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# THE POTENTIAL ROLE OF UROTHELIAL SENSORY PROTEINS IN PATIENTS WITH BLADDER OUTLET OBSTRUCTION AND DIFFERENT BLADDER DYSFUNCTIONS

### Hypothesis / aims of study

Bladder outlet obstruction (BOO) may be induced by specific functional and anatomic causes. The resulting obstruction can induce significant alterations in the morphology and physiology of urothelium and detrusor muscles, which manifest with various bladder dysfunctions. The underlying mechanisms responsible for the bladder dysfunction in BOO remain poorly understood. The purpose of this study is to investigate the role of sensory protein expressions in patients with BOO.

#### Study design, materials and methods

We prospectively enrolled patients who had undergone an urodynamic study for LUTS. Based on their results, it was categorized into 4 groups: Control (no BOO), BOO with detrusor overactivity (DO), BOO with detrusor underactivity (DU) and BOO with hypersensitive bladder (HSB). Bladder biopsies taken from each patient were examined using western blotting in search of sensory receptors included eNOS, iNOS, P2X3, M2, M3 and  $\beta$ 3. The results were compared against with urodynamic parameters.

#### Results

A total of 44 men were enrolled in this study. There were 34 patients presented with BOO (DO: 12, DU: 11 and HSB: 11). The expression of sensory proteins by western blotting was demonstrated in Fig.1. The distribution of M3 is significantly reduced in patients with BOO especially in those with HSB ( $0.685\pm0.333$ ) (p<0.000). The  $\beta$  3 signal in BOO with DU is significantly increased ( $1.353\pm0.499$ ) and lowest in BOO with HSB group (p<0.000) (Table 1). Higher number of  $\beta$ 3 expression is associated with delayed FSF and lower Qmax. A greater number in iNOS and eNOS is detected along with higher Pdet and larger PVR, respectively. P2X3 is associated with deferred FS whereas M2 is corresponding to increased Pdet.

#### Interpretation of results

LUTS secondary to BOO are associated with alteration in sensory receptors. Over expression or under expression of these sensory proteins could affect urodynamic parameters and might contribute to the pathophysiology of bladder dysfunction in BOO.

#### Concluding message

Alteration of urothelial sensory receptor expressions secondary to BOO may be responsible for the different bladder dysfunction.

#### Table 1. Expression of sensory proteins in control, BOO+HSB, BOO+DO and BOO+DU

	Control (N=10)	BOO+HSB (N=11)	BOO+DO (N=12)	BOO+DU (N=11)	P-value
eNOS	0.094±0.088	0.126±0.144	0.113±0.061	0.076±0.058	0.628
P2X3	0.097±0.109	0.287±0.175	0.211±0.106	0.275±0.299	0.103
M2	0.405±0.303	1.113±1.667	1.036±0.521	0.541±0.468	0.207
M3	1.593±0.708	0.685±0.333	0.721±0.298	1.025±0.315	0.000
iNOS	0.258±0.325	0.268±0.540	0.174±0.182	0.062±0.039	0.474
β3	0.878±0.584	0.820±0.316	0.904±0.225	1.353±0.499	0.024

Table. 2 Correlation between urodynamic parameters with sensory receptors expression

		β3	iNOS	eNOS	P2X3	M2	M3
FSF	Pearson	.616**	089	.027	031	.060	047
	р	.000	.623	.878	.864	.737	.791
FS	Pearson	.286	.000	002	.432*	188	.323
	р	.106	.999	.993	.011	.287	.063
Pdet	Pearson	131	.442*	.264	093	.499**	267
	р	.492	.015	.152	.620	.004	.146
Qmax	Pearson	380*	.054	.252	103	.111	347*
	р	.029	.765	.151	.563	.531	.044
Vol	Pearson	.288	114	098	237	205	.212
	р	.110	.535	.587	.185	.253	.237
PVR	Pearson	280	.186	.540**	032	.513**	029
	р	.114	.301	.001	.856	.002	.870

			Normal	BOO+DO	BOO+DU	BOO+HSB
eNOS	133 KDa	170KDa 130KDa	And part		-	-
M3	66 KDa	70KDa -				
M2	52 KDa	70KDa	-	==		==
P2X3	44 KDa	55KDa 40KDa	-		-	
GAPDH	37 KDa	35KDa	-			
iNOS	130 KDa		-		-	-
β3	45 KDa				-	

Fig.1. Expression of sensory proteins eNOS, iNOS, M3, M2, P2X3 and  $\beta$ 3 in BOO patients with different bladder dysfunctions.

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