

Noninvasive Method on ANS and 3D-Spectrogram Entropy Analysis

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Speaker : Liao Wen Chien M.D.PhD

Agenda





Introduction Autonomic Nervous System







External Sphincter m

Introduction What is Overactive Bladder ?

2002 ICS Terminology

Urgency, with or without urge incontience, usually with frequency and nocturia, in the absence of pathologic or metabolic factors that would explain these symptoms



ICS=International Continence Society Abrams P et al. Neuroyrol urodyn,2002,21,167-178 膀胱過動症自我檢查表 Self Screening for the Overactive Bladder

Frequent urination (> 8x per day)小便頻密 (每天八次以上)

Night-time urination (> 2x per night) 夜尿 (每晚兩次以上)

Sudden need to urinate.特發性尿意

Difficulty keeping it in.忍尿困難

May have "accidents" when laughing, coughing or sneezing.會因大笑、咳嗽、打噴嚏而漏尿 Incontinence: □ A lot □ A little 尿失禁: □ 大量 □ 少量



Hu T-W, Wanger TH, Bentkover JD et al. Estimated economic costs of overactive bladder in the United States . Urology 2003; to the estimated costs of osteoporosis and gyneacological 61:1123-1128





Urinary Diary

1.In the 1st column mark an (x)every

time you urinate into the toilet. 2.In the 2nd column, mark an (x) every time you accidentally leaked urine.

3.If an accident occurred, indicate the reason or circumstances surrounding the accident, for example, "coughed, bent over, sudden urge."

4.Under "Fluid Intake" describe the type (coffee, tea, juice, etc.)

and amount (a cup, 1 quart, etc). 5.Circle the time when you went to bed and when you got up in the

morning. 6.Record number and type of pads used.

7.Under Notes write any additional information you would like to include. For example, type and dose of medication you may be on for your urinary incontinence.

TIME	URINATE IN TOILET	LEAKING ACCIDENT	REASON FOR ACCIDENT	FLUID INTAKE TYPE AMOUNT
6 a.m.				
7 a.m.	<u>.</u>	-		3
8 a.m.				0
9 a.m.				·
10 a.m.				
11 a.m.	<u>.</u>			
12 Noon				
1 p.m.				¢
2 p.m.				
3 p.m.	2			
4 p.m.				
5 p.m.	× *			
6 p.m.				

Evaluation of OAB and the Use of Questionaire

OAB-q(2002) 4/25

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1

OAB Symptom Score(OABSS) Y,Homma(2006) (7)

3

OAB Symptom Score(OABSS) JG Blaivas(2006)

4

5

Patient Perception of Bladder Condition(PPBC) (2006)

OABSS in Taiwan Chinese (2008)

Material and Methods

ntroduction

Material Methods

The OAB group consisted of 33 OAB adult women 30–60 years, criteria (2003 International Incontinence Society), OABSS score >8. All patients :urination detailed the time and volume Medical history questionnaire

(1)medications that can disrupt the ANS function, (e.g. urinary incontinence medications, beta-receptor-specific drugs, or antidepressants),
(2)pathological conditions that may disrupt the ANS (e.g. coronary heart disease, neurological disease, diabetes and pelvic surgery, including hysterectomy).

OAB group :physical examination, ECG, urinalysis, uroflowmetry, and a pelvic ultrasound.



voiding (reflecting ANS control) was used.

500 to 1000cc water was orally consumed by each subject (both control and test groups). When patients expressed a need to urinate, they were requested to hold their urine for at least 3min, and later, to urinate in a standard urocytometry chair.

Continuous ECG recording was made while the patient was holding her urine, to 5 minutes immediately before voiding, during voiding, and then for 5 minutes post-voiding. 1000Hz ECG (P-QRS-T) wave pattern measurements were taken during urination.

The amount of urine voided and flow rate was recorded After voiding, a bladder scan was performed on each patient to determine the residual urine volume.

Material and Methods Material Methods ECG data was converted into HRV data (Matlab) Data was expressed as a mean \pm standard deviation(SD). The Mann-Whitney U test, Wilcoxon test, and Z tests were used to compare electrophysiological data values, between OAB and control subjects. (SPSS Inc, Chicago, III) software The relationship between various parameters was assessed using the Pearson correlation or Spearman rank correlation.

