Diagnostic value of median nerve shear wave ultrasound elastography in diagnosis and differentiation of carpal tunnel syndrome severity

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Abstract

Background: Carpal tunnel syndrome (CTS) is the commonest type of peripheral nerve entrapment syndromes. The study aimed at evaluation of diagnostic value of median nerve stiffness measured by shear wave ultrasound elastography for diagnosis and differentiation of CTS severity, correlated to electrophysiological studies. This case–control study involved 40 patients (56 wrists) with CTS of different severity and 40 controls (40 wrists). All participants underwent electrophysiological study to assess the CTS severity, high-resolution conventional B-mode ultrasound to assess cross-sectional area "CSA" of median nerve at carpal tunnel, ratio of median nerve CSA at carpal tunnel and forearm, and shear wave ultrasound elastography with measurement of median nerve mean stiffness and correlation to electrophysiological results as the reference standard.

Results: Mean median nerve stiffness by shear wave US elastography was increased in patients with CTS compared to controls and across the different CTS severity groups (P value < 0.001 & 0.001, respectively). The cutoff value by ROC curve analysis for median nerve stiffness to differentiate between patients with CTS and control group was 65.4 kPa (P value < 0.001, 94.6% sensitivity, 97.3% specificity). Higher diagnostic accuracy was noted with the combination of shear wave elastography and conventional B-mode US with improved AUC (B-mode + shear wave; 0.962, P value < 0.001).

Conclusions: Shear wave ultrasound elastography of median nerve was able to discriminate different severity subgroups of CTS with high sensitivity, while conventional US couldn't. The diagnostic accuracy of CTS was improved when combined high-resolution conventional B-mode US and complementary shear wave ultrasound elastography.

Keywords: Carpal tunnel syndrome, Mean stiffness, Median nerve, Shear wave elastography, Ultrasound elastography

Background

The most common type of peripheral nerve entrapment syndrome is carpal tunnel syndrome (CTS) and can affect one or both hands. The symptoms of CTS are pain, tingling, or numbness most notably in the radial three and a half fingers [1, 2].

Women have a threefold greater risk of carpal tunnel syndrome than men and is commonly diagnosed at age between 30 and 60 years old [2]. There are certain conditions associated with increased risk of carpal tunnel syndrome including arthritis, diabetes, hypertension, smoking, old wrist trauma, sedentary life style, high body mass index (BMI) and jobs requiring repetitive wrist movement [3–5].
CTS is typically a clinical diagnosis; however, the electrodiagnostic studies help to support the diagnosis and also aid in grading of CTS severity; that the electrodiagnostic studies give perspectives on peripheral nerve function, myelin dysfunction and axonal loss degree [1, 6, 7].

Ultrasound has played a growing role in the examination of peripheral nerves and gained popularity as a complementary method of investigation to electrophysiological studies [8, 9]. B-mode ultrasound and Doppler examination are used traditionally to examine peripheral nerves to gain information about nerve structure, cross-sectional area, echogenicity, vascularity and any anomalous anatomical structure [10, 11].

Ultrasound elastography is a noninvasive ultrasound-based imaging technique to examine and measure soft tissue stiffness. And depending on the physical quantity measured, the ultrasound elastography is divided into strain and shear wave elastography (SWE) [12–17].

Shear wave elastography measures the tissue stiffness by assessing the speed and the pattern of shear wave propagation in the target tissue. The lower speed of shear waves means soft tissue, while higher speed indicates stiff or hard tissue; the measured tissue stiffness is quantified on an absolute scale [18–21].

Ultrasound elastography is easy and fast technique and characterized by its feasibility and accessibility. And recently, there is an increasing interest within the use of ultrasound elastography for the evaluation of neuromuscular pathologies [22–24]; that ultrasound elastography has predominantly been used for evaluation of liver fibrosis and differentiation of malignant and benign neoplasms, especially breast neoplasms [25–28].

There are many histological changes that occur within the nerves in case of diseased peripheral nerves, so nerve ultrasound elastography can be retrieved as a noninvasive way to assess and evaluate nerve tissue composition changes as loss of more compliant myelin and its replacement with connective tissue [23, 29].

So, this study aimed at evaluation of diagnostic value of median nerve stiffness measured by shear wave ultrasound elastography for diagnosis and differentiation of carpal tunnel syndrome (CTS) severity, correlated to electrophysiological studies.

Methods
This is a case–control, prospective study approved by the local institutional ethics committee; written informed consent was obtained from all participants.

