

Efficacy of Education, Continuous Monitoring and Nutritional Care on Quality of Life of Cirrhotic Patients and Reducing of Liver Cirrhosis Complications

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Background and study aim: Liver cirrhosis is a complex chronic disease with a wide variety of morbidities and these complications directly affect the quality of life and result in significant morbidity and mortality. This study designed to determine the effects of an educational awareness program and continuous monitoring on quality of life (QOL) among cirrhotic patients and their complications.

Methods: Overall 18 cirrhotic patients included. An educational awareness program including a face-to-face training session, providing a guide booklet about caring of the liver cirrhosis, consulting with dietician and continuous monitoring of patients every other week was conducted and the patients followed for 6 months. The QOL, awareness score, complications of liver disease, severity of disease by using MELD score and laboratory parameters

were evaluated at baseline and at the end of intervention.

Results: At the end of the intervention, the prevalence of complications, including edema, ascites, hepatic encephalopathy, variceal bleeding and hospitalization rate were significantly decreased compare to baseline ($P < 0.05$). The average scores of awareness and QOL improved significantly (52.78 ± 20.36 to 88.89 ± 10.49 ; $P = 0.0001$ and 141.89 ± 20.40 vs. 182.72 ± 10.27 ; $P = 0.0001$ respectively). The laboratory parameters and the severity of the disease did not show any significant changes at the end of the study ($P > 0.05$).

Conclusion: Using a simple educational program can significantly improve not only cirrhotics awareness and QOL but also reduce complications and duration of hospitalization which could be cost effective and worth to try among cirrhotics.

INTRODUCTION

Liver cirrhosis by regeneration and fibrosis of the liver cells ultimately leads to portal hypertension and liver dysfunction [1]. This condition is a complex chronic disease with a wide variety of morbidities and estimated annual mortality rate about 5-10 cases per 100,000 people worldwide [2,3]. Nowadays, the most common causes of liver cirrhosis are alcohol-related liver disease (ARLD), non-alcoholic fatty liver disease (NAFLD) and viral hepatitis [4,5].

Hepatic encephalopathy, ascites, gastrointestinal bleeding and portal

hypertension are the most common complications of liver cirrhosis [6]. These complications directly affect the quality of life and result in significant morbidity and mortality [7]. In addition, malnutrition is one of the known complications of the liver cirrhosis and has significant prognostic effect on irreversibility of this condition and its complications [8]. Liver cirrhosis management methods, including screening of hepatocellular carcinoma (HCC) and bleeding from gastrointestinal varices often focus on risk reduction. These strategies mostly target clinical factors and do not consider other patient related factors such as

health-related quality of life (HRQOL). Measuring HRQOL can affect the disease management and is recognized as an important factor in chronic diseases such as liver cirrhosis [5].

Moreover, management of patients depends on their proper understanding of their condition and active participation in their care and decision making [9]. However, at the moment, patients' understanding and awareness about this condition is low and the reasons of this unawareness include: limited time for consultation in the clinic and the variable quality of educational resources such as websites and information leaflets [10,11]. The inherent complexity of liver cirrhosis and its complications may also limit access to information and understanding of the disease in some patients. However, there is little information and evidence about the patient's awareness about liver cirrhosis or the effectiveness of a routine educational intervention [13].

So it seems necessary to consider the importance of nutrition, counseling and awareness of patients about their disease, and its possible role for improving their quality of life. Moreover, its possible role in reducing the complications of liver cirrhosis should be emphasized. The current studies were limited to a specific range of cirrhotic patients with the same etiology as well as a certain stage of the disease according to the Child score, therefore, we conducted this study among patients who suffering liver cirrhosis with various causes and at various stages based on the MELD Score, to evaluate efficacy of close monitoring, education and nutritional support on the quality of life and their laboratory profile.

