COMMENTARY

The 3rd Canadian Symposium on Hepatitis C Virus: Expanding care in the interferon-free era

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Hepatitis C virus (HCV) currently infects approximately 250,000 individuals in Canada and causes more years of life lost than any other infectious disease in the country. In August 2011, new therapies were approved by Health Canada that have achieved higher response rates among those treated, but are poorly tolerated. By 2014/2015, short-course, well-tolerated treatments with cure rates >95% will be available. However, treatment uptake is poor due to structural, financial, geographical, cultural and social barriers. As such, ‘Barriers to access to HCV care in Canada’ is a crucial topic that must be addressed to decrease HCV disease burden and potentially eliminate HCV in Canada. Understanding how to better care for HCV-infected individuals requires integration across multiple disciplines including researchers, clinical services and policy makers to address the major populations affected by HCV including people who inject drugs, baby boomers, immigrants and Aboriginal and/or First Nations people. In 2012, the National CIHR Research Training Program in Hepatitis C organized the 1st Canadian Symposium on Hepatitis C Virus (CSCHCV) in Montreal, Quebec. The 2nd CSCHCV was held in 2013 in Victoria, British Columbia. Both symposia were highly successful, attracting leading international faculty with excellent attendance leading to dialogue and knowledge translation among attendees of diverse backgrounds. The current article summarizes the 3rd CSCHCV, held February 2014, in Toronto, Ontario.

Key Words: Biomedical; Clinical; Epidemiology; Hepatitis C; Public health; Social sciences

In 2013, it was estimated that there were >250,000 Canadians with chronic hepatitis C virus (HCV) infection (1). HCV infection causes progressive liver injury, and approximately 15% to 25% will develop cirrhosis, end-stage liver disease and liver cancer over a 30-year follow-up period (2). In August 2011, two protease inhibitors approved by Health Canada have led to higher response rates among those treated, but are poorly tolerated. By 2014/2015, short-course, well-tolerated treatments with cure rates >95% will be available. However, treatment uptake is poor due to structural, financial, geographical, cultural and social barriers. As such, ‘Barriers to access to HCV care in Canada’ is a crucial topic that must be addressed to decrease HCV disease burden and potentially eliminate HCV in Canada. Understanding how to better care for HCV-infected individuals requires integration across multiple disciplines including researchers, clinical services and policy makers to address the major populations affected by HCV including people who inject drugs, baby boomers, immigrants and Aboriginal and/or First Nations people. In 2012, the National CIHR Research Training Program in Hepatitis C organized the 1st Canadian Symposium on Hepatitis C Virus (CSCHCV) in Montreal, Quebec. The 2nd CSCHCV was held in 2013 in Victoria, British Columbia. Both symposia were highly successful, attracting leading international faculty with excellent attendance leading to dialogue and knowledge translation among attendees of diverse backgrounds. The current article summarizes the 3rd CSCHCV, held February 2014, in Toronto, Ontario.

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Extending care to HCV-infected individuals requires a coordinated effort by clinicians, nurses, researchers and policy makers, and requires consideration of the unique needs of specific Canadian populations affected by HCV including people who inject drugs (PWID) and

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HCV-infected Aboriginal persons, with both groups being identified by the Public Health Agency of Canada (PHAC) as being at increased risk for new HCV infections (3,4). In February 2012, the Canadian Institutes of Health Research (CIHR)-based National CIHR Research Training Program in Hepatitis C (NCRTP-HepC) organized the 1st Canadian Symposium on Hepatitis C Virus (CSHCV) in Montreal, Quebec. The 2nd CSHCV was held in March 2013 in Victoria, British Columbia (5). These symposia were well attended and attracted leading international experts, leading to dialogue and knowledge translation among attendees of diverse backgrounds. The current article summarizes the 3rd CSHCV, held February 2014, in Toronto, Ontario, where new Canadian and international HCV research findings were presented. It also gives an overview of the multidisciplinary discussions centred on improving access to care for HCV-infected Canadians. Specifically discussed were the challenges of delivering care to special populations of HCV-infected persons. Also highlighted was the need for a Canadian HCV action plan that would guide the provision of prevention, care and treatment for all Canadians infected with or at risk for acquiring HCV. The absence of such a plan remains a major obstacle that limits the marshalling of resources needed for HCV-infected Canadians to benefit from these potent, curative antivirals.

