

How Non Pharmacological Procedures Can Help to Reduce Drugs Need and Healthcare Costs in OVB -Urgency Micturition

Research article

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Abstract

Aim of this work is to verify the efficacy of the KEGEL procedure used in the micturition urge dysfunction and the contribute that this can provide in reducing-avoiding pharmacological therapy.

- i. This approach can be useful to reduce healthcare costs (direct and indirect) and in order to increase the patients quality of life.
- ii. The social implications are relevant.
- iii. The complexity in physiology and pathology in this condition need an adequate treatment:
- iv. Not only pharmacological approach or physiotherapeutic but also psychological and of lifestyle modification.
- v. This multidisciplinary approach can produce global benefit.
- vi. The more severe cases are not considered in this work because required treatment like Botulin toxin injection or surgery procedure.

Keywords: Overactive Bladder, Anatomy, Physiology, Nervous Control, Muscle, Pelvic floor Dysfunctions, Non Pharmacological Actions, Kegel Exercise, Bladder Training, Psychology, Pharmacology, Drug Need, Avoided Costs, Quality of life Urology, Micturition Urgency, Caffeine, Pharmacoeconomy

Introduction

Overactive bladder is a condition with specific symptoms like micturition urgency, increase of micturition frequency, incontinence, nocturia.

Other imply bladder involuntary detrusor contractions.

This condition imply great quality of life reduction (Figure 1)

Various conditions are involved like diabetes, reduced renal function, bladder cancers, IPB, increase bladder wall sensibility, weakened of pelvic musculature, pregnancy – birth, neurological condition like Parkinson disease, ICTUS, Multiple sclerosis, CNS or spinal cord or nerve damages, urological surgery or radiotherapy.



According Tammyr is Helena Rebecchi Silveira et al:

“It is now recognized that DM causes tissue damage by altering redox signaling in target organs. NADPH oxidase, whose sole function is the production of reactive oxygen species, plays a pivotal role in other well-known and bothersome diabetic complications” (Figure 2).

Also the ageing is a factor involved

Other iatrogenic factors can be diuretics, high amount of coffee, alcoholic drinks, wine, IVU, high body weight, Menopausa with low estrogen level.

The diagnosis: general physicians or urologic evaluation, anamnesis, imaging, laboratory test

PSA test in men, fluxometry, cysto- retrograde uretrography, cystoscopy

Urine analysis and uroculture. to exclude infections, blood traces

Neurological exam to detect sensorial or anomalous reflex

Test urodynamics: the bladder functionality and its capacity to fill in right way. If not correctly empty during the urination the residual urine can produce similar symptoms like OVB.

In order to measure the amount of urine not emitted the physician can carry out a bladder ecographic analysis or to put a catheter through uretra in order to drain and measure the post. Micturitional Residue still in bladder.

Urofloussometry, functional test to measure the volume and velocity of urinary flux.

Cistometry: to search involuntary contraction or if the bladder is not able to store urine correctly.

Uretrocistoscopia: to exclude cancers and kidney stones.

Various are the possibilities of Intervention

Behavioral: Reducing body weight, normalize fluid intake, avoid dietetic irritants like spices, alcohol, coffee, artificial sweeteners, etc.

Stop smoking, because irritant bladder and due to the fact that cough can produce urine loss.

Pelvic floor rehabilitation, Kegel exercise, double micturition, bladder training.

Other imply intermittent urologic catheterism.

Use of absorbent pads for social life, good diabetes control.

Related drugs: anticholinergic for the detrusor hyperactivity (contraindicated if IPB, Glaucoma, Myasthenia gravis), agonists of beta 3 adrenoceptor (Mirabegron), botulinic toxin A injection in refractory cases (MS and spinal cord lesions).

Sacral neuromodulation with pacemaker, surgery in particular patients,

The specialist involved: Urologist, Gynecologist, Neurologists, Physiotherapist, Geriatrics, General Physicians, Nurse, Pharmacist, Psychologist, Nutrition Specialist, Radiologists and laboratory medicine.

Systematic Review 10 November 2022

Healthcare and economic burden of anticholinergic use in adults with overactive bladder: a systematic literature review. Corinne Duperrouzel, Coby Martin, Ari Mendell, Megan Bourque, Adam Carrera, Alicia Mack and Jeffrey Nesheim.

“Overactive bladder is associated with increased incidence of various comorbidities, including depression, anxiety, falls and reduced health-related quality of life. OAB is also associated with an high economic burden EB, including medical costs, lost productivity and healthcare resource utilization; average healthcare costs (including inpatient, emergency room, outpatient, pharmacy and other costs) for patients with OAB have been estimated about at \$3003 per patient per month vs \$1123 per patient per month for a matched control group without OAB”

In this work are analyzed the Pelvic floor dysfunction in men especially related to the micturition urgency.

This situation implies heavy social implications related to the high frequency in the micturition with reduction of the quality of life of the patients. In the physiology of this function various muscles are involved and related nervous system: sympathetic, parasympathetic and also somatic-voluntary. Micturition is thus under cortical control, mediated by the spinal reflex arc which inhibits the pontine center until it is deemed appropriate to void. The motor cortex controls the voluntary muscle of the external urethral sphincter.

“The pelvic floor PF plays an integral part in lower urinary tract storage and evacuation. Normal urine storage necessitates that continence be maintained with normal urethral closure and with urethral support. The endopelvic fascia of the anterior vaginal wall, its connections to the arcus tendineus fascia pelvis and the medial portion of the levator ani muscles must remain intact to provide a normal urethral support. Normal pelvic floor function PFF is required for urine storage. Normal urine evacuation involves a series of coordinated events, the first of which involves complete relaxation of the external urethral sphincter and levator ani muscles. Acquired dysfunction of these muscles will initially result in a sensory urgency and detrusor overactivity; With time the acquired voiding dysfunction can result in intermittent urine flow and incomplete bladder emptying, progressing to urinary retention in the severe cases.”

“The urethral sphincter US is a muscular structure that regulates the outflow of urine from the bladder into the urethra. There are 2 urethral sphincters, the external and internal urethral sphincters. When these muscles contract, the urethra narrows, and urination stops or slows. The urethral sphincter US is critical for the maintenance of urinary continence. Urinary incontinence is often associated with the pathology of the urethral sphincter and is a very common problem worldwide.

The urethral sphincter muscles are located within the deep perineal pouch (Figures 3-8).”

“The act of micturition is an autonomic reflex at the level of the spinal cord SC. This reflex also helps to complete micturition when the act is voluntarily initiated, or when it follows a period of inhibition by the brain, by relaxing external sphincter. The control of this process is mediated via afferent signals AS from stretch and volume receptors in the bladder as well as from the muscles of the pelvic floor the vagina/penis, and the rectum, which informs the brain about the extent of filling, initiating several spinal reflexes. These serve to inhibit micturition until filling is complete while activating the voluntary external urethral sphincter EUS via the pudendal nerve. At the same time, detrusor activity is inhibited and the internal urethral sphincter is stimulated via sympathetic activity. Impulses from the filling bladder are carried to the spinal cord SC via the pelvic and hypogastric nerves, whereas the pudendal and hypogastric nerves carry impulses from the neck of the bladder and the urethra.”

