

BACKGROUND

In mammals, the heart arises from the differentiation of 2 sources of multipotent cardiovascular progenitors (MCPs). *Mesp1* (mesoderm posterior 1) is a key regulator of cardiovascular progenitors in vertebrates. Lineage tracing in mice demonstrated that *Mesp1* represents the earliest marker of cardiovascular progenitors, tracing almost all the cells of the heart including derivatives of the primary and second heart fields. *Mesp1* is a basic helix-loop-helix transcription factor that is transiently expressed by the earliest progenitors of the cardiovascular system from E6.5 to E7.5. During gastrulation, progenitors of cardiogenic mesoderm arise at E6.5 in the posterior lateral epiblast and migrate to form the cardiac crescent at E7.5, when regionalized cell fates are first delineated. Lineage tracing and heterotopic transplantation studies suggest that precursors in the earliest heart field possess potential to generate myocardium, endocardium, and epicardium, but subsequently become restricted as lineage-specific regulatory programs are activated.¹ *Mesp1* and *Mesp2* are closely related members but share significant sequence homology only in their bHLH regions. The inactivation of *Mesp1/2* indicated that *Mesp* genes are essential for early cardiac mesoderm formation and MCP migration. Several recent studies have demonstrated that *Mesp1* massively promotes cardiovascular differentiation during embryonic development and pluripotent stem cell differentiation and indicated that *Mesp1* resides at the top of the cellular and transcriptional hierarchy that orchestrates MCP specification.² Defining how *Mesp1* regulates the earliest step of MCP specification and controls their migration is essential to understand the root of cardiovascular development and how the deregulation of these processes can lead to congenital heart diseases. In addition, these findings will be very useful to boost the production of cardiovascular cells for cellular therapy, drug and toxicity screening.³

For the cellular and the molecular mechanism by which *Mesp1* acts during cardiac specification, it was shown that *Mesp1* acts as a cardiovascular master regulator during specification of MCPs during ESC differentiation. Embryonic stem cell (ESC) differentiation is a good model to study the cardiovascular progenitor specification. *Mesp1* both directly activated many key genes belonging to the core cardiac transcriptional machinery and directly repressed genes promoting early mesoderm and endoderm cell-fate specification. *Mesp1* first transiently stimulated its own endogenous expression through a direct positive autoregulatory loop and then inhibited its own expression, therefore, acting as a molecular switch during cardiac specification. Altogether, our results provide compelling evidence that *Mesp1* acts as a cardiovascular master regulator during specification of MCPs during ESC differentiation.⁴ In addition, it was further demonstrated that

Mesp1 promoted mesoderm development independently of Wnt signaling. Transient *Mesp1* expression in ESCs promotes changes associated with epithelial-mesenchymal transition (EMT) and induction of *Snai1*, consistent with a role in gastrulation. *Mesp1* expression also restricted the potential fates derived from ESCs, generating mesoderm progenitors with cardiovascular, but not hematopoietic, potential. Thus, in addition to its effects on EMT, *Mesp1* may be capable of generating the multipotent cardiovascular progenitor from ESCs *in vitro*.⁵

References:

1. Saga, Y. et al: Development 126:3437-47,1999
2. Takahashi, Y. et al: Develop. 135:787-96, 2008
3. Bondue, A. & Blanpain, C.: Circulat. Res. 107:1414-27, 2010
4. Bondue, A. et al: Cell Stem Cell 3:69-84, 2008
5. Lindsley, R.C. et al: Cell Stem Cell 3:55-68, 2008

TECHNICAL INFORMATION

Source:

Mesp1 Antibody is a mouse monoclonal antibody raised against recombinant human *Mesp1* proteins expressed in *E. coli*.

Specificity and Sensitivity:

This antibody detects endogenous *Mesp1* proteins without cross-reactivity with other family members.

Storage Buffer: PBS and 30% glycerol

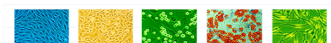
Storage:

Store at -20°C for at least one year. Store at 4°C for frequent use. Avoid repeated freeze-thaw cycles.

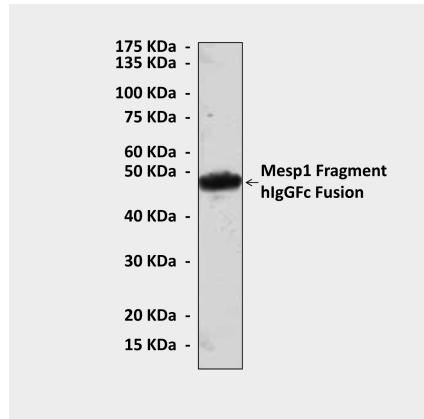
APPLICATIONS

Application:	*Dilution:
WB	1:1000
IP	n/d
IHC	n/d
ICC	n/d
FACS	n/d

*Optimal dilutions must be determined by end user.



QUALITY CONTROL DATA



Western Blot detection of Mesp1 proteins in 293 cell lysate containing human Mesp1-hlgGfc fusion proteins using Mesp1 Antibody.