THE NCRTP-HepC

The NCRTP-HepC is a CIHR-supported Strategic Training Initiative in Health Research that was first funded in 2003 (www.ncrtp-hepc.ca). The NCRTP-HepC is a partnership funded by public groups including the CIHR, PHAC and the Canadian Liver Foundation, and which receives private funding from both industry and community organizations. The main goal of the NCRTP-HepC is to build translational HCV research capacity, and to prevent, care and treat HCV with the desire to eliminate HCV disease in Canada within the next 10 to 15 years. The training program uses the knowledge of 35 leading researchers from universities across Canada, who act as mentors for the trainees and who advance Canadian HCV research. The NCRTP-HepC is interdisciplinary, with members representing diverse research backgrounds, including social and behavioural sciences, epidemiology and public health, clinical and biomedical sciences. Since 2003, the NCRTP-HepC has supported 71 trainees (11 MSc, 35 PhD, three MD and 22 postdoctoral) and 45 summer students. This has significantly enhanced Canadian HCV research capacity, knowledge translation and interdisciplinary collaboration.

The NCRTP-HepC is a leader in facilitating HCV research in Canada (5). In response to feedback from the 1st and 2nd CSHCV, and questionnaires completed by community groups including the Canadian Liver Foundation’s ‘Living with Liver Diseases’ support group and the Toronto Community Hepatitis C Program, the specific aims of the 3rd CSHCV were:

1. To facilitate transdisciplinary knowledge exchange and collaborations among Canadian trainees, established researchers, health care practitioners, health policy makers and community-based groups working in the field of HCV.
2. To discuss approaches that would lead to increased treatment uptake across the country with a focus on expanding care and access to difficult-to-reach populations with the ultimate goal of eliminating HCV in Canada.
3. To implement, plan and deliver knowledge exchange and dissemination of symposium findings to support practice change, community awareness, harm reduction and policy development.
4. To implement a long-term plan for sustaining an annual CSHCV.

THE 3rd CSHCV

Providing better care for HCV-infected individuals requires integration between researchers and policy makers across multiple disciplines. The theme of the 3rd CSHCV was “Expanding care for HCV-infected Individuals” and the program was divided into four sessions focusing on biomedical sciences, clinical sciences, social sciences, and epidemiology and public health. The primary audience included Canadian investigators, research trainees (including MSc, PhD and postdoctoral fellows), health care practitioners, policy makers, industry representatives and members of the affected community. The sessions, summarized below, consisted of an international plenary speaker, a Canadian speaker and peer-reviewed original oral abstracts.

Biomedical sciences

Advances in drug discovery and development have led to the approval of the first generation of direct-acting antivirals (DAAs) for the treatment of HCV (1,6). Triple therapy with first-generation protease inhibitors significantly improved sustained virological response rates compared with pegylated interferon (IFN) alpha-2a and ribavirin alone. However, first-generation protease inhibitors are associated with additional side effects and complicated regimens. This has made treatment more successful but more difficult than previous standard therapy with pegylated-IFN/ribavirin (6). At the end of 2013, Health Canada approved the second-wave protease inhibitor (simeprevir) and, most significantly, the first nucleotide inhibitor of the viral polymerase (sofosbuvir). These developments will pave the way for increased availability of IFN-free anti-HCV drug regimens for Canadians infected with HCV.

Although IFN will largely be replaced by IFN-free DAA oral regimens in the near future, IFN-based therapies may still have a role due to the prohibitively high cost of the new treatments. Hence, strategies to predict or enhance the success rate of IFN-based therapies will remain relevant for patient selection and, furthermore, understanding how IFN acts will still be of importance given its central role in the innate antiviral immune response. Dr Markus Heim (University Hospital Basel, Switzerland), a world leader on IFN as an antiviral, presented data from his laboratory on IFN responsiveness and discussed the future role of IFN in a world of highly effective DAAs (7).

He suggested that most countries will develop algorithms to ensure that DAAs are used to treat individuals in greatest need but perhaps not all patients. If new treatments continue to be very expensive, he projected that IFN-based regimens will still have an important role, particularly for those who are likely to respond to therapy. Thus, it is critical that we gain a better understanding of why some, but not all, patients clear chronic HCV infections with IFN-based therapy.

