Severe (complicated) hepatitis A in Cotonou (Benin) concerning an incompletely vaccinated Spanish expatriate

Těžký (komplikovaný) průběh virové hepatitidy A v Cotonou (Benin) u nedostatěčně očkováného pacienta španělského původu

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Summary: Viral hepatitis A is a major cause of acute hepatitis in the world. Its progression can be fulminant. Generally, it does not usually pose a major health problem among people living in endemic areas, but can be an important source of morbidity for unvaccinated subjects from western countries travelling in highly endemic areas. We report a severe case of viral hepatitis A concerning a patient of Spanish origin, incompletely vaccinated and recently immigrated in Benin. Two months before hospitalisation, she had an elective cholecystectomy for symptomatic cholecystolithiasis.

Key words: jaundice – hepatitis A – expatriate – incomplete vaccination

Souhrn: Virová hepatitida A je hlavní příčinou akutní hepatitidy na světě. Její průběh může být fulminantní, hlavně u dospělých. Tato choroba většinou nepředstavuje zásadní zdravotní problém u lidí žijících v endemických oblastech, ale může mít těžký průběh u neočkovaných subjektů ze západních zemí, kteří do takových oblastí cestují. Popisujeme těžký průběh virové hepatitidy A u pacientky španělského původu, která byla nedostatečně očkována proti hepatitidě A a nedávno se přistěhovala do Beninu. Dva měsíce před hospitalizací měla elektivní cholecystektomii pro symptomatickou cholecystolitiázu.

Klíčová slova: žloutenka – hepatitida A – přistěhování – neúplné (částěčné) očkování

Introduction

Viral hepatitis A (HAV) is a major cause of acute hepatitis in the world [1]. Each year, there are more than 1.5 million new infections with hepatitis A worldwide [2]. The virus is transmitted via the faecal-oral route, usually through inter-human contact or from ingestion of food or contaminated water. The HAV is almost always asymptomatic in children under three years, but usually symptomatic in adults and may be severe (complicated). Very rarely, the HAV may have a fulminant course which may lead to liver failure and death. In Sub-Saharan

Africa (SSA), the HAV is endemic with more than half of all children (under five years) contaminated [3]. In endemic areas, the risk of contracting HAV is 90%, usually in childhood. Subjects who once have contracted HAV are immune for life. This is why the World Health Organization recommends no systematic vaccination in people living in areas of high endemicity such as Africa [2,4]. However, subjects travelling from Western countries to SSA have a high risk of contracting the disease in the absence of vaccine protection. We report a severe case of HAV in a Spanish-born patient who was incompletely vaccinated and recently immigrated in Benin.

Observation

S. E., a 54-year-old woman was admitted in October 1st, 2013 to a hospital in Cotonou for abdominal pain, vomiting, fatigue and low-grade fever lasting for one week. Her history indicated there had been a laparoscopic cholecystectomy in Madrid in September 2013 for symptomatic cholecystolithiasis. She had no history of liver disease or metabolic disease. She didn't take any drug regularly. The Spanish-born woman who had been living in Benin

Tab. 1. Laboratory results during the hospitalization.

Tab. 1. Laboratorní výsledky během hospitalizace.

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	1. 11. 2013	5. 11. 2013	10. 11. 2013	15. 11. 2013	12. 12. 2013	10. 1. 2014	12. 3. 2014
ALT (UI/L)	3,800	3,200		501	305	124	33
AST (UI/L)	1,800	1,400		133	74	32	30
GGT (UI/L)	450	350					
ALP (UI/L)	750	570					
TP (%)	47	59	77	80	82	86	88
total bilrubin (mg/L)	73	78		42			19
conjugated bilirubine (mg/L)	57.2	62.2		34			5
HBsAg	negative						
anti-HBs (mUI/mL)	411						
anti-HBc	negative						
HAV-Ab IgM	positive						
HIV	negative						
anti-HCV	negative						
CRP (mg/L)	32.10						6.4
glycemia (g/L)	0.92						0.9
creatinine (mg/L)	4.10						4
thick blood	negative						
hemoglobin (g/dL)	12.5						13
leukocytes (g/L)	3.75						3.8
platelets (g/L)	199						250
		_					

 $ALT-alanine\ aminotransferase,\ AST-aspartate\ aminotransferase,\ GGT-\gamma-glutamyl\ transpeptidase,\ ALP-alkaline\ phosphatase,\ TP-prothrombin\ time,\ HAV-Ab\ IgM-viral\ hepatitis\ A\ antibody\ Immunoglobulin\ M,\ HCV-hepatitis\ C\ virus,\ CRP-C-reactive\ protein$