Micturition is a SPINAL REFLEX facilitated or inhibited by higher brain centre, subjected to voluntary/involuntary facilitation (Figure 9).



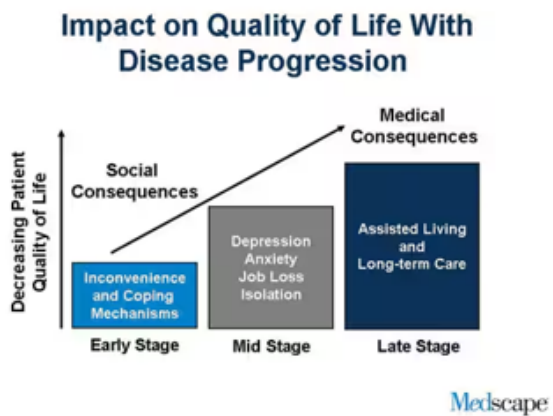


Figure 1: From Diane K. Newman, Mescap Urology 2024.

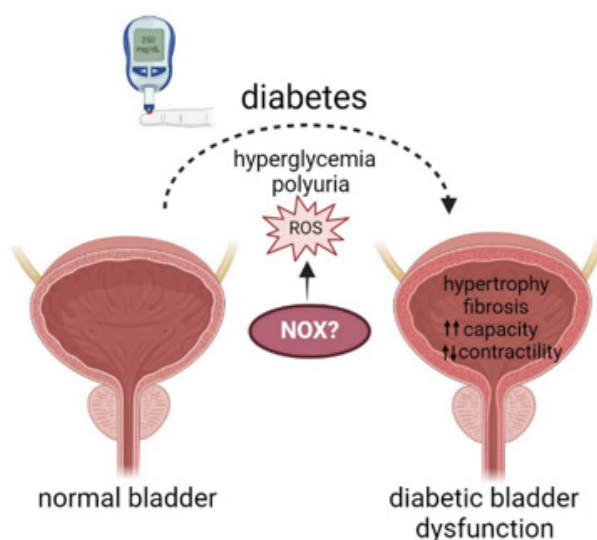


Figure 2: From <https://doi.org/10.3390/antiox13101155>.

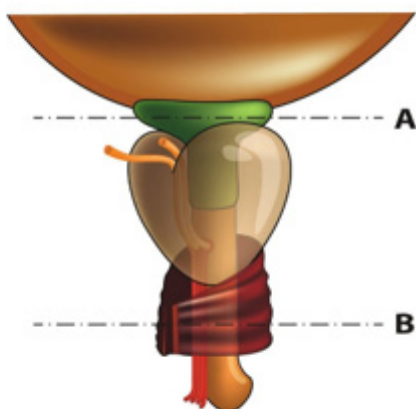


Figure 3: Male urethral sphincter image showing: A) the internal urethral sphincter. B) the external urethral sphincter.



Figure 4: From <https://cioffredi.com/belly-care-overview/>.

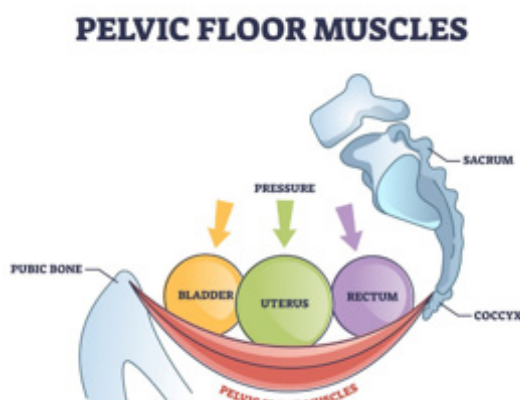


Figure 5: From the Pelvic clinic.

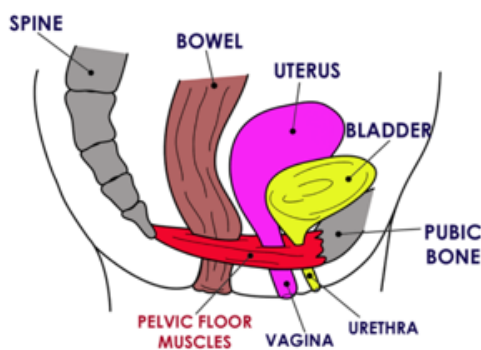


Figure 6: <https://norfolkandwaveneycommunityhealth.nhs.uk/msk/self-help/women-s-men-s-health-pregnancy/the-pelvic-floor/>.

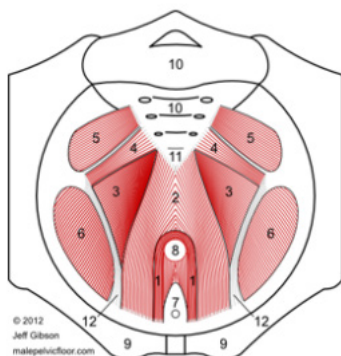


Figure 7: The pelvic diaphragm viewed from above. From <http://www.malepelvicfloor.com/anatomy.html>

1. Puborectalis muscle, 2. Pubococcygeus muscle, 3. Iliococcygeus muscle, 4. Coccygeus muscle Other muscles, 5. Piriformis, 6. Obturator Internus, Other landmarks, 7. Urethral opening, 8. Anal opening, 9. The paired pubic bones in the front of the pelvis, 10. The sacrum, 11. Coccyx (tail bone), 12. Tendinous arch



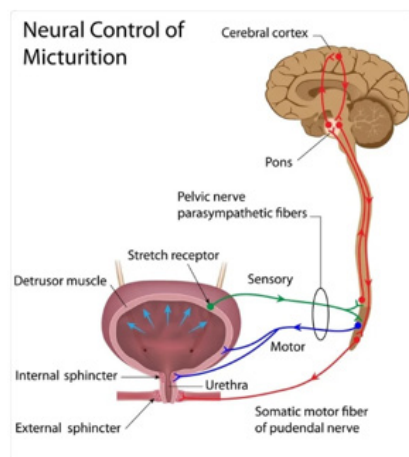


Figure 8: From <https://www.news-medical.net/health/Micturition-Reflex-Neural-Control-of-Urination.aspx>.

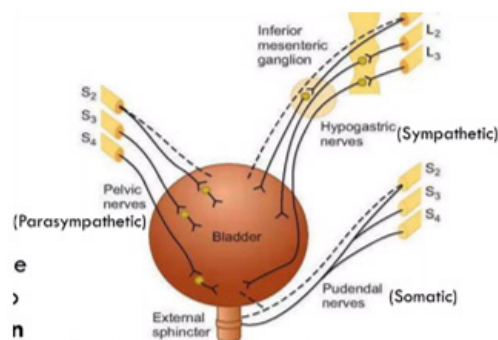


FIGURE 38–20 Innervation of the bladder. Dashed lines indicate sensory nerves. Parasympathetic innervation is shown at the left, sympathetic at the upper right, and somatic at the lower right.