P values lower than 0.05 were considered significant.

Material and Methods

ntroduction





Future Work

(MatlaThe Matlab software program was used to calculate Power spectrum density(PSD) Multiscale entropy graphs 3D-spectrograms. $Z(t) = \sum_{j=1}^{n} a_j(t) e^{i2\pi \int f_j(t) dt}$

Each of these resulting data sets represents a different mathematical approach to quantify and display HRV data, and therefore ANS activity.

Each provides a specific fingerprint that distinguishes HRV between normal and OAB patients.

The neuron signal normalization entropy graphs were explained using the Hilbert Hwang Transform (HHT) equations;

$$h(\omega) = \frac{1}{T} \int_{0}^{T} H(\omega, t) dt$$

Stastatics Analysis





Dr

JEC

Micturition 3D-Spectrogram Normal VS OAB



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CLINICAL ARTICLE

A noninvasive evaluation of autonomic nervous system dysfunction in women with an overactive bladder

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ABSTRACT

Objective: To verify the hypothesis that a connection exists between overactive bladder (OAB) syndrome and a bladder-specific dysfunction of the autonomic nervous system (ANS). Method: An electrocardiogram recorded heartheat cycles from the onset of urinary urgency to 5 minutes after voiding in 33 women with an overactive bladder and 176 controls. Power spectral density (PSD) analysis allowed to quantify heart rate variability (HRV), which is in relation to ANS function. Three-dimensional spectrograms and multiscale entropy graphs were used to display HRV values. Results: The differences between patients and controls were all significant in the time and frequency domains of HRV (P<0.05), which suggests disturbances in bladderspecific ANS activity in women with OAB. Conclusion: By quantifying HRV data, PSD analysis provides a simple, noninvasive method of assessing disturbances in ANS activity and monitoring treatment in women with OAB. It can also be used to evaluate other neuronal conditions.

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popiny sive method of measuring the activity of the perve fibers that

1. Introduction

1. Ind could but	control the urge to urgest (or prigturition) and urgestion
The overactive bladder (OAB) syndrome was defined by the	A direct relationship between heart rate variability (HRV) and
International Continence Society as "urinary urgency, with or without	parasympathetic effects has been reported [11.12] including in
utinary incontinence, usually with frequency and nocturia " in the	studies focusing on women with OAB or uninary stress incontinence
absence of local infection or other nathologic changes [1]. This	[12] or children with monosymptomatic poctumal enurgis [14]
	Contracting the second se
	and when the first one can be evaluated by performing
	constrait density (ISD) and sees of HRV. The sim of the present
or the and in treatment bries an care 141. There are several theories for	study use to: (1) grantify HBV/ data my manne of DCD an always for all
the owners of (12) which prove not be multiply and the life in the life of the	study was to. (1) quantify the attained values by manyses of 2 D
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activity, whether a set of the interpretation of the particular	
which can release acetychome mappiophately, of in the balasym-	would indicate disturbances in Arts activity in women with GAB.
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	A Cremary and methods
stimulated alter the bladder has emptied, which caused the bladder to	The study was carned out from January 2004 through December
contract again before it is full; and these noers are part of the	2006 with 33 women between 30 and 60 years of age seen for OAB at
automic and ho how is that your	Couparies of the Department of Obstetrics and Gynecology
Quantifying the Events of the LEVING Steele downer the	of P Whateven it is contained. If the gematched women without
difficult with metalen oralities are assaulte based ormptore	CA Swith a Contact of the Contact of the Contact of CAB
score (OABSS), which goes from 0 to 15, is subjective and therefore	as defined by the 2003 International Incontinence Society criteria [1]
has limitations, and grodynamic investigations can be invasive and	or had an OABSS score greater than 8. All had previously provided a
are time construct of a region of the role of ective, and	urination pattern detailing the time and volume of each urination over
	at least 3 days. Further screening was the same for study patients and
 Corresponding author, Mailing address; Taiwan Adventist Hospital, No. 424, Sec2. 	controls. It included answering a medical history questionnaire
Ba Da Road, Taipei, Taiwan, Tel.: +86 2 27718151.	designed to exclude women treated with medications for urinary
E-mail address: dr.liaowc@msa.hinet.net (WC. liao).	incontinence, drugs targeting beta receptors, antidepressants, or any

