Persistent risk of tuberculosis in migrants a decade after arrival in Australia

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n 2005, an estimated 8.8 million new cases of tuberculosis (TB) worldwide caused 1.6 million deaths. However, Australia and other developed countries have relatively low TB incidence rates, with most new cases occurring in migrants. The most reliable predictor of TB incidence in Australia is the TB prevalence in countries from where migrants are accepted. TB incidence in people migrating to developed countries is highest in the first 2 years after migration, although it remains high for up to a decade after arrival. 5-10

Control of TB in Australia is facilitated through entry screening of migrants using chest x-ray, and contact tracing of people with active TB. ¹¹ However, this strategy is inadequate for detecting latent and non-pulmonary disease and is unlikely to identify migrants whose TB will be diagnosed many years after their arrival.

We aimed to examine the risk of TB in migrants a decade after their arrival in Australia, and to determine characteristics of migrants that may be associated with ongoing risk, as these could have an impact on TB control policies in Australia.

METHODS

Study population

People born overseas and residing in Victoria with a laboratory-confirmed diagnosis of TB made between 1990 and 2004 were included in this study. Data on date of diagnosis, age at diagnosis, sex, clinical site of infection and drug resistance were obtained from the state TB reference laboratory at the Victorian Infectious Diseases Reference Laboratory. The laboratory database did not include all demographic variables, so patients were matched against notification data from the Victorian Department of Human Services to obtain further details, including country of birth and year of arrival in Australia. Patients for whom data on country of birth and arrival year were not available were excluded from the

We chose 10 years after arrival as the cutoff point for analysis, as TB may be less likely to be considered in the differential diagnosis in migrants a decade after arrival.

ABSTRACT

Objective: To examine the risk of tuberculosis (TB) in migrants a decade after their arrival in Australia.

Design, setting and patients: Retrospective review of laboratory-confirmed cases of TB in migrants diagnosed between 1990 and 2004 by the state TB reference laboratory in Victoria, analysed by a multivariate model comparing migrants diagnosed 10 or more years after arrival with those diagnosed within 10 years of arrival.

Main outcome measures: Time to diagnosis; characteristics of migrants diagnosed with TB, including age, sex, region of birth, site of infection, and drug resistance.

Results: A third of migrants (734/1924) were diagnosed with TB 10 or more years after arrival in Australia. This group was more likely to be European-born (adjusted odds ratio [AOR], 3.4; 95% CI, 2.4–5.0) and older than 34 years (35–49 years: AOR, 3.8; 95% CI, 2.0–7.0), reflecting the longer time European migrants have been in Australia. There were two distinct European groups: European Union (EU)/Western and Central/Eastern. The Central/Eastern group were from countries with current high TB rates and, compared with the EU/Western group, were younger (mean age, 50 v 64 years) and more likely to be diagnosed within 10 years of arrival (47% v 14%; P < 0.001).

Conclusion: European migrants were more likely to be diagnosed a decade or more after arrival in Australia. Once migrants from the currently high TB incidence areas of Asia and Africa have been in Australia for a similar period of time, their timing of diagnosis may resemble that for migrants from Europe. The current screening policy should be complemented with more sensitive techniques to detect latent TB.

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This cut-off also allowed comparison with other studies. ^{6,8-9}

Statistical analysis

Time between arrival in Australia and diagnosis was examined for migrants from specific regions, and a multivariate logistic regression model was constructed to compare those diagnosed 10 or more years after arrival with those diagnosed within 10 years of arrival. Independent variables in the model were year of diagnosis, sex, age group, clinical site (pulmonary or non-pulmonary), drug resistance, and region of birth, all of which were associated with time to diagnosis at the univariate level (*P*<0.1) (Box 1).

Data were analysed using Stata, version 8.0 (StataCorp, College Station, Tex, USA).

Ethical approval

Ethics approval was not required for this study because analysis and dissemination of notifiable diseases data is a core public health responsibility under the *Health Act* 1958 (Vic).

RESULTS

Of 3191 laboratory-confirmed TB cases in Victoria between 1990 and 2004, 2769 (87%) had the patient's country of birth recorded. Most of this group (2315; 84%) were migrants, with 45% (1047) coming from South-East Asia, 17% (400) from southern Asia, 15% (346) from Europe and 13% (299) from the Middle East or Africa.

Time between arrival in Australia and diagnosis of TB was available for 1924 of the patients born overseas (83%). The median time to diagnosis was 6 years (interquartile range, 2–14 years), but 38% (734) of all migrants with complete data were diagnosed 10 or more years after arrival. Most patients born in the Middle East/Africa and Asia (86% and 71%, respectively) were diagnosed within 10 years of arrival, compared with 21% of Europeans (Box 2).