Study population
Our study involved 80 subjects: 40 patients (56 wrists) with carpal tunnel syndrome of different severity and 40 controls (40 wrists) of matched ages and sex (mean age of 41 years and age range of 25–66 years for patient group and mean age of 43 years and age range of 25–66 years for control group). They were sampled in a simple random manner over a period from December 2018 to January 2020.

Inclusion criteria of patients group involved patients with clinical diagnosis of CTS based on the clinical diagnostic criteria of the American Academy of Neurology and American Association of Electrodiagnostic Medicine [30, 31]. These criteria include: swelling, pain, paresthesia, numbness or weakness of the hand worsened by sleep or sustained arm or hand position, repetitive action of the hand or wrist that is relieved by shaking the hand or changing posture, sensory deficit or atrophy of the thenar muscles. Exclusion criteria of patients group involved (a) patients < 18 years, (b) past history of wrist fracture or surgery, (c) previous carpal tunnel release surgery, (d) clinical evidence of other forms of neuropa-thies as mono- or poly-neuropathies and cervical radiculopathy, (e) history of diabetes or autoimmune disease, (f) patients refused to participate in the study.

Inclusion criteria of control group involved healthy volunteers of matched ages, sex and free of clinical signs of CTS and without past history of wrist trauma or surgery, any clinical signs of neuropathy, diabetes or autoimmune disease.

All participants underwent revision of medical history, clinical examination of both wrists, electrophysiological tests, high-resolution conventional B-mode ultrasound and shear wave ultrasound elastography (SWE) of median nerve, as the following:

History and clinical examination
The following points were assessed: (a) age, (b) clinical presentation, (c) history of diabetes, autoimmune disease, previous wrist surgical maneuvers, or previous wrist fractures. Clinical examination of both wrists was done by referring clinician.

Electrophysiological diagnostic study
A standard electrophysiological diagnostic examination was done for all participant subjects on the basis of the American Association of Electrodiagnostic Medicine recommendations [32]. Correlation between electrophysiological examination results, as reference standard tool for diagnosis and assessment of CTS severity, and median nerve mean stiffness value measured by ultrasound elastography, was done.

Technique of electrophysiological test was as in the following [32]; it was done at room temperature. The examiner was blinded to the clinical diagnosis of both patients and volunteers.
Antidromic sensory median nerve and ulnar nerve conduction studies were recorded from digit II and digit V, respectively. Motor median nerve conduction studies with stimulation at the wrist and elbow and motor ulnar nerve conduction studies with stimulation at the wrist, above and below elbow, were recorded, both including F-waves.

Positive results for the presence of CTS were considered if ≥0.4 ms difference between median and ulnar sensory peak latencies and/or a median distal motor latency (DML) ≥0.4 ms [32].

CTS severity was graded as mild, moderate, and severe on the basis of the electrophysiological diagnostic study results.

Ultrasound imaging of the median nerve

Conventional high-resolution B-mode ultrasonography and shear wave elastography of the median nerve were done for all subjects by using Logic P9 (GE healthcare medical system, USA) with high-frequency linear transducer L3-12 (12-MHz).

Ultrasound examinations were carried out by a radiologist of 15-year experience and who was blinded to the electrodiagnostic test results of the participants.

The participants were asked to lie supine with relaxed upper and lower limbs and no movement of their wrists or fingers during the examination. Images of shear wave elastography were also acquired in the same plane and position.

In control participants, the Rt. wrist was standardized in US examination.

Conventional high-resolution B-mode ultrasound imaging of the median nerve

The examination was done by using the musculoskeletal preset, highest frequency resolution mode with the depth of 2 cm and adjusted focus at the level of the median nerve. Identification of the median nerve in the transverse view at the carpal tunnel was done. The median nerve CSA was measured at carpal tunnel inlet, with reference to bony landmarks of the scaphoid and pisiform bones and at the forearm (12 cm above the carpal tunnel inlet). CSA was obtained using freehand boundary tracing technique, just inside the hyperechoic rim of the nerve. The transducer was kept perpendicular to the nerve to obtain accurate CSA with no additional applied force other than the transducer to avoid any artificial nerve deformity.

Measurement was repeated for three times, and the mean of the three values was analyzed. Ratio of median nerve CSA at carpal tunnel to CSA at forearm was also calculated.

Normal median nerve CSA at the distal wrist crease ranges from 7.2 to 9.8 mm², according to previous literature [33, 34].

Power Doppler examination was also done to assess whether it increased the intra-neural blood flow or not.