METHODS

This interventional cross sectional pilot study conducted on patients with liver cirrhosis who referred to the hepatology outpatient clinic of Ahvaz Imam Hospital in 2017. After being approved by the Ethics Committee of Ahvaz Jundishapur University of Medical Sciences (IRAJUMS. REC.1396.811), all over-18-year-old patients suffering from non-alcoholic liver cirrhosis were invited to participate without exclusion based on various stages of the Child Score. Then, 25 patients were invited to enter the study and informed consent was obtained from all of the participants. Exclusion criteria included: simultaneous medical conditions and synchronous diseases such as

uncontrolled cardiopulmonary or renal disease, presence of any kind of malignancy such as hepatocellular carcinoma, pregnancy, and non-referring or failure to follow after 6 months of intervention. This interventional study was performed on 25 cirrhotic patients, of which 7 were excluded from the study because of various reasons (death of the patient and lack of follow-up and or timely referral), and the final analysis were performed on 18 subjects. The study protocol conforms to the ethical guidelines of the 1975 Declaration of Helsinki as reflected in a priori approval by the institution's human research committee.

Before the intervention, the demographic information of each patient (including age, sex, education, marriage, etc.) and his or her clinical profile (including family history of liver disease, duration of illness, cause of illness plus results of laboratory parameters, complications of disease, quality of life, and knowledge of patients about their condition) were evaluated and recorded in each patient's special file.

The educational and awareness program delivered to all of the participants which include one-hour face-to-face awareness training session in a safe and relaxed environment. Educational content included some information about liver function, cirrhosis development with the progression of inflammation, risk factors, disease complications and management strategies. In addition, the patients were provided a booklet containing information about liver cirrhosis. Training about cirrhosis of the liver and its complications, diet, observing nutritional guidelines for reducing complications of the illness and hygiene was given to all of them. In this stage, a diet was also given to patients. In addition, they were contacted every two weeks and their questions and concerns about the disease were answered.

Patient evaluation

In all of the patients, the level of knowledge, quality of life, severity of illness, laboratory parameters (Na, K, BUN, Cr, CBC, ALT, AST, ALK P, INR, TSH, BIL) were evaluated at the beginning of the study and six months after the intervention (end of study). Body mass index (BMI) determined as Kg/m^2 . The frequency of hospitalization and complications such as incidence of gastrointestinal bleeding, hepatic encephalopathy, severity of ascites and edema, and other complications such as electrolyte and

laboratory abnormalities over a 6 months period were investigated, and they were compared with their same results during 6 months before of the study period.

Quality of life and disease severity assessment

Severity of the liver cirrhosis was determined based on the Model for End-Stage Liver Disease (MELD) scale. The chronic liver disease questionnaire (CLDQ) was used to assess the quality of life of the patients [14]. This questionnaire has been widely used as a health-related quality of life assessment (HRQOL) tool and its validity has been confirmed in chronic liver disease with different etiologies and severity and in different languages [14-16]. The questionnaire consists of 29 questions in 6 sections: 8 questions about emotional functioning, 3 questions about abdominal symptoms, 3 questions about activity, 5 questions about systemic symptoms, 5 questions about fatigue, and 5 questions about anxiety. The questions are measured on a 7-point Likert-scale: Always 1; Very often 2; Mostly 3; Sometimes 4; Rarely 5; hardly ever 6; Never 7 scores. The final score of QOL obtained by calculating the total score of each individual. In general, each individual's score range varies between 29 and 203, and higher scores represent better QOL [14]. The validity of this questionnaire has been confirmed in previous studies in Iran [15] and in this study, the reliability of the quality of life questionnaire was assessed by calculating Cronbach's alpha (0.84).

Awareness level

A researcher-made questionnaire was used to measure patient's awareness about liver cirrhosis, consisting of 8 two-choice questions (yes 1; no 0). The awareness level of each person was obtained by calculating the total score of each individual. This level varies from 0 to 8 for them, and a higher score indicates more awareness among patients. Content validity of the questionnaire was evaluated by supervising of two well experienced hepatologists. Reliability of the questionnaire was calculated as 0.86 by using Cronbach's alpha.

