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ORIGINAL RESEARCH

Efficacy of direct-acting antiviral therapy for hepatitis C viral infection. Real-life experience in Bahrain

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Purpose: The introduction of direct-acting antivirals (DAAs) has revolutionized the treatment of chronic hepatitis C viral (HCV) infection. This study aims to establish real-world treatment efficacy of Sofosbuvir-based (SOF-B) and Ombitasvir/Paritaprevir/Ritonavir-based (OPR-B) regimens.

Patients and methods: This prospective, non-randomized observational real-life study was conducted in Salmaniya Medical Complex, Bahrain, and included consecutive patients with chronic HCV infection (genotypes 1–4) who were treated with direct-acting antivirals. Sustained virologic response to therapy was assessed at week 12 post end of treatment (SVR12).

Results: Of the 167 patients included, 60.5% (n=101) were treated with SOF-B and 39.5% (n=66) with OPR-B regimens for 12 weeks (n=148; 88.6%) or 24 weeks (n=19; 11.4%). SVR12 was achieved in 156 (93.4%) patients, 4 patients failed to achieve SVR despite completion of treatment, and 7 patients discontinued treatment due to non-compliance and were included in the analysis on an intention-to-treat basis. There was no difference between SOF-B and OPR-B regimens (95/101; 94.1%) and (61/66; 92.4%), respectively (p=0.68). However, SVR12 rates were significantly higher in patients without liver cirrhosis (103/104; 99.0%) compared to patients with cirrhosis (53/63; 84.1%; p<0.001), and in patients who received 12-week-regimen (141/148; 95.3%) compared to those who received 24-week regimen (15/19; 78.9%; p<0.024). However, logistic regression analysis identified cirrhosis at baseline to be the only independent predictor of non-SVR12 (OR: 16.1, 95% confidence interval 1.96–131.91, p=0.01). Apart from Hb, INR, and ALP, all other laboratory parameter improved following treatment (p<0.05).

Conclusion: Both SOF-B and OPR-B regimens achieved high SVR12 rates in this real-life cohort of patients with chronic HCV infection, similar to what is reported in other real-world studies. Cirrhosis was the only independent predictor of poor response.

Keywords: HCV, DAAs, treatment, sustained viral response, cirrhosis, liver disease

Introduction

Hepatitis C virus (HCV) infection represents a serious challenge to global health, with an estimated worldwide prevalence of 71.1 million[1] even though it has dropped significantly from the estimated 170 million a decade ago.[2] A substantial percentage of patients with chronic HCV infections develop significant complications, mainly chronic hepatitis C (CHC), liver cirrhosis, liver cell failure, and hepatocellular carcinoma (HCC).[3–7]

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The primary goal of HCV treatment is to achieve a sustained virologic response (SVR), which is defined as undetectable HCV RNA levels at 12 weeks (SVR12) or 24 weeks (SVR24) after the completion of treatment.[8] The achievement of SVR in patients with HCV infection is associated with infection eradication, improvement in their quality of life, and a reduced risk of complications including cirrhosis and HCC.[9,10]

Pegylated interferon-based therapy was the standard-of-care (SOC) therapy for HCV infection for nearly 2 decades. However, the introduction of 2nd generation direct-acting antivirals (DAAs) and interferon-free regimens represents the beginning of a new era and a revolution in the treatment of HCV. In a systematic review and a network meta-analysis of 27 randomized controlled trials (RCTs) involving 3415 patients with CHC treated with different DAA regimens, the SVR ranged from 94% to 99%, the greatest rates for patients without cirrhosis being estimated for those receiving sofosbuvir + velpatasvir with ribavirin for 12 weeks (99%; 95% Credible Intervals, 98–100%), and those with cirrhosis receiving sofosbuvir + velpatasvir for 24 weeks (96%; 95% CrI, 92-–99%). Ribavirin increased efficacy in patients with and without cirrhosis (Odds Ratio, 2.6–4.5).[11] However, real-life results concerning the efficacy of such therapies for HCV are still scarce. In fact, efficacy rates reported in RCTs can be lower in community-based practice settings due to concomitant diseases or constitutional factors. Knowledge of these factors is valuable for the future management of affected patients.[12]

In this study, we share our clinical experience in treating patients with chronic HCV infection and evaluate treatment efficacy on a real-life practical ground. The objectives were to ascertain the SVR12 in consecutive patients treated at our facility and identify which factors are associated with better sustained virologic rates.