Shear wave ultrasound elastography imaging of the median nerve

SWE images required no external compression or transducer pressure. After few seconds of no movement that enables stabilization of the SWE image, freeze, then an ideal image of the median nerve at the longitudinal view was obtained. An automated fixed-sized 3-mm circular region of interest (ROI) in diameter was placed on the median nerve directly at the level of lunate bone. Automatically calculated quantitative stiffness (elasticity) value was obtained in kPa. The measurement was repeated for three times, and the mean stiffness or elasticity value was analyzed. Quantitative stiffness values ranged from 0 to 180 kPa and displayed tissue stiffness on a color scale from dark blue (lowest stiffness) to red (highest stiffness) as seen in Figs. 1, 2, 3, 4, 5, 6 [18, 20–22].

Statistical analysis

Statistical analysis and tests were determined according to the variable type. IBM Statistical Package for Social Sciences software (SPSS), 21st edition, IBM, USA, was used for data analysis. Continuous data were expressed as mean ± standard deviation (SD); categorical data were expressed as a percentage. Comparison of conventional high-resolution B-mode US results, mean stiffness value, and correlation to results of electrophysiological studies were done. The sensitivity and specificity were determined. Chi-squared and Spearman correlation tests were specified for qualitative data comparison. ROC curves for determination of cutoff values of mean stiffness value were defined with area under curve (AUC), specificity, and sensitivity.

Results

The study involved 80 participants: 40 patients with CTS of different severity and 40 controls of matched age and sex to the involved patients. Fifty-six wrists were examined from 40 patients (24 females and 16 males; mean age of 41 ± 1.1 years; age range of 25–66 years) and 40 wrists from 40 control participants (24 females and 16 males; mean age 43 ± 2.3 years; age range of 25–66 years). Sixteen patients had bilateral disease (10 females and 6 males), and 6 patients with clinical CTS displayed normal electrophysiological results. Mild CTS was found in 16 wrists, while 19 wrists displayed moderate CTS and 15 wrists displayed severe CTS.
Fig. 1 A 32-year-old normal female free from symptoms of carpal tunnel syndrome with normal electrophysiological diagnostic studies. (Left) High-resolution conventional B-mode ultrasound normal median nerve thickness (≈ 6 mm). (Right) Shear-wave elastography displayed homogenous blue color of the median nerve and mean median nerve stiffness value = 42 kPa.

Fig. 2 A 45-year-old female patient with moderate grade of carpal tunnel syndrome by electrophysiological diagnostic studies. (Left) High-resolution conventional B-mode ultrasound showed thickened median nerve with increased CSA (12.8 mm²), and wrist/forearm ratio (2.7). (Right) Shear-wave elastography of the median nerve displayed increased mean median nerve stiffness value = 94.6 kPa.

Fig. 3 A 58-year-old male patient with severe grade of carpal tunnel syndrome by electrophysiological diagnostic studies. (Left) High-resolution conventional B-mode ultrasound showed thickened median nerve with increased CSA (13.6 mm²) and calculated wrist/forearm ratio (2.9). (Right) Shear-wave elastography of the median nerve displayed increased mean median nerve stiffness value = 102.7 kPa.
Fig. 4 A 37-year-old female patient with mild grade of carpal tunnel syndrome by electrophysiological diagnostic studies. (Left) High-resolution conventional B-mode ultrasound showed mildly thickened median nerve with increased CSA (10.4 mm²) and wrist/forearm ratio (1.9). (Right) Shear-wave elastography of the median nerve displayed mildly increased mean median nerve stiffness value = 68.9 kPa

Fig. 5 A 40-year-old female patient with moderate grade of carpal tunnel syndrome by electrophysiological diagnostic studies. (Left) High-resolution conventional B-mode ultrasound showed thickened median nerve with increased CSA (12.5 mm²) and wrist/forearm ratio (2.4). (Right) Shear-wave elastography of the median nerve displayed increased mean median nerve stiffness value = 89.4 kPa