Statistical analysis

SPSS version 22 was used for statistical analysis. Data were analyzed by descriptive statistics including mean, standard deviation, frequency and percentage. The normality of data was assessed

by Kolmogorov-Smirnov test and homogeneity of variances by Leven test. Moreover, Wilcoxon singled-rank test was used to compare the mean of variables before and after intervention and chi-square test was used to compare qualitative variables. The significance level in the tests was considered to be 0.05.

RESULTS

Average age of the participants was 47.00 ± 17.42 y (18 to 83 years), which included 4 women and 14 men. The average duration of the disease from first diagnosis to participation in study was 2.52 ± 2.32 months (1 month to 9 years). The baseline characteristics of the patients are presented in Table (1).

After the completion of the study, the average time of participants hospitalization significantly decreased (Before the study 1.28 ± 1.22 days (0 to 4) and at the end of the study 0.33 ± 0.59 (0-2) days, $P = 0.001$). Before study, 55.6% of patients had edema while after 6 months of close monitoring, this symptom decreased to 16.7%. In addition, at the end of the study, the prevalence and incidence of bleeding from gastrointestinal varices and hepatic encephalopathy were also decreased. Hypothermia and serum potassium disturbance were not observed in any of the patients at the beginning and at the end of the intervention. Comparison of the liver cirrhosis complications before and after intervention is presented in Table (2).

As presented in Table 3, the laboratory values at the end of the study did not show any significant changes ($P > 0.05$). The severity of liver cirrhosis based on MELD score did not change significantly at the end of the study ($P > 0.05$) (Table 4).

As presented in Table 5, the level of patient's awareness before and after the intervention was 52.78 ± 20.36 and 88.89 ± 10.40 , respectively (average 36.11% improve, $P < 0.0001$). The quality of life of the patients before and after the intervention was $69.9\% \pm 10.05$ and 90.01% , respectively (20.11% improvement). The dimensions of QOL including emotion, systemic and abdominal symptoms, activity, fatigue, and anxiety significantly improved at the end of the study ($P < 0.0001$, Fig. 1).

Table (1) : Basic characteristics of the participants (BMI: Body Mass Index)

Characteristics	Frequency (percent)
Sex	4 (22.2 %)
Female	14 (77.8 %)
Male	
Marital status	3 (16.7 %)
Single	15 (83.3 %)
Married	
Education	
Illiterate	2 (11.1 %)
Sub diploma	9 (50.0 %)
Diploma	4 (22.2 %)
Academic education	3 (16.7 %)
Weight (kg)	68.09 ± 16.04 (45 -110)
Height (cm)	170.00 ± 10.49 (152 – 187)
BMI (Kg / m ²)	24.06 ± 5.07 (16.46- 35.51)
History of alcohol consumption	2 (11.1 %)
Smoking history	6 (33.3 %)
Family history of liver disease	3 (16.7 %)
Hospitalization history (yes)	9 (50%)
Background of the underlying disease	
Diabetes Melitus	3 (16.7 %)
Chronic kidney disease	2 (11.1 %)
Others	3 (16.7 %)
Cause of disease	
Hepatitis C	8 (44.4%)
Autoimmune hepatitis	3 (16.7)
Hepatitis B	2 (11.1)
Fatty Liver	1 (5.6)
Unknown	4 (22.2)

Table (2) : Comparison of complications before and after intervention

Complication of liver cirrhosis	Before intervention	After intervention	P-value
Ascites			
Yes	10 (55.6 %)	2 (11.1 %)	0.005
No	8 (44.4 %)	16 (88.9 %)	
Edema			
No	8 (44.4 %)	15 (83.3 %)	0.002
Intensity of edema (+1)	7 (38.9 %)	3 (16.7 %)	
Intensity of edema (+2)	3 (16.7 %)	0	
Bleeding from gastrointestinal varices			
Yes	2 (11.1 %)	0	0.157
No	16 (88.9 %)	18 (100 %)	
Hepatic encephalopathy			
Yes	2 (11.1 %)	0	0.157
No	16 (88.9 %)	18 (100 %)	