Figure 9: The parasympathetic nerve (pelvic) are motor to the detrusor and inhibitory to the internal urethral sphincter. The somatic nerve (pudendal) involved in the voluntary control of the external urethral sphincter.

Sensory nerve: involved in cortical sensation

The detrusor muscle is mainly responsible for emptying the bladder during micturition. The internal urethral sphincter is a smooth muscle, the external sphincter is a skeletal muscle. So it is possible to see in the nervous control of micturition various centers:

- i. Cortical center involved: inhibitory to the pontine center
- ii. Brain stem center (pons) facilitatory micturition
- iii. Sacral spinal cord - parasympathetic (reflex evacuation)

The cortical center controls the pontine in order to have a socially acceptable micturition frequency.

The Pontine is a Coordination Center

Involved in synchronization for a complete evacuation.

The spinal center: involved in the reflex. An inefficient contraction of the detrusor produces incomplete evacuation. By Dr. Liji Thomas, MD

“Micturition is thus under cortical control (CC) as well as mediated by the spinal reflex arc, which inhibits the pontine center until it is deemed appropriate to void. In addition, the motor cortex controls the voluntary muscle of the external urethral sphincter (EUS).” Related lifestyle and medication involved: Pelvic Floor Dysfunction. Grimes; Michael Stratton.

June 26, 2023.

“Therapeutic interventions for patients with pelvic floor dysfunction (PF) should be tailored to their specific needs. A multidisciplinary approach is many times necessary. Patients with a history of sexual, physical, or emotional abuse should have the information relayed to the entire treatment team to facilitate modifications of therapy to accommodate the patient’s needs.

From medline website: How to do Kegel Exercises

Once you know what the movement feels like, do Kegel exercises 3 times a day:

- i. “Make sure your bladder is empty, then sit or lie down.
- ii. Tighten your pelvic floor muscles. Hold tight and count 3 to 5 seconds.
- iii. Relax the muscles and count 3 to 5 seconds.
- iv. Repeat 10 times, 3 times a day (morning, afternoon, and night).
- v. Breathe deeply and relax your body when you are doing these exercises.
- vi. Kegel exercises can be done any time you are sitting or lying down. You can do them when you are eating, sitting at your desk, driving, and when you are resting or watching television”.

Lifestyle Modifications

Diet: avoidance of alcohol, caffeine (cola, tea, coffee), acidic foods/



beverages, including citrus and tomatoes, concentrated sugar, artificial sweeteners, like aspartame, spicy foods and cigarettes for urinary frequency and incontinence. These changes have overlapping benefits for anorectal symptoms, including incontinence.

Weight loss

a 3% to 5% weight reduction can decrease urinary incontinence episodes by about 50%.

Pelvic floor exercises (Kegel): to strengthen the pelvic floor PF.

Core exercises: to strengthen the pelvic floor and support.

Medications

Topical vaginal estrogen for overactive bladder, vaginal thinning and dyspareunia.

Anticholinergics (fesoterodine, tolterodine) for OVB.

Beta3 agonists (mirabegron) for OVB.”

Material and Methods

With an observational point of view various relevant literature is reported related the scope of this work.

The reported figure helps to better explain the concepts. An experimental project is submitted.

After evaluation of all this a global conclusion is provided.

Results

“In 1948, Kegel exercises were first described by A. Kegel for pelvic floor muscle PFM strengthening. The perineometer, also called the vaginal manometer, has been designed to record the contraction strength of pelvic floor muscles and can be used to guide the participants to conduct the exercises correctly. Dr. Kegel’s study showed that the exercises could help to prevent cystocele, rectocele or urinary stress incontinence [1].”

“We conducted a prospective cohort work of women with urinary urgency and frequency symptoms. Participants underwent PFPT once or twice per week for 10 weeks. Symptom improvement was assessed using validated questionnaires, voiding diaries, subjective measures. 57 participants enrolled; 21 (36.8%) withdrew or completed less than 5 weeks of PFPT. Thirty-one (54.4%) of the remaining 36 participants completed 10 weeks of PFPT. The mean age of the study group (n = 36) was 48.9 ± 15.0 years. The primary diagnoses were overactive bladder OB syndrome (n = 24, 66.7%) and painful bladder syndrome PBS (n = 12, 33.3%). Women attended a median of 14.0 (interquartile range [IQR], 8.0-16.0) PFPT visits over a median of 11.9 weeks (IQR, 10.0-18.1). At baseline, the median Pelvic Floor Distress PFD Inventory-Short Form 20 score was 79.2 (IQR, 53.1-122.9), and decreased to 50.0 (IQR, 25.0-88.5; P < 0.001) after PFPT; the urinary and prolapse symptom subscales both decreased significantly. Participants reported a decrease from a median of 10.0 voids per day to 8.0 (P < 0.001).

On the Patient Global Impression of Improvement, 62.5% of women reported : they were "much better" or "very much better."

The PFPT with myofascial release techniques improves urinary symptoms while avoiding the medications and more invasive therapies. The high dropout rates suggest that the motivation or logistic factors may play a significant role in the utilization and success of this treatment option [2].” “Pelvic floor muscle training consists in the repetition of one or more sets of voluntary contractions of the pelvic muscles. By building muscles volume, PFMT elevates the pelvic floor PF and the pelvic organs, closes the levator hiatus, reduces pubovisceral length and elevates the resting position of the bladder [3].”

Related the pharmacology involved

“Urinary antispasmodic agents UAA are medications prescribed to treat bladder overactivity which can cause urinary urgency and incontinence. Urinary antispasmodic agents increase bladder capacity, delay the desire to void, decrease the frequency and urgency to urinate, and reduce pain caused by difficulty in the urination.

UAA relax the detrusor smooth muscle which contracts to release urine and prevent spasms of the muscles around the urinary tract. Urinary antispasmodic agents work in the following ways to decrease uninhibited the bladder contractions:

Block acetylcholine (ach) from stimulating muscarinic receptors, which are protein molecules on smooth muscle cells in the bladder lining. ACH is a natural chemical (neurotransmitter) that nerve endings secrete to make the muscles contract. Block the activity phosphodiesterase, an enzyme that breaks down signaling molecules known as cyclic adenosine monophosphate. An increase in cAMP concentration results in the relaxation of smooth muscles.

Urinary antispasmodic agents include:

oxibutinin oral, topical, transdermal, flavoxate

Other

Tolterodin Detrusitol antimuscarinic

Genurin flavoxate

Anticholinergic-parasympatholytic action reduces the tonus of smooth muscle in the bladder, Solifenacin antagonizes the M2 and M3 muscarinic receptors in the bladder to treat an overactive bladder.