While there have been great advances in our understanding of the innate immune response to HCV infection, it remains unclear why some patients have a relatively high level of baseline activation of IFN-responsive genes whereas others have minimal or no activation of these genes despite persistent HCV infection (8). The baseline expression of IFN-responsive genes is the strongest predictor of treatment outcome with IFN-based therapy (9). The gene expression pattern in a given individual is at least partially genetically determined, with strong associations with variants near the IFN lambda-3 (IFNλ3) gene (formerly known IL-28B) as well as with the recently identified IFNλ4 (10-15). Patients with the treatment-unfavourable variants tend to have activated IFN systems before receiving treatment and tend to be refractory to further stimulation with IFN. Overcoming IFN refractoriness by combining DAAs with IFN, using alternative dosing strategies or other IFNs, such as IFNa, could enhance IFN responsiveness, allowing for shorter, more effective treatments that may be an acceptable alternative to cost-prohibitively IFN-free DAA regimens.

In addition to its association with treatment outcome, the IFNλ3 polymorphism is also associated with spontaneous resolution of acute HCV infection (15,16). Dr Naglaa Shoukry (Université de Montréal, Montreal, Quebec) discussed recent work that indicated increases in IFNλ3 in plasma occurring in acute HCV infection correlated with natural killer cell markers and degradation (17). Natural killer cell-mediated degradation also correlates with the magnitude of the HCV-specific T cell responses, suggesting crosstalk between innate and adaptive immunity in acute HCV infection (18). A pilot transcriptome analysis of peripheral blood mononuclear cells from acute HCV patients of the Montreal Acute Hepatitis C Cohort (HEPICO) using RNA sequencing will hopefully shed some light on this important interaction (17).

Preliminary results indicate differential expression of...
numerous genes, many of which function in innate and adaptive immunity. A robust induction of IFN-stimulated gene expression was demonstrated during acute HCV and, surprisingly, despite spontaneous clearance of the virus, peripheral blood mononuclear cells from resolver patients continued to demonstrate altered expression of immune genes for >1 year after clearance. Future studies will correlate parameters of HCV disease progression and compare these findings with data from patients infected with other viruses, vaccine studies and in vitro models. This will provide further insight into the interplay between innate and adaptive immunity on the outcome of HCV infection.

Clinical sciences
Access to care and treatment for marginalized and remotely located patients living with HCV is a critical issue along the cascade of care, Dr Sanjeev Arora, Director of Project ECHO (Extension for Community Healthcare Outcomes) at the University of New Mexico (Albuquerque, USA), described the objectives, structure, operation, successes and obstacles encountered with this innovative program (19). The mission of Project ECHO is to develop capacity to safely and effectively treat chronic, common and complex diseases in rural and underserved areas, and to monitor the outcomes of these treatments (20). HCV lends itself well to this model of care. Expert interdisciplinary specialist teams are linked with primary care clinicians via tele-ECHO clinics (21). Experts comanage patient cases and impart their expertise by way of mentorship, guidance, feedback and didactic education. In the case of HCV, primary care physicians retain responsibility for patient management, with the ultimate goal of developing independent expertise allowing them to manage future patients and become ‘local experts’ for their peers, thereby reducing travel costs, wait times and improving health care quality (21). HCV care and expertise has been dramatically expanded to remote regions and marginalized populations including prisoners who historically had never received HCV treatment in New Mexico. Evaluation of this care model demonstrates health care provider and patient satisfaction, as well as cure rates that are as good, if not better, than those achieved within an academic setting. This approach to HCV care appears to be ideal for Canada but will require a willingness to demonopolize specialist knowledge, and to task shift from services traditionally provided by physicians to other health care providers, as well as convincing public payers that investing in this type of care delivery system is cost-effective and sustainable.

In an extension of this topic, Drs Curtis Cooper and Julie Bruneau engaged in a lively and humorous debate entitled “Treatment should be strictly delivered by specialists rather than by primary care practitioners in the community” (22). Dr Cooper was given the difficult task of representing the ‘pro’ side and presenting arguments in favour of specialists strictly providing HCV care (when in fact he supported a broader approach of primary care physician involvement). Quoting Hamlet as well as research on HIV and oncology care, Dr Cooper argued that specialists were more likely to initiate treatment and choose appropriate regimens when facing complex conditions. He also pointed out that general practitioners may lack training, expertise and experience to treat specific conditions for which treatment evolves rapidly, such as with HIV and cancer survivorship (23,24). He suggested that, similar to HIV and cancer, HCV treatment is quite complex and management by specialists may be required.

