

IMPACT OF RECTAL DISTENSION ON PRIMARY BLADDER MECHANOSENSITIVE AFFERENT NERVE ACTIVITIES IN THE RAT: MECHANISM OF A PERIPHERAL AFFERENT ORIGIN OF PELVIC ORGAN CROSS-SENSITIZATION

Hypothesis / aims of study

Cross-sensitization among pelvic structures may contribute to chronic pelvic pain of unknown etiology and involves convergent neural pathways of noxious stimulus transmission from two or more organs (1). It has been speculated that convergence of sensory information from discrete pelvic structures may occur at different levels of nervous system hierarchy; locally (peripherally) via axon collaterals, in the spinal cord (dorsal root reflexes), and/or in the central nervous system (1, 2). In the present study, we aimed to directly test the hypothesis that rectal distension (RD) can alter the mechanosensitive properties of urinary bladder afferents at a peripheral afferent nerve level by using established techniques of measurement of primary bladder single unit afferent activities (SAAs) and artificial stable RD.

Study design, materials and methods

Female Sprague-Dawley rats were used. Under urethane anesthesia (1.5 g/kg intraperitoneally), a catheter and a balloon-attached catheter were inserted into the bladder and the rectum, respectively (Figure 1). Both L6 dorsal roots were cut and fine filaments were dissected from the left L6 dorsal roots and placed across a bipolar electrode for monitoring SAAs. SAAs of the nerves primarily originating from the bladder were identified by electrical stimulation of the pelvic nerve and by bladder distension. Nerves with conduction velocities (CV) more than 2.5 m/sec were designated as A δ -fibers and those with CV less than 2.5 m/sec as C-fibers. RD was applied by filling the balloon with saline. First, to exclude the direct mechanical effect of RD on bladder pressure, intravesical pressure was assessed under empty and saline-filled (at 30 cmH₂O of intravesical pressure) states of the bladder with application of RD. To evaluate the effect of continuous RD on bladder SAAs during bladder filling, first, SAAs were recorded during cystometry with filling saline into the bladder at a rate of 0.08 ml/min until an intravesical pressure of 30 cmH₂O without RD as the baseline, and then the SAA measurement repeated 3 times under different RD conditions at the rectal balloon pressure of 20, 40, and 60 mmHg. SAAs were averaged at 5 cmH₂O interval of pressure or at equally divide into five parts of volume in the filling phase, and the average unitary activity was totalled as an integrated activity during the whole filling phase, based on pressure and volume, respectively. For comparison of these integrated activities, the values are expressed as a percentage of the base-line activity.

Results

Nineteen rats were used in this study. No significant direct mechanical influence of RD on the bladder pressure either empty or full bladder state was confirmed in 4 rats (data not shown). Thirty single unit SAAs (A δ -fibers: n=11, CV=5.20 \pm 0.71 m/sec., C-fibers: n=19, CV=1.28 \pm 0.12 m/sec.) were isolated from 15 rats. RD facilitated bladder SAAs of both A δ - and C-fibers during bladder filling in a strength-dependent manner (Figures 2 and 3).

Interpretation of results

The present results indicate that SAAs of both A δ - and C-fibers conveying sensation of bladder filling can be enhanced by rectal distension in the rat through a mechanism other than direct mechanical influence of RD on the bladder. Peripheral convergent mechanisms at an afferent nerve level may be involved in this cross-sensitization, since no reflex arc through the L6 dorsal roots was preserved in the present experimental set-up. Such cross-organ afferent pathways may originate, at least partly, directly from the rectum via antidromic axon reflexes from a single dichotomizing primary afferent supplying two structures (prespinal convergence).

Concluding message

The present study demonstrates that rectal distension can increase bladder mechanosensitive afferent activities of both A δ - and C-fibers as a consequence of peripheral neural-cross-talk.