Material and methods

Study design

This was a prospective, non-randomized, observational single-center cohort study.

Patients

All consecutive patients who started treatment for HCV infection (genotypes 1–4) at the Salmaniya Medical Complex hospital were included in this study and followed up from January 2016 to September 2017. Patients who had not received any prior treatment (treatment-naïve) and those who had (treatment-experienced) were both included, as well as patients with hepatocellular carcinoma. Patients with concomitant hepatitis B virus and/or HIV infections were excluded.

Methods

All patients were subjected to thorough history talking (age, sex, history of diabetes mellitus, hypertension, liver transplantation, hyperlipidemia, hypothyroidism, end-stage renal disease, renal transplantation, sickle-cell disease, other systemic comorbidities) and full clinical examination.

At baseline (pre-treatment) and 12 weeks after the end of therapy, the following laboratory investigations were performed: HCV antibody, HCV RNA PCR quantitation, complete blood count, international normalization ratio (INR), partial thromboplastin time (PTT), serum creatinine, serum albumin, total serum bilirubin, alanine aminotransferase (ALT), alkaline phosphatase (ALP), and gamma-glutamyl transferase (γ-GT). HCV genotype was performed only at baseline.

HCV genotyping and consolidated HCV viral load estimation were performed using a fully automated Abbott m2000 machine along with the manufacturer supplied reagent kits (Abbott Molecular, Abbott Park, IL, USA). This assay quantifies HCV RNA using in vitro reverse transcription-polymerase chain reaction (PCR) method, and it has a sensitivity of 12 IU/mL for 0.5 mL and 30 IU/mL for 0.2 mL sample volume with a detection range of 12 IU/mL (log 1.08 IU/mL) to 100 million IU/mL (log 8.0 IU/mL). It detects genotypes 1–6 with a specificity of ≥99.5%. Genotyping was performed using standard oligonucleotide-specific primers through PCR.

A baseline abdominal ultrasound was performed to evaluate the presence of cirrhosis and its complications (shrunken liver, coarse echotexture, irregular surface, dilated portal vein, ascites, splenomegaly).

Efficacy and safety assessment

Sustained virologic response to therapy was assessed at week 12 post the completion of treatment (SVR12) by HCV RNA PCR quantitation. Patients were followed up regularly for adverse events or abnormal findings on physical examination and clinical laboratory tests. They were seen fortnightly during the first 4 weeks of treatment, then every 4 weeks till the end of treatment, and 12 weeks after end-of-treatment.

Ethical considerations

The study was approved by the local institutional research ethics and scientific committees of Salmaniya Medical Complex Hospital. This work was conducted in accordance with the Declaration of Helsinki (2013) and the International Conference on Harmonization Guidelines for Good Clinical Practice (ICHG-GCP). A written informed consent was obtained from all participants, and their data sheets were coded to ensure anonymity and confidentiality.

Statistical analysis

Data were collected, revised, coded, and analyzed with the statistical software SPSS (Statistical Package for Social Sciences) Version 16.0 (SPSS, Chicago, IL, USA). Descriptive analysis of data was in the form of percentages, mean, or medians, and data are expressed as mean ±standard deviation (SD) or number and percentages (%) as appropriate. An intention-to-treat analysis was performed. For numerical data, univariate analysis was performed for all independent variables using two sample *t*-tests, Wilcoxon Signed Ranked, or Mann–Whitney U tests as appropriate. For categorical data, univariate binary logistic regression analysis was performed for all independent variables using Chi-Square or Fisher's exact test as appropriate. Based on the variables that showed statistical significance in the univariate analysis, multiple logistic regression analysis with the forward stepwise variable selection was used to identify the independent predictors impacting response to treatment. A *p*-value of <0.05 was set as a level of significance.

