Re: Declining trend in incidence of tuberculosis in adolescents and young adults in Taiwan

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20 May 2019

Dr Martin Kolb
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442 Glossop Road
Sheffield S10 2PX
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Re: Declining trend in incidence of tuberculosis in adolescents and young adults in Taiwan

Snow K¹, Sawyer S², Denholm J³, Graham S¹.

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To the Editor,

We read with interest the recent publication by Chen et al regarding the burden of notified tuberculosis (TB) among adolescents and young adults in Taiwan[1]. The authors report that 10-24 year olds constituted only 5.2% of all newly diagnosed TB patients in Taiwan from 2005 to 2016, and that this percentage declined from around 6% in 2005 to around 4% in 2016. They contrast these figures with our estimate that young people constitute 17% of all people developing incident TB globally in 2012[2]. There are several factors that explain this apparently large discrepancy.

Firstly, our figure of 17% is a global average, covering settings that are demographically and epidemiologically diverse with regard to both population age structure and TB epidemiology. The age distribution of TB in any setting depends on three factors: the age structure of the population, the intensity of TB transmission, and the success of TB prevention efforts in different age groups. The proportion of TB cases which affect young people will be lower than average in less intense TB epidemics, in countries with older populations, and in countries with more successful prevention programs. At one extreme, in Western Europe, most nations have TB incidence below 20 cases per 100,000 person years (py)[3], and young people constitute only 14-18% of the total population[4]. At the other extreme, in southern Africa estimated TB incidence exceeds 300 per 100,000py[3] and fully one third of the population are aged 10-24[4], resulting in an epidemic concentrated among those in young and middle adulthood (see Figure 1). In Taiwan, young people constitute around 16% of the population, similar to Europe and other high income settings[5], and as noted by the authors, reported TB incidence is 44 cases per 100,000py. As such, it is to be expected that the burden of disease among young people in Taiwan is below the global average.
The proportion of cases among young people will be highest in countries with high TB incidence and young populations, as well as high rates of TB transmission have a disproportionate impact on children and adolescents[6]. Furthermore, high rates of TB transmission have a disproportionate impact on children and adolescents[6]. As TB incidence declines, the burden of disease shifts into progressively older age groups, reflecting a greater burden due to reactivation of infection acquired in periods of higher transmission early in life[7]. Chen et al describe impressive declines in reported tuberculosis in Taiwan, from 73 per 100,000py in 2005 to 44 per 100,000py in 2016, a decline of approximately 3.6% per year[1]. Consequently, it is not surprising that the proportion of disease among young people has declined at the same time, and that there were greater declines in TB among young people than older people. This is consistent with observations in Japan during a period of substantial decline in TB incidence[7], and with other countries in the Western Pacific region (Figure 1).

![Figure 1. Estimated burden of disease (outlines) and case notifications (female in red and male in teal) by age and sex in Africa, South East Asia, and the Western Pacific regions. Reproduced with permission from the WHO Global TB Report 2018[3].](image)

Secondly, there will usually be a discrepancy between notification data, which show only diagnosed and reported cases, and estimates of all incident disease which account for imperfect case-detection[3] (see the white gaps in Figure 1). Evidence is emerging that young people may be at higher risk of misdiagnosis and under-detection than adults[8]. This is consistent with the reality that in most settings, young people have reduced access to health services due to limited financial resources, the demands of school and work, family control over their ability to seek care, and lower health literacy than adults[9]. Our estimates were for all incident TB, including undiagnosed and unreported TB[2]. Consequently, our estimates are higher than what is shown in notification data, because we adjusted for the case detection gap.

Chen et al describe a strong TB control response in Taiwan over recent decades, including an expansion of contact tracing to cover almost 100% of close contacts in 2017[1]. This is a major achievement. We would add a note of caution, however, in response to the assertion that “young patients [...] may have compromised treatment adherence in some areas. Therefore, adolescents and young adults with TB clearly benefit more from the DOTS [Directly Observed Treatment, Short Course] programme”. Medication adherence is a well-recognised challenge for adolescents with any chronic health condition, including TB, due to complex developmental and health system factors[9]. However, the traditional approach of clinic-based DOTS-direct observation of therapy
(DOT) can majorly disrupt participation in education and employment, which will already be threatened by the diagnosis of a stigmatised condition like TB. Unless health clinics are co-located or close to schools, adolescents who are required to attend a daily clinic will miss many hours of education, a risk factor for poor learning, disengagement from schooling and early school leaving. The World Health Organization has de-emphasised clinic-based DOT in the End TB Strategy[10], and is encouraging TB services to develop and implement patient-centred models of care. Young people with TB may indeed require more intensive support to achieve consistent adherence to medications, especially with second-line drugs or when complex comorbidities such as HIV are present, and it poorly supported by family. However, this support should be delivered in ways that respect their developing autonomy, their desire for privacy, and the importance of their continued engagement in education or employment. We in the TB community have much to learn from colleagues in HIV and other chronic disease services with regard to providing high-quality, patient-centred care to young people with TB.

Yours sincerely,

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Centre for International Child Health
Department of Paediatrics
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On behalf of Prof Susan Sawyer, A/Prof Justin Denholm, Prof Stephen Graham

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Firstly, our figure of 17% is a global average, covering settings that are demographically and epidemiologically diverse. The age distribution of TB in any setting depends on three factors: the age structure of the population, the intensity of TB transmission, and the success of TB prevention efforts in different age groups. The proportion of TB cases which affect young people will be lower than average in less intense TB epidemics, in countries with older populations, and in countries with more successful prevention programs. At one extreme, in Western Europe, most nations have TB incidence below 20 cases per 100,000 person years (py)[3], and young people constitute only 14-18% of the total population[4]. At the other extreme, in southern Africa estimated TB incidence exceeds 300 per 100,000py[3] and fully one third of the population are aged 10-24[4], resulting in an epidemic concentrated among those in young and middle adulthood (see Figure 1). In Taiwan, young people constitute around 16% of the population, similar to Europe and other high income settings[5], and as noted by the authors, reported TB incidence is 44 cases per 100,000py. As such, it is to be expected that the burden of disease among young people in Taiwan is below the global average.

Furthermore, high rates of TB transmission have a disproportionate impact on children and adolescents[6]. As TB incidence declines, the burden of disease shifts into progressively older age groups, reflecting a greater burden due to reactivation of infection acquired in
periods of higher transmission early in life[7]. Chen et al describe impressive declines in reported TB in Taiwan, from 73 per 100,000py in 2005 to 44 per 100,000py in 2016, a decline of approximately 3.6% per year[1]. Consequently, it is not surprising that the proportion of disease among young people has declined at the same time, and that there were greater declines in TB among young people than older people. This is consistent with observations in Japan during a period of substantial decline in TB incidence[7], and with other countries in the Western Pacific region (Figure 1).

Figure 1. Estimated burden of disease (outlines) and case notifications (female in red and male in teal) by age and sex in Africa, South East Asia, and the Western Pacific regions. Reproduced with permission from the WHO Global TB Report 2018[3].

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Snow, KJ; Sawyer, SM; Denholm, JT; Graham, SM

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