

*Research Article***Role of Musculoskeletal Ultrasound in the Diagnosis of Entrapment Neuropathies of Median and Ulnar Nerves****Faten I. Mohamad, Shereen R. Kamel, Mona H. Mahmoud, Fatma A. Mohamed and Ahmed E. Hafez,**

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Abstract

Objective: the aim of the study was to evaluate the role of musculoskeletal ultrasound in the assessment of entrapment neuropathies of median nerve at the wrist and ulnar nerve at the elbow; and to determine the relationships of ultrasound findings with the clinical severity of carpal tunnel syndrome (CTS) and cubital tunnel syndrome (CuTS) and the electrophysiological grading scale. **Methods:** one hundred hands with CTS, twenty elbows with CuTS and one hundred twenty asymptomatic controls were assessed by nerve conduction studies and underwent ultrasonography of wrists and elbows. Data from patients and controls were compared to determine the diagnostic relations in patients with CTS and CuTS and the grade of severity. **Results:** There was a high degree of correlation between NCS of the median and ulnar nerves, clinical parameters and variable ultrasound measurements. A cut-off point of 1.0 mm² for the cross-sectional area (CSA) of the median nerve at the tunnel inlet was found to be 94% sensitive and 76% specific. Palmer bowing (PB) of the flexor retinaculum was found to have 91% sensitivity and 90% specificity at cut-off point 2.00mm, while flattening ratio (FR) at 3.71 cut-off point was less sensitive and highly specific (21% and 99% respectively). The CSA of the ulnar nerve was the most sensitive parameter and a cut-off point of 1.0 mm² behind medial epicondyle was found to be 100% sensitive and 87% specific. The ulnar nerve ratios (UNR) had a diagnostic accuracy of 90% with 80% specificity. **Conclusion:** Ultrasonographic measurements of median and ulnar nerves CSAs, FR, PB and UNR have a comparable diagnostic value as a non-invasive and an alternative modality for the evaluation of CTS and CuTS.

Keywords: Ultrasonography, nerve conduction, entrapment neuropathies.**Introduction**

Carpal Tunnel Syndrome (CTS) is the commonest neuropathy of the upper extremities and accounts for 90% of all entrapment neuropathies⁽¹⁾. The Cubital tunnel syndrome (CuTS) is the second most common compressive neuropathy of the upper limb following carpal tunnel syndrome⁽²⁾ and is the most common site for entrapment for the ulnar nerve⁽³⁾. Neuromuscular ultrasound (NMUS) has been introduced into electrodiagnostic laboratories as a complement to nerve conduction studies and electromyography for the diagnosis of a variety of nerve and muscle conditions⁽⁴⁾.

Characteristic symptoms and signs, and electrophysiological studies are the cornerstones in the diagnosis of CTS^(5,6).

False negative cases can be seen in about 10-20% of patients⁽⁷⁾.

The electrophysiological studies usually show the level of the lesion, but do not provide anatomical information about the nerve or its surroundings^(8,9). In the last few years, NMUS; being inexpensive and non-invasive imaging modality; has been shown to be useful diagnostic tools in CTS, providing information on the median nerve and surrounding structures^(10,11).

The aim of the current study was to evaluate the role of neuromuscular ultrasound in the assessment of idiopathic entrapment neuropathies of median nerve at the wrist and ulnar nerve at the elbow; and to determine the relationships of ultrasound findings with the clinical severity of CTS and cubital tunnel syndrome (as assessed by

validated clinical scale) and the electro-physiological grading scale.

Patients and Methods

Patients:

Between May 2015 and January 2016, Sixty five patients (53 females and 12 males) with 100 diseased hands and clinically suspected to have idiopathic CTS and seventeen patients (9 females and 8 males) with 20 diseased elbows and clinically suspected to have idiopathic CuTS were included. One hundred twenty (100 for CTS and 20 for CuTS) healthy age and sex matched individuals were served as a control group. All patients and controls were underwent nerve conduction studies and subsequent sonographic evaluation.

Excluded from the study patients with history of wrist/elbow trauma, local joint injection, fracture or surgery, Clinical, electrophysiological or radiological evidence of proximal median or ulnar neuropathy, cervical radiculopathy or polyneuropathy, history of underlying disorders (physiologic/pathologic) that can be associated with CTS: physiological e.g pregnancy, drugs e.g hormonal contraception, neuropathic causes e.g diabetes mellitus, alcoholism, endocrinal e.g

hypothyroidism, acromegaly, wrist and / or elbow arthritis due to any cause e.g rheumatoid arthritis, renal failure, congestive heart failure.