Table (3) : Laboratory values before and at the end of the study

Variable	Before intervention	After the intervention	*P-value
BUN (mg / dL)	16.12±10.5	17.48±8.11	0.397
Cr (mg / dl)	0.93±0.37	1.09±0.36	0.164
Na (mEq/L)	138.71±2.8	139.58±2.71	0.246
K (mEq/L)	4.02±0.38	4.11±0.35	0.135
Hb (g / dL)	11.73±2.36	11.47±1.41	0.34
Ca (mg / dL)	8.84±0.86	9.08±0.24	0.854
AFP (µg / L)	10.45±8.16	11.33±7.61	0.735
ALT (IU / L)	52.11±29.96	43.46±20.40	0.473
AST (IU / L)	64.72±36.45	53.92±18.68	0.397
ALK P (IU / L)	311.44±215.50	271.5±138.35	0.831
INR	1.34±0.42	1.29±0.29	0.656
TSH (µg / mL)	4.51±2.92	4.14±3.85	0.893
Bil (total) (mg / dL)	1.89±1.33	1.60±0.83	0.138
Bil (direct)	0.59±0.58	0.61±0.35	0.326

(BUN: Blood Urea Nitrogen; Cr: Creatinine; Hb: hemoglobin; AFP: alpha-fetoprotein; ALT: Alanine aminotransferase; AST: Aspartate aminotransferase; ALK P: Alkaline Phosphatase; INR: International Normalized Ratio; TSH: Thyroid-Stimulating Hormone; BIL: Bilirubin); Numbers are presented as mean ± standard deviation.

* Wilcoxon signed-rank test was used to compare the mean before and after intervention

Table (4) : Severity of liver cirrhosis before and after intervention based on MELD scoring system (MELD: model for end-stage liver disease)

MELD Score	Before intervention	After intervention	*P-value
Average	11.14±3.59	12.17±4.84	0.552
Median	11.5	11	
Range (Min-Max)	6 - 17	6 - 21	

Table (5) : Mean score of awareness and quality of life of participants before and after intervention (All of the variables are presented as Mean ± standard deviation (Min-Max))

	Before intervention	After intervention	Difference	P-value
Awareness level	4.22±1.62 (0-7)	7.11 ± 0.83 (5-8)	2.88±0.96 (1-5)	0.0001>
Quality of Life	141.89±20.40 (102-170)	182.72±10.27 (167-198)	40.83 ± 17.73 (6-71)	0.0001>

