Abnormal Ultrasonographic Findings in Patients with Chronic HCV: An Egyptian Experience

MOHAMED A.R. MOSTAFA, M.Sc.*; TAHER M.H. EL ZANATY, M.D.**; AYMEN M.R. FODA, M.D.** and NEHAD M. TAWFIK, M.D.**

The Department of Internal Medicine, Faculty of Medicine, Misr University for Science & Technology* and Department of Internal Medicine & Gastroenterology, Faculty of Medicine, Cairo University**

Abstract

Background: Ultrasound plays a crucial role in the noninvasive evaluation of chronic HCV patients, particularly in limited resources setting where it can provide an easy-to-use and cheap imaging modality. Ultrasound can evaluate the liver size, texture, liver edge, hepatic parenchyma, presence of liver nodules, and hepatic vascularity, as well as the affection of spleen and venous supply. In most hepatic centers, ultrasound is a part of the routine evaluation of chronic HCV to evaluate the stage of disease and the severity of fibrosis. The present study assessed the frequency of abnormal ultrasonographic findings in Egyptian patients with chronic HCV infection.

Aim of Study: This study assessed the fruency of abnormal ultrasonographic findings in Egyptian patients with chronic hepatitis C virus (HCV) infection.

Patients and Methods: A cross-sectional study was conducted on HCV patients presented to the Cairo University hospitals through the period from July 2014 to July 2015. All examinations were performed by the same operator.

Results: Three hundred patients were included in the present study. The mean age of the included patients was 48.1 \pm 13 years old and nearly 60% of the patients were males. Overall, the prevalence of abnormal ultrasound findings during abdominal examination was 60.7% (n=180). Besides, 20% of the patients had enlarged spleen, with an average size of 13.9 \pm 3cm.

Conclusion: In conclusion, our findings highlight the high prevalence of abnormal ultrasound findings amongst Egyptian patients with chronic HCV. We found that more than 60% of the chronic HCV patients had variable degree of abnormalities during abdominal ultrasound findings. Thus, ultrasound examination should be implemented as routine investigation during evaluation of chronic HCV patients, which can provide an easy-to-use, reliable, and cheap modality in limited settings.

Key Words: Chronic HCV – Ultrasound – Chronic liver disease – Abnormalities.

Introduction

THE endemic of the hepatitis C virus (HCV) infection is a major public health problem with devastating long-term consequences. According to recent epidemiological figures, the seroprevalence of HCV is still increasing, especially in the Eastern Mediterranean and European Regions, with an estimated global prevalence of 2.5% [1,2]. Egypt has the highest prevalence rate of HCV worldwide with approximately 2 to 6 new cases per 1000 population every year [3]. Based on viral genome sequences, there are currently seven recognized genotypes of HCV; though HCV genotype 1 and 3 are the most prevalent worldwide, genotype 4 is the most commonly identified genotype in the Middle East [1,4]. HCV infection is transmitted mainly through blood contact and to a lesser extent by sexual contact or perinatally. Despite the fact that HCV infection is largely asymptomatic in the acute stage, chronic HCV is associated with a significant risk of mortality and liver-related complications [5]. Recently, it was reported that up to 30% of patients with chronic HCV had liver cirrhosis, while the prevalence of decompensated liver failure is 11 % [6]; moreover, the current body of evidence shows that chronic HCV is a major risk for the development of hepatocellular carcinoma (HCC) [7]. In addition, the mortality rates secondary to chronic HCV showed a notable increase from 2006 to 2010 [6]. Chronic HCV represents a financial burden, both in national and global levels, as well; according to previous reports, it was estimated that the treatment expenses of HCV infection were \$6.5 billion in 2011 in the United States alone [8].

Ultrasound plays a crucial role in the noninvasive evaluation of chronic HCV patients, particularly in limited resources setting where it can

Correspondence to: Dr. Mohamed A.R. Mostafa,

The Department of Internal Medicine, Faculty of Medicine, Misr University for Science & Technology

provide an easy-to-use and cheap imaging modality [9]. Ultrasound can evaluate the liver size, texture, liver edge, hepatic parenchyma, presence of liver nodules, and hepatic vascularity, as well as the affection of spleen and venous supply. In most hepatic centers, ultrasound is a part of the routine evaluation of chronic HCV to evaluate the stage of disease and the severity of fibrosis [10,11]. The present study assessed the frequency of abnormal ultrasonographic findings in Egyptian patients with chronic HCV infection.

Material and Methods

The study was initially approved by the local ethics committee of Cairo University Hospital, and run in compliance with regulatory laws and the Declaration of Helsinki [12]. The manuscript was prepared per the recommendations of the STROBE guidelines [13]. All patients signed the written informed consent before enrollment.

