Kenis, Y., Dustin, P., jun., and Peltzer, T. (1958). *Acta Haematologica*, 20, 329.
 Kissmeyer-Nielsen, F., et al. (1973). *Histocompatibility Testing 1972*, ed. J. Dausset and J. Colombani. Copenhagen, Munksgaard.
 Kristensen, M. (1938). *Acta Scandinavica Pathologica et Microbiologica*, suppl. 37, 399.
 Levine P. H., et al. (1971). *Cancer*, 27, 416.
 Levine, P. H., Stevens, D. A., Coccia, P. F., Dabish, L., and Roland, A. (1972). *Cancer*, 30, 875.

Massey, F. C., Lane, L. L., and Imbriglia, J. E. (1953). *Journal of the American Medical Association*, 151, 994.
 Miller, G. (1971). *Yale Journal of Biology and Medicine*, 43, 358.
 Miller, R. W., and Beebe, G. W. (1973). *Journal of the National Cancer Institute*, 50, 315.
 Rosdahl, N., Olesen Larsen, S., and Thamdrup, A. B. (1973). *Scandinavian Journal of Infectious Diseases*, 5, 163.
 Smithers, D. W. (1967). *British Medical Journal*, 2, 337.

Prevalence of Hepatitis B Antigen and Antibody in Prostitutes

G. PAPAEOVANGELOU, D. TRICHOPOULOS, T. KREMASTINO, G. PAPOUTSAKIS

British Medical Journal, 1974, 2, 256-258

Summary

We investigated the prevalence of hepatitis B antigen (HBaG) and antibody (HBaB) in 293 prostitutes and in 379 pregnant women of similar age and of low socio-economic level, who served as controls. HBaG was found in 4.4% of prostitutes and 3.4% of controls. The prevalence of HBaB was significantly higher ($P < 0.001$) in prostitutes (56.7%) than in controls (24.5%). The prevalence of HBaB was clearly age-dependent in both groups. Evidence of hepatitis B virus infection significantly increased with the number of years in prostitution. The evidence of increased infection rates among prostitutes and their distribution support the hypothesis that hepatitis B infection is sexually transmitted.

Introduction

Recent studies suggest that hepatitis B may be transmitted by non-parenteral or at least inapparent routes (Cossart, 1971) and that sexual transmission may play an important role in the spread of the disease (Jeffries et al., 1973). Thus a high rate of infection might be expected among prostitutes. We therefore decided to test this hypothesis by studying the prevalence of hepatitis B antigen (HBaG) and antibody (HBaB) in prostitutes in Athens, where the incidence of hepatitis B virus infection is high (Papaevangelou et al., 1971). We report the results of the study in this paper.

Subjects and Methods

The study sample consisted of 293 registered prostitutes in the Greater Athens area. Their age, length of time in prostitution, and history of syphilis were ascertained by direct questioning and from the special records of regular medical examinations. An unselected sample of 379 pregnant women who were delivered in hospital served as controls. Nearly all of them were in social class IV or V (General Register Office, 1951). The age range of the prostitutes was 20-51 years (mean 30.1), which

was slightly but not significantly ($P > 0.1$) higher than that of the controls (mean age 27.7 years).

Blood samples were allowed to clot and the separated serum was stored at -20°C . Screening for HBaG was done by counter-immunoelectrophoresis (Pesendorfer et al., 1970) and immunodiffusion test (Prince, 1968); for HBaB by passive haemagglutination assay (Vyas and Shulman 1970); for syphilis by Wasserman as well as by R.P.R.* card test; and for SGPT by a modified method of Reitman-Frankel (1957). Differences between frequency rates were tested by χ^2 test with Yates's correction.

Results

HBaG was present in 13 (4.4%) of the prostitutes and in 13 (3.4%) of the controls (table I, fig. 1). This difference was not statistically significant ($P > 0.1$). In two of the HBaG-positive prostitutes and one of the controls SGPT levels were above normal but not higher than 100 Karmen units. HBaG was more prevalent in prostitutes than in controls in almost all age groups. There was also a definite tendency for the HBaG prevalence to decrease with age in both prostitutes and controls. HBaG when found was still present in all cases on retesting at least six months later, but in no case was HBaB present.