Results

Baseline (pre-treatment) patients' characteristics

Baseline demographic, comorbidities, virologic, and laboratory characteristics of the cohort of the study are shown in Table 1. A total of 167 patients were included in this study. Their mean age was 50.9±12.4 years, 91 (54.5%) were males, 55 (32.9%) were diabetic, 41 (24.6%) were hypertensive, 63 (37.7%) were cirrhotic, 21 (12.6%) had liver transplantation, 14 (8.4%) had hypothyroidism, 14 (8.4%) had sickle cell disease, 31 (18.6%) had hyperlipidemia, 8 (4.8%) had end-stage renal disease, and 3 (1.8%) had renal transplant. One patient had HCC and he achieved SVR. High baseline (pre-treatment) viral RNA load ≥400,000 IU/L was detected in 115 (68.9%) of patients. Genotype 1, 2–3, and 4 were detected in 122 (73.1%), 22 (13.1%), and 23 (13.8%) patients, respectively.

Table 1 Baseline (pre-treatment) patients' characteristics (n=167)

Variable	Unit or category	Result
Age	Years	50.9±12.4
	<40 years	34 (20.4)
	≥40 years	133 (79.6)
Sex	Male:Female	91 (54.5):76 (45.5)
Cirrhosis		63 (37.7)
Diabetes mellitus		55 (32.9)
Hypertension		41 (24.6)
Liver transplant		21 (12.6)
ESRD		8 (4.8)
Renal transplant		3 (1.8)
Hyperlipidemia		31 (18.6)
Hypothyroidism		14 (8.4)
Sickle-cell disease		14 (8.4)
Viral load	(IU/mL)	1.35E+6±1.9E+6
	Log10 viral load	5.81±0.66
	<400,000	6.52 (31.1)
	≥40,000	115 (68.9)
HCV genotype	G1	122 (73.1)
	G2-3	22 (13.1)
	G4	23 (13.8)
WBCs	(x109/L)	5.81±2.16
Hemoglobin	(gm/dL)	12.78±2.10
Platelets	(x109/L)	191.83±95.49
INR		1.29±2.10
PTT	Second	26.10±3.68
Serum creatinine	(µmol/L)	88.05±140.73
Serum albumin	(gm/L)	38.79±5.85
Total bilirubin	(µmol/L)	18.73±17.86
ALT	(IU/L)	61.00±40.35
ALP	(IU/L)	102.66±44.47
γGT	(IU/L)	97.69±89.27

Note: Data expressed as mean±SD or number (%) as appropriate.
Abbreviations: ALP, alkaline phosphatase; ALT, alanine aminotransferase; ESRD, end-stage renal disease; HCV, hepatitis C virus; INR, international normalization ratio; γGT, gamma-glutamyl-transferase; PTT, partial thromboplastin time.

Types, duration, and combinations of treatment regimens

Ombitasvir/Paritaprevir/Ritonavir-based (OPR-B) and Sofosbuvir-based (SOF-B) regimens were given to 66 (39.5%) and 101 (60.5%) patients, respectively (Table 2). A total of 148 (88.6%) patients received therapy for 12 weeks and the remaining 19 (11.4%) patients were treated for 24 weeks. In addition, 120 (71.9%) patients were treated

Table 2 Types, duration, and combinations of treatment regimes

Type	OPR-based SOF-based	66 (39.5) 101 (60.5)
Duration	12 weeks 24 weeks	148 (88.6) 19 (11.4)
Combination	Ombitasvir/Paritaprevir/Ritonavir + Dasabuvir Ombitasvir/Paritaprevir/Ritonavir + Dasabuvir + Ribavirin Ombitasvir/Paritaprevir/Ritonavir + Ribavirin Sofosbuvir + Daclatasvir Sofosbuvir + Daclatasvir + Ribavirin Sofosbuvir + Pegylated Interferon + Ribavirin Sofosbuvir + Ledipasvir + Ribavirin Sofosbuvir + Ledipasvir Sofosbuvir + Simeprevir + Ribavirin Sofosbuvir + Ribavirin	27 (16.2) 28 (16.8) 11 (6.6) 5 (3.0) 26 (15.6) 1 (0.6) 50 (29.9) 15 (9.0) 1 (0.6) 3 (1.8)
Ribavirin included?	**Yes** With SOF-based regimens With OPR-based regimens **No**	120 (71.9) 81 (67.5) 39 (32.5) 47 (28.1)

Note: Data expressed as n (%).
Abbreviations: SOF; sofosbuvir. OPR; Ombitasvir/Paritaprevir/Ritonavir.

concomitantly with ribavirin, 81 (67.5%) of those received the SOF-B regimen and 39 (32.5%) the OPR-B regimen.