Fig. 6 A 48-year-old male patient with severe grade of carpal tunnel syndrome by electrophysiological diagnostic studies. (Left) High-resolution conventional B-mode ultrasound showed thickened median nerve with increased CSA (13.2 mm²) and calculated wrist/forearm ratio (2.7). (Right) Shear-wave elastography of the median nerve displayed increased mean median nerve stiffness value = 105.2 kPa
Table 1  Quantitative analysis of CSA of median nerve, wrist/forearm ratio and mean stiffness (elasticity) by ROC analysis between CTS patients and controls

| Variable               | Mean stiffness value (at cutoff = 65.4 kPa) | CSA by US (at cutoff = 8.5 mm²) | Wrist/forearm ratio (at cutoff = 1.3) |
|------------------------|---------------------------------------------|---------------------------------|---------------------------------------|
| Control (40 wrists)    | 38.13±23.56                                 | 7.2±1.82                        | 1.6±0.41                              |
| Patients with CTS (56 wrists) | 95.73±27.58                               | 11.7±2.93                       | 2.5±0.81                              |
| P value                | <0.001                                      | 0.01                            | 0.01                                  |
| Sensitivity (%)        | 94.6                                        | 81                              | 89                                    |
| Specificity (%)        | 97.3                                        | 83                              | 87                                    |
| AUC (area under curve) | 0.901                                       | 0.721                           | 0.734                                 |

Fig. 7  A Receiver operating characteristic curve analysis showed optimal cutoff value for mean stiffness value of median nerve, CSA and wrist/forearm ratio. Green line for mean median nerve stiffness, yellow line for CSA, blue line for wrist/forearm ratio

Receiver operating characteristic curve analysis revealed that the optimal cutoff values for median nerve stiffness, median nerve CSA, and wrist/forearm median nerve CSA ratio were (65.4 kPa, 8.5 mm², and 1.3, respectively) to differentiate between patients with CTS and control group with sensitivity and specificity for median nerve stiffness of 94.6% and 97.3%, respectively (Table 1; Fig. 7).

The median nerve CSA and wrist/forearm median nerve CSA ratio were found to be larger in patients with CTS compared to control group (mean 11.7±2.9 mm² vs. 7.2±1.8 mm²; P=0.01, and mean 2.5±0.8 vs. 1.6±0.4; P=0.01, respectively). Also, the mean stiffness value of the median nerve was greater in patients with CTS than in controls (mean 95.73 kPa±27.58 vs. 38.13 kPa±23.56; P<0.001) (Table 1).

Median nerve stiffness, CSA, and wrist/forearm CSA ratio were examined in control group and across different CTS severity subgroups. The study revealed that CSA and wrist/forearm CSA ratio of median nerve were significantly different between controls and CTS wrists, but there was no role of CSA or CSA ratio of median nerve to differentiate between different CTS severity groups. On
the contrary, mean median nerve stiffness was found to be helpful in differentiation of different CTS severity subgroups (*P* value = −0.001) (Table 2).

Patients with CTS were grouped into two subgroups (CTS with normal NCS/mild and moderate/severe) with subsequent analysis, which revealed that mean median nerve stiffness was significantly different between two subgroups as: CTS with normal NCS/mild CTS group (78.71 kPa ± 16.58) and moderate/severe CTS group (114.03 kPa ± 15.51) with a *P* value of 0.001 (Table 3).

As regards the power Doppler examination, none of controls showed positive power Doppler signals, but it was seen in 16 wrists with CTS (0% vs. 28%, *P* = 0.001).

No statistically significant difference was found in mean median nerve stiffness (90.45 kPa ± 16.7 vs. 85.68 kPa ± 13.2), CSA (12 ± 1.9 vs. 10 ± 1.4) or CSA ratio (1.9 ± 0.4 vs. 1.7 ± 0.3) in CTS wrists with positive versus negative power Doppler signals (*P* value = 0.06).

Quantitative analysis of median nerve CSA, forearm/wrist CSA ratio and mean median nerve stiffness were done for both CTS patients and controls according to age group, body mass index and sex. In both groups of controls and patients with CTS, there was no significant difference in mean median nerve CSA, forearm/wrist median nerve CSA ratio and median nerve stiffness between younger (< 55 years old) versus older (≥ 55 years old) patients, and non-overweight patients (BMI < 25) versus overweight (BMI ≥ 25) patients. But the mean median nerve stiffness was significantly higher in males compared to females either in CTS patients or controls as: (109.78 kPa ± 13.73 vs. 78.35 kPa ± 19.58, *P* = 0.01, and 45.91 kPa ± 16.29 vs. 18.84 kPa ± 13.67, *P* = 0.01) (Tables 4, 5).

The combined conventional high-resolution B-mode US and shear wave elastography examinations revealed significantly higher diagnostic performance and accuracy than each single examination; that the AUC for combined examination had been improved (conventional US + shear wave, AUC = 0.962 and conventional US, AUC = 0.721; *P* < 0.001) (Table 6).