Methodology

The clinical diagnosis of CTS was based on Azami et al., 2013 diagnostic criteria⁽¹⁷⁾. A modified Arabic version of the Boston Carpal Tunnel Questionnaire (BCTQ) was used to obtain a patient-oriented measurement⁽¹⁷⁾. The Historical-Objective distribution based (Hi-Ob-Db) scale was also used to assess the condition regarding subjective symptoms as well as objective signs⁽¹⁴⁾. All neurophysiological studies were done using *Neuropack S1, MEB-9400K, 8 channels EMG/EP Measuring System; Nihon Kohden; Japan*. An electrodiagnosis grading scale for CTS was introduced by Bland 2000⁽¹⁵⁾. Musculoskeletal ultrasound scans were performed using *Siemens ACUSON P500 Ultrasound System (Siemens Healthcare, Boulevard, Malvern, USA) multi-frequency 1-14 MHz linear transducer*. Subjects were seated facing the examiner with arms extended and wrists resting on the examination couch, forearms supinated, and the fingers semi-extended⁽¹⁶⁾ as shown in fig (1).



Figure (1): patient positioning during neuromuscular ultrasound of the median nerve at the wrist

The CSA of the median nerve was measured by tracing method at the tunnel inlet⁽¹⁶⁾; and outlet^(17,18). The median nerve then imaged in cross-section at mid

forearm, then the wrist forearm ratio (WFR) was calculated^(14,15). The median nerve flattening ratio (FR) (at the pisiform or hamate level) was calculated by dividing

the major transverse axis of the nerve by its minor longitudinal axis⁽¹¹⁾. Flexor retinaculum bowing was defined as a measurement at 90° from a line drawn from the hook of the hamate bone to the tubercle of the trapezium bone⁽¹¹⁾. The median nerve mobility (transverse sliding) in the carpal tunnel was observed dynamically during flexion/extension of the fingers and wrist. An imaginary, transverse line was drawn bisecting the levels of the pisiform and the hook of the hamate. The mobility of the MN was evaluated on axial plane at this level⁽¹¹⁾. and finally blood flow in the MN sheath was then detected around 2 cm above the carpal tunnel using color and power Doppler sonography⁽¹⁷⁾.

The clinical diagnosis of cubital tunnel syndrome was based on signs and symptoms of ulnar nerve distribution⁽¹⁵⁾. Ulnar neuropathy at the elbow questionnaire (UNEQ) for symptoms severity was used⁽¹⁶⁾. Electrodiagnostic studies were done according AAEM 1999⁽¹⁷⁾. Patients then divided into five grades of severity on the basis of neurophysiological

classification⁽¹⁷⁾. Neuro-muscular ultrasound evaluation, using the same machine; patient was in supine position and arm was abducted and flexed 90° at the elbow for evaluation of the nerve from the wrist to the mid-arm⁽¹¹⁾ fig. (2). To obtain a sagittal view, the transducer was placed in the ulnar groove. Then, rotated 90° degrees to obtain a cross sectional view at the elbow. The transducer was then advanced distally to the mid-forearm and proximally to the mid-arm, for imaging of the nerve in the cubital tunnel, at the level of the medial epicondyle, and in the supracondylar region⁽¹⁵⁾. On transverse scans, the CSA of the ulnar nerve was determined using direct tracing method. Nerve size was measured at the following levels: the medial epicondyle, 2cm proximal 2cm distal to this level and in the middle of the upper arm and forearm.

Ulnar nerve ratios; upper arm and forearm swelling ratios were calculated by dividing the maximum ulnar nerve CSA at the elbow by the ulnar nerve CSA at the middle of the upper arm and forearm respectively⁽¹⁸⁾.

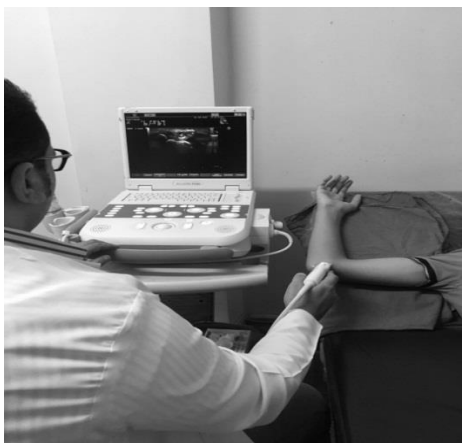


Figure (2): Positioning for ulnar nerve imaging

Statistical analysis

Analysis of data was done by personal computer using SPSS (Statistical program for social science) version 16. Data were expressed as mean \pm SD for parametric variables and as number and percent for non-parametric variable. Comparison between groups for parametric data was done by independent samples t-test (unpaired t-test). Chi – square (χ^2) test was

used to compare qualitative variables. The difference was expressed as probability of value (P value). The difference was considered significant if $P < 0.05$.