Response to therapy

SVR was achieved in 156 (93.4%) patients, and only 11 (6.7%) did not. The actual characteristics of the 11 patients with no response to therapy are shown in Table 3. Of them, 7 patients had adherence issues although they did not report any side effects, and 4 failed to achieve SVR despite completion of their regimens. Among the 7 who stopped therapy and did not complete their course, 4 were on SOF-B regimens and 4 were on a 24-week regimen. Two of the patients who completed treatment were on a SOF-B regimen while the other 2 were on an OPR-B regimen. A higher proportion of patients who achieved SVR12 had a genotype 1&4 infection (87.2%) compared to those who did not respond (81.8%); however, this was not statistically significant (p=0.611, Table 4).

Comparison between patients according to response to therapy

As shown in Figures 1 and 2 and Table 4, patients who achieved SVR (n=156) were similar to those who did not achieve SVR (n=11) regarding all demographics, comorbidities, virologic parameters, regimens used, and laboratory parameters apart from the rate of cirrhosis (p<0.001) and the duration of therapy (p=0.024). Using logistic regression analysis, only cirrhosis was found to independently predict SVR (OR: 16.09; 95% confidence interval 1.96–131.09; p=0.01) (Table 5).

Comparison between patients according to treatment regimen

As shown in Figure 3, patients who received SOF-based therapy (n=101) and those who received OPR-based (n=66) were similar regarding the rate of SVR (94.1% and 92.4%, respectively; p=0.754).

The impact of treatment on laboratory results

As shown in Table 6, successful completion of the treatment regimen with achievement of SVR in 93.4% of cases lead to a significant improvement in the mean total WBCs, platelets, and albumin levels, with significant reduction in the mean serum bilirubin, ALT, and γGT levels. However, Hb, INR, and ALP levels did not change.

Safety

The 7 patients (4.2%) who could not adhere to treatment did not report any side effects. The reduction in Hb with ribavirin was insignificant and did not lead to need for transfusion, dose reduction, nor cessation of therapy. There were no mortalities.

Discussion

Even though phase III randomized controlled trials offer robust evidence of the efficacy of drug treatments, real-life therapeutic studies, like the present study, are invaluable. The conditions in RCTs are tightly controlled, and the results may not necessarily translate into real-world outcomes where there is inability to control over patients' adherence, there is absence of inclusion and exclusion criteria, presence of variable comorbidities, and other factors which may affect the rate of SVR. In this study, we have shown that DAAs, irrespective of the type of or length of regimen used, are highly effective in achieving SVR, and the only independent predictor of response is presence of cirrhosis.

Although our experience indicates higher but not statistically significant SVR in patients infected with genotype 1&4 compared to those infected with other genotypes,

Table 3 Baseline characteristics of 11 patients with failure to achieve SVR12

Characteristics	Patient 1	Patient 2	Patient 3	Patient 4	Patient 5	Patient 6	Patient 7	Patient 8	Patient 9	Patient 10	Patient 11
Age (years)	53	61	68	61	37	52	32	52	34	47	79
Sex	M	F	F	M	M	M	M	M	F	M	F
Cirrhosis	Yes	Yes	Yes	Yes	Yes	Yes	No	Yes	Yes	Yes	Yes
DM	Yes	No	No	Yes	No	No	No	No	Yes	Yes	No
Hypertension	Yes	No	No	Yes	No	No	No	No	No	No	No
Liver Tx.	No	No	No	No	No	No	No	No	No	No	No
Renal Tx.	No	No	No	No	No	No	No	No	Yes	No	No
ESRD	No	No	No	No	No	No	No	No	No	No	No
Hypothyroidism	No	No	No	No	No	No	No	No	No	No	No
Hyperlipidemia	No	No	No	Yes	No	No	No	No	Yes	No	Yes
SCD	No	No	No	No	No	No	No	No	No	No	Yes
Genotype	3	1b	4	1a	4	1b	1a	1a	1b	1a	1b
Log_{10} viral load	6.17	5.47	6.05	5.56	6.66	5.61	6.1	*5.25*	5.40	4.17	5.84
Regimen type	SOF-B[A]	SOF-B[B]	OPR-B[C]	OPR-B[D]	SOF-B[E]	SOF-B[E]	OPR-B[F]	SOF-B[E]	SOF-B[E]	OPR-B[F]	OPR-B[F]
RBV used	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	No	Yes
Fail OR Stop	Stop	Stop	Fail	Stop	Fail	Stop	Fail	Stop	Stop	Fail	Stop
Therapy period	24	24	24	24	12	12	12	12	12	12	12