### Table 3

| Variable | Mean stiffness value (at cutoff = 95.8 kPa) | CSA by US (at cutoff = 16.2 mm²) | Wrist/forearm ratio (at cutoff = 3.5) |
|----------|------------------------------------------|----------------------------------|-----------------------------------|
| CTS with normal NCS/mild CTS (n = 22) | 78.71 ± 16.58 | 14.53 ± 1.6 | 2.73 ± 1.2 |
| Moderate/severe CTS (n = 34) | 114.03 ± 15.51 | 18.9 ± 2.2 | 4.1 ± 0.6 |
| *P* value | 0.001 | 0.065 | 0.074 |
| Sensitivity (%) | 96.7 | 46 | 45 |
| Specificity (%) | 97.8 | 43 | 42 |
| AUC (area under curve) | 0.947 | 0.398 | 0.331 |
| NPV (%) | 98 | 43.2 | 41.7 |
| PPV (%) | 96 | 46.3 | 44.5 |

### Table 4

| Variable | CSA (mm² ± SD) | Wrist/forearm ratio (± SD) | Mean stiffness (kPa ± SD) |
|----------|----------------|--------------------------|--------------------------|
| Gender   |                |                          |                          |
| Male (n = 16, 22 wrists) | 12.6 ± 2.26 | 2.3 ± 0.43 | 109.78 ± 13.73 |
| Female (n = 24, 34 wrists) | 10.3 ± 2.52 | 2.1 ± 0.62 | 78.35 ± 19.58 |
| *P* value | 0.05 | 0.652 | 0.01 |
| Age group |                |                          |                          |
| < 55 (n = 30, 38 wrists) | 10.8 ± 2.54 | 2.1 ± 0.71 | 99.78 ± 23.27 |
| ≥ 55 (n = 10, 18 wrists) | 11.3 ± 2.73 | 1.9 ± 0.52 | 111.61 ± 12.43 |
| *P* value | 0.053 | 0.468 | 0.283 |
| BMI      |                |                          |                          |
| < 25 (n = 19, 26 wrists) | 11.2 ± 3.63 | 2.02 ± 0.84 | 94.75 ± 29.12 |
| ≥ 25 (n = 21, 30 wrists) | 11.6 ± 2.91 | 2.3 ± 0.53 | 103.98 ± 18.86 |
| *P* value | 0.414 | 0.637 | 0.591 |
Table 5: Quantitative analysis of CSA of median nerve, wrist/forearm ratio and mean stiffness (elasticity) between controls (n=40, 40 wrists)

| Variable          | CSA (mm²) | Wrist/forearm ratio | Mean stiffness (kPa) |
|-------------------|-----------|---------------------|---------------------|
| **Gender**        |           |                     |                     |
| Male (n=16)       | 6.3±2.83  | 1.4±0.52            | 45.91±16.29         |
| Female (n=24)     | 5.7±2.51  | 1.2±0.61            | 18.84±13.67         |
| P value           | 0.048     | 0.587               | 0.01                |
| **Age group**     |           |                     |                     |
| < 55 (n=29)       | 6.9±1.74  | 0.9±0.85            | 29.37±24.43         |
| ≥ 55 (n=11)       | 7.3±1.69  | 1.2±0.81            | 31.74±25.71         |
| P value           | 0.067     | 0.453               | 0.251               |
| **BMI**           |           |                     |                     |
| < 25 (n=24)       | 5.6±1.93  | 1.6±0.39            | 30.96±21.75         |
| ≥ 25 (n=16)       | 6.4±2.58  | 1.5±0.48            | 37.21±22.93         |
| P value           | 0.597     | 0.711               | 0.443               |

Table 6: Diagnostic performance of combination of conventional high-resolution ultrasonography and shear wave elastography of median nerve using determined cutoffs for elastographic values calculated from ROC curves

| Modality                | Sensitivity | Specificity | AUC     | P value |
|-------------------------|-------------|-------------|---------|---------|
| Conventional US         | 81          | 83          | 0.721   | <0.001  |
| + Shear wave elastography | 97.8         | 98          | 0.962   | <0.001  |

Discussion

Ultrasound elastography is a noninvasive imaging tool, which identifies tissue elasticity (stiffness) [1]. The use of ultrasound elastography for evaluation of neuromuscular pathologies has recently gained increased popularity [22–24].