Study design and patients:

In this study conducted a cross-sectional study that recruited Egyptian patients with chronic HCV who were being followed at the outpatient clinics of Cairo University hospitals through the period from July 2014 to July 2015. Patients were deemed eligible if they aged >18 years old and had a laboratory-confirmed chronic HCV, regardless of the state of the cirrhosis. Patients were excluded if they had concurrent hepatitis B virus (HBV), autoimmune disease, malignancy, or cryoglobulinemia.

Data collection and ultrasound evaluation:

Patients were assessed for demographic characteristics, routine laboratory assessment, and abdominal ultrasound (US) examination. The ultrasound examination was performed using Bmode ultrasound (Philibis infinity 70) with a 7.5 MHz linear array transducer. A single ultrasound operator performed all ultrasound examinations The primary outcome of the present study was to assess the frequency of abnormal ultrasonographic findings in Egyptian patients with chronic HCV infection.

Statistical analysis:

The statistical software MINITAB (16.0) was used for data processing and analysis. According to the normality of data distribution, the central tendency and variability of the numerical data were presented in the form of mean \pm standard deviations (SD) or median with interquartile range (IQR). Frequency counts and percentages summarized categorical variables.

Results

Three hundred patients were included in the present study. The mean age of the included patients was 48.1 ± 13 years old and nearly 60% of the patients were males. The mean and standard deviation of the laboratory and imaging findings were estimated as shown in Table (1). The mean hemoglobin level was 11.5 ± 2.1 g/dL, while the mean of white blood cells and platelet count was $5.4 \pm 2.5 \times 10^3$ and $147.5 \pm 80 \times 10^6$, respectively. The mean direct bilirubin level was 1.4 ± 1.6 mg/dL. The liver enzymes (ALT and AST) were mostly within the average range, as well as the serum albumin level. The mean alpha fetoprotein (AFP) was 8.6 ± 4.3 mg/dL.

Overall, the prevalence of abnormal ultrasound findings during abdominal examination was 60.7% (n=180), Fig. (1). Besides, 20% of the patients had enlarged spleen, with an average size of 13.9 ± 3 cm.

Table (1): Characteristics of the studied groups.

te (1): Shurdeteristies of the studied groups.		
Variables	Patients (N=300)	
Age (years):		
Mean \pm SD	48.1 ± 13	
Range	18-85	
Sex (n, %):		
Male	182 (60.7%)	
Female	118 (39.3%)	
Laboratory findings:		
HB	11.5 ± 2.1	
WBCs	5.4±2.5	
Platelets	147.5 ± 80	
Total bilirubin	2.2±2.3	
Direct bilirubin	1.4 ± 1.6	
ALT	57±23	
AST	60±26	
Albumin	3,2±0.9	
PT	15.2±4	
INR	$1.4{\pm}0.3$	
AFP	8.6±4.3	

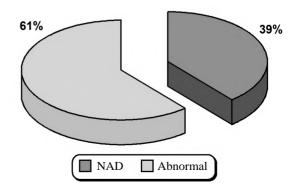


Fig. (1): Pelvi-abdominal ultrasonography findings.

Discussion

Ultrasound plays a crucial role in the noninvasive evaluation of chronic HCV patients, particularly in limited resources setting where it can provide an easy-to-use and cheap imaging modality [9]. Ultrasound can evaluate the liver size, texture, liver edge, hepatic parenchyma, presence of liver nodules, and hepatic vascularity, as well as the affection of spleen and venous supply. In most hepatic centers, ultrasound is a part of the routine evaluation of chronic HCV to evaluate the stage of disease and the severity of fibrosis [10,11]. The present study assessed the frequency of abnormal ultrasonographic findings in Egyptian patients with chronic HCV infection.

We found that the prevalence of abnormal ultrasound findings during abdominal examination was 60.7% (n=180). Besides, 20% of the patients had enlarged spleen, with an average size of 13.9 ± 3 cm. In a previous report by Màaji et al., the rate of abnormal ultrasound findings amongst patients with chronic liver disease was nearly 43%. Besides, nearly 93% of the cohort showed hepatic edge irregularities [9]. In another report, the rate of abnormal ultrasound findings was 32% [14]. In Weickert et al., study, the rate of abnormal ultrasound findings was 55% [15]. Other reports showed similar findings [16].

The current body of evidence shows that ultrasound examination can yield high sensitivity and reliability in assessing the severity of liver fibrosis. The multidimensional value of ultrasound cannot be overstated. In Choong et al., it was confirmed that routine abdominal ultrasound was sensitive and reliable for differentiating early stage of liver fibrosis, with nodularity being the most distinguished feature [17]. Other reports demonstrated that abnormal hepatic surface is a useful indicator of liver fibrosis and the need for liver biopsy [18]. Abnormalities in ultrasound was even found to have higher diagnostic yield when they combined with abnormal liver functions (albumin level <3.5g/dL) [19]. Martinez et al., recommended noninvasive tests can be combined with abdominal ultrasound to differentiate clinically-significant fibrosis [20].

