## Investigating endothelial polarization and vascular diseases using zebrafish embryos

Hyouk-Bum Kwon<sup>1\*</sup>, Shengpeng Wang<sup>1</sup>, Christian S.M. Helker<sup>1</sup>, S. Javad Rasouli<sup>1</sup>, Hans-Martin Maischein<sup>1</sup>, Stefan Offermanns<sup>1</sup>, Wiebke Herzog<sup>2</sup>, <sup>3</sup> and Didier Y.R. Stainier<sup>1\*</sup>

## Address

 Max Planck Institute for Heart and Lung Research, Department of Developmental Genetics, Ludwigstraße 43, 61231 Bad Nauheim, Germany
University of Münster, Schlossplatz 2, 48149 Münster, Germany
Max Planck Institute for Molecular Biomedicine, Röntgenstraße 20, 48149 Münster, Germany

\* Correspondence should be addressed to Hyouk-Bum Kwon (email: Hyouk-Bum.Kwon@mpi-bn.mpg.de) or Didier Stainier (email: Didier.Stainier@mpibn.mpg.de)

## Abstract

Endothelial cells (ECs) have the ability to align in the direction of flow in response to shear stress. However, how ECs respond to flow in complex *in vivo* environments is less clear. To analyze endothelial polarization *in vivo*, we generated an endothelial-specific transgenic line to mark the Golgi apparatus. We find that most ECs polarize within 4.5 hours after the onset of vigorous blood flow and, by manipulating cardiac function, observed that flow-induced EC polarization is a dynamic and reversible process. Based on its role in EC migration, we analyzed the role of Apelin signaling in EC polarization, and find that it is critical for this process. Knocking down Apelin receptor function in primary human ECs also affects their polarization. Our studies provide new tools to analyze the mechanisms of EC polarization *in vivo*, and reveal an important role in this process for a signaling pathway implicated in cardiovascular disease.