According to our study results, the median nerve CSA and wrist/forearm median nerve CSA ratio were found to be larger in patients with CTS compared to control group (mean 11.7±2.9 mm² vs. 7.2±1.8 mm²; P=0.01, and mean 2.5±0.8 vs. 1.6±0.4; P=0.01, respectively). The cutoff values for median nerve CSA and wrist/forearm median nerve CSA ratio were (8.5 mm² and 1.3, respectively) to differentiate between patients with CTS and control group with sensitivity and specificity of 81% and 83% for CSA of median nerve, and 87% and 89%, respectively, for wrist/forearm CSA ratio. These results showed comparable results to those mentioned by Simon et al. [1], Lin et al. [29] and Wee et al. [2, 35].

Many studies were performed about diagnosis of CTS by ultrasound elastography; regardless of US elastography technique either strain or shear wave. These studies evoked a significant difference in median nerve mean stiffness measured by SWE between patients with CTS and controls [18, 19, 23, 34–38]. This trend was similarly observed in our study to those recorded by the former studies. That the mean stiffness value of the median nerve was greater in patients with CTS than in controls (mean 95.73±27.58 kPa vs. 38.13±23.56 kPa) with a statistically significant difference in differentiation between CTS patients and controls (P<0.001).

There are various cutoff values that have been proposed for the diagnosis of CTS with reasonable respective sensitivity and specificity. In our study, the cutoff value of mean median nerve stiffness measured by SWE was 65.6 kPa (P value <0.001, AUC=0.910, sensitivity =94.6%, specificity =97.3%); these findings were matched with the previous results recorded by many studies, as the range of cutoff values was between 38.25–79 kPa and 3.0–3.23 m/s [18, 19, 35–38], as the study by Cingoz et al. [18], which involved 59 patients and 18 controls, reported that the best cutoff value of median nerve stiffness was 38.25 kPa, with 78.6% sensitivity, and 62.5% specificity.

We agreed to a large extent with the results observed by Paluch et al. [19] who studied 87 patients and 34 controls and reported cutoff value of mean median nerve stiffness of 79 kPa, with 96.6% sensitivity and 100% specificity, and by Kantarci et al. [36] who studied 60 patients and 36 controls and recorded cutoff value of median nerve stiffness of 40.4 kPa, 93.3% sensitivity, and 88.9% specificity.

But the demonstrated cutoff value of mean median nerve in this study was found to be lower than had been reported by Wee et al. [35], which was 84 kPa. And this could be explained by the younger age of participants involved in our study compared to older patients included in a study by Wee et al. [35]. This may be the cause of increased mean median nerve stiffness. Few authors demonstrated that carpal tunnel syndrome in elderly patients is a separate entity [39, 40] due to nerve microstructure changes in the aging process and that the nerve stiffness become more stiffer and smaller in size, and subsequently, the measured mean median nerve stiffness increased, while on the contrary the CSA of median nerve tends to be smaller, so it is less sensitive indicator for CTS.

Our study revealed increased mean median nerve stiffness correlated to increased severity grade of CTS demonstrated by electrophysiological diagnostic studies. It was found that mean median nerve stiffness was significantly different between two subgroups as: CTS with normal NCS/mild CTS group (78.71 kPa±16.58) and moderate/severe CTS group (114.03 kPa±15.51) with a P value of 0.001. These findings matched and agreed with formerly described and recorded results by other studies [18, 35–38]. As Kantarci et al. [36] recorded a difference in mean median nerve stiffness between mild and severe...
groups of CTS as (55.1 kPa ± 12.9 vs. 101.4 kPa ± 26.7), respectively, also Cingoz et al. [18] found a difference between mild and moderate groups as (44 kPa vs. 82 kPa), respectively. In other study by Wee et al. [35], the mean median nerve stiffness of median nerve across EDx/mild CTS and moderate/severe groups was (107.6 kPa ± 71.6 vs. 159.3 kPa ± 68.8), respectively, while a study done by Zhang et al. [38] and Arslan et al. [37] recorded mean median nerve stiffness of (3.25 m/s ± 0.41 vs 5.24 m/s ± 0.55 and 3.65 m/s ± 0.53 vs 5.09 m/s ± 0.57), respectively, between mild and severe CTS groups.

The combined conventional high-resolution B-mode US and shear wave elastography examinations revealed significantly higher diagnostic performance and accuracy than each single examination.

Our study had some limitations. Firstly, ultrasound technique is operator dependent and needs experienced radiologists to avoid misinterpretation and reduce the false-positive and false-negative outcomes. Secondly, a need for additional prospective studies with larger size is necessary to provide stronger results, conclusions and standardization of elastographic protocols and cutoff values with more realistic representative external validity.