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Micturition 3D-Spectrogram Normal VS OAB





Normal Micturition 3D-Spectrogram

Project





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UESUIT

Micturition MSE Normal VS OAB





Introduction 市場地位



MarketingStrategy



Publication

Journal paper

- 1 <u>Liao Wen Chien</u> and Jaw F-S. "Non-invasive Evaluation of Autonomic Nervous System Dysfunction in Female Patients with Overactive Bladder INTERNATIONAL JOURNAL OF GYNECOLOGY & OBSTETRICS Volme110,Issue1 Page12-17,July 2010 (SCI)
- 2 <u>Liao Wen Chien</u>, and Jaw FS. "Noninvasive Impedance Analysis to Measure Human Urinary Bladder Volume" The Journal of Obstetrics and Gynaecology Research Accept Nov 2010 (SCI)
- 3 <u>Liao Wen Chien</u>, and Jaw FS. "A Noninvasive Evaluation Analysis of Amniotic Fluid Embolism and Disseminated Intravascular Coagulopathy" The Journal of Maternal-Fetal & Neonatal Medicine Accepted: Dec 17 2010 (SCI)
- 4 <u>Liao Wen Chien</u> Yili Tseng, Yu-Rong Liang, Jaw F-S. "Portable Electrical Stimulator For Urinary Incontience Biomedical Engineering: Applications, Basis and Communications Vol: 20, Issue: 1(2008) pp. 61-64 (SCI)

Paper

1 <u>Liao W-C</u>, Jaw FS. Non-Invasive Impedance Measurement of Urinary Bladder Volume TUGA & TCS 2007; 12: P67

2 <u>Liao Wen Chien</u>. Evaluation on the Autonomic Function of Normal Physiologic Storage and Micturition International Conference on Preventive Medicine Mar 25 2007

3 2009專利發明人1. 廖文劍 2. 高瑀絜 3. 趙福杉 4. 林啟萬

台灣大學專利案件發明名稱:[自律神經狀態的偵測裝置、分析裝置及分析方法]

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4 <u>Liao W-C</u>, Jaw FS. Non-Invasive Impedance Measurement Human Urinary Bladder Volume The 9th TCS & TUGA 2010; 12: P5

5 <u>Liao W-C</u>, Jaw FS. Non-Invasive Evaluation of Autonomic Nervous System Dysfunction in Female Patients with Overactive BladderThe 9th TCS & TUGA 2010; 12: P4

$$\cos\frac{n2\pi t}{T} \Rightarrow \cos\omega_{0}t \quad \sin\frac{n2\pi t}{T} \Rightarrow \sin\omega_{n}t \qquad \omega_{0} = \frac{2\pi}{T} \qquad n\frac{2\pi}{T} = n\omega_{0} = \omega_{n}$$

$$n=1, n\omega_{0} = 1\omega_{0} = \omega_{1} \qquad n=2, n\omega_{0} = 2\omega_{0} = \omega_{2}$$

$$e^{j\omega_{n}t} = \cos\omega_{n}t + j\sin\omega_{n}t \qquad (e^{i\theta} = \cos\theta + i\sin\theta) \qquad (e^{-i\theta} = \cos\theta - i\sin\theta)$$

$$\cos\omega_{n}t = \frac{e^{j\omega_{n}t} + e^{-j\omega_{n}t}}{2} \qquad \sin\omega_{n}t = \frac{e^{j\omega_{n}t} - e^{-j\omega_{n}t}}{2j}$$

$$f(t) = a_{0} + \sum_{n=1}^{\infty} [a_{n}\cos\omega_{n}t + b_{n}\sin\omega_{n}t] \qquad f(t) = a_{0} + \sum_{n=1}^{\infty} \left[a_{n}\left(\frac{e^{j\omega_{n}t} + e^{-j\omega_{n}t}}{2}\right) + b_{n}\left(\frac{e^{j\omega_{n}t} - e^{-j\omega_{n}t}}{2j}\right)\right]$$