This study has some limitations including the small sample size and the single-center setting of the study that may hinder the generalizability of our data. In addition, we could not assess the patterns of abnormalities due to the insufficiency of the data. Moreover, the implication of these abnormalities on HCV condition in form of clinical severity and response to treatment was not assessed. In conclusion, our findings highlight the high prevalence of abnormal ultrasound findings amongst Egyptian patients with chronic HCV. We found that more than 60% of the chronic HCV patients had variable degree of abnormalities during abdominal ultrasound findings. Thus, ultrasound examination should be implemented as routine investigation during evaluation of chronic HCV patients, which can provide an easy-to-use, reliable, and cheap modality in limited settings. Further research should be performed to examine whether high frequency probe can enhance the sensitivity of ultrasound examination, and whether ultrasound examination can replace liver biopsy.

References

- 1- PETRUZZIELLO A., MARIGLIANO S., LOQUERCIO G., COZZOLINO A. and CACCIAPUOTI C.: Global epidemiology of hepatitis C virus infection: An up-date of the distribution and circulation of hepatitis C virus genotypes. World Journal of Gastroenterology, 22: 7824-40, 2016. https://doi.org/10.3748/wjg.v22.i34.7824.
- 2- COOKE G.S., LEMOINE M., THURSZ M., GORE C., SWAN T., KAMARULZAMAN A., et al.: Viral hepatitis and the Global Burden of Disease: A need to regroup. Journal of Viral Hepatitis, 20: 600-1, 2013. https://doi.org/ 10.1111/jvh.12123.
- 3- MOHAMOUD Y.A., MUMTAZ G.R., RIOME S., MILL-ER D. and ABU-RADDAD L.J.: The epidemiology of hepatitis C virus in Egypt: a systematic review and data synthesis. BMC Infectious Diseases, 13: 288, 2013. https://doi.org/10.1186/1471-2334-13-288.
- 4- ELGHARABLY A., GOMAA A.I., CROSSEY M.M.E., NORSWORTHY P.J., WAKED I. and TAYLOR-ROBINSON S.D.: Hepatitis C in Egypt - past, present, and future. International Journal of General Medicine, 10: 1-6, 2017. https://doi.org/10.2147/IJGM.S119301.
- 5- DITAH I., DITAH F., DEVAKI P., EWELUKWA O., DITAH C., NJEI B., et al.: The changing epidemiology of hepatitis C virus infection in the United States: National health and nutrition examination survey 2001 through 2010. Journal of Hepatology, 60: 691-8, 2014. https://doi.org/10.1016/j.jhep.2013.11.014.
- 6- LU M., LI J., RUPP L.B., ZHOU Y., HOLMBERG S.D., MOORMAN A.C., et al.: Changing trends in complications of chronic hepatitis C. Liver International, 38: 239-47, 2018. https://doi.org/10.1111/liv.13501.
- 7- ANDRADE L. DE O, D'OLIVEIRA A., MELO R., DE SOUZA E., SILVA C.C. and PARANA R.: Association between hepatitis C and hepatocellular carcinoma. Journal of Global Infectious Diseases 2009. https://doi.org/ 10.4103/0974-777X.52979.
- 8- RAZAVI H., ELKHOURY A.C., ELBASHA E., ESTES C., PASINI K., POYNARD T., et al.: Chronic hepatitis C virus (HCV) disease burden and cost in the United States. Hepatology 2013. https://doi.org/10.1002/ hep.26218.
- 9- MAÀJI S.M., YAKUBU A. and ODUNKO D.D.: Pattern of abnormal ultrasonographic findings in patients with

clinical suspicion of chronic liver disease in Sokoto and its environs. Asian Pacific Journal of Tropical Disease, 3: 202, 2013. https://doi.org/10.1016/S2222-1808 (13)60041-9.