Logistic regression analysis indicated that, compared with migrants from South-East Asia (the largest group of migrants), migrants diagnosed 10 or more years after arrival were more likely to be diagnosed

1 Comparison between migrants diagnosed with tuberculosis less than 10 years after arrival with those diagnosed 10 or more years after arrival in Australia, Victoria, 1990–2004

Characteristic $(n = 1860)$ *	Patients diagnosed < 10 years after arrival	Patients diagnosed ≥ 10 years after arrival	Crude odds ratio [†] (95% CI)	Adjusted odds ratio [†] (95% CI)	P (from model)
Year of diagnosis					0.001
1990–1994	315 (27.3%)	181 (25.6%)	1.0	1.0	
1995–1999	320 (27.7%)	160 (22.7%)	0.9 (0.7–1.1)	1.1 (0.9–1.5)	
2000–2004	519 (45.0%)	365 (51.7%)	1.2 (1.0–1.6)	1.9 (1.4–2.6)	
Sex					0.845
Male	576 (49.9%)	368 (52.1%)	1.0	1.0	
Female	578 (50.1%)	338 (47.9%)	0.9 (0.8–1.1)	1.0 (0.8–1.3)	
Age group (years)					< 0.001
0–19	79 (6.8%)	14 (2.0%)	1.0	1.0	
20–34	647 (56.0%)	107 (15.2%)	1.0 (0.5–1.8)	0.9 (0.5–1.6)	
35–49	234 (20.3%)	183 (25.9%)	4.5 (2.5–8.1)	3.8 (2.0–7.0)	
50–64	101 (8.8%)	135 (19.1%)	8.0 (4.3–15.0)	5.5 (2.9–10.6)	
≥ 65	93 (8.1%)	267 (37.8%)	17.3 (9.4–32.0)	10.3 (5.4–19.7)	
Clinical site					0.751
Pulmonary	618 (53.6%)	415 (58.8%)	1.0	1.0	
Non-pulmonary	536 (46.4%)	291 (41.2%)	0.7 (0.7–1.0)	1.0 (0.9–1.2)	
Drug resistance					0.018
Not resistant	1006 (87.2%)	655 (92.8%)	1.0	1.0	
Resistant	148 (12.8%)	51 (7.2%)	0.5 (0.4–0.7)	0.7 (0.5–1.1)	
Region of birth					< 0.001
South-East Asia	528 (45.8%)	301 (42.6%)	1.0	1.0	
Europe	56 (4.9%)	206 (29.2%)	6.5 (4.7–8.9)	3.4 (2.4–5.0)	
Middle East/Africa	213 (18.5%)	36 (5.1%)	0.3 (0.2–0.4)	0.3 (0.2–0.5)	
North-eastern Asia	74 (6.4%)	41 (5.8%)	0.9 (0.6–1.4)	0.5 (0.3–0.8)	
Southern Asia	249 (21.5%)	94 (13.3%)	0.7 (0.5–0.9)	0.5 (0.4–0.7)	
Other	34 (2.9%)	28 (4.0%)	1.4 (0.8–2.3)	1.1 (0.6–2.0)	

^{*} Patients with missing data for any of the variables were excluded. $\uparrow \ge 10$ years versus < 10 years. All independent variables were associated with time to diagnosis at the univariate level (P < 0.1).

during 2000–2004, to be aged \geq 35 years (with the risk increasing with age), and to be born in Europe. They were less likely to have been born in the Middle East/Africa, north-eastern Asia or southern Asia. There was no association between years to diagnosis and sex, site of infection or drug resistance (Box 1).

There were 267 migrants from Europe, forming two distinct groups: those from countries in the European Union (EU)/Western Europe and those from Central/Eastern Europe. The EU/Western group formed the majority (207; 78%), and 60% (125) of them were aged ≥65 years (mean, 64 years). Patients in the Central/Eastern group (60; 22%), from countries where TB prevalence is higher, were younger, with 32%

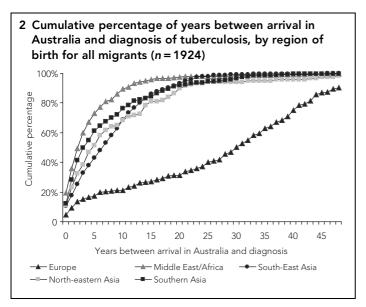
(19) aged \leq 34 years (mean, 50 years). Of all 267 migrants from Europe, 56 (21%) were diagnosed within 10 years of arrival in Australia — 28 from EU/Western Europe and 28 from Central/Eastern Europe, representing 14% and 47%, respectively, of the total patients from each region (P<0.001).

