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# Must countries shoulder the burden of mesothelioma to ban asbestos? A global assessment

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# ABSTRACT

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Professor Ken Takahashi: ken.takahashi@uwa.edu.au Introduction Mesothelioma is a key asbestos-related disease (ARD) but can be difficult to diagnose. Countries presumably ban asbestos to reduce future ARD burdens, but it is unknown if countries ban asbestos as a

consequence of ARD burdens. We assessed if and to what extent mesothelioma burden has an impact on a country banning asbestos and obtaining targets for preventative strategies.

Methods We analysed the status of asbestos ban and mesothelioma burden during 1990-2019 in 198 countries. We assessed mesothelioma burden by age-adjusted mortality rates (MRs) estimated by the Global Burden of Disease Study (GBD) and mesothelioma identification by the WHO mortality database. For GBD-estimated mesothelioma MR, the pre-ban period in the asbestosbanned countries was compared with the 1990-2019 period in the not-banned countries. For mesothelioma identification, the 1990-2019 period was applied to both banned and not-banned countries.

Results The association of mesothelioma MR with ban status increased as the ban year approached. Logistic regression analyses showed that the odds of a country banning asbestos increased 14.1-fold (95% CI 5.3 to 37.9) for mesothelioma identification combined with a 26% (12% to 42%) increase per unit increase of mesothelioma MR (one death per million per year) during the period 1-5 year before ban (model p<0.0001).

Conclusion Mesothelioma burden had an impact on, and together with its identification, explained the banning of asbestos in many countries. Asbestos-banned countries likely learnt lessons from their historical policies of using asbestos because mesothelioma burden and identification follow historical asbestos use. Prevention targets for ARD elimination should combine asbestos ban with mesothelioma identification.

# INTRODUCTION

Mesothelioma is a fatal malignancy with at least 80% attributable fraction to asbestos exposure.<sup>1</sup> The disease can be difficult to diagnose and only approximately 50% of countries report mesothelioma deaths to the WHO.<sup>2</sup> Nonetheless, due to its importance

### WHAT IS ALREADY KNOWN ON THIS TOPIC

- $\Rightarrow$  Six papers analysed the possible effect of a ban on the subsequent mesothelioma burden, that is, the post-ban effect: Four assessed individual countries and two studied the issue on a global scale.
- $\Rightarrow$  Another two global assessment studies investigated the possible effect of pre-ban factors on subsequent asbestos bans: one included an analysis of mesothelioma burden and another assessed the ratification status of international conventions.
- $\Rightarrow$  The former study had limited generalisability regarding the possible effect of mesothelioma on an asbestos ban because it covered relatively few countries using country-reported data in a study interwoven with several hypotheses.

## WHAT THIS STUDY ADDS

- $\Rightarrow$  Pre-ban mesothelioma mortality rates increased incrementally towards the ban year in asbestosbanned countries, and the difference to that of the not-banned countries widened as the ban year approached.
- $\Rightarrow$  Asbestos bans can be explained by pre-ban mesothelioma burden and by mesothelioma identification, independently and in combination.

### HOW THIS STUDY MIGHT AFFECT RESEARCH, **PRACTICE OR POLICY**

- $\Rightarrow$  Our study adds evidence that the burden and identification of mesothelioma are important drivers for banning asbestos.
- $\Rightarrow$  Asbestos-banned countries likely learnt lessons from their historical policies of using asbestos because mesothelioma burden and identification follow historical use of asbestos.
- $\Rightarrow$  International cooperation to eliminate asbestosrelated disease should integrate technologies to identify mesothelioma in the roadmap to ban asbestos.

as a key asbestos-related disease (ARD), the burden of disease has been estimated for many countries of the world.<sup>3-5</sup> The Global Burden of Diseases Study (GBD) recently

estimated that 29300 people worldwide die each year from mesothelioma.  $\!\!\!\!^4$ 

The WHO has been calling on countries to cease the use of asbestos in order to eliminate ARD.<sup>6</sup> Since the mid-1980s, around 60 countries have adopted asbestos bans,<sup>7</sup> but the pace of new bans has recently slowed.<sup>8</sup> The current global consumption of raw asbestos lingers at around 1 million metric tonnes per year.<sup>9</sup>

At the national level, the course of ARD burdens closely follows that of asbestos consumption after a period of several decades,<sup>10–12</sup> which is reflected in geographical correlations between disease burdens and historical asbestos use.<sup>13 14</sup> These relationships add to the scientific basis of policies to justify asbestos bans. In our recent literature review,<sup>8</sup> we addressed the question of whether mesothelioma burdens decreased *after* asbestos bans and concluded that the timing of studies was too early to demonstrate the full effect of bans. Our review also emphasised the importance of asking the alternative question of whether high mesothelioma burdens burdens *preceded* asbestos bans. This is part of a larger question as to why countries ban asbestos.