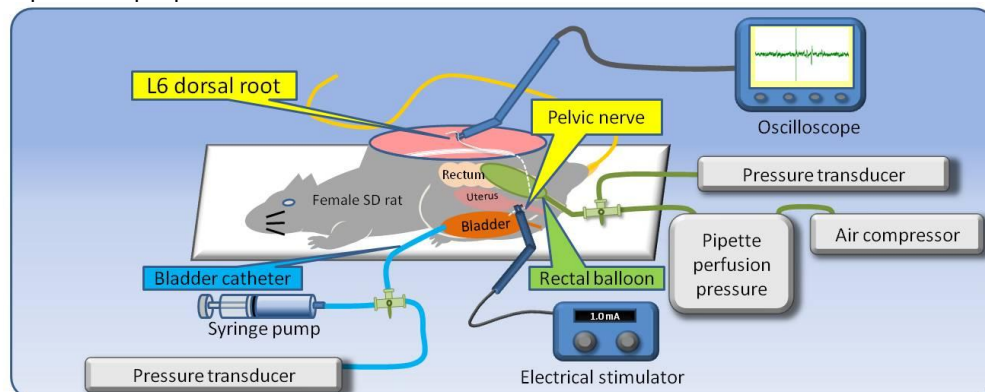


Figure 1 Experimental setup

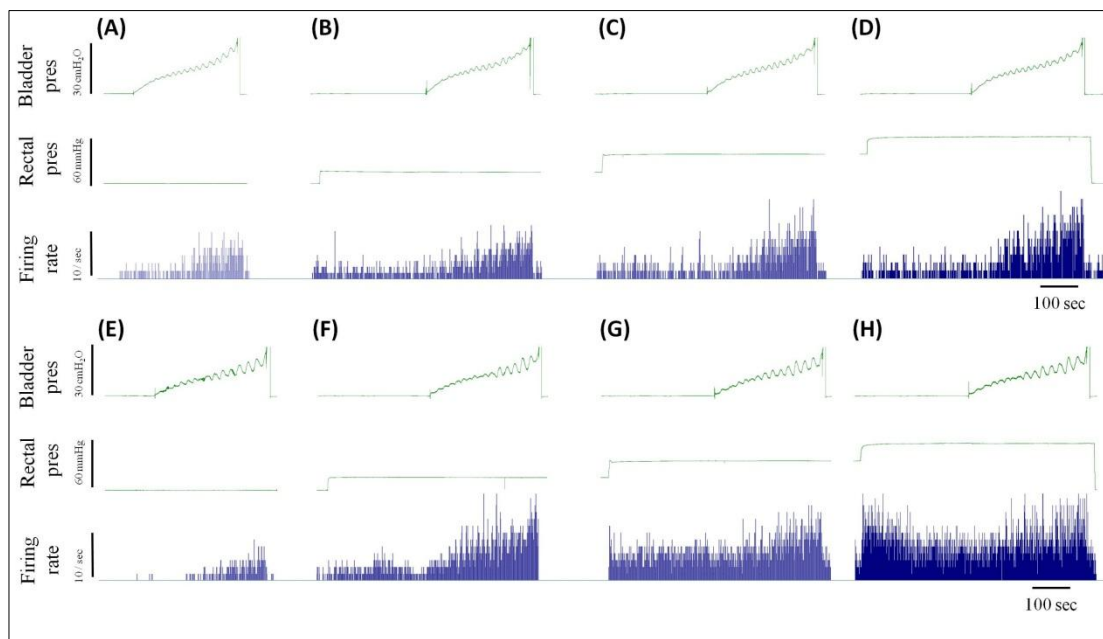


Figure 2 Typical tracings of the bladder pressure (upper traces), the rectal pressure (middle traces), and bladder single unit afferent activities (SAAs) (lower traces) of A δ -fiber (A, B, C, D), and C-fiber (E, F, G, H) during bladder filling with saline. Bladder SAAs were measured 4 times repeatedly during bladder filling under different strengths [baseline, 0 mmHg (A, E), 20 mmHg (B, F), 40 mmHg (C, G), and 60 mmHg (D, H)] of rectal distension (RD).

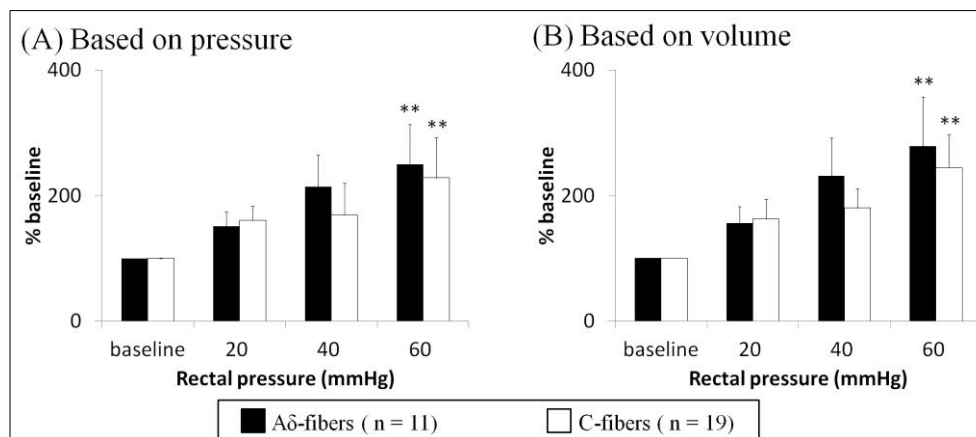


Figure 3 Integrated responses of the afferent activities of A δ -fiber and C-fiber, during the whole filling phase, based on pressure (A) and volume (B). The values are expressed as a percentage of baseline activity (mean \pm SEM). SAAs of the bladder A δ -fibers and C-fibers during bladder filling were increased with rectal distension (RD) in a strength-dependent manner (** P < 0.01, one- and two-way ANOVA followed by Tukey's test).

References

1. Malykhina AP. Neural mechanisms of pelvic organ cross-sensitization. *Neuroscience*. 2007;149(3):660-72.
2. Ustinova EE, Fraser MO, Pezzone. Cross-talk and sensitization of bladder afferent nerves. *MA. Neurourol Urodyn*. 2010;29(1):77-81.

Disclosures

Funding: None **Clinical Trial:** No **Subjects:** ANIMAL **Species:** Rat **Ethics Committee:** The Animal Ethics Committee of the University of Antwerp, Faculty of Medicine