FIG. 1—Prevalence of hepatitis B antigen (HBaG) and antibody (HBaB) among prostitutes (P) and pregnant women (C) in relation to age.

Department of Hygiene and Epidemiology, School of Medicine, University of Athens, Athens 609, Greece

G. PAPAEOVANGELOU, M.D., Associate Professor of Hygiene and Epidemiology

D. TRICHOPOULOS, M.D., Professor and Chairman, Department of Hygiene and Epidemiology

T. KREMASTINO, M.D., Lecturer in Hygiene and Epidemiology

G. PAPOUTSAKIS, M.D., Lecturer in Hygiene and Epidemiology

*Hynson, Westcoff and Dunning, Inc., Baltimore, Maryland, U.S.A.

TABLE I—Prevalence of Hepatitis B Antigen (HBAG) and Antibody (HBAb) among Prostitutes and Controls

Age Group	Controls								Prostitutes							
	No.	HBAG		HBAb		HBAG or HBAb		No.	HBAG		HBAb		HBAG or HBAb		No.	%
		Positive	%	Positive	%	Positive	%		Positive	%	Positive	%	Positive	%		
<19	33	1	3.0	4	12.1	5	15.2	—	—	—	—	—	—	—	—	—
20-24	115	6	5.2	24	20.9	30	26.1	18	2	11.1	9	50.0	11	61.1	18	15.7
25-29	106	4	3.8	28	26.4	32	30.2	73	4	5.5	32	43.8	36	49.3	73	69.8
30-34	81	1	1.2	23	28.4	24	29.6	57	3	5.3	33	57.9	36	63.2	57	63.2
≥35	44	1	2.3	14	31.8	15	34.1	138	2	1.5	91	65.9	93	67.4	138	67.4
Total	379	13	3.4	93	24.5	106	28.0	293*	13	4.4	166	56.7	179	61.1	293*	61.1

*Age unknown in seven (two HBAG positive and one HBAb positive).

TABLE II—Prevalence among Prostitutes of Hepatitis B Antigen (HBAG) and Antibody (HBAb) by Years in Prostitution

Years in Prostitution	No.	HBAG		HBAb		HBAG or HBAb	
		Positive	%	Positive	%	Positive	%
<1	29	1	3.5	11	37.9	12	41.4
1-2	30	4	13.3	13	43.3	17	56.7
3-4	26	3	11.5	16	61.5	19	73.1
≥5	178	3	1.7	115	64.6	118	66.3
Total	293*	13	4.4	166	56.7	179	61.1

*Duration in prostitution unknown in 30 (two HBAG positive and 11 HBAb positive).

HBAb was present in 166 of the prostitutes (56.7%) and in 93 (24.5%) of the controls (table I, fig. 1). The difference was significant ($\chi^2=70.6$; $P<0.001$). The prevalence among controls under 20 years old was 12.1%. It increased rapidly with age, reaching 31.8% among women over 35 years old. The increase with age was less appreciable in prostitutes, the corresponding rates being 50% in the 20-24 age group and 65.9% in those over 35 years old. Thus in both groups the prevalence of HBAb was age dependent.

When the prevalences of HBAG and HBAb were related to years in prostitution a substantially higher rate of HBAG was noted in prostitutes in their first five years of prostitution (table II, fig. 2). It decreased substantially in those who had been in prostitution more than five years. The prevalence of HBAb was directly related to the duration of prostitution. It was present in 37.9% of those who had been in prostitution for less than one year and in 64.6% of those who had been in it for over five years. This difference was slightly accentuated after standardization for age (corresponding rates 28.1% and 72.4%).

FIG. 2—Prevalence among prostitutes of hepatitis B antigen (HBAG) and antibody (HBAb) according to years in prostitution.

Serological evidence of syphilis was present in or recorded in the history of 45 prostitutes. HBAG was detected in three (6.7%) and HBAb in 27 (60%) of them. These rates were not statistically higher ($P>0.1$) than those of HBAG (4%) or HBAb (56%) in prostitutes without evidence of syphilis.