On the ‘con’ side, Dr Bruneau pointed out that, to reduce HCV morbidity and mortality, we must first and foremost substantially increase treatment uptake among those who are infected. Currently, <10% of all infected people have access to treatment (25). Conceding that patients with severe liver conditions or multiple comorbidities should be treated in specialized academic settings with skilled and experienced specialists, she stressed the importance of having primary care physicians engaged in treating HCV-infected individuals. Not only are primary care physicians qualified to treat HCV, they can provide testing, counselling, and treat other medical conditions that often coexist with HCV, such as addiction and psychiatric disorders, ultimately improving the overall health outcomes (21). In the end, both debaters amicably concluded that HCV treatment should be widely available to patients according to best practices via the collaboration of primary care physicians and specialists in the best-adapted environment to meet individualized needs.

In further discussion regarding access to care, there was an emphasis on the population of HIV/HCV coinfected individuals. This group is a key population in Canada at risk for accelerated disease progression, and which faces multiple barriers in access to care and successful HCV treatment. Based on information generated from the Canadian Co-Infection Cohort, Dr Marina Klein (McGill University, Montreal, Quebec) described the current characteristics of HIV/HCV coinfected individuals in Canada, as well as the modifiable risk factors and interventions that could improve health outcomes (26). From 2003 to 2013, this population had a standardized mortality rate that was >12-fold that of the Canadian population (27). This is driven primarily by end-stage liver disease, drug overdose and malignancy. Use of current IFN-based HCV antiviral and HIV antiretroviral therapies reduces this risk for the individual but not at the general population level. Although HCV antiviral therapy initiation rates are higher than in other cohorts, females, Aboriginals and those with a history of crack/cocaine use were less likely to initiate treatment. HCV antiviral initiation rates differed considerably among the 18 sites contributing data to this cohort, suggesting that standardizing HCV care in coinfection should be prioritized. The value of enhanced access to HCV antiviral therapy was also discussed. Benefits include improved patient quality of life scores and reduced health care utilization following achievement of a sustained virological response. However, the multiple socioeconomic barriers faced by the HIV-HCV coinfected population will need to be addressed to ensure maximum benefit.

Social and behavioral sciences
In Canada, the majority of new and existing cases of HCV occur among PWID, either currently or in the past (1,28). Dr Jason Grebely (University of New South Wales, Australia) discussed a community-based study of PWID in Vancouver (British Columbia) documenting that HCV treatment uptake was only 2% per year in 2010 (29), despite universal access to health care. Treatment among PWID has added benefits, including the potential to reduce HCV transmission. Treatment as prevention has the potential to markedly reduce incidence and, ultimately, prevalence across the country (25,30). Clearly, strategies are needed to enhance HCV treatment uptake, particularly among PWID.

Barriers to HCV treatment are multifactorial and include issues of access to therapy and barriers at the level of the patient, practitioner and the health care system (31-33). Dr Carla Treloar (University of New South Wales, Australia), explored how stigma, social exclusion, symbolic violence and trust could be better understood and used to develop more effective therapeutic encounters to address barriers to HCV care, particularly among PWID (34). Drawing from social theory and research from a number of qualitative studies with clients and practitioners (35-39), she suggested some pragmatic strategies of how practitioners could develop inclusive, nondiscriminatory, trust-building HCV services to enhance HCV assessment and treatment.

In addition to enhancing engagement to HCV care and treatment, strategies to enhance responses to HCV therapy are also needed. Dr Louise Balfour (University of Ottawa, Ottawa, Ontario), highlighted the importance of adherence to HCV therapy (40). Specifically, Dr Balfour emphasized the important role that psychologists can play in helping to address barriers to care at the level of the patient, and helping to better prepare HCV patients for therapy, which may lead to improved adherence and, ultimately, improved outcomes in HCV therapy (41). Adherence will be particularly important with the new oral DAA regimens given the very high costs of failed therapy. Adherence strategies are particularly important for PWID or individuals with serious mental health issues (42).
One collaborative community-based group model of care that has been incredibly successful for treating a highly marginalized population was presented by Dr Jeff Powis from the Toronto Community Hep C program (43). This model has demonstrated that PWID can be engaged in HCV treatment (44) and treatment responses resemble those observed in clinical trials, despite the presence of multiple comorbidities (44,45).