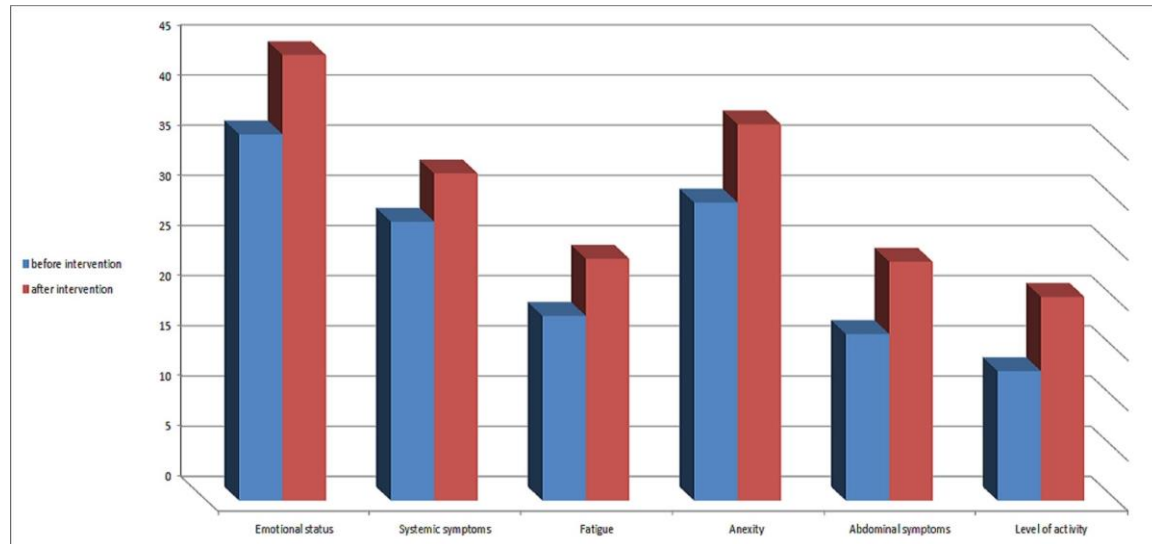


Figure (1) : Comparison of dimensions of quality of life in cirrhotic patients before and after intervention

DISCUSSION

Patients with chronic liver disease could experience different conditions and need to be aware of disease management course and treatment strategy. Usually in clinical practice, it is not easy for physicians to provide the patients enough information during their routine visiting sessions, and the time allocated to each patient is very limited [17]. The aim of the current study was to investigate the effect of awareness, continuous monitoring and nutritional care on QOL and rate of the liver cirrhosis complications.

The results of this study showed that at the end of the intervention, liver cirrhosis complications, including the rate and severity of edema, the incidence of ascites, bleeding from gastrointestinal varices and hepatic encephalopathy decreased compared to the beginning of the study. Moreover, after the completion of the study, the average time of hospitalization decreased significantly. The increase in the level of patient's awareness consequently resulted in the reduction of complications and duration of hospital stay which could be cost effective.

Volk et al. [18] reported that improving the patient's understanding of the condition could significantly reduce the hospital readmissions among patients with decompensated cirrhosis. In addition, by increasing the level of awareness of patients, they may benefit from better clinical outcomes. For example, Serper et al. [19] found that patients who receive liver transplant had more adherences to treatment and had less need

to refer to the hospital and be hospitalized with better knowledge about the treatment regimen.

On the other hand, Wigg et al. [20] investigated the efficacy of a chronic disease management model and demonstrated that the intervention had no significant effect on reducing the number of hospitalization days and the severity of the disease or improving the QOL in patients with chronic liver failure which contradict with the findings of the present study. The reason of this difference could be different study population as well as the type of intervention. In the study by Wigg et al. [20] the intervention was conducted in areas of self-management support, decision support, and clinical information systems; whereas, in the present study, increasing patients' awareness about the disease, nutritional support and continuous monitoring of the patients were conducted for six months.

In the current study, laboratory parameters and severity of liver cirrhosis did not change significantly based on MELD score at the end of intervention though the lack of laboratory improvement should be attributed to the progressive nature of the liver cirrhosis. Moreover, patients' awareness about liver cirrhosis was low at the beginning of the study and after educating and informing, the level of awareness of participants increased significantly (36.11% rising in awareness). During the study period, patients had been permitted to obtain information from other sources about their condition, which may contribute to increasing the knowledge of

patients. Nevertheless, from a practical perspective, this factor has no significant role.

There are limited studies about patients' awareness about liver cirrhosis or the efficacy of a routine educational intervention. In a study by Volk et al. [21], patients' awareness about self-management of liver cirrhosis was reviewed at a U.S.-based health facility. The results indicated a poor awareness level at the beginning of the study and 26% improvement of knowledge of the patients after a simple nutritional and non-nutrition education intervention for 3 months by using a brief booklet. This increase in awareness about the management of liver cirrhosis is valuable because this intervention was done by using very few resources. However, it was unclear whether this level of awareness would lead to improved outcomes. In another study, Goldsworthy et al. [13] also showed understanding and awareness of patients about liver cirrhosis was poor at the beginning of the study and after using multimedia education and informing patients, the score of patients' awareness questionnaire increased significantly compared to the beginning of the intervention by 41.7%. Therefore, the conducted intervention in this study was an effective way to empower patients with liver cirrhosis. In addition, Kadokawa et al. [17] investigated the effectiveness of conducting chronic liver disease education classes for informing patients with chronic hepatitis and liver cirrhosis. The level of awareness of the participants in the study improved significantly after participating in the classes, moreover, the recovery rate depended on the number of class attendance. These results are consistent with the findings of the present study.

