Role of adipokines in enhanced pain and inflammation in a rodent model of obesity

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Obese individuals are more likely to be affected by chronic pain, however, the biological mechanisms underpinning this comorbidity are not known. A causal link may be dysregulated secretion of inflammatory adipokines both from expanding adipose tissue and centrally. The aim of this study was to characterize altered pain processing and changes in inflammatory cytokine expression in spinal cord in rodent models of obesity. Responses to thermal and mechanical stimulation of the hind paw were assessed in adult male Wistar rats fed a high fat diet (HFD; 22%) or normal diet for 16 weeks (n=6/group) in absence of inflammation, and then in response to intradermal hind paw injection of carrageenan (3%; 50μl), a model of acute inflammation. Spinal cord was collected and adipokine mRNA expression, cholesterol and triglycerides (TAGs) measured using real-time PCR and ELISA. Rats fed a HFD gained significantly more weight than controls (502 ± 12g vs. 444 ± 7g; P<0.01), and displayed plasma hyperinsulaemia and hypercholesterolaemia (both P<0.05 vs. controls) but normoglycaemia. Acute nociceptive responses were unchanged in obese rats but they displayed potentiated mechanical and thermal hyperalgesia and increased paw edema (all P<0.05 vs. lean controls) in response to carrageenan. Significant changes in levels of resistin C reactive protein, TGFβ and visfatin (but not IL1β or TNFβ) were detected in obese rat spinal cord. The increased pain and inflammation in obese rats fits with the hypothesis that obesity is a chronic low-grade inflammatory disorder, producing a state where responses to inflammatory challenge are potentiated. The altered adipokine profile observed suggests adipokines may be useful biomarkers for monitoring initiation and progression of pain with obesity, or even be involved in the development of co-morbid pain in obese individuals.

Real-time PCR analysis of adipokine mRNA in spinal cord from HFD fed rats and control rats (n=6/group). Target mRNA are expressed relative to housekeeping gene cyclophilin. Data are expressed as mean ± SEM. * p <0.05.

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Biography

Sharron Dolan is a Senior Lecturer in Pharmacology and Assistant Head of the Department of Life Sciences at Glasgow Caledonian University. After completing her PhD in Neuropharmacology at University of Stirling with Dr Peter Cahusac, she took up a BBSRC funded Post-Doctoral position with Professor Andrea Nolan at Glasgow University’s Veterinary School, working to characterize the spinal mechanisms of inflammatory pain and analgesia. She took up a tenured post as lecturer at GCU in 2004. Her research over the past 20 years has focused on understanding the neuronal mechanisms of pain and inflammation and more recently focused on mechanisms underlying co-morbid pain with diabetes and obesity.

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