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## ULTRASTRUCTURAL AND FUNCTIONAL ANALYSIS OF WEIBEL-PALADE BODIES

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## Abstract:

The adhesive glycoprotein von Willebrand factor (VWF) is stored for secretion in specialized endothelial cell secretory organelles called Weibel-Palade bodies (WPBs). VWF plays a crucial role in hemostasis and defects in the protein result in the bleeding disorder Von Willebrands disease. Previously we used Fourier analysis of cryo-EM images and tomographic reconstructions of vitrified human umbilical vein endothelial cells (HUVEC) to show that VWF is stored as a series of helical tubules that tight pack to a paracrystalline matrix in the WPB lumen (Berriman et al 2009, 2009;106:17407-17412). WPB fusion with the plasma membrane is associated with disassembly of the paracrystalline array of tubules and appearance of a membranous extension of the WPB that we speculated might represent a connection to the plasma membrane. Here we extend our ultrastructural studies of WPBs in situ in vitrified HUVEC to reveal a novel structural feature of the organelle. Using light microscopy of live and fixed cells we identify a marker of this novel structural feature. Direct visualization of these features during cell stimulation sheds new light on the nature of the morphological changes during WPB exocytosis.

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