Discussion

Transmission of hepatitis B to sexual partners has been reported (Hersh *et al.*, 1971, Heathcote and Sherlock, 1973). HBAG is found more often in young adults (Prince *et al.*, 1970) and the carrier rate is high in homosexuals (Vahrman, 1973), in patients attending venereal disease clinics (Jeffries *et al.*, 1973), and in prostitutes (Papaevangelou *et al.*, 1973). HBAG carrier rates are not an index of cumulative past experience of hepatitis B (Blumberg *et al.*, 1969, Szmuness *et al.*, 1973). In contrast HBAb prevalence rates reflect past exposure to hepatitis B virus and therefore they provide reliable information from which to draw conclusions. Fulford *et al.* (1973) reported that promiscuity increases the incidence of HBAb. Their sample consisted of patients attending a clinic for sexually transmitted diseases and HBAb was detected by the insensitive immunodiffusion technique.

In the present study the highly sensitive method of passive haemagglutination assay was used to show the presence of HBAb in prostitutes, who may be considered as the group with the highest risk of contracting sexually transmitted diseases. The findings were compared with a sample of pregnant women of a low socioeconomic level and of roughly comparable age. The results showed that evidence of hepatitis B virus infection (prevalence of HBAG or HBAb) was significantly higher in the prostitutes. As expected, evidence of infection was related to age in both groups. The higher prevalence among the younger prostitutes may be explained by, among other things, the practice of illegal prostitution, which is usual for a time before registering as a prostitute.

Age and duration of prostitution are closely correlated. Therefore the increase seen in the hepatitis B infection rates with the duration of prostitution could be partly attributed to differences in the ages of the prostitutes concerned. Standardization for age scarcely affected the duration gradient of the infection rates. Therefore the duration effect is probably independent of the age effect. These findings are further evidence for the sexual transmission of hepatitis B.

Increased exposure to hepatitis B virus by other than the sexual route may be expected in prostitutes. Transmission by the parenteral route as a result of drug addiction may be more common in prostitutes. None in our sample had a history of parenteral drug abuse, which is very rare in Greece (Adjutantis, 1973). Treatment for syphilis, gonorrhea, and other infections may be other vehicles for parenteral transmission, but evidence of hepatitis B infection was no greater among the prostitutes with syphilis in our sample. Thus these factors do not seem to explain the great difference in prevalence rates between the prostitutes and controls in our study. We believe that the distribution of evidence of hepatitis B virus infection in the prostitutes in our study is compatible with a venereal mode of transmission. Other carefully controlled studies would, we hope, confirm our findings and provide further support for our belief.

This study was supported in part by a grant from the office of Scientific Research and Development, Ministry of Culture and Sciences.

References

- Adjutantis, G. (1973). *Minerva Medica Greca*, 1, 119.
 Blumberg, B. S., Sutnick, A. T., and London, T. (1969). *Journal of the American Medical Association*, 207, 1895.
 Cossart, Y. (1971). *Journal of Clinical Pathology*, 24, 394.
 Fulford, K. W., Dane, D. S., Catterall, R. D., Woof, R., and Denning, J. V. (1973). *Lancet*, 1, 1470.
 General Register Office. (1951). *Classification of Occupations*, 1950. London, H.M.S.O.
 Heathcote, J., and Sherlock, S. (1973). *Lancet*, 1, 1468.
 Hersh, T., Melnick, J., Goyal, R. K., and Hollinger, F. B. (1971). *New England Journal of Medicine*, 285, 1363.
 Jeffries, D. J., James, W. H., Jefferiss, F. J. G., Macleod, K. G., and Willcox, R. R. (1973). *British Medical Journal*, 2, 455.
 Papaevangelou, G., Kourea, T., and Tsoukas, S. (1971). *Pathologia et Microbiologia*, 37, 361.
 Papaevangelou, G., Trichopoulos, D., Papoutsakis, G., Kremastinou, T., and Pavlides, E. (1973). Ninth International Congress on Tropical Medicine and Malaria. *Abstracts*, 2, 166.
 Pesendorfer, F., Krassnitzky, O., and Wewalka, F. (1970). *Klinische Wochenschrift*, 48, 58.
 Prince, A. M. (1968). *Proceedings of the National Academy of Sciences of the United States of America*, 60, 814.
 Prince, A. M., et al. (1970). *New England Journal of Medicine*, 282, 987.
 Reitman, S., and Frankel, S. (1957). *American Journal of Clinical Pathology*, 20, 56.
 Szmunes, W., Prince, M., Brotman, B., and Hirsch, R. (1973). *Journal of Infectious Diseases*, 127, 17.
 Vahrman, J. (1973). *Lancet*, 2, 157.
 Vyas, G. N., and Shulman, N. R. (1970). *Science*, 170, 332.

