

CLINICAL SPECIFICATIONS

ACTOMYOSIN NETWORK

Function:

Gastrointestinal microfilaments of the actomyosin network are critical for apical junctional complex biogenesis and function.¹ The apical junctional complex, made up in part by tight junction proteins zonulin and occludin, is responsible for preventing antigen invasion and preservation of the biochemical homeostasis within the gastrointestinal tract.³ The actomyosin network can signal tight junction contractions and give structure to their assembly.

Antibodies Appear:

Autoimmune liver disorders² Celiac disease^{1, 4} Chronic hepatitis² Crohn's disease⁴ Myasthenia Gravis⁵

Known Cross-Reactions: Giardia lamblia,⁶ Entamoeba histolytica⁷

Clinical Significance:

Many environmental factors such as bacterial toxins can affect the stability of the actomyosin network and occludin/zonulin. Antibodies to the actomyosin network are therefore biomarkers of intestinal barrier dysfunction, either via bacterial infiltration or by an autoimmune mechanism aimed at the gastrointestinal tract. For the best clinical value, antibodies against the actomyosin network should be measured in conjunction with lipopolysaccharide (LPS) and occludin/zonulin proteins. When antibodies are detected against the actomyosin alone, it is an indication of autoimmunity against the mucosal epithelium and other tissue cell cytoskeleton of the intestinal barrier. When antibodies are detected against the actomyosin network and LPS, but none are detected for occludin/zonulin, this indicates a breakdown in intestinal barrier integrity by bacterial antigens through the transcellular pathway. The detection of antibodies against actomyosin, LPS, and occludin/zonulin indicates that there has been both transcellular and paracellular penetration of the intestinal barrier.

References:

- 1. Clemente, et al. Enterocyte actin autoantibody detection: a new diagnostic tool in Celiac disease, results of a multi-center study. Am Gastroenterol, 2004; 99: 1551-1556.
- 2. Gröschel-Stewart and Doniach. Immunological evidence for human myosin isoenzymes. Immunology, 1969; 17:991-994.
- 3. Ivanov, et al. Differential roles for actin polymerization and a myosin II motor in assembly of the epithelial apical junctional complex. Mol Biol Cell, 2005; 16:2636-2650.
- 4. Magalhaes, et al. Studies on the nature and significance of connective tissue antibodies in adult coeliac disease and Crohn's disease. Gut, 1974; 15:284-288.
- 5. Romi, et al. Striational antibodies in Myasthenia Gravis: reactivity and possible clinical significance. Arch Neural, 2005; 62:442-446.
- 6. Peattie, et al. Ultrastructural localization of giardins to the edges of disk microribbons of *Giardia lamblia* and the nucleotide and deduced protein sequence of alpha giardin. J Cell Biol, 1989; 109(5):2323-2335.
- 7. Edman, et al. Genomic and cDNA actin sequences from a virulent strain of *Entamoeba histolytica*. Proc Natl Acad Sci U S A, 1987; 84(9):3024-3028.