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## **Title**

MAp44

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## **Authors**

Chandrasekhar, Anjana Dinasarapu, Ashok Reddy Matsushita, Misao et al.

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Review Article Open Access

# MAp44

Anjana Chandrasekhar<sup>1</sup>, Ashok Reddy Dinasarapu<sup>1</sup>, Misao Matsushita<sup>2</sup>, Shankar Subramaniam<sup>3</sup>

MAp44 is a ~44 kDa alternate splice product of *MASP1* and is mainly expressed in the heart. Mannose/mannan binding lectin (MBL) associated serine proteases, MASP-1 and MASP-3 are other products of *MASP1*. Similar to MASP-1 (isoform 1 of *MASP1*, which represents the longest transcript), MAp44 has a C1r/C1s/Uegf/bmp1 (CUB) domain, calcium-binding EGF-like domain and complement control protein (CCP) domains. However, it lacks the serine protease domain of MASP-1 and therefore cannot perform MASP-1's functions. MAp44 binds to multimeric pathogen receptors such as MBL and the three ficolins, and is believed to play a regulatory role in the lectin pathway of complement activation.

#### **KEYWORDS**

Mannan-binding lectin serine peptidase 1 (C4/C2 activating component of Ra-reactive factor); Mannose-binding lectin-associated protein of 44 kDa; Mannose-binding lectin-associated serine protease 1; Mannose-binding protein-associated serine protease; MAP-1; MAP1; Map44; MASP; MASP1

#### **IDENTIFIERS**

Molecule Page ID:A008392, Species:Human, NCBI Gene ID: 5648, Protein Accession:NP\_001027019.1, Gene Symbol:MASP1

#### PROTEIN FUNCTION

MASP-1 (*MASP1* Isoform 1), MASP-3 (*MASP1* Isoform 2) and MAp44 (*MASP1* Isoform 3) are splice products of *MASP1* (Degn *et al.* 2009). MAp44 is also known as MAP-1. Similar to MASP-1, MAp44 circulates in the serum in complex with mannose/mannan-binidng lectin (MBL), L-ficolin or H-ficolin. However, unlike MASP-1 and MASP-3, MAp44 lacks serine protease domain (Skjoedt *et al.* 2010, Skjoedt *et al.* 2011, Skjoedt *et al.* 2012). The only known function of MAp44 is to down-regulate C4 activation by competing with MASP-1 and MASP-2 to bind to pattern recognition molecules such as MBL and ficolins. This further affects downstream complement pathway activation. MAp44 can also disrupt MASP-1-MASP-2 heterodimer and thereby down-regulate complement pathway (Degn *et al.* 2013).

### REGULATION OF ACTIVITY

null

#### INTERACTIONS

MAp44, like other MASP proteins forms a head-to-tail homodimer (Skjoedt *et al.* 2012). The homodimer interacts with MBL, L-ficolin and H-ficolin, with H-ficolin being its preferred partner (Degn *et al.* 2009, Skjoedt *et al.* 2010, Skjoedt *et al.* 2011). MAp44 also interacts with the recently discovered novel collectin, CL-11 (CL-K1) (Ma *et al.* 2013).

The experimental methods used to characterize these interactions are documented in CMAP, a complement map database (Yang *et al.* 2013).

#### **PHENOTYPES**

No SNPs or disease associations have been documented so far.

## MAJOR SITES OF EXPRESSION

MAp44 is mainly expressed in the myocardial fibers followed

by the skeletal muscle, unlike other MASP proteins (MASP-1, MASP-2 and MASP-3), which are mainly expressed in the liver (Degn *et al.* 2009, Skjoedt *et al.* 2010).

#### SPLICE VARIANTS

MAp44 transcript (MASP1 isoform 3) is an alternative splice variant of MASP1 (Degn et al. 2009). MASP1 also encodes for two more proteins, MASP-1 (MASP1 isoform 1) and MASP-3 (MASP1 isoform 2) (Dahl et al. 2001). MASP1 encodes for six domains: two C1r/C1s/Uegf/bone morphogenetic protein 1 (CUB), an epidermal growth factor (EGF)-like, two complement control proteins (CCPs) and a serine protease domain. The first five domains (encoded by exons 1-11) together form the heavy (or 'A') chain, while the serine protease domain forms the light (or 'B') chain (encoded by exons 12-18) (Fujita et al. 2002). MAp44 is formed by alternative splicing at the ninth exon of MASP1. Thus, MAp44 has two CUB domains, EGF, one CCP domain, a unique C-terminal domain of 17 a.a (encoded by exon 9) and importantly lacks the serine protease domain (Degn et al. 2009, Skjoedt et al. 2010). Please refer to MASP-1 and MASP-3 Molecule Pages at www.signalinggateway.org for more information.

#### REGULATION OF CONCENTRATION

MAp44 is found in human serum at 0.24  $\mu$ g/ml (Skjoedt et al. 2011) or 1.4  $\mu$ g/ml in Ca<sup>2+</sup>-dependent complexes with MBL and ficolins (Degn *et al.* 2009).

## ANTIBODIES

Commercial antibody specific to MAp44 is available from Hycult Biotech.

# Table 1: Functional States

STATE DESCRIPTION	LOCATION	REFERENCES
MAp44	extracellular space	
2(MAp44)	extracellular space	Skjoedt MO et al. 2012
2(MAp44) - glycosylated	extracellular space	Skjoedt MO et al. 2011
2(MAp44)/MBL	extracellular space	Skjoedt MO et al. 2011; Skjoedt MO et al. 2010; Skjoedt MO et al. 2012
2(MAp44)/L-FCN	extracellular space	Degn SE et al. 2009; Skjoedt MO et al. 2011; Skjoedt MO et al. 2010
2(MAp44)/H-FCN	extracellular space	Degn SE et al. 2009; Skjoedt MO et al. 2011; Skjoedt MO et al. 2010
2(MAp44)/CL-K1	extracellular space	Ma YJ et al.

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#### SUPPLEMENTARY

Supplementary information is available online.

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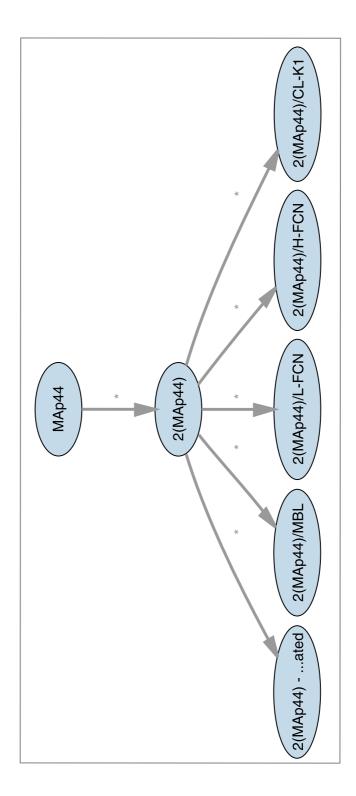
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This molecule exists in 7 states , has 6 transitions between these states and has 0 enzyme functions.(Please zoom in the pdf file to view details.)



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