for 10 months did not have a significant alcohol consumption. She had been vaccinated against hepatitis A (single dose) three years previously on an earlier trip to South America and had also been immunised against hepatitis B. She lost the international vaccination card and didn't remember the name of the vaccine she took. The symptoms had motivated an antimalarial treatment with the combination of artesunate 200 mg, sulfamethoxazole 500 mg and pyrimethamine 25 mg - normal dose (one tablet per day for three days), without success. During the admission in hospital, the patient was afebrile (after paracetamol), the general condition had altered (because of severe asthenia), her body mass index was 22.9 (weight 64 kg), umbilical perimeter was 81 cm, blood pressure 123/75 mm Hg and heart rate 88/min. It was noted a conjunctival jaundice and tenderness of the epigastrium. The remainder of the examination was normal, there was no sign of hepatic encephalopathy. Laboratory tests (tab. 1) showed a negative thick blood, normal blood count as well as renal function tests, blood electrolytes and creatinine. We noted cytolysis (alanine aminotransferase (ALT) 95N (N – times the upper limit of normal), aspartate aminotransferase (AST) 45N), cholestatic jaundice (total bilirubin 7.5N, conjugated bilirubin to 23N, alkaline phosphatase (ALP) 7N, y-glutamyl transpeptidase (GGT) 10N). The rate of prothrombin was low at 47% and the C-reactive protein amounted to 32 mg/L. Seeing this picture of cholestatic jaundice in a patient shortly after cholecystectomy for stones, the first diagnostic hypothesis raised was residual stones in the bile ducts (cholelithiasis/choledocholithiasis). The second etiological hypothesis was acute viral hepatitis. Abdominal ultrasound showed a homogeneous hepatomegaly without bile duct dilatation of intra- and extrahepatic, and prothrombin time (TP) did not go back after the administration of vitamin K₁. Thus, the diagnosis was moving more towards acute viral hepatitis. Serology for hepatitis B immunisation showed anti--HBs antibodies to 411 mIU/mL, the serology for hepatitis C (HCV antibodies) was negative, immunoglobulin M type of anti-HAV antibodies (IgM) was positive. The control in another laboratory confirmed acute HAV. The diagnosis of acute HAV with severe hepatic impairment was retained. The symptomatic treatment was favourable with slow normalisation of hepatic tests and TP. TP was normal after 10 days and bilirubin after three weeks. The normalisation of transaminases was slow (the

control after three months showed normal AST but ALT to 3N, after five months transaminases were completely normalised), (tab. 1).

Discussion

This clinical case is interesting for several reasons:

- 1. The background the patient had undergone a recent cholecystectomy for symptomatic gallstones; and therefore seeing the picture of febrile jaundice and abdominal pain, the hypothesis of residual stones in the common bile duct was primary evocated. But this assumption was quickly ruled out in the absence of bile duct dilatation from the abdominal ultrasound, despite not performing any additional radiological examination such as magnetic resonance cholangiopancreatography. The positivity of IgM HAV led to the diagnosis of acute HAV.
- 2. The subject affected an incompletely HAV-vaccinated immigrant. Although the HAV does not pose a health problem for people in endemic areas [4], it can be a source of morbidity in travellers from low-prevalence countries [5]. Vaccination is recommended for these subjects. The doses and vaccination schema depend of the type of used vaccines and traveller's age. For the isolated vaccines againt HAV (for example Havrix®), it is recommended the first dose of vaccination at least 15 days before departure to the endemic area with a booster 6-12 months later [4,6]. Better is two full dose vaccination for adults before traveling to endemic area. The scheme for this vaccination is a 1,440 IU dose in adults (720 IU in children) at months 0 and 6 to 12 [4]. The vaccine is effective against HAV and protection is durable (12-25 years) [6]. This vaccination scheme differs in case of combined vaccines (for example boosters 2 months (M2) and 6-12 months (M6–M12) after the first dose in case of vaccines against HAV + HBV and another schema in case of vaccine against HAV + typhoid fever) [4,6]. Our

patient had received only one dose of unknown vaccine during a previous trip, so anyway was incompletely protected. In the literature, cases have been reported in incompletely HAV-vaccinated patients [7] and also in fully vaccinated patients reflecting the possible failures of the vaccine [6]. It seems that the vaccine efficacy decreases with age. Wolters et al. had found that vaccination against HAV was only 65% effective in subjects over 40 years of age [8]. They advocated, therefore, to verify vaccine protection in patients over 40 years of age before contact with the virus.

3. The last interesting point of this case is its clinical severity, shown by the presence of hepatocellular insufficiency. Indeed, HAV is usually a benign affection with 0.3-1.5% mortality [6]. The adult patients usually present with symptomatic forms. Morbidity and mortality increase with age [9]. In subjects aged 40 and over, mortality exceeds 2% [10]. The liver failure and mortality risk factors are age, co-infection with hepatitis B or C and sex (women being at a higher risk) [6]. Our patient had two of these factors (female sex and age) as well as being an incompletely vaccinated immigrant. Fortunately, evolution was spontaneously favourable.

Conclusion

This case demonstrates the importance and the necessity of full immunisation against HAV in patients from Western countries who are traveling to highly endemic areas. It would be useful to check the effectiveness of vaccination in subjects over 40 years of age before contact with the virus.

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The authors declare they have no potential conflicts of interest concerning drugs, products, or services used in the study. Autoři deklarují, že v souvislosti s předmětem studie nemají žádné komerční zájmy.

The Editorial Board declares that the manuscript met the ICMJE "uniform requirements" for biomedical papers.

Redakční rada potvrzuje, že rukopis práce splnil ICMJE kritéria pro publikace zasílané do biomedicínských časopisů.

> Submitted/Doručeno: 14. 3. 2016 Accepted/Přijato: 24. 5. 2016

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