One of the reasons for the poor level of awareness about liver disease in this study, as well as other studies, is difficulty in remembering the information given by the responsible doctors in routine visits; or insufficiency of the provided information. However, these two reasons are not exclusive. Information given in routine clinical visits usually focuses on commonly used treatment regimens as well as alternative therapies that may be implemented in the near future, and less attention is paid to issues such as long-term care and or other details. This is partly due to the limited time assigned for counseling into the routine clinical program. In this situation, patients and their family members may not be aware about the importance of the information presented to them. It should also be noted that it is difficult to

measure and correct or change patients' awareness or beliefs about their disease and treatment due to several interfering factors such as health literacy, beliefs about health, medications and treatment, the relationship between the patient and the physician (such as the quality of education and communication), self-efficacy, and the impact of the other internal and external barriers [22-24]. These barriers may include social or economic factors, physician-related or healthcare-related factors, and patient-related factors, and could potentially affect adherence to treatment [25]. Therefore, further studies are needed to identify the factors affecting patients' awareness and following of the given instructions.

Another finding of this study was improving of the QOL of cirrhotic patients which increased by 11.1% after intervention. Various studies have shown that protein-energy malnutrition is associated with reduced survival in patients with liver cirrhosis [7,26]. Optimal nutrition status is essential in managing patients with advanced liver disease. Patients with liver cirrhosis are suffering from malnutrition which result in increased morbidity and reduced QOL due to adverse side effects [7]. In a randomized controlled study in India by Maharshi et al. (2016), patients with liver cirrhosis were evaluated in two groups of nutrition support and control for 6 months. The results showed that the QOL associated with health in the intervention group was significantly improved compared to the control group [27]. In another study, Zandi et al. [15] showed that implementing self-care education programs and continuous monitoring of cirrhotic patients for three months would significantly improve the QOL of these patients. These results are consistent with the findings of the present study.

In sum, the results obtained in this study and similar studies show that patients' QOL improves significantly after the intervention by education and informing. Therefore, educational interventions can be used as a useful way to increase the QOL of cirrhotics in clinical setting.

One of the main limitations of this study is the using self-assessment method and no objective measurement to assess the level of awareness. Other limitations of this study include being a single-center study and low number of participants, being a pilot study, lack of long-term follow-up and not considering the control group to compare the effectiveness of intervention results due to

ethical issues. Among the uncontrollable limits of this study, we can mention the previous knowledge and experience, and the motivation and interest of patients who are effective in implementing the program.

CONCLUSION

The results of this study showed that patients' awareness about liver cirrhosis could be improved through a simple educational intervention and continuous monitoring for 6 months and would result in fewer side effects, reduced the number of hospitalization days and improved QOL of the patients. By improving the patients' QOL, they would encourage to prevent leaving of treatment. More studies are required to determine the exact impact of more intensive educational interventions on improving results, adhering to the treatment, and lowering medical costs in patients with liver cirrhosis.

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Conflict of interest

The authors declare to have any conflict of interest. While this research involving Human Participants, we did not prescribed them any medicine or supplementary material with potential hazards and all of the participants requested to sign an informed consent before inclusion in study. This study has been done under supervision of Alimentary Tract Research Center without any financial funding and approved by Ahvaz Jundishapur University of Medical Sciences ethical committee (IRAJUMS. REC.1396.811).

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