Note: The following treatments were given: [A]Sofosbuvir + daclatasvir + ribavirin for 24 weeks. [B]Sofosbuvir/Ledipasvir + Ribavirin for 24 weeks. [C]OPR + Ribavirin for 24 weeks. [D]OPR + Dasabuvir + Ribavirin for 24 weeks. [E]Sofosbuvir/Ledipasvir + Ribavirin for 12 weeks. [F]OPR + Dasabuvir + Ribavirin for 12 weeks.

Abbreviations: DM, diabetes mellitus; M, male; F, female; SCD, Sickle cell disease; ESRD, end-stage renal disease; RBV, ribavirin; SOF-B, sofosbuvir-based; OPR-B, Ombitasvir/Paritaprevir/Ritonavir based; Tx, transplantation; Stop, patient stopped therapy; Fail, patient failed to achieve SVR12.

Table 4 Comparison between patients according to response to therapy

Variable	SVR (n=156)	No SVR (n=11)	*P*-value
Age	50.81±12.31	52.36±14.56	0.882
Age ≥40 years	125 (80.1)	8 (72.6)	0.556
Male gender	84 (53.8)	7 (63.6)	0.529
Cirrhosis	53 (34.0)	10 (90.9)	<0.001
Diabetes mellitus	52 (33.3)	3 (27.3)	0.679
Hypertension	39 (25.0)	2 (18.2)	0.612
Liver transplant	21 (13.5)	0 (0.0)	0.193
End-stage renal disease	8 (5.1)	0 (0.0)	0.441
Renal transplant	2 (1.3)	1 (9.1)	0.661
Hyperlipidemia	29 (18.7)	2 (18.2)	0.965
Hypothyroidism	14 (9.0)	0 (0.0)	0.299
Sickle-cell disease	13 (8.3)	1 (9.1)	0.930
Regimen (SOF-Base)	95 (60.1)	6 (54.5)	0.677
Duration of therapy 12-weeks	141 (90.4)	7 (63.6)	0.007
Presence of Ribavirin	110 (70.5)	10 (90.9)	0.148
Log_{10} viral load	5.82±0.60	5.66±0.64	0.375
Viral load	109 (69.9)	6 (54.5)	0.520
Genotype 1 and 4	136 (87.2)	9 (81.8)	0.611
Genotype 4	21 (13.5)	2 (18.2)	0.661
Genotype 1	115 (73.7)	7 (63.6)	0.466
WBC ($x10^9$/L)	5.76±2.15	6.49±2.31	0.281
Hemoglobin (gm/dL)	12.77±2.10	12.90±2.13	0.950
Platelets ($x10^9$/L)	191.38±89.50	198.09±164.45	0.324
PT (Second)	13.31±1.84	13.61±1.66	0.399
PTT (Second)	26.06±3.75	26.77±2.58	0.437
INR	1.30±2.18	1.16±0.15	0.399
Serum creatinine (μmol/L)	89.63±145.42	65.64±19.62	0.834
Serum albumin (g/L)	38.85±5.92	38.00±4.94	0.468
Total bilirubin (μmol/L)	18.93±18.38	16.00±6.84	0.727
ALT (IU/L)	80.19±39.98	72.45±45.79	0.391
ALP (IU/L)	103.03±45.40	97.45±29.10	0.900
γGT (IU/L)	94.94±88.09	136.73±100.96	0.125

Note: Data expressed as mean±SD or n (% of SVR group) as appropriate.
Abbreviations: ALP, alkaline phosphatase; ALT, alanine aminotransferase; INR, international normalization ratio; γGT, gamma-glutamyl-transferase; PTT, partial thromboplastin time; SVR; SOF, sofosbuvir sustained virologic response; WBCs, white blood cells.

another real-life study from Hawaii contradicts these findings with lower overall SVR rates for genotypes 1 and 4 (75%) compared to genotype 3 (81%).[13] The authors attributed this to the inclusion of patients who may be excluded from clinical trials, such as those with prior treatment history, nonadherence issues, or with comorbidities that could result in discontinuation of treatment or loss to follow-up. Higher rates of noncompliance were noted among genotype 3 patients because of the longer duration of the regimen (24 weeks vs 12 weeks). In contrast, compliance was high in our cohort, and only 4 patients failed to achieve SVR despite completion of their course.