Results

Concerning CTS group; their ages ranged from 18 to 60 years with a mean of 30.86 ± 9.74 and their illness duration was ranged from 2 to 20 weeks with a mean of 7.24 ± 3.39 .

Measures of CSA of the MN at the inlet, CSA at the outlet, palmer bowing and inlet/forearm ratio in the CTS patients were significantly different compared to control group ($p < .001$ for all) and flattening ratio $p = .02$ (table 1). The accuracy of ultrasonographic measurements was evaluated by using cut off points of ROC curve. The area under the curve (AUC) of CSA was $.90$, indicating a sensitivity and specificity of 94% and 76% respectively, at cut off value of 4.0 mm^2 . The AUC of inlet/forearm ratio was $.91$ at cut off value of 2.18 indicating a sensitivity and specificity of 90% and 92% respectively. The AUC of FR was $.90$ at cut off value of 2.7 indicating a sensitivity and specificity of 91% and 99% respectively. The AUC of PB was $.90$ at cut off value of 2.00 mm indicating a sensitivity and specificity of 91% and 90% respectively (table 2).

Our results showed that CSA and Palmer bowing were the most statistically significant parameters that had correlation with the clinical parameters. There was positive correlation between symptom severity score and US parameters as regards CSA at the inlet, inlet/forearm ratio, palmer bowing and mobility ($r = .30, p = .003, r = .21, p = .004, r = .22, p = .003$ and $r = .26, p = .008$ respectively) moreover there was positive correlation between functional score and US parameters as regards CSA at the inlet, CSA at the outlet, inlet/forearm ratio, flattening ratio and palmer bowing ($r = .27, p = .007, r = .28, p = .004, r = .23, p = .001, r = .27, p = .008$ and $r = .20, p = .004$ respectively).

The Hi-Ob-Db. score showed positive correlation with US parameters as regards CSA at the inlet, CSA at the outlet, inlet/forearm ratio, palmer bowing, mobility

and doppler signals ($r = .29, p = .003, r = .20, p = .004, r = .31, p = .002, r = .29, p = .004, r = .28, p = .000$ and $r = .29, p = .004$ respectively).

Concerning CuTS group; their ages ranged from 20 to 70 years with a mean of 38.20 ± 9.90 and disease duration ranged from 2 to 34 weeks with a mean of 13.10 ± 10.04 .

CSA of the ulnar nerve at the medial epicondyle ranged from 1 to 20 mm^2 with a mean of $13.90 \text{ mm}^2 \pm 2.90$, while the CSA 2 cm proximal to the medial epicondyle ranged from 6 to 11 mm^2 with a mean of $8.0 \text{ mm}^2 \pm 1.80$. The CSA 2 cm distal to the medial epicondyle ranged from 6 to 10 mm^2 with a mean of $8.00 \text{ mm}^2 \pm 1.39$. The medial epicondyle /mid arm CSA ratio was found to be from 1.20 to 2.81 with a mean of 2.08 ± 0.50 while the medial epicondyle /mid forearm CSA ratio as found to be from 1.00 to 4.50 with a mean of 2.28 ± 0.84 . Power Doppler signals was found to be positive in 11 (100%) of diseased elbows. There was a difference in the CSA of the ulnar nerve at the medial epicondyle in controls (8.70 mm^2) compared to those with UNE (13.90 mm^2), which was statistically significant ($P = .001$) (fig.3). The ulnar nerve ratios at the medial epicondyle and mid arm and mid forearm were also statistically significant with $p < .001$ (table 3).

CSA at the medial epicondyle was found to be the most sensitive parameter in diagnosis of CuTS. It showed the highest sensitivity 100% and specificity 80% with 98.0% for the AUC at cut off point 9.0 mm^2 , compared to other US parameters. Combined CSA at the medial epicondyle and ulnar nerve ratio add no more values to sensitivity or specificity (table 4).

