Colonic wall remodeling in patients with short- and long-lasting ulcerative colitis

¹ Chiara Ippolito - ² Cristina Segnani - ² Erika Tirotta - ³ Rocchina Colucci - ² Carolina Pellegrini - ² Corrado Blandizzi - ¹ Sauro Dini - ⁴ Mariella Errede - ⁴ Francesco Girolamo - ⁴ Daniela Virgintino - ¹ Amelio Dolfi - ¹ <u>Nunzia Bernardini</u>

¹University of Pisa, Department of Clinical and Experimental Medicine, Unit of Histology and Medical Embryology, Pisa, Italia - ²University of Pisa, Department of Clinical and Experimental Medicine, Division of Pharmacology, Pisa, Italia - ³University of Padova, Department of Pharmaceutical and Pharmacological Sciences, Padova, Italia - ⁴University of Bari, Department of Basic Medical Sciences, Neurosciences and Sensory Organs, Unit of Human Anatomy and Histology, School of Medicine, Bari, Italia

Inflammatory bowel disease (IBD) are chronic and progressive pathologies associated with invalidating abdominal symptoms and increasing incidence in Western countries. Although fibrostenosis is a rare event in ulcerative colitis [UC], it may evolve to fibrosis in the later stages (1, 2). In the present study we examined the histopathological remodeling of the colonic wall from short- and long-lasting (SL and LL) UC patients. Full-thickness left colonic surgical samples were obtained from nonstenotic SL (\leq 3 years) and LL (\geq 10 years) UC patients with a severe exacerbation of colitis and active disease, without clinical signs of fibrostenosis. Collagen and elastic fibres, vascular networks and parameters of fibrosis have been evaluated by histochemistry, immunohistochemistry, confocal immunofluorescence and/or western blot. For comparison, normal colonic control samples from subjects who underwent surgery for uncomplicated colon cancer and without IBD and diverticular disease, were considered.Both SL- and LL-UC showed a thickening in tunica muscularis and activation of transmural neovessels, with proliferating CD105-positive endothelial cells and activated nestin-positive pericytes, as compared with controls. The colonic wall of LL-UC displayed a significant increase in collagen deposition and fibrotic markers (type I and III collagen, fibronectin, vimentin, RhoA), an enhancement of proliferation (PCNA), a decrease in elastic fibres, together with a fibrotic rearrangement of the tunica muscularis smooth muscle cells. In conclusion, a significant full-thickness remodeling of the colonic wall has been documented in the colonic tissues from both SL-UC and LL-UC, as compared to controls: transmural fibrotic thickening and angiogenesis were found in both UC groups, together with a cellular fibrotic/proliferative switch in the tunica muscularis which occurs in the later stages. These histopathological remodeling may contribute to the reduced elastic properties and tissue stretch capability of the wall, thereby impairing the occurrence of a normal motor activity in these subjects.

References

[1] Bush et al. Curr Gastroenterol Rep 2014.

[2] Torres et al. Inflamm Bowel Dis 2012.

Keywords

Ulcerative colitis; active disease; colonic muscle remodeling; intestinal angiogenesis/fibrosis.