SVR of patients on SOF-based regimens was not affected by age, high viral load, or advanced fibrosis in the present cohort. Presence of comorbidities like diabetes mellitus, essential hypertension, hypothyroidism, hyperlipidemia, and sickle cell disease also did not affect the overall SVR, a similar conclusion to the Hawaii study;[13] however, the authors identified that male gender was a statistically significant factor for failure to achieve SVR, something that did not hold true in our cohort.

In another large real-world cohort study (n=485), patients on sofosbuvir and daclatasvir therapy, with or without ribavirin, achieved high SVR12 (91%) and

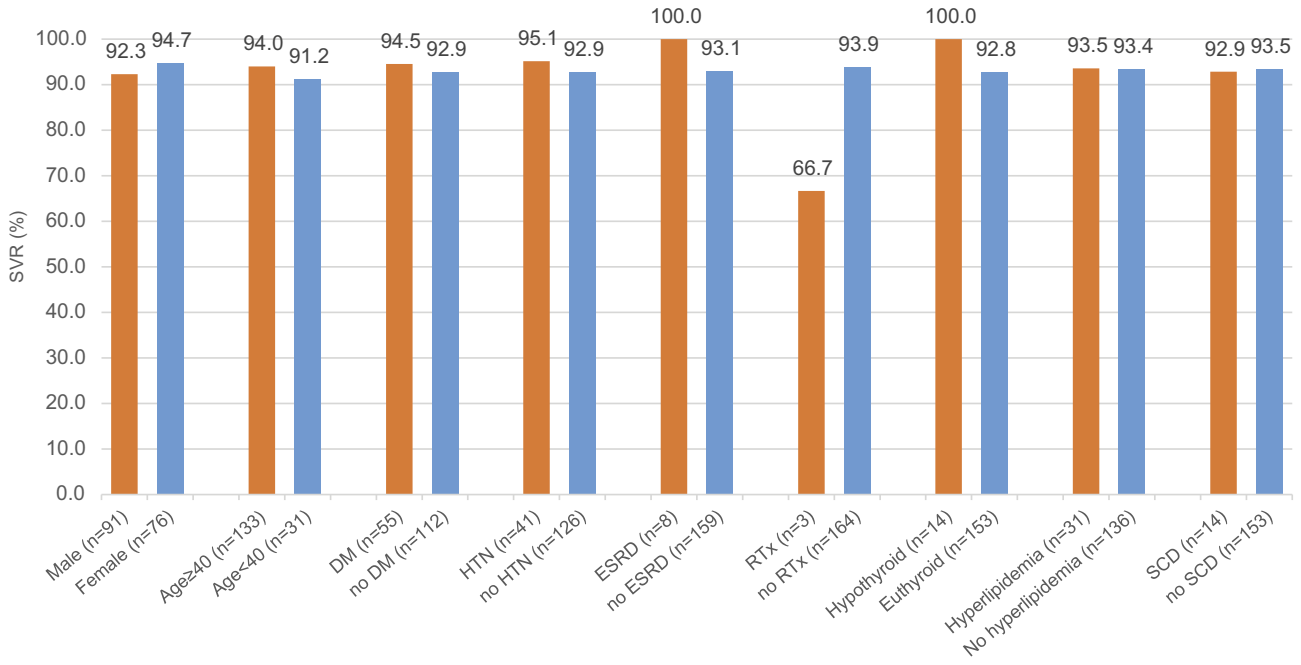

Figure 1 Sustained virologic response (SVR) rate based on different baseline patient demographic data and comorbidities.
Abbreviations: DM; diabetes mellitus. ESRD; end-stage renal disease. HTN; hypertension. RTx; renal transplant. SCD; sickle-cell disease. *P*-values are >0.05 in all by Chi-Square.

