scientific correspondence

High prevalence of non-virus/non-alcohol-related alanine-aminotransferase increase in blood donors

A slight increase of alanine-aminotransferase (ALT) levels is frequently observed (23.7%) in blood donors negative for hepatitis viruses (B and C) and alcohol abuse. The highest rate of subjects with ALT abnormalities was observed in men 20–40 years old, with a body mass index \geq 25. Non-alcoholic fatty liver disease (NAFLD) is probably the most frequent cause of hypertransaminasemia in Italian blood donors.

Hepatitis viruses (HAV, HBV, HCV, HDV) and alcohol intake have played a major role in causing acute and chronic liver damage in Italy during the last decades.¹ However, these factors do not account for all cases of hypertransaminasemia observed in clinical practice. Screening of the general population in northern Italy during the years 1991 to 1993² revealed that a high proportion of subjects showed elevated transaminase levels despite being negative for hepatitis virus markers and denying excessive alcohol intake.

We evaluated the alanine-aminotransferase (ALT) levels in 9,495 blood donors negative for both hepatitis B surface antigen and antibody to hepatitis C virus. The population consisted of 7,527 men and 1,968 women (median age 35.4 years; range 16–65) who had all undertaken screening procedures for blood donation at a blood bank in southern Italy. Diabetes, history of drug addiction, alcohol abuse, recent use of potentially hepatotoxic drugs or parenteral exposure were excluded using a specific questionnaire. Alanine-aminotransferases were measured by an ultraviolet kinetic test (Boehringer-Mannheim, Italy) using the optimized method recommended by the International Federation of Clinical Chemistry. The upper limit of normal was 40 U/L as indicated by the manufacturer.

Overall, 23.7% of subjects (2252/9495) presented abnormal ALT levels. The ALT levels were above the normal limit in 423/3,451 (12.2%) of subjects with a body mass index (BMI) < 25 and in 1,829/6,044 (30.2%) of subjects with BMI \ge 25 (p< 0.05). The highest rate of subjects with altered ALTs was observed in men 20-40 years old (Table 1). ALT abnormalities are generally slight, usually twice as high as the maximum normal value (MNV) (Table 2). These findings suggest that: 1) non-virus/non-alcohol-related hypertransaminasemia is a major epidemiological problem; 2) overweight is a frequent factor associated with ALT abnormalities. As these findings agree with corresponding figures from studies conducted in northern Italy,² this phenomenon is widespread throughout our country.

Non-alcoholic-fatty liver disease (NAFLD) – steatosis and steatohepatitis frequently present in overweight or obese patients, is probably the most common cause of abnormal transaminase levels in blood donors³ although the role played by other, rarely explored, factors of liver injury – i.e. celiac disease, occult infection, etc.⁴⁻⁸ has to be assessed.

> Filomena Morisco,* Abele Di Lonardo,° Tommaso Stroffolini,* Danilo Leone,@ Nicola Caporaso*

*Department of Food Science, University of Naples "Federico II"; °Immunohematology and Transfusion Center, "G. Rummo"

Hospital, Benevento; #Laboratory of Epidemiology, Istituto Superiore di Sanità, Rome; @Department of Mathematics and

Statistics, University of Naples "Federico II", Italy

Correspondence: Prof. Nicola Caporaso, M.D., Department of Food Science, University of Naples "Federico II, via Caravaglios, 36, 80125 Naples, Italy. Phone: international +39.081.2395535. Fax: international +39.081.5932575. E-mail: nicola.caporaso@tin.it

Age groups (years)	Men		Women	
	Ν	%	Ν	%
0 - 20	38/246	15.44	5/89	5.61
20 - 40	1447/4786	30.23	45/1185	3.79
40 - 60	653/2437	26.79	56/679	8.24
> 60	5/58	8.62	3/15	20.00
Total	2143/7527	28.47	109/1968	5.53

Table 1. Prevalence of ALT abnormality by age and sex among blood donors negative for hepatitis virus markers and denving

Table 2. Prevalence of different levels of ALT abnormality by age and sex among our population of blood donors in southern Italy, 1994 – 1998.

Age Groups (years) 0 - 20	Men < 2 × MNV% > 2 ×MNV %		Women <2 × MNV > 2 × MNV%	
	12.60	2.84	5.61	0
20 - 40	26.57	3.65	3.45	0.33
40 - 6	25.19	1.60	7.80	0.44
> 60	8.62	0	20.00	0
Total	25.53	2.93	5.18	0.35

References

- Stroffolini T. Alcohol, HCV infection, and liver cirrhosis: is the cup half full or half empty? J Hepatol 1998; 28:728-30.
- Bellentani S, Tiribelli C, Saccoccio G, et al. Prevalence of chronic liver disease in the general population of northern Italy: the Dionysos Study. Hepatology 1994; 20:1442-9.
- Bellentani S, Saccoccio G, Masutti F, et al. Prevalence of and risk factors for hepatic steatosis in northern Italy. Ann Intern Med 2000; 132:112–7.
- Volta U, De Franceschi L, Lari F, Molinaro N, Zoli M, Bianchi FB. Coeliac disease hidden by cryptogenic hypertransaminasaemia. Lancet 1998; 352:26-9.
- Berasain C, Betes M, Panizo A, et al. Pathological and virological findings in patients with persistent hypertransaminasaemia of unknown aetiology. Gut 2000; 47:429-35.
- Fargion S. Dysmetabolic iron overload syndrome. Haematologica. 1999; 84:97–8.
- Pisani G, Antigoni I, Bisso G, Wirz M, Iudicone P, Miceli M, Gentili G. Prevalence of TT viral DNA in italian blood donors with and without elevated serum ALT levels: molecular characterization of viral DNA isolates. Haematologica 2000; 85:181-5.
- Pisani G, Cristiano K, Bisso G, Wirz M, Gentili G. Further evidence on the underestimation of the prevalence of TT viral DNA in blood donors. Haematologica 2000; 85:1218-9.