Tropio clorure: antimuscarinic.

“OVB is a symptom complex consisting of bothersome storage urinary symptoms that is highly prevalent among both sexes and has a significant impact on the quality of life. Various antimuscarinic agents and the beta-3 agonists mirabegron and vibegron BETMIGA are currently available for the treatment of OAB [4]”

“ The human bladder expresses various sympathetic and parasympathetic receptors to help regulate The micturition. Activation of the parasympathetic system will result in bladder contraction. Receptors positively affecting bladder contraction and micturition include the M2 and M3 receptor subtypes, which function via the parasympathetic nervous system PNS . These receptors work to increase intracellular calcium ICA and down-regulate cyclic-adenosine, respectively, which will increase muscle contraction. Negatively affecting micturition predominantly includes the sympathetic beta-3 adrenergic receptors. Mirabegron is a beta-3 rec. agonist which will cause detrusor muscle relaxation. Animal studies have shown that the beta-3 receptor agonists exhibit a dose-dependent detrusor relaxation DR (mediated via up-regulation of cyclic-adenosine) during the storage phase of micturition. Mirabegron can aid in the symptomatic relief of OAB and symptoms of urge urinary incontinence, urgency, and urinary frequency.

Contraindications

Previous hypersensitivity reaction to mirabegron or any excipients of tablet or oral suspension. Mirabegron use correlates with hypertension, and its contraindication includes severe un-controlled hypertension. Blood pressure BP optimization should be considered before initiating mirabegron therapy [5].”

“Pelvic floor exercises PFE are also known as Kegel, or childbirth, exercises. Kegel exercises are commonly used and have a 30% to 90% success rate in women with stress incontinence.” [6] Generally for iperreflexia are more used anticholinergic drugs instead the alfa -litics if functional obstruction.



Related Oxibutin

“Oxibutinina tablets XL is contraindicated in the patients with urinary retention, gastric retention and other severe decreased gastrointestinal motility conditions, uncontrolled narrow-angle glaucoma.”

“Tamsulosin is a α -1A and α -1D blocker, 70% of the α -1 adrenoceptors in the prostate are of the α -1A subtype. Label By blocking these adrenoceptors, smooth muscle in the prostate is relaxed, urinary flow is improved. Label The blocking of α -1D adrenoceptors relaxes the detrusor muscles of the bladder which prevents storage symptoms. The specificity of tamsulosin focuses the effects to the target area while minimizing effects in other areas.”

Buscopan HYOSCINE N-Butylbromide (antimuscarinic and anticholinergic) is a muscle relaxant prescribed by doctors to reduce spasms of the gastrointestinal tract, and/or urinary-genital tract

Silodyx Silodosin

α 1A adrenoceptor, indication for IPB symptoms. Blockade of these α 1A-adrenoceptors causes smooth muscle in these tissues to relax, thus decreasing bladder outlet resistance BOR, without affecting detrusor smooth muscle contractility. This causes an improvement of both storage (irritative) and voiding (obstructive) symptoms (Lower urinary tract symptoms, LUTS) associated with benign prostatic hyperplasia.

Other Useful Drugs Used : duloxetine (CYMBALTA), TCA, SSRI.

Relevant are also the effect played by stress, anxiety and depression of the overactive bladder.

Third line treatments for Overactive Bladder

Botulinum toxin injections into the bladder Neuromodulation – altering nerve messages or signals to the bladder using electrical stimulation by:

Posterior tibial nerve stimulation (PTNS) or Sacral neuromodulation using InterStim®.

Between other kind of factor to be taken in consideration

Related alimentary behaviour: drink at least 1000-1500 ml water die (if physician allow) day in order to dilute urine : too much concentrated irritate the bladder mucosa. It is necessary not to take in a single way by refracted during the day.

Not drink water the two hour before to go to sleep in order to reduce the need to wake up to go in bath.

Not intake irritant food like caffeine and ethil alcohol or species because irritate the bladder.

No smoke to avoid smoke (nicotin and the chemical substance inhaled can irritate the bladder).

Related psychological aspects: When there is an episode of bladder urgency it is useful also to think to other things in order to have distraction (Mindfulness or other technique).

“The objective of the work was to evaluate if a mindfulness-based stress reduction program is a viable treatment worthy of further evaluation for the treatment of urinary urge incontinence.

This was a single-arm pilot study of 7 women who participated in an 8-week mindfulness-based stress reduction program MSRP to evaluate its effectiveness in reducing urinary leakage episodes. Improvement was measured by 3-day bladder diary, OVB Symptom and Quality of Life-Short Form, Health-Related Quality of Life, and Patient Global Impression of Improvement. Mean incontinence episodes per day decreased from 4.14 (SD, 1.96; range, 2.67-7.67) at baseline to 1.23 (SD, 0.93; range, 0.33-2.67) after the treatment ($P = 0.0005$

for change). After treatment, 5 of 7 women were improved based on the Patient Global Impression of Improvement PGII, and 2 were unchanged. Significant improvements on both Health-Related Quality of Life and OVB Symptom and Quality of Life-Short Form were reported after treatment. Results were sustained at 1 year in 4 of 7 women. Two women sought treatment, and one was lost to follow-up at 12 months [7].”

“Pelvic floor muscle training involves the contraction of the puborectal, anal sphincter and external urethral muscles, inhibiting the detrusor contraction, what justify its use in the treatment of OAB symptoms.

Prospective clinical trial with 27 women with mixed urinary incontinence (MUI), with predominance of OAB symptoms and loss ≥ 2 g in the pad test. It was evaluated: pelvic floor muscles function (digital palpation and manometry); urinary symptoms (nocturia, frequency and urinary loss); degree of discomfort of OAB symptoms; and quality of life (Incontinence Quality-of-Life Questionnaire). The PFMT program consisted of 24 outpatient sessions (2x/week + home PFMT). There was a significant improvement of urinary symptoms to the pad test (5.8 ± 9.7 , $p < 0.001$), urinary loss (0.7 ± 1.1 , $p = 0.005$), nocturia (0.8 ± 0.9 , $p = 0.011$). Reduction in the degree of discomfort of urinary symptoms was observed according to OAB-V8 questionnaire (10.0 ± 7.7 , $p = 0.001$). There were significant results in PFMTs function: Oxford (3.6 ± 0.9 , $p = 0.001$), endurance (5.2 ± 1.8 , $p < 0.001$), fast (8.9 ± 1.5 , $p < 0.001$) and manometry (26.6 ± 15.8 , $p = 0.003$). Quality of life had a significant improvement in the 3 domains evaluated by I-QoL.