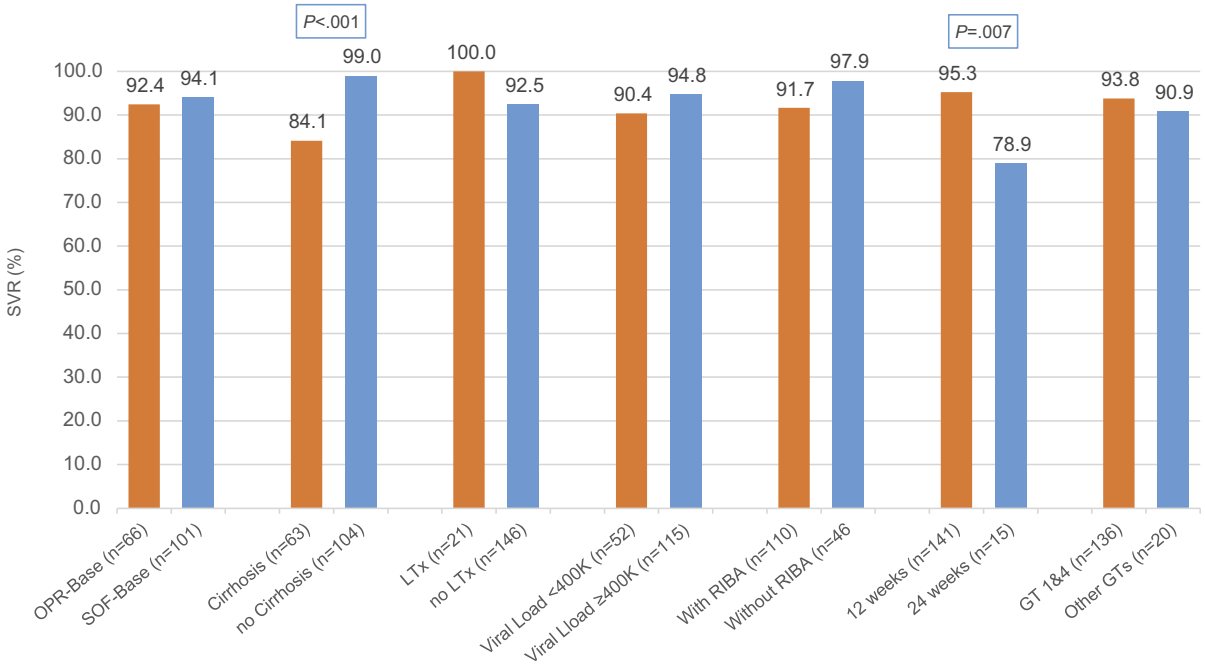

Figure 2 Sustained virologic response (SVR) rate based on viral, hepatic, and regimen parameters. SOF; sofosbuvir.
Abbreviations: OPR; Ombitasvir/Paritaprevir/Ritonavir. LTx; liver transplantation. 400 K; 400,000 IU/L. GT; genotype. RIBA; ribavirin. *P*-value by Chi-Square test.

tolerated the treatment well, regardless of HCV genotype or cirrhosis, liver transplant or HIV/HCV coinfection status, and only 28 patients discontinued treatment.[14] Our results are similar, with the exception of the influence of cirrhosis which appears to influence SVR. What is encouraging for patients with HCV is the large number of recent real-world studies from around the world that show similar results with high SVRs with different regimens, different genotypes, different durations of therapy (from 8 to 24 weeks), with or without ribavirin, whether

Table 5 Results of the multivariate logistic regression analysis for the independent predictors of sustained virologic response

Variable	Odd ratio	*P*-value	95% Confidence interval	
			Minimum	Maximum
Presence of cirrhosis	16.091	0.010	1.963	131.905
Duration of therapy (12 weeks)	2.379	0.127	0.731	12.280