The impressive advances in the development of DAA-based therapy, particularly IFN-free regimens, provide the potential to cure HCV infection in the vast majority of treated individuals. However, as shown in Figure 1, there is a considerable disparity between the potential treatment efficacy (>90%) and the current HCV treatment effectiveness (<10%), given that <10% of people have been treated in most countries, including Canada. Unless the proportion of individuals screened, assessed and treated for HCV infection is substantially increased, these anticipated therapeutic advances will have limited impact at the population level (46).

**Epidemiology and public health**

More than one-half of all existing HCV infections are estimated to be in current or former PWID (2,25). HCV treatment uptake in this population is low overall, and is especially low among current PWID (47-49). Barriers to HCV treatment are multifactorial and include issues of access to therapy and barriers at the level of the patient, practitioner and the health care system (25). Active drug use and mental health problems are often key factors affecting HCV treatment decision by both patients and providers (25). Some of these barriers are associated with medication side effects, factors that affect therapy adherence and concerns regarding reinfection. However, there is ample evidence that, with appropriate care and support, PWID can be effectively treated for HCV, with international guidelines recommending HCV treatment for PWID on a case-by-case basis (25,50). Dr Kim Page (University of San Francisco, California, USA), discussed HCV ‘treatment cascade’ in PWID and individuals with substance use and mental health issues, how these issues impact care delivery and reviewed strategies that may impact successful delivery of care in this population (51). She spoke of the ‘incredible shrinking patient’. After clinicians apply a series of criteria to exclude patients (eg, AUDIT score for alcohol use, depression scores, laboratory criteria and active injection drug use) as few as 4% of patients are deemed eligible for current HCV treatments. Dr Page described the UFO “You Find Out” study, which provides an opportunity for PWID younger than 30 years of age in San Francisco to undergo testing and learn if they are infected with HCV or HIV. The UFO model was presented as an example of a successful HCV education and prevention program, demonstrating how integration of harm reduction, outreach, testing, education and onsite primary care can engage and care for young injection drug users.

Another significant at-risk population with a high prevalence of HCV and low treatment uptake is the First Nations community (52). Dr Kathy Pouteau (Family Physician, Sioux Lookout and Kasabonika Lake First Nation, Ontario) shared her experiences and model for delivering HCV prevention and care to a remote Aboriginal community in Northern Ontario (53). First Nations communities in Northwestern Ontario have seen markedly increased rates of HCV infection. Risk factors for infection include injection drug use and, in 2009, the Chiefs of the Nishnawbe Aski Nation declared a state of emergency around prescription drug use in their communities. Dr Pouteau discussed the challenges, opportunities and collaborative response of health service providers and First Nations in rural and remote communities in the Sioux Lookout area. Barriers to HCV prevention, diagnosis and care in these communities include fragmented primary and public health systems, limited access to harm reduction programs, difficulties in confirming HCV diagnosis and accessing post-test counselling, limited access to HCV support and treatment, barriers due to as well as ongoing inequities in social, economic and political determinants of health. These barriers are being addressed by health care providers in First Nations communities using a collaborative approach. This approach centres around five main priorities: building political advocacy and support; encouraging communication and collaboration with health services agencies; establishing meaningful partnerships based in First Nations communities (ie, opioid substitution programs); building First Nations HCV capacity, local ‘expert’ resources and tertiary partnerships; and promoting culturally safe and trusting relationships between health care providers and community members.

A critical player in the national response to HCV is PHAC. One of the challenges facing policy makers is the limitations in the comprehensiveness of HCV surveillance data in Canada, which makes it highly challenging to estimate the HCV disease burden. Dr Maxim Troubnikov from PHAC presented an updated model for determining estimates of the prevalent and undiagnosed HCV infections in Canada in 2011 (54). This model followed approaches used by the Center for Communicable Diseases and Infection Control (CCDIC) to estimate the 2011 HCV disease burden using the workbook and the back-calculation modelling methods. Using the workbook method, preliminary results, based on a Canadian population of 34,483,975, estimated that the anti-HCV prevalence for 2011 was 0.96% (95% CI 0.61 to 1.34), corresponding to 331,046 persons (95% CI 210,352 to 462,085 persons), while the chronic HCV prevalence was 0.71% (95% CI 0.45% to 0.99%), corresponding to 244,836 persons (95% CI 155,177 to 341,391 persons). It was further estimated that 44% of anti-HCV-positive persons were likely undiagnosed in 2011, although before having a final estimate, input from the back-calculation process and validation is required.