The PFMT without any additional guidelines improves the symptomatology, the function of PFMTs and the quality of life of women with OAB symptoms [8]. “

“Lifestyle and behavioral factors play a role in OAB as well. A body mass index (BMI) of >30 kg/m² is another risk factor for the OAB symptoms. Studies have found smoking to be risk factor for increased urgency, but this has not been consistent with other studies. High caffeine intake (>400 mg/d) has also been associated with OAB. There are also behaviors that can increase the likelihood of OAB. These include both inadequate /excessive fluid intake, caffeine, carbonated beverages, spicy food, artificial sweeteners, and alcohol. Identifying modifiable risk factors earlier on can aid in the treatment of OAB [9].”

“Caffeine may promote incontinence through its diuretic effect, particularly in individuals with underlying detrusor overactivity, in addition to the increasing muscle contraction of the bladder smooth muscle. Caffeine may affect bladder function via central micturition centers, including the medial preoptic area, ventrolateral periaqueductal gray, pontine micturition center. The results indicated that caffeine consumption increased contraction pressure CP and time significantly ($P < 0.05$, vs. control). c-Fos expression levels in the central micturition areas were increased by the caffeine administration ($P < 0.05$, vs. control), and the lowest dose of caffeine (10 mg/kg) enhanced c-Fos expression to the greatest extent ($P < 0.05$, 50 and 100 mg/kg vs. 10mg/kg).

NGF expression in the central micturition areas CMA increased with caffeine administration ($P < 0.05$, vs. control), and the lowest dose of caffeine (10 mg/kg) enhanced NGF expression to the greatest extent ($P < 0.05$, 50 and 100 mg/kg vs. 10 mg/kg).” (Figure 10 & Figure 11) [10]. “Caffeine at 4.5 mg/kg caused diuresis and decreased the threshold of sensation at filling phase, with an increase in flow rate and voided volume. So, caffeine can promote early urgency and frequency of urination. Individuals with LUTS should avoid or be cautious in consuming caffeine containing foodstuffs. “

“OVB and IBS are pathologically characterized by overactivity (irritability) of the bladder and bowel, respectively, previous studies have shown that they frequently occur concurrently [11].”



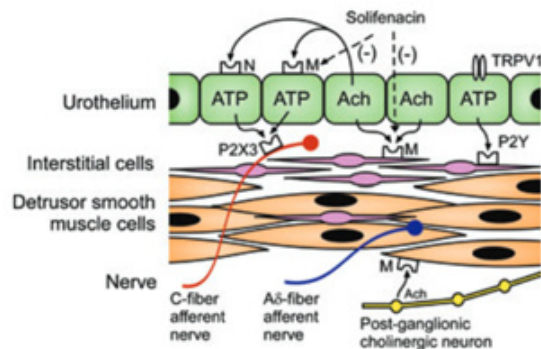


Fig. 2 Urothelium may synthesize and release multiple neurotransmitters which consequently alter the excitability of afferent nerves and affect the detrusor muscle contractility. Ach, acetylcholine; ATP, adenosine triphosphate; M, muscarinic receptors (M₂, M₃); N, nicotinic receptors; TRPV1, transient receptor potential vanilloid 1.

Figure10: Srom Pathophysiology of Overactive Bladder February 2012 Lower Urinary Tract Symptoms 4(s1) En Meng et al.

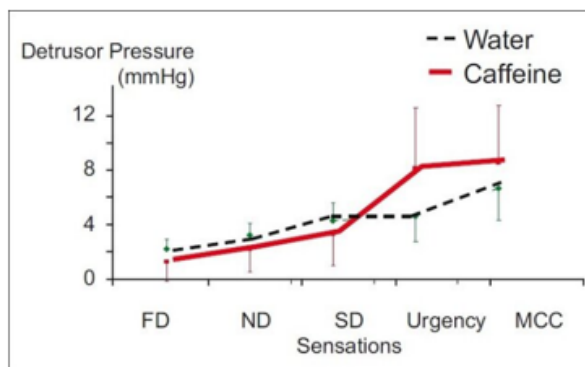


Figure 11: Detrusor pressure at filling phase (mean±SE) in mmHg after water and caffeine ingestions (FD=first desire to void, ND=normal desire to void, SD=strong desire to void, MCC=maximal cystometric capacity). From DOI: 10.4103/0974-7796.75862.

“Functional constipation is associated with OVB and its dry subtype, particularly in the younger population. In addition, this association is responsible for lower quality of life scores, especially when urinary incontinence is present [12].”

“When compared to never to <1 serving/week, women consuming ≥1 serving/day of artificially sweetened beverages had 10% greater odds of reporting mixed urinary incontinence MUI after adjustments. Amount of artificially sweetened beverage consumption was not associated with stress or urgency urinary incontinence symptoms [13].”

“Diabetic cystopathy is a well-recognized complication of DM, which usually develops in middle-aged or elderly patients with long-standing and poorly controlled disease. It may have broad spectrum clinical presentations. Patients may be asymptomatic, or have a wide variety of voiding complaints from OVB and urge incontinence to decreased bladder sensation and overflow incontinence [14].”

Between the mechanism involved in poliuria : osmotic property of glucose, oxidative stress , nerve damages (neurogenic bladder), diabetes cause excessive thirst with increase water intake. Diabete and methabolic syndrome increase body weight and this increase intraabdominal pressure . “Sodium glucose cotransporter 2 inhibitors are the newest class of drugs for the treatment of diabetes mellitus type II and off-label use for heart failure and other diagnoses has been expanding. SGLT2i use results in glucosuria which has a diuretic effect. Diuresis and glucosuria together may provoke / exacerbate OVB symptoms such as urinary frequency, urgency, and urgency incontinence. Addi-

tionally, higher rates of infection may be seen withglucosuria (Figure 12-15) [15].

When you inhale, your diaphragm rises up, which allows your pelvic floor to move upward with a gentle contraction. Breathing out contracts your pelvic floor.

From Hinge Health

“Slow, deep breathing allows your pelvic floor PF to relax and stretch, which helps the muscles remain strong and flexible. sneeze This can cause tightness and weakness and, over time, may lead to issues like urinary incontinence (or leakage), pelvic pain, prolapse, and other pelvic symptoms.”

“The Knack maneuver KM training with vaginal palpation and verbal instruction improved MVC of PFM.” [16] The Knack Manoeuvre is a term coined by Prof. J. Ashton-Miller from the University of Michigan. It describes the skill of intentionally contracting the pelvic floor muscles PFM to prevent urine loss just before and throughout a sudden rise in the intra-abdominal pressure associated with forceful actions such as a cough, or sneeze (Figure 16). “In the management of OAB symptoms, patients can be taught to control urgency by performing general relaxation techniques, including slow deep breathing exercises ,to decrease the intensity of the urgency and allow the patient to delay voiding and distraction techniques in which patients get involved in tasks that involve mental concentration (Figure 17)”(17).