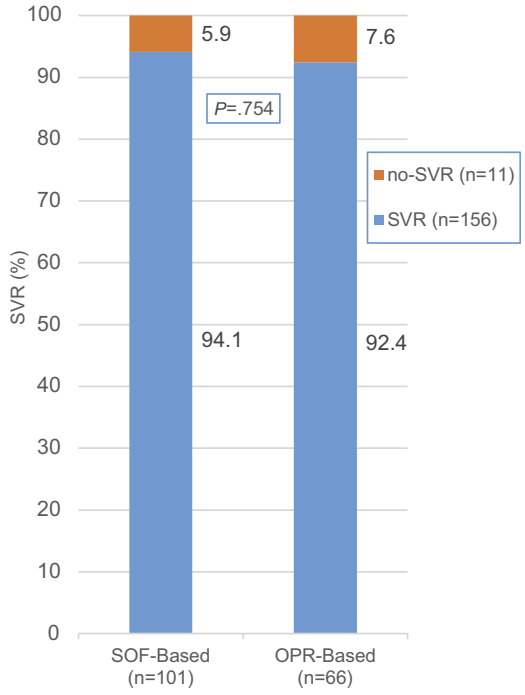

Figure 3 Percentage Sustained virologic response (SVR) and non-sustained virologic response (No SVR) in patients who received the SOF-based (n=101) and OPR-Bree (n=66) regimens. *P*=0.754 by Fisher's exact test. SOF= sofosbuvir.

patients are treatment-naïve or treatment-experienced, and whether the DAAs are original or generic.[15–29]

The present study failed to demonstrate a difference in SVR12 between treatment regimes, whether they were SOF-based or OPR-based. Although this may be interpreted as non-SVR being probably related to host, disease, or viral factors rather than regimen-related factors such as type and length of treatment, this is likely a type 2 error. In the literature, for example, ribavirin has a positive additional effect on SVR in certain regimens[11] while in others, such as daclatasvir plus sofosbuvir, it does not.[18]

Patients with cirrhosis had significantly lower SVR12 compared to those without cirrhosis, and in multivariate

Table 6 The impact of treatment on the laboratory results

Variable	Pre-treatment	Post-treatment	*P*-value
WBC (x10^9/L)	5.81±2.16	6.07±2.38	0.025
Hemoglobin (gm/dL)	12.78±2.10	12.82±2.26	0.671
Platelets (x10^9/L)	191.83±95.49	215.70±99.09	<0.001
INR	1.29±2.10	1.12±0.17	0.320
PTT (second)	26.10±3.68	26.07±4.02	0.981
Serum creatinine (μmol/L)	88.05±140.73	90.40±133.79	0.625
Serum albumin (g/L)	38.79±5.85	41.15±5.35	<0.001
Total bilirubin (μmol/L)	18.73±17.86	15.24±13.68	0.003
ALT (IU/L)	61.00±40.35	27.42±22.72	<0.001
ALP (IU/L)	102.66±44.47	93.03±71.87	0.061
γGT (IU/L)	97.69±89.27	47.06±55.88	<0.010

Note: Data expressed as mean±SD or n (%) as appropriate.
Abbreviations: ALP, alkaline phosphatase; ALT, alanine aminotransferase; INR, international normalization ratio; WBCs, white blood cells.

analysis, cirrhosis was found to be the only independent predictor of non-SVR in our cohort. This is supported by a recent study from Egypt involving 2400 patients with HCV-related cirrhosis, where sofosbuvir and ribavirin therapy lead to SVR in only 71.2%, with more than 5% of patients discontinuing therapy due to adverse effects.[21] In a Chinese study, patients with HCV-related decompensated cirrhosis achieved 90% SVR;[23] however, this likely relates to the small number of patients (n=30).

One of the interesting findings in this real-world cohort is the high rate of response in the 21 patients who received treatment post-liver transplant (SVR12=100%). A recent Swedish study also identified that SVR12 was achieved in 91/93 (97.8%) of patients who relapsed post-liver transplantation, with 100% rates for genotype 2, 3, and 4, and a 96% rate for genotype 1.[29]

The main limitation of this study is its sample size, which limits any subgroup comparisons. In addition, since the allocation of patients to treatment was not randomized, direct comparisons between regimens, even as broadly as SOF-B and OPR-B, is limited. This is also confounded by the fact that there is great heterogeneity in the regimens used, where there is also one patient who received pegylated interferon in addition to sofosbuvir. Despite that, our results are comparable to other real-life studies of DAAs, where SVRs in excess of 90% are demonstrated. Finally, we did not evaluate the rapid viral response (RVR) with viral kinetics at 4 weeks. However, detecting a difference in RVR would be unlikely due to the high SVR.

Conclusion

Patients with chronic HCV irrespective of genotype, viral load, age, gender, and other medical comorbidities benefit greatly from SOF-B and OPR-B regimens. Both 12 and 24-week treatments are effective and well tolerated. However, cirrhosis influences the rate of SVR adversely.

Disclosure

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