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HIV associated culture proved tuberculosis has increased in north central London from 1990 to 1996

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Objectives: To examine rates of culture proved tuberculosis in HIV infected patients in three specialist centres in north central London.**Methods:** Cases of tuberculosis in patients with previously documented HIV infection from 1990 to 1996 were identified retrospectively from microbiological/clinical records at Chelsea and Westminster, St Mary's, and University College London Hospitals.**Results:** Between 1990 and 1996 202 cases of culture proved tuberculosis were identified at the three centres. Of these, 132/202 (65.3%) occurred in homosexual/bisexual men, 41/202 (20.3%) were in patients with heterosexual contact in sub-Saharan Africa, and 29/202 (14.4%) were in "others." Overall 148/202 (73.3%) had pulmonary tuberculosis. The total number of HIV infected individuals seen at the three centres increased from 4298 in 1990 to 5048 in 1996. Rates of tuberculosis in the three centres increased from 0.46% in 1990 to 0.83% in 1996. Part of this increase was due to an increase in tuberculosis among Africans from 1993 to 1996.**Conclusions:** Rates of HIV associated tuberculosis increased in these three centres in north central London between 1990 and 1996. In part this was due to an increase in the number of African patients with HIV infection attending the three centres. In addition, there was circumstantial evidence of recent transmission among homosexual men with HIV infection. Prospective "real time" surveillance of tuberculosis in HIV infected patients is needed in order to detect case clustering and to improve tuberculosis control.

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Introduction

Before 1988 there was a pattern of declining incidence of pulmonary tuberculosis in England and Wales. Since 1988 this decline has reversed such that between 1988 and 1993 approximately 8000 more cases of tuberculosis were notified than would have been anticipated had the previous secular trend continued.¹ Increasing socioeconomic deprivation, changes in patterns of immigration, and improved notification of tuberculosis have all been suggested as causes of this increase in incidence of tuberculosis. HIV infection is also a possible factor. The rise in tuberculosis in the United States in the mid 1980s was due in part to the high prevalence of HIV infection.² Previous studies in the United Kingdom have shown little evidence of a major overlap between HIV infection and tuberculosis.^{3,4} In this study we examined rates of culture proved tuberculosis in HIV infected patients attending three specialist centres in north central London.

Methods

We retrospectively studied rates of tuberculosis in adults with HIV infection between 1990 and 1996 at three specialist centres in north central London (Chelsea and Westminster, St Mary's, and University College London Hospitals). In order to eliminate bias due to changes in notification practices, microbiological records at each hospital were used to identify cases of *Mycobacterium tuberculosis* from whom the

organism was cultured from any site, rather than records of notified cases. These records were then cross referenced with HIV clinic records to identify patients known to be HIV infected at the time of diagnosis of tuberculosis. Risk factors for HIV infection in each patient were recorded.

Records from each centre were cross checked to eliminate duplication: patients seen at more than one centre during a course of treatment for tuberculosis were counted only once. Patients re-presenting with culture proved tuberculosis after completion of a course of chemotherapy lasting ≥ 6 months were regarded as having two separate episodes of infection.

The total number of HIV infected patients attending the three centres in each year of the study was ascertained. This figure may overestimate the number of patients seen as some patients are known to attend more than one centre in any given year. We assumed that transfers between centres remained stable over the study period. Data on the total reported cases of AIDS from all three centres were obtained from the Public Health Laboratory Service Communicable Disease Surveillance Centre.

Results

During the study period 202 cases of culture proved tuberculosis were diagnosed in HIV infected patients at the three centres, of whom

Table 1 Annual rates of tuberculosis in HIV infected patients attending the three centres from 1990 to 1996

Year	No with culture proved tuberculosis	No with HIV attending the three centres	TB rate in those with HIV infection (%)
1990	20	4298	0.46
1991	30	4649	0.64
1992	31	4854	0.64
1993	19	4776	0.40
1994	25	4777	0.52
1995	35	4821	0.72
1996	42	5042	0.83

Figure 1 Number of cases of culture positive tuberculosis each year, by risk factor for HIV infection.