Table (1): US findings in CTS patients and control

	Patients (n=100)	Control (n=100)	P value
CSA inlet	11.89 ± 2.64	7.80 ± 1.08	$<.001^*$
CSA outlet	10.34 ± 2.37	7.76 ± 1.29	$<.001^*$
Inlet/forearm ratio	2.03 ± 0.79	1.77 ± 0.29	$<.001^*$
Flattening ratio	3.10 ± 0.83	2.88 ± 0.48	$.002^*$
Palmer bowing	4.27 ± 1.37	1.90 ± 0.37	$<.001^*$

Independent sample t test *Significant p-value $<.05$, CSA; Cross Sectional Area

Table (γ): accuracy of ultrasonographic parameters in CTS group:

	AUC	sensitivity	specificity	Cut off value	P value
CSA inlet	90%	94%	77%	8.0	<0.001*
CSA outlet	92%	80%	89%	8.0	<0.001*
Inlet/ forearm ratio	90%	70%	92%	2.08	<0.001*
Flattening ratio	00%	21%	99%	3.71	0.267
Palmar bowing	90%	91%	90%	2.00	<0.001*

ROC curve, CSA; Cross Sectional Area, *Significant p-value < 0.05

Table (ϛ): Ultrasonographic findings in cubital tunnel syndrome group and control

		Elbows (No = 20)	control	p-value
CSA at the medial epicondyle	Range	10 - 20	8-10	<0.001*
	Mean± SD	13.90 ± 2.90	8.70 ± 0.79	
CSA 2 cm proximal to media epicondyle	Range	6 - 11	6-8	0.001*
	Mean± SD	8.0 ± 1.80	7.90 ± 0.79	
CSA 2 cm distal to media epicondyle	Range	6 - 10	0-7	<0.001*
	Mean± SD	8.00 ± 1.39	0.90 ± 0.79	
medial epicondyle CSA/ arm ratio	Range	1.20 - 2.81	0.80 - 1.33	<0.001*
	Mean± SD	2.08 ± 0.50	1.10 ± 0.19	
medial epicondyle CSA/ forearm Ratio	Range	1.00 - 4.50	1-2	<0.001*
	Mean± SD	2.78 ± 0.84	1.02 ± 0.29	

Table (ε): Diagnostic value of ultrasonographic parameters in cubital tunnel syndrome:

	AUC	Sensitivity	Specificity	Cutoff value	P value
CSA at the medial epicondyle	98.0%	100%	80%	9.0	<0.001*
CSA 2 cm proximal to medial epicondyle	76%	70%	70%	7.0	0.0005*
CSA 2 cm distal to medial epicondyle	94%	90%	80%	6.0	<0.001*
medial epicondyle CSA/arm ratio	90.0%	90%	80%	1.29	<0.001*
medial epicondyle CSA/forearm ratio	90%	90%	80%	1.7	<0.001*
Combined CSA at the med. Epicondyle and delta arm	99%	100	80%	384.64	<0.001*

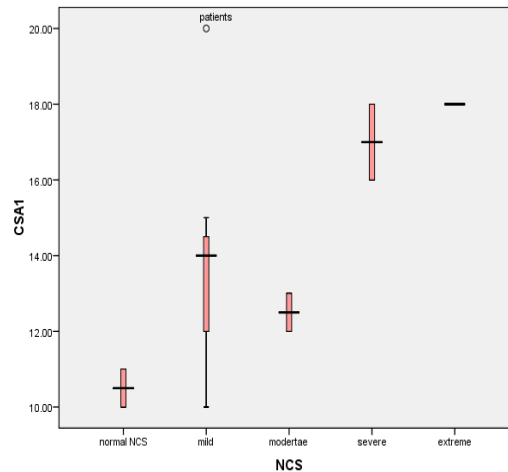


Fig. (3): Range of CSA in CuTS group

Discussion

In most instances, the value of complementary testing should be determined by the extent to which it affects the probability of the patient having the diagnosis that had been established clinically. Clearly, there are diagnoses that cannot be well established on the basis of clinical criteria alone and appropriate testing. Confirmation of CTS is usually evaluated by electrophysiological study⁽¹³⁾. However, sometimes, it is difficult to diagnose CTS using only this modality, early cases and even severe CTS that show no response to the stimulation, elderly patients and associated peripheral polyneuropathy patients⁽¹⁴⁾.

Recently, US techniques came into advancement as a tool to complement the diagnosis of CTS⁽¹⁵⁾.

In the present study, the efficacy of ultrasound for the assessment of CTS and CuTS was evaluated. Electrodiagnostic studies and clinical criteria were used as gold standard diagnostic procedures.