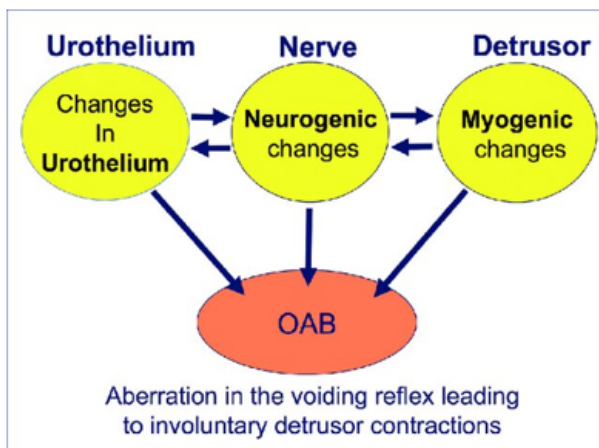


Figure 12: Mechanisms underlying overactive bladder. From DOI: 10.5489/cuaj.11181.

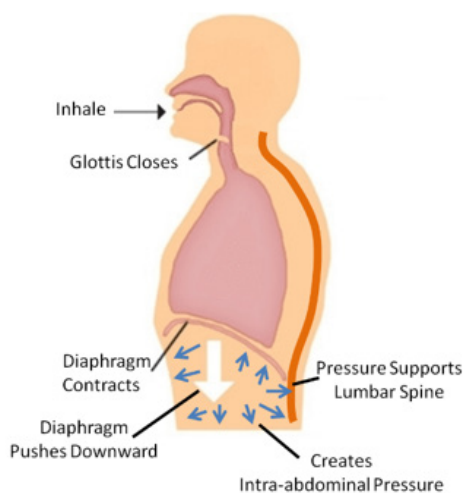


Figure 13: From Pinfree Health clinic.

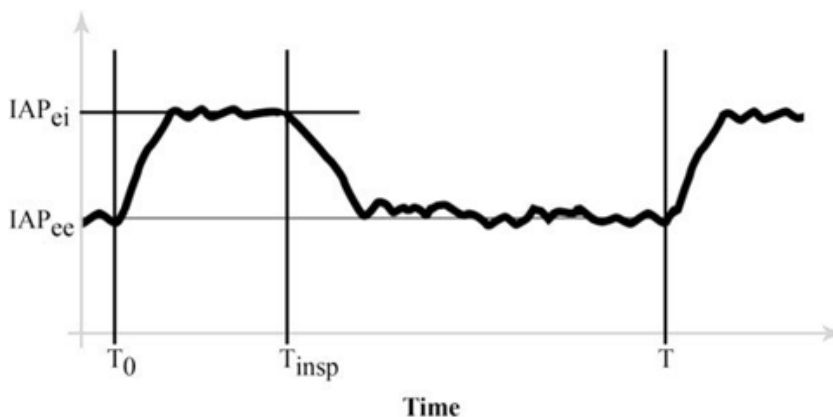


Figure 14: Effect of respiration on intra-abdominal pressure (IAP). T₀, start of inspiration; T_{insp}, inspiratory time; T, total respiration time; IA-Pee, end-expiratory IAP; IAPei, end-inspiratory IAP. From DOI: 10.1186/2110-5820-2-S1-S18.



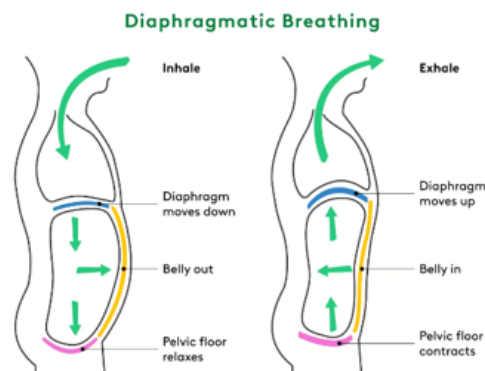


Figure 15: From Hinhe Health Breathing in stretches your pelvic floor. Breathing out contracts your pelvic floor.

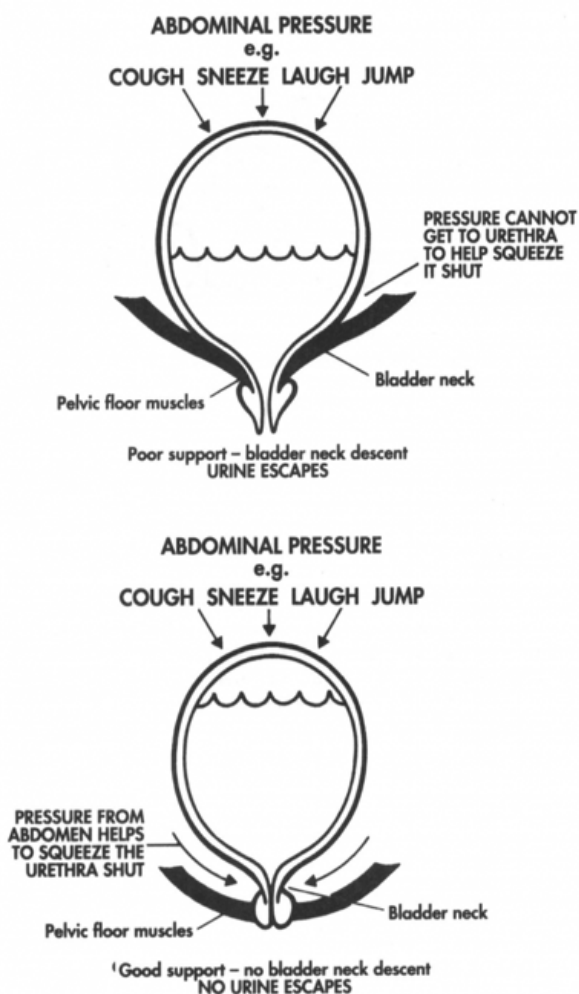


Figure 16: From <https://i.innerstrength.com.au/pelvic-floor-dynamic-bracing-knack-female>

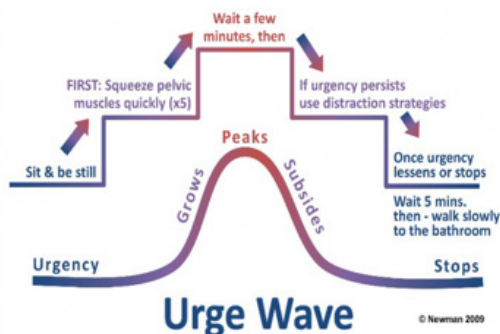


Figure 17: From Urotoday.com.



From Urotoday.com Diane Newman

“urgency follows a wave pattern; it starts, grows, peaks, and then subsides until it stops. The key to controlling bladder urgency BU is by practicing “bladder training” BT that uses methods like “urge suppression” which are calming messages to your brain to stop you peeing so often. When you feel a sudden, urgent need to pee, do not rush or run to the bathroom. Rushing will jiggle your bladder and increase the feeling of urgency and may cause bladder leaks.