$$c_{0} = a_{0} \qquad c_{n} = \frac{a_{n} - jb_{n}}{2} \qquad c_{-n} = \frac{a_{n} + jb_{n}}{2} \qquad f(t) = C_{0} + \sum_{n=1}^{\infty} \left[C_{n}e^{j\omega_{n}t} + C_{-n}e^{-j\omega_{n}t}\right] = \sum_{n=\infty}^{\infty} C_{n}e^{j\omega_{n}t}$$

$$f(t) = \frac{1}{T}\int_{-\frac{T}{2}}^{\frac{T}{2}} f(t)e^{j\omega_{n}t}dt + C_{0}e^{-j\omega_{n}t} + C_{1}e^{-j(\omega_{1} - \omega_{n})t} + \cdots + C_{n}e^{-j(\omega_{1} - \omega_{n})t}$$

$$F(\omega) = \int_{-\infty}^{\infty} f(t)e^{-j\omega_{n}t}dt$$

Power Spectrum Density





Boltzaman

Total $u^* = u + u'$ if $u = u_r \rightarrow u = u_s \leftrightarrow$

$$\Omega^* = \Omega + \Omega' \checkmark$$

$$\frac{p(u = u_s)}{p(u = u_r)} = \frac{p(u_s)}{p(u_r)} = \frac{\Omega^*(u = u_s)}{\Omega^*(u = u_r)} = \frac{\Omega(u_s)\Omega'(u^* - u_s)}{\Omega(u_r)\Omega'(u^* - u_r)} = G \bullet R$$

$$\frac{p(u_s)}{p(u_r)} = GR \qquad G = \frac{\Omega(u_s)}{\Omega(u_r)} \qquad R = \frac{\Omega'(u^* - u_s)}{\Omega'(u^* - u_r)} \checkmark$$

$$\frac{1}{\Omega} \left(\frac{d\Omega'}{du'}\right) = \frac{1}{K_B T'} \qquad \qquad \frac{d\Omega'}{du'} = \left(\frac{1}{K_B T'}\right)\Omega' \checkmark$$

$$\Omega'(u) = const \times e^{\frac{u'}{K_B T}}$$

$$\frac{\Omega'(u^* - u_s)}{\Omega'(u^* - u_r)} = R = \frac{const \times e^{\frac{u^* - u_s}{K_B T'}}}{const \times e^{\frac{u^* - u_s}{K_B T'}}} = e^{\frac{-(u_s - u_s)}{K_B T}} = e^{\frac{-(u_s - u_s)}{K_B T}} \checkmark$$



Electrical Stimulation

Boltzman factor.

Nernst Equation∉ $\frac{C_2}{C_k} = \frac{P(2)}{P(1)} \qquad u = E_k + E_p \qquad E_p = ZeV +$ Total $u^* = u + u'$ if $u = u_r \rightarrow u = u_s \leftrightarrow$ $\Omega^* = \Omega + \Omega' +$ $\frac{p(u=u_s)}{p(u=u_r)} = \frac{p(u_s)}{p(u_r)} = \frac{\Omega^*(u=u_s)}{\Omega^*(u=u_r)} = \frac{\Omega(u_s)\Omega'(u^*-u_s)}{\Omega(u_r)\Omega'(u^*-u_r)} \frac{P_2}{P_1} = \frac{C_2}{C_1} = \frac{\Omega(2)}{\Omega(1)}e^{\frac{-(u_2-u_1)}{K_sT}} \qquad \frac{\Omega(2)}{\Omega(1)} = \frac{\alpha\Delta x\Delta y\Delta z}{\alpha\Delta x\Delta y\Delta z} = 1$ $\frac{p(u_s)}{p(u_s)} = GR \qquad G = \frac{\Omega(u_s)}{\Omega(u_s)} \qquad R = \frac{\Omega'(u^* - u_s)}{\Omega'(u^* - u_s)} \cdot u_2 - u_1 = E_k(2) - E_k(1) + Ze(V_2 - V_1)$ $\frac{d\Omega'}{du'} = \left(\frac{1}{K_B T'}\right) \Omega_{c} \frac{C_2}{C_c} = e^{\frac{-Z_2(V_2 - V_1)}{K_B T}}$ $\frac{1}{\Omega} \left(\frac{d\Omega'}{du'} \right) = \frac{1}{K_* T'}$ $\Omega'(u') = const \times e^{\frac{u}{K_s T}}$ $\ln(\frac{C_2}{C_1}) = \frac{-Ze}{K_BT}(V_2 - V_1) \Longrightarrow V_2 - V_1 = \frac{K_BT}{Ze}\ln(\frac{C_2}{C_1}) + V_2 = \frac{K_BT}{Ze}\ln(\frac{C_2}{C_1}) + \frac{K_BT}{ZE}\ln($ $\frac{\Omega'(u^* - u_s)}{\Omega'(u^* - u_r)} = R = \frac{const \times e^{\frac{u^* - u_s}{K_B T}}}{const \times e^{\frac{u^* - u_s}{K_B T}}} = e^{\frac{-(u_s - u_s)}{K_B T}} = e^{\frac{-(u_s - u_s)}{K_B T}}, \quad N_A \cdot K_B = R = 8.31451J \cdot mol^{-1} \cdot K^{-1}, \quad N_A \cdot e = F = 96485.31C \cdot mol^{-1}$ $\frac{p(u_s)}{p(u_s)} = G \bullet R = Ge^{\frac{-(u_s - u_s)}{K_s T}} = \left[\frac{\Omega(u_s)}{\Omega(u_s)}\right]e^{\frac{-(u_s - u_s)}{K_s T}} \qquad \qquad \frac{K_B T}{Ze} = \frac{RT}{ZF} \Longrightarrow V_2 - V_1 = \frac{RT}{ZF}\ln(\frac{C_1}{C_s})$