- 10-CELLE G., SAVARINO V., PICCIOTTO A., MAGNOLIA M.R., SCALABRINI P. and DODERO M.: Is hepatic ultrasonography a valid alternative tool to liver biopsy? Report on 507 cases studied with both techniques. Digestive Diseases and Sciences, 33: 467-71, 1988. https://doi.org/10.1007/BF01536033.
- 11-FONTANA R.J. and LOK A.S.F.: Noninvasive monitoring of patients with chronic hepatitis C. Hepatology (Baltimore, Md), 36: s57-64, 2002. https://doi.org/10.1053/ JHEP.2002.36800.
- 12- JAVA: Declaration of Helsinki World Medical Association Declaration of Helsinki. Bulletin of the World Health Organization, 79: 373-4, 2013. https://doi.org/S0042-96862001000400016 [pii].
- 13- VON ELM E., ALTMAN D.G., EGGER M., POCOCK S.J., GØTZSCHE P.C. and VANDENBROUCKE J.P.: The strengthening the reporting of observational studies in epidemiology (STROBE) statement: Guidelines for reporting observational studies. International Journal of Surgery 2014. https://doi.org/10.1016/j.ijsu.2014.07.013.
- 14- CHOONG C-C., VENKATESH S.K., SIEW E.P.Y., CHOONG C-C., VENKATESH S.K. and SIEW E.P.Y.: Accuracy of Routine Clinical Ultrasound for Staging of Liver Fibrosis. Journal of Clinical Imaging Science, 2: 58, 2012. https://doi.org/10.4103/2156-7514.101000.

- 15- WEICHERT U., BUTTMANN A., JAKOBS R., SCHILL-ING D., EICKHOFF A. and RIEMANN J.F.: Diagnosis of liver cirrhosis: A comparison of modified ultrasound and laparoscopy in 100 consecutive patients. Journal of Clinical Gastroenterology, 39: 529-32, 2005. https://. doi.org/10.1097/01.MCG.0000165669.17649.20.
- 16- NISHIURA T., WATANABE H., ITO M., MATSUOKA Y., YANO K., DAIKOKU M., et al.: Ultrasound evaluation of the fibrosis stage in chronic liver disease by the simultaneous use of low and high frequency probes. The British Journal of Radiology, 78: 189-97, 2005. https://doi.org/ 10.1259/BJR/75208448.
- 17- CHOONG C-C., VENKATESH S.K. and SIEW E.P.Y.: Accuracy of routine clinical ultrasound for staging of liver fibrosis. Journal of Clinical Imaging Science, 2: 58, 2012. https://doi.org/10.4103/2156-7514.101000.
- 18- ALLAN R., THOIRS K. and PHILLIPS M.: Accuracy of ultrasound to identify chronic liver disease. World Journal of Gastroenterology, 16: 3510-20, 2010. https:// doi.org/10.3748/WJG.V16.128.3510.
- 19- LEE H.S., KIM J.K., CHEONG J.Y., HAN E.J., AN S.Y., SONG J.H., et al.: Prediction of compensated liver cirrhosis by ultrasonography and routine blood tests in patients with chronic viral hepatitis. The Korean Journal of Hepatology, 16: 369-75, 2010. https://doi.org/ 10.3 3 50/KJHEP.2010.16.4.3 69.
- 20- MARTÍNEZ S.M., CRESPO G., NAVASA M. and FORNS X.: Noninvasive assessment of liver fibrosis. Hepatology (Baltimore, Md), 53: 325-35, 2011. https://doi.org/10.1002/HEP.24013.

التغيرات التي تحدث في الموجات الصوتية على البطن في مرضى الالتهاب الفيروسي الكبدي سي المزمن

الهدف من هذه الدراسة هو تحديد التغيرات التى تحدث فى الأشعة الموجات الصوتية على البطن فى مرضى التهاب الفيروس الكبدى سى المزمن وتعتبر الموجات الصوتية من أرخص وأسرع الطرق التى تستخدم لأكتشاف التغييرات التى تطرأ فى مرضى فيروس سى المزمن ومن هذه التغيرات التى تطرق على الكبد (دهون الكبد – تليف الكبد – أورام الكبد).

التغيرات التي تطرق على الطحال من تضخم وارتفاع ضغط الوريد البابي الكبدي والاستصقاء الغشاء البرويتوني.

هذه دراسة مسحية مسبقة تم اجرائها من الفترة يونيو ٢٠١٤ – يونيو ٢٠١٥ فى مستشفيات كلية طب القصر العينى وقد شملت ثلاثمائة مريض مصاب بالتهاب الكبد الفيروسى نوع سى جرى تعريضهم لسلسلة من الفحوصات السريرية والمناعية بالاضافة إلى التشخيص بالموجات الصوبية.

وتعتبر أيضاً الأشعة التلفزيونية من الفحوصات الدورية التى يتم عملها لمرضى التهاب الفيروس الكبدى سى وتليف الكبد كل ٦ شهور لمتابعة ظهور أى أورام بالكبد وأى مضاعفات من التليف.

وتبين من هذه الدراسة أوضحت تغيرات في أكثر من ٦٠٪ من مرضى التهاب الفيروس الكبدي سي لذا يعتمد عليها في التشخيص والمتابعة.