functional grade biosimilar antibodies may be stored sterile as received at 2-8°C for up to<br/>one month. For longer term storage, aseptically aliquot in working volumes without diluting<br/>and store at -80°C. Avoid Repeated Freeze Thaw Cycles.