

## **Recombinant Rat TIM-3 Fc Chimera**

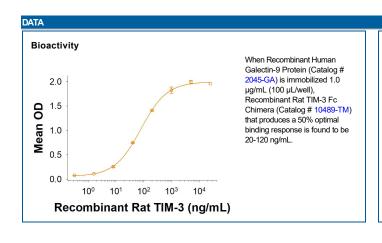
Catalog Number: 10489-TM

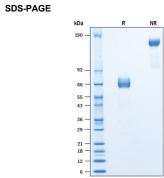
DESCRIPTION				
Source	Mouse myeloma cell line, NS0-derived rat TIM-3 protein			
	Rat TIM-3 (Leu22-Ala194) Accession # P0C0K5.2	IEGRMD	Mouse IgG <sub>1</sub> (Pro100-Lys330)	
	N-terminus		C-terminus	

N-terminal Sequence Analysis	Leu22
Structure / Form	Disulfide-linked homodimer
Predicted Molecular	46 kDa

SPECIFICATIONS		
SDS-PAGE	60-75 kDa, under reducing conditions	
Activity	Measured by its binding ability in a functional ELISA.  When Recombinant Human Galectin-9 Protein (Catalog # 2045-GA) is immobilized at 1.0 μg/mL (100 μL/ well), Recombinant Rat TIM-3 Fc Chimera (Catalog # 10489-TM) that produces a 50% optimal binding response is found to be 20-120 ng/mL	
Endotoxin Level	<0.10 EU per 1 µg of the protein by the LAL method.	
Purity	>95%, by SDS-PAGE visualized with Silver Staining and quantitative densitometry by Coomassie® Blue Staining.	
Formulation	Lyophilized from a 0.2 µm filtered solution in PBS with Trehalose. See Certificate of Analysis for details.	

PREPARATION AND STORAGE			
Reconstitution	Reconstitute at 250 µg/mL in PBS.		
Shipping	The product is shipped at ambient temperature. Upon receipt, store it immediately at the temperature recommended below.		
Stability & Storage	Use a manual defrost freezer and avoid repeated freeze-thaw cycles.  12 months from date of receipt, -20 to -70 °C as supplied.  1 month, 2 to 8 °C under sterile conditions after reconstitution.  3 months, -20 to -70 °C under sterile conditions after reconstitution.		





2 µg/lane of Recombinant Rat TIM-3 Fc Chimera Protein (Catalog # 10489-TM) was resolved with SDS-PAGE under reducing (R) and non-reducing (NR) conditions and visualized by Coomassie® Blue staining, showing bands at 60-75 kDa and 120-150 kDa, respectively.

Rev. 5/20/2020 Page 1 of 2





## Recombinant Rat TIM-3 Fc Chimera

Catalog Number: 10489-TM

## BACKGROUND

T cell immunoglobulin and mucin domain-3 (TIM-3) also known as HAVCR2, is a member of the TIM family of immune regulating molecules. TIMs are type I transmembrane glycoproteins with one Ig-like V-type domain, a Ser/Thr-rich mucin stalk region (1, 2). While lacking a specific immunoreceptor tyrosine-based inhibition motif (ITIM), the cytoplasmic domain of TIM-3 contains a conserved region of five tyrosine residues important for downstream signaling (3). A soluble form of TIM-3 lacking the mucin stalk and transmembrane domains is formed as either a result of alternative splicing or metalloproteinase-dependent cleavage. Within the ECD, mature rat TIM-3 shares 60% and 69% amino acid sequence identity with human and mouse TIM-3, respectively. TIM-3 is up-regulated on several populations of activated myeloid cells (macrophage, monocyte, dendritic cell, microglia, mast cell) and T cells (Th1, CD8+, NK, Treg) (3-10). Its binding to Galectin-9 induces a range of immunosuppressive functions which enhance immune tolerance and inhibit anti-tumor immunity (3,11). TIM-3 ligation attenuates CD8+ and Th1 cell responses (11-13) and promotes the activity of Treg and myeloid derived suppressor cells (8, 11, 13, 14). In addition, dendritic cell-expressed TIM-3 dampens inflammation by enabling the phagocytosis of apoptotic cells and the cross-presentation of apoptotic cell antigens (4). It also binds the alarmin HMGB1, thereby preventing the activation of TLRs in response to released tumor cell DNA (7). TIM-3 interactions with Galectin-9 can alternatively trigger immune stimulatory effects, such as the coactivation of NK cell cytotoxicity (10).

## References:

- 1. Sakuishi, K. et al. (2011) Trends Immunol. 32:345.
- 2. Anderson, A.C. (2012) Curr. Opin. Immunol. 24:213.
- 3. Wolf, Y. et al. (2020) Nature Reviews Immunology 20:173.
- 4. Nakayama, M. et al. (2009) Blood 113:3821.
- 5. Anderson, A.C. et al. (2007) Science 318:1141
- 6. Wiener, Z. et al. (2007) J. Invest. Dermatol. 127:906.
- 7. Chiba, S. et al. (2012) Nat. Immunol. 13:832.
- 8. Sanchez-Fueyo, A. et al. (2003) Nat. Immunol. 4:1093.
- 9. Ndhlovu, L.C. et al. (2012) Blood 119:3734.
- 10. Gleason, M.K. et al. (2012) Blood 119:3064.
- 11. Zhu, C. et al. (2005) Nat. Immunol. 6:1245.
- 12. Sakhdari, A. et al. (2012) PLoS ONE 7:e40146.
- 13. Sabatos, C.A. et al. (2003) Nat. Immunol. 4:1102.
- 14. Dardalhon, V. et al. (2010) J. Immunol. 185:1383.