Ways to Take Control of Urgency

Focus on another body sensation. Deep breathing . Sit down and take five slow, deep breaths. Think about the air moving in and out of your lungs instead of how your bladder feels. Squeeze your pelvic muscles. So when urgency comes on, sit and do a couple of gentle pelvic muscle PM squeezes, just 3 to 5 squeezes. This will calm your bladder and the urgency will go away. You can also try holding one strong squeeze of your pelvic muscles.

Distract yourself. If you get your mind off your bladder long enough, the feeling of urgency will often pass. One way is by focusing on a mental activity like counting backward from 100 by 7s.

Do a task that requires a lot of thought, play a game on your phone, write a letter.

Use self-talk or good self-statements. “I am the boss, not my bladder.”

“Compared to the lowest level of occupational PA Physical activity, the multivariate ORs for BPH for the heavy/strenuous level were 0.6 (95% CI, 0.4–0.8) at age 15–19, 0.6 (95% CI, 0.4–0.8) at age 30–39 and 0.7 (95% CI, 0.5–0.9) at age 50–59. Compared to <2 hr/week of recreational PA, the ORs for BPH for the highest level (≥ 5 hr/week) were 0.5 (95% CI, 0.4–0.7) at the age 15–19, 0.6 (95% CI, 0.5–0.8) at age 30–39, and 0.7 (95% CI, 0.5–0.8) at age 50–59. All inverse trends in risk were significant, no heterogeneity was found by reason of BPH-diagnosis, age at diagnosis, and BMI (Figure 18) [18].

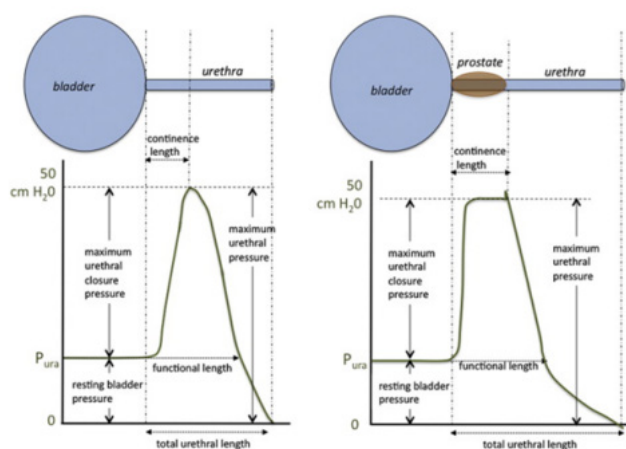


Figure 18: Urethral pressure profiles of the female (left) and male (right) lower urinary tract. From Animal Models of Lower Urinary Tract Dysfunction Rita I. Jabr, Christopher H. Fry, in Animal Models for the Study of Human Disease, 2013.

Related the pressure in uretra

“The patients were tested after 7 days and also after 14 days of yoga practice. In case 1, it was seen that the prostate size, on sonography SG readings, was reduced from 144 to 37 cc by just 1 month of yoga practice. Prevoid urine volume was 230 ml and postvoid was 180 ml, and after 1 month of yoga practice, it was 694 ml and 19 ml, respectively. Serum PSA initially was 25.24 ng/ml, while at the time of posttesting, it was 0.82 ng/ml. In case 2, the postvoid urine volume PVV was 50 ml initially. On the 14th day of yoga practice, it was nil. Serum PSA was 128.5 ng/ml at the time of pretesting, while on the 14th day, it was 16.35 ng/ml. The prostate size was 91 cc on sonography SG examination at pre yoga testing, while on 14th day of yoga practice, it was 23.6 cc. The yoga module, used in this study, is recommended to be employed to help patients of enlarged prostate [19].”

“postvoid residual urine PVRU significantly decreased in the treatment arms containing dutasteride, but not with tamsulosin alone. Table 2 displays the key RCTs with 5ARIs.” [20] “The decreases in the PVR volume were significantly ($P < 0.01$) greater with alfuzosin than with the placebo” [21].

“By intention- to-treat analysis ITT, at the end of 6-month trial, 232 (81%) of 287 patients in the Urtica dioica group reported improved LUTS compared with 43 (16%) of 271 patients in the placebo group ($P < 0.001$). Both IPSS and Qmax showed greater improvement with drug than with the placebo. The IPSS went from 19.8 down to 11.8

with Urtica dioica and from 19.2 to 17.7 with placebo ($P = 0.002$). Peak flow rates improved by 3.4 mL/s for placebo recipients and by 8.2 mL/s for treated patients ($P < 0.05$) [22].”

“The Lotus Pose also engages the muscles of the lower back and the pelvic floor [23].”

Experimental project hypothesis

In order to evaluate the efficacy of the KEGEL exercise it can be performed an practical experience:

20 patients with simpthoms of overactive bladder – urgency

They must to be divided into 2 groups

i.group that use KEGEL exercise (That can be used alone by the patients without physician intervention)

ii.group using 1 cp of ioscin as antispatic (OCT over the counter) when needed.

iii.At all it must to be subministrated a quality of life questionnaire to cover 4 hours later this two strategies

Time of observation: 30-60 days

Statistical analisis: To be verified if there is significative difference using this 2 strategy or not in order to to verify The effect of the KEGEL exercise.



Discussion

In actual health care systems is needed to avoid to spend too much money or to reduce the working days lost or to increase the quality of life (good social events participations) if alternatives not pharmacological can help. Overactive bladder and micturition urgency are universally treated with various approach because

A complexity in muscle and nervous fiber, spinal cord and upper SNC control centers are involved.

(Autonomous or voluntary), other factor can be diabetes, BPH, urethral restriction.

Pharmacological, rehabilitative or surgery approach are adopted by physicians and other healthcare professionals in relation to the severity of the pathology (urologist, physiotherapists, surgeon, geriatrics and other) The pharmacology of the substance used is clear and also the contraindication or side effect.

It is clear the effect played by bladder irritants like spices, alcohol, and coffee and the by the BMI role. The same the efficacy of the KEGEL procedure (see literature) and the easy way to be autosubministered as daily practice.

The physicians and the healthcare professionals involved in this field can help the patients.

The body weight must be controlled as well as the diabetes condition (Today there are various kind of drugs that make possible under physician control to fine regulate diabetes in some population). The right level of hydration (water drinking) and the normal functionality of the intestine (colon, rectus) as well as the right amount of physical activity are factors to be reached.

A special consideration to the stress level, anxiety or depressive status for the implication in this system and the physical activities to be performed in order to reduce symptoms.(see also literature about the specific deep breath exercise or Yoga for the prostate enlargement : in order to reduce abdominal pressure AP on the bladder). The stress condition increase a wrong way breath (more rapid) that contribute to increase the AP that act on the bladder.

Diabetes and the glycosuria acts with various mechanism in the bladder pathology. (like neuronal damage, oxidative stress, osmotic way).

To be considered the pharmacological effects played by the drugs in use for this condition (systemic or local): antimicrobials for bacterial prostatitis (ciprofloxacin and other), antimicrotics (antifungine), NSAIDS, local cortisones, 5 -alfa reductase, alfa1- antagonist, antimuscarinics, and other.