148 (73.3%) had a positive culture from a pulmonary specimen. Three patients had a second episode of tuberculosis during the study period. The number of cases of tuberculosis increased from 20 in 1990 to 42 in 1996 (table 1, fig 1). The total number of HIV infected individuals seen at the three centres increased from 4298 in 1990 to 5048 in 1996 (table 1). The rate of tuberculosis in HIV infected individuals was 0.46% in 1990 and increased to 0.83% by 1996 (table 1). Overall, 132 (65.3%) cases of tuberculosis occurred in homosexual/bisexual men, 41 (20.8%) in patients with heterosexual contact in sub-Saharan Africa, and 29 (13.9%) in "others" (including 11 (5.4%) in injection drug users). There was an increase in the proportion of Africans with tuberculosis between 1993 and 1996 (fig 1). There was an increase in cases of tuberculosis in homosexual men in the three centres in 1991 and 1992 (fig 1).

Twelve patients with multidrug resistant (MDR) tuberculosis were seen during this study. Eight were part of a previously reported nosocomial outbreak which occurred in an inpatient unit, one other patient presented with MDR tuberculosis despite directly observed therapy and the remaining three cases acquired MDR tuberculosis outside the United Kingdom. No secondary cases linked to these three cases were identified.

During the study 3397 new AIDS cases were notified by the three centres, representing 31% of all cases notified in England and Wales over this period.

Discussion

We found a significant increase in rates of tuberculosis over the period of this study. We believe that this represents a true rise in rates of tuberculosis in patients with HIV infection attending these three centres, but considered several other explanations for the increase in cases. Notification of cases of tuberculosis from HIV centres has previously been poor.⁵ Improved notification may cause an apparent increase in tuberculosis in patients with HIV infection, and may also have led to more widespread HIV testing of patients with tuberculosis.⁵ Indeed, notifications of tuberculosis in North Thames rose from 1310 cases in 1990 to 1795 cases in 1995 (J Watson, personal communication). In order to minimise bias due to changes in notification practice we included in our analysis only cases of tuberculosis proved by culture. Another explanation for our data is that an increased number of patients were seen each year in the three centres. Since the total number of HIV infected patients at these centres increased by <18% during the entire study period, this accounts for only a small part of the increase in cases.

Others have shown that, in London, Africans with HIV infection have a high incidence of tuberculosis.⁶ We found an increase in the number of Africans with HIV and tuberculosis from 1993 and 1996. This explains part of the overall increase of tuberculosis; nevertheless, only 20% of the cases of tuberculosis in our study occurred in Africans. Some of the observed increase in tuberculosis in HIV infected Africans may be due to more frequent HIV testing in patients presenting with tuberculosis. In contrast, almost two thirds of cases of tuberculosis in our study occurred in homosexual men. Studies from the United States which used restriction fragment length polymorphism (RFLP) analysis and epidemiological case finding show that over half the cases of tuberculosis in patients with HIV infection were clustered and, thus, were due to recently acquired infection.^{7,8} Although we did not perform routine RFLP typing of isolates of *M. tuberculosis* we are aware of several outbreaks of tuberculosis that occurred in our units during the study period.⁹⁻¹¹ The increase in tuberculosis in 1995 was in part due to an outbreak of nosocomial MDR tuberculosis. The peak of tuberculosis in homosexual men in 1991 and 1992 may also have been due to undetected recent transmission of tuberculosis. This peak was seen in all three units, so nosocomial transmission in a single centre would not account for the higher rate of tuberculosis observed in these years. Of note, the increased rate of tuberculosis in HIV infected patients was not recognised in any of the three centres until this study was performed. This highlights the need for better systems of surveillance of HIV associated tuberculosis.

Rates of tuberculosis in patients with HIV infection may reflect the effectiveness of general tuberculosis control programmes, providing an "early warning" system of poor tuberculosis control.¹² There is a need for regional databases of patients with tuberculosis, including data on

HIV status and drug resistance patterns linked to prospective molecular typing of isolates to enable rapid detection of case clustering/nosocomial transmission and to contain the threat of MDR tuberculosis.

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Contributors: RC initiated this project; DC, MH, and RM were responsible for data collection; DC wrote the first draft; and RM wrote the final draft of the paper. All authors were involved in design of the study and commented on interim drafts: all reviewed the final manuscript.

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