Conclusions
Shear wave ultrasound elastography of median nerve is a noninvasive procedure with high sensitivity. It was able to discriminate different severity subgroups of CTS (CTS with NCS/mild CTS versus moderate/severe CTS groups), while conventional US could not. It was observed that the diagnostic accuracy of CTS was improved when combined high-resolution conventional B-mode US and complementary shear wave ultrasound elastography of the median nerve.

Abbreviations
BMI: Body mass index; CSA: Cross-sectional area; CTS: Carpal tunnel syndrome; NCS: Nerve conduction study; SWE: Shear wave elastography.

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Authors’ contributions
HR was involved in formulation of the study, preparation of methodology, data collection, analysis of the data, and writing the paper. The author has read and approved the manuscript.

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Availability of data and materials
The dataset used and/or analyzed during the current study are available from the corresponding author on reasonable request.

Declarations
Ethical approval and consent to participate
Approved by the local institutional ethics committee (Committee of Scientific Research Ethics (CSRE), Suez Canal University, Egypt); written informed consent was obtained from all patients. The reference number is not applicable and/or not available.

Consent for publication
Consent for publication was obtained from the patients and controls.

Competing interests
The author of this manuscript declares no relationships with any companies, whose products or services may be related to the subject matter of the article.

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References
1. Simon NG, Talbott J, Chin CT et al (2016) Peripheral nerve imaging. Handb Clin Neurol 136:811–826
2. Wee TC, Simon NG (2019) Ultrasound elastography for the evaluation of peripheral nerves: a systematic review. Muscle Nerve. https://doi.org/10.1002/mus.26624
3. Hobson-Webb LD, Padua L (2016) Ultrasound of focal neuropathies. J Clin Neuropathophysiol 33(2):94–102
4. Boitie AA, Arnold R, Russell BA et al (2018) Effects of hemodialysis on intraneural blood flow in end-stage kidney disease. Muscle Nerve 57(2):287–293
5. Boitie AA, Issar T, Kwai NC et al (2018) Correlation between markers of peripheral nerve function and structure in type 1 diabetes. Diabetes Metab Res Rev 34(7):e3028
6. Alanyaz MH (2017) Clinical and electrophysiological evaluation of carpal tunnel syndrome: approach and pitfalls. Neurosciences 22(3):169–180
7. Bourque PR, Brooks J, Mobach T, Gammon B, Papp S et al (2020) Systematic prospective electrophysiological studies of the median nerve after simple distal radius fracture. PLoS ONE 15(12):e0243490. https://doi.org/10.1371/journal.pone.0243490
8. Georgiev GP, Karabinov V, Kotov G, Iliev A (2018) Medical ultrasound in the evaluation of the carpal tunnel: a critical review. Cureus 10(10):e3487
9. Kuo TT, Lee MR, Liao YL et al (2016) Assessment of median nerve mobility by ultrasound dynamic imaging for diagnosing carpal tunnel syndrome. PLoS ONE 11(1):e0147051
10. Chen YT, Williams L, Zak MJ et al (2016) Review of Ultrasonography in the diagnosis of carpal tunnel syndrome and a proposed scanning protocol. J Ultrasound Med 35(11):2311–2324
11. Boitie AA, Visser LH, Padua L et al (2017) Utility of maximum perfusion intensity as an ultrasonographic marker of intraneural blood flow. Muscle Nerve 55(1):77–83
12. Nowicki A, Dobruch-Sobczak K (2016) Introduction to ultrasound elastography. J Ultrasound 16(65):113–124
13. Gdaci Ayasov R, Erdal A, Bu O et al (2018) The effectiveness of ultrasonography and ultrasonographic elastography in the diagnosis of carpal tunnel syndrome and evaluation of treatment response after steroid injection. Eur J Radiol 108:172–176
14. Taljanovic MS, Gimber LH, Becker GW et al (2017) Shear-wave elastography: basic physics and musculoskeletal applications. Radio Graph 37(3):855–870
15. Martin MJ, Cartwright MS (2017) A pilot study of strain elastography in the diagnosis of carpal tunnel syndrome. J Clin Neuropathophysiol 34(2):114–118