PRELIMINARY COMMUNICATIONS

"Anticonvulsant Action" of Vitamin D in Epileptic Patients? A Controlled Pilot Study

CLAUS CHRISTIANSEN, PAUL RØDBRO, OLE SJØ

British Medical Journal, 1974, 2, 258-259

Summary

The frequency of epileptic seizures was observed in a controlled therapeutic trial on 23 epileptic inpatients before and after treatment with vitamin D, or placebo in addition to anticonvulsant drugs. The number of seizures was reduced during treatment with vitamin D, but not with placebo. The effect was unrelated to changes in serum calcium or magnesium. The results may support the concept that epileptics should be treated prophylactically with vitamin D.

Introduction

Minor degrees of hypocalcaemia (Richens and Rowe, 1970; Christiansen *et al.*, 1973) and hypomagnesaemia (Christiansen *et al.*, 1974) are known to occur in epileptics on anticonvulsant therapy. As a lowering of the concentration of these ions in extracellular fluid would tend to increase neuromuscular irritability it is conceivable that the effect of anticonvulsants could be partially counteracted. These biochemical changes are part of an osteomalacic state in such patients, and since small doses of vitamin D can increase the bone mineral content

(Christiansen *et al.*, 1973) it is possible that vitamin D might also influence the number of epileptic fits. For practical purposes the problem might be reformulated in the following way: Does vitamin D reduce the number of seizures in epileptic patients on anticonvulsant therapy? We report here the results of a controlled pilot study which seem to indicate that this may be the case.

Patients and Methods

Altogether 23 epileptic inpatients who had relatively frequent fits were selected for the study (table I). All the patients originally had typical grand mal or focal motor attacks and several of them also had other types of seizure. Since they were receiving various combinations of phenytoin, phenobarbitone, and primidone, for the purpose of analysis dosage (table 1) was calculated according to the data of Richens and Rowe (1970).

The number and type of seizures were recorded by experienced nurses who were unaware whether the patient was receiving vitamin D or placebo. Grand mal, abortive forms of grand mal as seen in patients on anticonvulsant treatment, and focal motor seizures were recorded.

TABLE I—Details of Patients and Dose and Duration of Anticonvulsant Treatment. Patients were divided into Two Treatment Groups (see text)

Group	No. of Patients	Sex		Age (Years)		Daily Dose of Anticonvulsant (Units*)		Duration of Anticonvulsant Treatment (Years)	
		M.	F.	Mean	Range	Mean	Range	Mean	Range
A	9	4	5	15	9-27	6.9	2-11	11	3-23
B	14	11	3	13	6-25	6.0	3-12	11	5-17

*Dose calculated in units according to data of Richens and Rowe (1970).

Blood samples were drawn for the determination of serum calcium, serum magnesium, and serum alkaline phosphatase. These determinations, all in duplicate, were performed as previously reported (Christiansen *et al.*, 1973, 1974).

Procedure.—The patients were allocated to two treatment groups (A and B, see table I) by means of random sampling numbers (Bradford Hill, 1967). The duration of the trial was 84 days and it was divided into three periods of 28 days. The

Department of Clinical Chemistry, Glostrup Hospital, 2600 Glostrup, Denmark

CLAUS CHRISTIANSEN, CAND. MED., Assistant Physician
 OLE SJØ, CAND. MED., Assistant Physician

Department of Clinical Physiology, Aalborg Sygehus Syd, 9000 Aalborg, Denmark

PAUL RØDBRO, M.D., Physician-in-Chief