The second PHAC presentation, by Dr Margaret Gale-Rowe of the CCDIC, outlined the actions and the current federal response to HCV prevention and control (50). The current PHAC response includes HCV research collaboration and support, active disease surveillance and epidemiology, as well as the development of professional guidelines and resources for the management of HCV (50). Workshops and studies held by PHAC over the past year include discussion of birth cohort screening for HCV, a topic that was debated in the 2nd CSHCV (5). As a result, there is an interim statement that is under development on HCV screening based on the birth cohort approach in addition to current risk-based screening. Programs and partnerships...
that PHAC has fostered include promoting public and professional awareness campaigns including World Hepatitis Day and funding for the expansion of HCV knowledge including funding for the NCRT- HepC and the Global Hepatitis C Technical Network. The current PHAC response to HCV was outlined and presently exists as part of an integrated response to sexually transmitted and blood-borne infections including HIV, HCV and other infections. In the discussion period following these presentations, there was discussion about the need for a unified national strategy for hepatitis C in Canada. Although the PHAC representatives acknowledged the value of such an approach, no concrete plans to develop a national strategy currently exist.

OUTCOMES OF THE 3rd CSHCV
Although current treatments for HCV are effective, the severe side effect profile, prolonged duration, need for specialized support and the high cost, greatly limit treatment uptake. By 2014/2015, it is likely that all-oral, well-tolerated HCV treatment will be available with cure rates >95% in treated individuals. These new treatments will dramatically improve the ability to initiate treatment and cure infected individuals. However, translating these remarkable therapeutic improvements into tangible reductions in disease morbidity and mortality will require significant improvements in treatment uptake. With this in mind, the major focus on HCV in Canada will likely shift from the drug discovery to developing strategies to engage people into care and treatment. The 3rd CSHCV highlighted strategies that have proven effective in increasing treatment uptake internationally and nationally, with a focus on practical implementation of successful approaches in Canada.

The challenges of increasing HCV treatment uptake in Canada are structural, financial, geographical, cultural and social. It is time to develop achievable goals of treating the four major population groups in Canada that are most affected – PWID, baby boomers, immigrants and Aboriginal/First Nations people – so that HCV in Canada can be eliminated within the next 10 to 15 years. Among those at risk, PWID will require a specific focus for harm reduction to prevent reinfection. Without a screening program to identify those who remain undiagnosed, disease elimination will not be achieved. It will be critical to confirm whether the preliminary PHAC estimate that 44% of Canadians are unaware of their infection is correct. The key topic of discussion at this symposium was whether HCV care should be strictly provided by subspecialist clinicians or whether primary care physicians should treat HCV-infected individuals. The issues raised were the complexity of HCV treatment regimens in addition to the fact that, if all HCV-infected individuals were to be treated, there would not be a sufficient number of subspecialists to fill the need. The broad consensus was that a blended model of care, including many health care disciplines, will be required to maximize identification, treatment initiation and HCV cure. The notion of employing the ECHO model of telemedicine to reach geographically isolated areas was presented, a model that has been especially effective in providing HCV care in remote areas of the United States (21).

This symposium highlighted that given the unique challenges of the HCV epidemic and the imminent potential for curability, there is a clear need for national coordination of HCV prevention, care and treatment. National strategies have been adopted in countries, such as Australia, Scotland and the United States, which have resulted in improvements in HCV prevention, care and treatment (5). Canada needs a national strategy or action plan so that Canadians can also benefit from the imminent curability of HCV so that disease elimination within the next 10 to 15 years can become a realized dream.

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REFERENCES
57. NCHECR. Epidemiological and economic impact of potential increased hepatitis C treatment uptake in Australia: National Centre in HIV Epidemiology and Clinical Research, The University of New South Wales, 2010.