16. Tatar IG, Kurt A, Yavasoglu NG et al (2016) Carpal tunnel syndrome elastosonographic strain ratio and cross-sectional area evaluation for the diagnosis and disease severity. Med Ultrasound 18(3):305–311
17. Yoshii Y, Tung WL, Ishii T (2017) Strain and morphological changes of median nerve after carpal tunnel release. J Ultrasound Med 36(8):1153–1159
18. Cingoz M, Kandemirli SG, Alis DC et al (2018) Evaluation of median nerve by shear wave elastography and diffusion tensor imaging in carpal tunnel syndrome. Eur J Radiol 101:59–64
19. Paluch L, Pietruski P, Walecik J et al (2018) Wrist to forearm ratio as a median nerve shear wave elastography test in carpal tunnel syndrome diagnosis. J Plast Reconstr Aesthet Surg 71:1146–1152
20. Kesikburun S, Adiguzel E, Kesikburun B et al (2016) Sonoelectrostatic assessment of the median nerve in the longitudinal plane for carpal tunnel syndrome. PM R 8(2):183–185
21. Greening J, Dilley A (2017) Posture-induced changes in peripheral nerve stiffness measured by ultrasound shear-wave elastography. Muscle Nerve 55(2):213–222
22. Zhu B, Yan F, He Y et al (2018) Evaluation of the healthy median nerve elasticity: feasibility and reliability of shear wave elastography. Medicine (Baltimore) 97(43):e12956
23. Harmon B, Wells M, Park D et al (2019) Ultrasound elastography in neuro-muscular and movement disorders. Clin Imaging 53:35–42
24. Kubo K, Zhou B, Cheng YS et al (2018) Ultrasound elastography for carpal tunnel pressure measurement: a cadaveric validation study. J Orthop Res 36(1):477–483
25. Sigrist RMS, Liau J, Kaffas AE et al (2017) Ultrasound elastography: review of techniques and clinical applications. Theranostics 7(5):1303–1329
26. Dietrich CF, Bamber J, Berzigotti A et al (2017) EFSUMB guidelines and recommendations on the clinical use of liver ultrasound elastography, update 2017 (Long Version). Ultraschall Med 38(4):e48
27. Ferraioli G (2019) Review of liver elastography guidelines. J Ultrasound Med 38(1):9–14
28. Lower L, Beaver A, Cox K et al (2018) Evolving imaging techniques for staging axillary lymph nodes in breast cancer. Clin Radiol 73(4):396–409
29. Lin CP, Chen LJ, Chang KV et al (2019) Utility of ultrasound elastography in evaluation of carpal tunnel syndrome: a systematic review and meta-analysis. Ultrasound Med Biol 45(11):2855–2865
30. The American Association of Electrodiagnostic Medicine (2013) Practice parameter for electrodiagnostic studies in carpal tunnel syndrome. Muscle Nerve 25:912–922
31. Tengfei Fu, Cao M, Liu F et al (2015) Carpal tunnel syndrome assessment with ultrasonography: value of inlet-to-outlet median nerve area ratio in patients versus healthy volunteers. PLoS ONE. https://doi.org/10.1371/journal.pone.0116777
32. Werner RA, Andary M (2011) Electrodiagnostic evaluation of carpal tunnel syndrome. Muscle Nerve 44:597–607
33. Bathala L, Kumar P, Kumar K et al (2013) Ultrasoundographic cross-sectional area normal values of the ulnar nerve along its course in the arm with electrophysiological correlations in 100 Asian subjects. Muscle Nerve 47:673–676
34. Won SJ, Kim BJ, Park KS et al (2013) Reference values for nerve ultrasonography in the upper extremity. Muscle Nerve 47:864–871
35. Wee TC, Simon NG (2020) Shear wave elastography in the differentiation of carpal tunnel syndrome severity. Am Acad Phys Med Rehabilitation. https://doi.org/10.1002/prmr.12334
36. Kantarcı F, Ustubasögüloğlu FE, Delil S et al (2014) Median nerve stiffness measurement by shear wave elastography: a potential sonographic method in the diagnosis of carpal tunnel syndrome: Eur Radiol 24(2):434–440
37. Arslan H, Yavuz A, Iğen F et al (2018) The efficiency of acoustic radiation force impulse (ARFI) elastography in the diagnosis and staging of carpal tunnel syndrome. J Med Ultrason 45(3):453–459
38. Zhang C, Li M, Jiang J et al (2017) Diagnostic value of virtual touch tissue imaging quantification for evaluating median nerve stiffness in carpal tunnel syndrome. J Ultrason Med 36(9):1783–1791
39. Gregoris N, Bland J (2019) Is carpal tunnel syndrome in the elderly a separate entity? Evidence from median nerve ultrasound. Muscle Nerve 60:217–218
40. Mulroy E, Pelosi L (2019) Carpal tunnel syndrome in advanced age: a sonographic and electrodiagnostic study. Muscle Nerve 60:236–241

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