The sensitivity of the CSAs for diagnosis of CTS was ranged from 88% to 89%^(16, 17, 18, 19) and the CSA cutoff at which the values were considered abnormal, varied from 9-10 mm^(16, 17, 18, 19, 20). Our study showed 94% sensitivity and 76% specificity at 8.0 mm⁽¹⁾ cut off value for the mean CSA at the inlet which was the same cut off value for the study of Mohammadi et al., 2010⁽²¹⁾ with near equal sensitivity (94%) and different specificity (98%). This higher specificity

probably was due to higher number of patients included in that study (164 wrists vs. 100 wrists in our study). Kim et al., 2014 results showed a higher sensitivity 88.0% probably due to less number of their control (30 wrists vs. 100 wrists in our study) and higher sample size (247 wrists) and showed a higher specificity 90% probably due to higher cutoff value of 10mm⁽²⁾.

The sensitivities of increased palmer bowing of the flexor retinaculum varied from 80% to 89.2%^(3, 4, 5, 6) and sensitivities of flattening ratio ranged from 37% to 100%^(7, 8, 9, 10). Our results showed a sensitivity of 91% and 90% specificity at cut off value of 2.00 mm for the PB. These data of sensitivities corresponds with the findings reported in earlier studies. Sensitivity of FR was found to be 21% at cut off value of 3.7 mm which is less than previous data by kim et al., 2014 (59.8% vs. 21% in this study) due to higher cutoff value (3.8 vs. 3.7 respectively). FR had a poor predictive value as found by^(11, 12).

In our study, CSA and FR, PB of US were significantly increased in the CTS group than the control group. Among them, CSA at the inlet and PB were found to have a relatively higher accuracy than FR according to the ROC curve. Therefore, measurement of CSA at the inlet and/or PB can be considered as an alternative modality to distinguish CTS patients from asymptomatic controls. These data found to be in concordant with previous data^(13, 14, 15, 16, 17).

Mild CTS cannot show abnormal findings on US study in previous studies^(3,21). In our results, US could detect abnormalities as regards CSA at the inlet and PB (80% and 90% of patients in subgroup 1 respectively) with a significant difference in comparison to control group which is in concordant with Mhoon et al., 2012.

Results from Keles et al., 2000 and Kim et al. 2014, could not show a significant difference of US parameters regards CSA in patients with normal NCS that is different from our results; as their cut off value was higher compared to our results (9.3 mm² and 10 mm² vs. 8.0 mm² respectively) as regards CSA and 3.7 mm and 3 mm vs. 2.00 mm respectively as regards PB.

The diagnosis of ulnar neuropathy at the elbow is usually made by clinical neurologic examination and standard nerve conduction studies⁽²²⁾. In the majority of cases it is usually easily diagnosed by means of NCS; however, clinical examination is often non-localizing, the role of provocative tests only marginal, while electrophysiological tests may be normal or non-localizing with sensitivities ranging from 37% to 86%^(23, 24).

Our finding from the ultrasound measurement at different levels around the elbow showed positive correlation with NCS in concordant with Volpe et al., 2009; Omejec and Podnar, 2010.

The CSA behind the medial epicondyle was found to be the site of maximum enlargement (CSA max.) with the highest sensitivity and specificity (100% and 80% respectively) at 9.0 mm² cut off value and the AUC was 98.0%. This cut off value was in concordant with Pompe and Beekman 2013⁽²⁵⁾.

In a study by Simon et al, 2010⁽²⁶⁾; the CSA max was comparable to our results (13.0±1.4 vs. 13.90±2.90 respectively) and they found a significant difference to their control (p<0.01).

The range of CSA max was 10-20 mm² which was the same range of our study. Also data from Omejec and Podnar 2010⁽²⁷⁾

found that CSA max was comparable to our results and ranged from 10-14 mm².

As regards the ulnar nerve ratios; Simon et al., 2010 found that mid-arm ratio to be 2.2±0.2 which was significant regarding their control. These data was comparable to our results.

In terms of sensitivity and specificity the ulnar nerve ratio was found to be 90% and 80% respectively at cut off value 1.29 which was comparable to results gained from Omejec and Podnar, 2010. Pompe and Beekman, 2013 concluded that ROC-analysis of their results did not show a cut-off point for the swelling ratio with much higher sensitivity (without loss of too much specificity). The lower specificity found in their study compared to our results may be explained by their control group which consisted of disease not healthy controls.

The AUC of The forearm ratio was found to be 90%. The sensitivity and specificity was 90% and 80% respectively, which was more than those found by Bayrak et al., 2011⁽²⁸⁾; 88% and 77% respectively probably due to lower number of controls compared to their patients (21 vs. 41 healthy controls).

In conclusion, US is considered a new diagnostic modality in entrapment neuropathies of the median and ulnar nerves and several cut off values are considered nowadays as a diagnostic criteria for CTS and CuTS.

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