de Broglie, *Recherches sur la théorie des quanta* (Researches on the quantum theory), Thesis (Paris), 1924; L. de Broglie, *Ann. Phys.* (Paris)

de Broglie Hypothesis Lightwave=== \rightarrow ?Photon MatterWave?<===Particle (1) $E = h \upsilon$

(h:Plank's Const v:Frequency) (2) $P = \frac{h}{\lambda}$ (3) $\lambda = \frac{h}{P}$ (h:Plank's constant

=6.626x10-34Js)

Hie, Ann. Phys. (Paris)

$$V = \frac{\lambda}{T} = \lambda \upsilon \frac{1}{T} = \upsilon \upsilon = \frac{c}{\lambda}$$

$$V = \frac{\lambda}{T} = \lambda \upsilon \frac{1}{T} = \upsilon \upsilon = \frac{c}{\lambda}$$

$$E = mc^{2} \quad E = h\upsilon = h\frac{c}{\lambda}$$

$$mcc = h\frac{c}{\lambda} \quad mc = \frac{h}{\lambda}$$

$$p = \frac{h}{\lambda} \quad mc = p$$

$$\frac{h}{\lambda} = \frac{h}{\mu}$$

Quantum Theory



Heisenberg uncertain Principle2

$$\lambda_q \lambda_l = (\frac{h}{mv})(CT_l) \quad C = \frac{\lambda_l}{T_l} \quad \lambda_q \lambda_l = \frac{h}{p}(CT_l)$$

$$P\lambda_{q}(\frac{\lambda_{l}}{C}) = h(T_{l}) \qquad P(\frac{\lambda_{q}}{T_{l}})(\frac{\lambda_{l}}{C}) = h \qquad T_{l} = \frac{\lambda_{l}}{C}$$
$$\Delta v = \frac{\lambda_{q}}{T_{l}} \qquad T_{l} = \frac{\lambda_{l}}{C} \qquad P = mV$$
$$P\Delta vT_{l} = h \qquad (\therefore \quad \Delta P = m\Delta v)$$
$$\frac{Pm\Delta v}{m}T_{l} = h \qquad (\therefore \quad \Delta P = m\Delta v)$$

$$\frac{P\Delta P}{m}T_{l} = h \qquad P = mv \quad E = \frac{1}{2}mv^{2} \quad E = \frac{1m^{2}v^{2}}{2m}$$

$$E = \frac{P^2}{2m}$$
 $\Delta E = \frac{2P\Delta P}{2m}$ $\Delta E = \frac{P\Delta P}{m}$ $\Delta T = T_l$

 $\Delta E \Delta t = h$ indeterminacy relation

 $\Delta t = h$

Tower Chen et al The Heisenber Uncertainty Relation Derived By Multiplying Matter Wavelength And Lightl Wavelength Concepts of Physics Vol. III (2006)