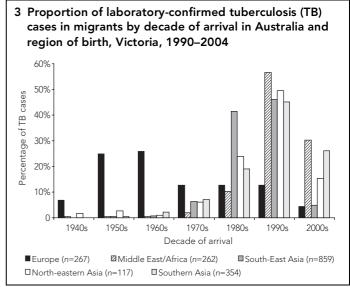
European patients in the study population have been arriving in Australia since the 1940s, when Europeans represented the majority of Australia's immigrants (Box 3). By contrast, patients from other migrant groups have been arriving mostly since the 1970s or 1980s, when immigration policies changed in Australia. To determine whether these changes in immigration may explain the pattern of diagnosis for European migrants, we compared time between arrival

and diagnosis for migrants by region of birth for the total study population (Box 2), and for only those migrants who arrived in Australia after 1980 (Box 4). Only 21% of European migrants in the total study population were diagnosed within a decade of arrival, compared with 73% of all other migrant groups combined (P<0.001) (Box 2). However, of migrants arriving in Australia after 1980, 68% of Europeans were diagnosed within a decade of arrival, similar to the proportion of all other groups (76%) (P=0.151) (Box 4).

DISCUSSION

We found that over a third of TB cases in migrants were diagnosed a decade or more





after their arrival in Australia. This proportion represents an increase from the 20% previously reported for cases diagnosed between 1948 and 1992 in Victoria. This finding is consistent with the experience in other developed countries, where the risk of active TB continues beyond the immediate post-migration period. 5-8,10

Overseas-born patients diagnosed a decade or more after arrival in Australia were more likely to be from Europe, mainly from countries in the EU and Western Europe. Similar to Australian-born TB patients² and patients diagnosed in the EU/Western Europe in 2004, 12 they were also more likely to be older. The older generations of European and Australian-born populations have a higher relative prevalence of latent

infection because they were born at a time when the risk of TB infection was relatively high, and they have had a long cumulative probability of acquiring infection.³ This is also true for people born in North America.¹³

The second group of European migrants, those from Central or Eastern Europe, were younger at the time of diagnosis with active TB and formed the majority of European patients diagnosed within 10 years of arrival in Australia. This is similar to patients born in Asia and Africa and reflects the current higher incidence of TB in their home countries. ¹³

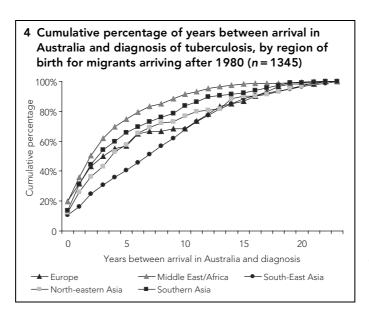
Relatively little TB transmission from migrants to the

broader community has been demonstrated in Australia and other low-incidence countries, and most TB diagnoses are due to reactivation of latent TB infection. In Victoria, genetic typing of all new TB isolates has occurred since 1993, and, apart from limited transmission within immigrant family groups and domestic contacts (including first-generation Australian-born children), most cases of TB in migrants who have been in Australia for 10 or more years have involved unique strains (DL, unpublished data).

Given that European migrants have been in Australia longer than migrants from other regions, it is not surprising that TB in this group was found to be diagnosed longer after arrival. This has implications for future control of TB in Australia. When migrants from the currently high-incidence countries of Africa and Asia have been in Australia for longer periods, their timing of diagnosis may resemble that now seen for European migrants, resulting in an increase in the number of TB cases diagnosed in these groups. A similar pattern may be observed in other low-incidence countries with migration policies comparable to those in Australia.

Further efforts to improve detection and treatment of latent TB infection, along with the realisation that the risk of infection persists beyond the first decade after migration, may help to minimise the burden of TB in Australia. Additional strategies that include more sensitive screening tech-

niques for latent TB, such as the Mantoux test and the secondgeneration interferon-y release assays including QuantiFERON (Cellestis International, Melbourne, Vic) and enzyme-linked immunosorbent spot (ELIS-POT) tests, 16 should be introduced into the migrant screening program. Refugees and migrants from high-prevalence areas should be screened for latent TB on arrival, offered treatment as appropriate, and followed up beyond 10 years. A further control strategy, as recommended by the European framework for tuberculosis control and elimination in countries with low incidence, 3 is a global investment to improve TB control in



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high-incidence countries and thus reduce the international transfer of risk.

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COMPETING INTERESTS

None identified

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