To be considered also the effect of phytotherapy on BPH (URTICA DIOICA, SERENOA R) and related Suppository of Fenolmicin P3 (a polyphenol derivate from propoli) - Boisexil (Boswellia serrata Fitosom).

The psychological aspects are relevant : the distraction technique (Think to other during and urge bladder event can be useful if added to KEGEL exercise and to the deep breath expiration). In an evolutive point of view it is interesting to observe that humans under an evolutionary point of view was in the first Evolution not bipede (in vertical position the abdominal pressure more act on bladder) and not used to sitting in rigid chair.

The ancient Lotus Yoga position is of interest. (relaxation, breath control and for pelvic floor wellbeing).

Conclusion

As conclusion of this work it is possible to say that Kegel procedure because easy to adopt also in autonomous way, added to psychological strategy and alimentary specific indication can be an interesting

approach to reduce, when possible, various healthcare costs (direct or indirect like working days lost) or to improve social life of the patients in some pelvic floor dysfunction like Micturition urgency and OVB.

Obviously the goldens standard according literature are pharmacological, rehabilitative or surgery intervention (in some conditions) prescribed by the physicians but the same Kegel procedure can be added to better achieve good results as well as some YOGA and breath exercise. It is always better if under the control a specific physicians or other healthcare professional involved. A good use of the pharmacology (drugs) may imply to observe the efficacy also of non pharmacological intervention and in special way if this approach make possible to stop or to reduce the vicious cycle of the, IPB, OVB, urgency, prostatitis, cystitis, flogosys, linfocite infiltration, edema and other improving the urethral flux or the pelvic floor function.

Conflict of interests

No.

References

1. Yi CH, Ke VC (2024) Book Kegel Exercises Treasure Island (FL): Stat Pearls Jan.
2. Sonia RA, Sybil GD, Laura ED, Jessica LM, Michele RH, et al. (2015) Pelvic Floor Physical Therapy as Primary Treatment of Pelvic Floor Disorders With Urinary Urgency and Frequency-Predominant Symptoms. *Female Pelvic Med Reconstr Surg* 21(5): 252-256.
3. Ilaria S, Simona S, Maddalena M, Flavia N, Roberto M, et al. (2019) Pelvic floor muscle training for prevention and treatment of urinary incontinence during pregnancy and after childbirth and its effect on urinary system and supportive structures assessed by objective measurement techniques). *Arch Gynecol Obstet* 299(3): 609-623.
4. Evgenyi IK, Cristiano MG, Francisco C (2021) Current pharmacotherapy of overactive bladder. *Int Braz J Urol* 47(6): 1091-1107.
5. Omar D, Ahmed EZ (2023) *Mirabegron* August 28.
6. Kris RB, Sue BC, Jessica H (2019) *Occupational Therapy with Elders (Fourth Edition) Strategies for the COTA*, Chapter 17 - Strategies to Maintain Continence in Elders.
7. Jan B, Donna C, Ingrid N (2012) Mindfulness-based stress reduction for treatment of urinary urge incontinence: a pilot study. *Female Pelvic Med Reconstr Surg* 18(1): 46-49.
8. Fátima F, Marair S, Manoel JG (1992) Rodrigo Castro. Pelvic floor muscle training for overactive bladder symptoms - A prospective study. *Rev Assoc Med Bras* 63(12): 1032-1038.
9. Marcella GWG, Alexis AD, Elizabeth JG (2016) Evaluation and management of overactive bladder: strategies for optimizing care. *Res Rep Urol* 8:113-122.
10. Young SC, Il GK, Sung EK, Lakkyong H, Mal-SS, et al. (2014) Caffeine enhances micturition through neuronal activation in micturition centers. *Mol Med Rep* 10(6): 2931-2936.
11. Seiji M, Kazumi H, Naoki W, Jyunichi H, Gaku T, et al. (2012) Relationship between overactive bladder and irritable bowel syndrome: a large-scale internet survey in Japan using the overactive bladder symptom score and Rome III criteria. *BJU Int* 111(4): 647-652.
12. Glícia EA, Eneida RD, Danielle NA, Milly QA, Natália SPM, et al. (2018) functional constipation and overactive bladder in women: a population-based study. *Arq Gastroenterol* 55Suppl 1(Suppl 1): 35-40.
13. Nancy ER, Kathleen MH, Chris AA, Yasmin MR, Aladdin HS, et al. (2022) Artificially Sweetened Beverages and Urinary Incontinence- A Secondary Analysis of the Women's Health Initiative Observational Study (WHI-OS) *Menopause* 30(3): 283-288.
14. Saeid G, Ismail L (2010) Bladder Dysfunction in Diabetes Mellitus *Front Pharmacol* 16;1: 136.
15. Patrick JM, Taylor W, Erin G, Hafsa A, Dairon DD, et al. (2024) PD12-



- 04 SGLT2 Inhibitors Associate With Overactive Bladder Symptoms. *Journal of Urology* May 01.
16. Seda YY, Nuriye Ö, M Ata T (2022) Comparing the efficacy of the Knack maneuver on pelvic floor muscle function and urinary symptoms using different teaching methods: a prospective, nonrandomized study. *Int Urogynecol J* 33(10):2895-2903.
 17. JF Wyman, KL Burgio, DK Newman (2009) Practical aspects of lifestyle modifications and behavioural interventions in the treatment of overactive bladder and urgency urinary incontinence. *Int J Clin Pract* 63(8): 1177-1191.
 18. Luigino DM, Antonella Z, Alessandra T, Maurizio M, Valerio R, et al. (2006) Lifetime occupational and recreational physical activity and risk of benign prostatic hyperplasia. *International Journal of Cancer* 118(10): 2632-2635.
 19. Pathak SD, Rajbhoj PH, Bhogal RS (2017) Effect of a yoga module on an enlarged prostate in elderly patients: Two case studies. *Yoga Mimamsa* 49(1): 34-39.
 20. Stavros G, Matthias O (2010) Current status of 5 α -reductase inhibitors in the management of lower urinary tract symptoms and BPH. *World J Urol* 28(1): 9-15.
 21. S Alan MN, Timothy BH, Christine GR, Jean PS (2001) Claus G Roehrborn, Postvoid residual urine in patients with lower urinary tract symptoms suggestive of benign prostatic hyperplasia: pooled analysis of eleven controlled studies with alfuzosin. *Adult urology* 57(3): 459-465.
 22. Mohammad RS (2005) *Urtica dioica* for treatment of benign prostatic hyperplasia: a prospective, randomized, double-blind, placebo-controlled, crossover study *J Herb Pharmacother* 5(4): 1-11.
 23. Siddappa N, Vadiraja S (2023) The Profound Influence of Padmasana (Lotus Pose) on an Individual's Life. *International Journal of Novel Research and Development* 8(9).

