



Incidence of Hepatocellular Carcinoma After Direct Antiviral Therapy for HCV in Patients With Cirrhosis Included in Surveillance Programs.

Submitted by Beatrice Guillaumat on Tue, 02/05/2019 - 16:11

Titre	Incidence of Hepatocellular Carcinoma After Direct Antiviral Therapy for HCV in Patients With Cirrhosis Included in Surveillance Programs.
Type de publication	Article de revue
Auteur	Nahon, Pierre [1], Layese, Richard [2], Bourcier, Valérie [3], Cagnot, Carole [4], Marcellin, Patrick [5], Guyader, Dominique [6], Pol, Stanislas [7], Larrey, Dominique [8], de Ledinghen, Victor [9], Ouzan, Denis [10], Zoulim, Fabien [11], Roulot, Dominique [12], Tran, Albert [13], Bronowicki, Jean-Pierre [14], Zarski, Jean-Pierre [15], Riachi, Ghassan [16], Calès, Paul [17], Péron, Jean-Marie [18], Alric, Laurent [19], Bourlière, Marc [20], Mathurin, Philippe [21], Blanc, Jean-Frédéric [22], Abergel, Armand [23], Serfaty, Lawrence [24], Mallat, Ariane [25], Grangé, Jean-Didier [26], Attali, Pierre [27], Bacq, Yannick [28], Wartelle, Claire [29], Dao, Thong [30], Thabut, Dominique [31], Pilette, Christophe [32], Silvain, Christine [33], Christidis, Christos [34], Nguyen-Khac, Eric [35], Bernard-Chabert, Brigitte [36], Zucman, David [37], Di Martino, Vincent [38], Sutton, Angela [39], Roudot-Thoraval, Françoise [40], Audureau, Etienne [41], ANRS CO12 CirVir Group [42]
Editeur	Elsevier
Type	Article scientifique dans une revue à comité de lecture
Année	2018
Langue	Anglais
Date	2018 11
Pagination	1436-1450.e6
Volume	155
Titre de la revue	Gastroenterology
ISSN	1528-0012
Mots-clés	Aged [43], Antiviral Agents [44], Carcinoma, Hepatocellular [45], Female [46], Hepatitis C [47], Humans [48], Incidence [49], Interferons [50], Liver Cirrhosis [51], Liver neoplasms [52], Male [53], Middle Aged [54], Proportional Hazards Models [55], Retrospective Studies [56]

BACKGROUND & AIMS: Retrospective studies have found an unexpectedly high incidence of hepatocellular carcinoma (HCC) among patients with hepatitis C virus (HCV)-associated cirrhosis who received direct-acting antiviral (DAA) agents. We analyzed data from the ANRS CO12 CirVir cohort to compare the incidence of HCC in patients with cirrhosis who received DAA therapy vs patients treated with interferon (IFN).

METHODS: Data were collected from 1270 patients with compensated biopsy-proven HCV-associated cirrhosis recruited from 2006 through 2012 at 35 centers in France. For descriptive purpose, patients were classified as follows: patients who received DAA treatment (DAA group, n = 336), patients who achieved a sustained virologic response (SVR) following an IFN-based regimen (SVR-IFN group, n = 495), or patients who never received DAA treatment and never had an SVR following IFN therapy (non-SVR group, n = 439). The patients were included in HCC surveillance programs based on ultrasound examination every 6 months, and clinical and biological data were recorded. To account for confounding by indication due to differences in patient characteristics at treatment initiation, we constructed a time-dependent Cox regression model weighted by the inverse probability of treatment and censoring (IPTCW) to assess the treatment effects of DAA on time until HCC.

RESULTS: Compared with patients in the SVR-IFN group, patients in the DAA group were older, higher proportions had diabetes or portal hypertension, and liver function was more severely impaired. The crude 3-year cumulative incidences of HCC were 5.9% in the DAA group, 3.1% in the SVR-IFN group, and 12.7% in the non-SVR group (overall $P < .001$; unadjusted hazard ratio [HR] for HCC 2.03; 95% confidence interval [CI] 1.07-3.84; $P = .030$ for the DAA group vs the SVR-IFN group). HCC characteristics were similar among groups. Among patients with HCC, the DAA group received less-frequent HCC screening than the other 2 groups ($P = .002$). After Cox analyses weighted by the IPTCW, we found no statistically significant increase in risk of HCC associated with DAA use (HR 0.89; 95% CI 0.46-1.73; $P = .73$).

CONCLUSIONS: Analysis of data from the ANRS CO12 CirVir cohort reveals that the apparent increase in HCC incidence observed in patients with cirrhosis treated with DAAs compared with patients who achieved SVR following an IFN therapy can be explained by patient characteristics (age, diabetes, reduced liver function) and lower screening intensity.

Résumé en anglais

URL de la notice <http://okina.univ-angers.fr/publications/ua18781> [57]
DOI [10.1053/j.gastro.2018.07.015](https://doi.org/10.1053/j.gastro.2018.07.015) [58]
Autre titre Gastroenterology
Identifiant (ID) PubMed [30031138](https://pubmed.ncbi.nlm.nih.gov/30031138/) [59]

Liens

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