A series of articles aimed to elucidate how asbestosbanned countries transitioned from asbestos using to asbestos banned.<sup>15</sup> A myriad of pathways was unravelled, involving wide variations in sociopolitical, juridical, cultural, scientific and technological conditions and the interplay of various stakeholders, for example, patients, non-governmental organisations, workers/employers, administrators and the health/scientific sectors. This raised the question: what common factor incentivised stakeholders to ultimately ban asbestos? We hypothesise this may be the ARD burden experienced by a country in the period preceding a ban.

Countries presumably ban asbestos *for the purpose* of eliminating the future ARD burden. However, it is unknown if countries ban asbestos as a consequence of the ARD burden. We believe this question merits thorough examination. Estimated burdens of mesothelioma are now available for a wide range of countries, and can be used to represent the ARD burden at a relevant time. Here, we aimed to determine if and to what extent mesothelioma burden has an impact on a country banning asbestos and obtaining targets for preventative strategies.

#### **METHODS**

We analysed the status of countries with data available for both the mesothelioma burden and asbestos ban. Due to limited reporting of mesothelioma burden by countries, we used widely referenced data estimated by the GBD, which analyses the global situation of diseases and injuries for a wide range of countries.

Recent GBD covers the period from 1990 to 2019 and includes estimated age-adjusted mortality rates (MRs) for mesothelioma.<sup>4 16</sup> We employed the GBD-estimated age-adjusted MR of mesothelioma to represent the mesothelioma burden in each country. In the GBD and our study,

mesothelioma was defined as C45 including all its subcategories in accordance with the International Classification of Diseases and Related Health Problems, 10th Revision<sup>17</sup> (ICD-10). Our study subjects comprised 198 countries for which a GBD-estimated mesothelioma MR was available during our observation period of 1990–2019.

Mesothelioma MR data were downloaded from the global health data exchange of the Institute for Health Metrics and Evaluation,<sup>16</sup> University of Washington, USA, which states that data 'can be used, shared, modified or built on by non-commercial users via the Creative Commons Attribution-NonCommercial-NoDerivatives 4.0 International License'.<sup>18</sup>

We calculated the average and median of mesothelioma MR during the period before asbestos ban (pre-ban) in asbestos-banned countries and during the entire observation period in not-banned countries. Note that the latter period represents the period during which asbestos was not banned despite the accumulation of mortality in the not-banned countries.

Six countries (Austria, Denmark, Iceland, Norway, Sweden and Switzerland) banned asbestos in or before 1990 (ie, 'early banned') and thus lacked data on the preban mesothelioma MR. These countries were excluded from our main analysis of 198 countries and were instead analysed for their GBD-estimated mesothelioma MR during 1990–1994 (ie, the earliest period with available data on mesothelioma MR) to approximate pre-ban mesothelioma MR.

We defined mesothelioma identification as the entire process from clinical identification and attribution as cause of death to notification of statistics to the WHO. We refer to the country status of mesothelioma identification as 'mesothelioma identified' when data were present and 'not-identified' when data were absent from the WHO mortality database<sup>19</sup> during the entire observation period for both banned and not-banned countries. The first year of mesothelioma identification for a particular country was defined as the year that data on mesothelioma deaths first appeared for that country in the WHO mortality database. Note that mesothelioma was introduced into the WHO mortality database for the first time in 1994 and the mesothelioma identification may have been initiated before or after the asbestos ban in banned countries.

We defined asbestos ban as a country's banning of the production, import and new use of asbestos. We refer to a country's asbestos ban status as 'asbestos banned' or 'not-banned' and the year of asbestos ban adoption based on the database of the International Ban Asbestos Secretariat.<sup>7</sup>

 $\chi^2$  test was used to examine the association between the country status of asbestos ban and mesothelioma identification, each as a binary variable (yes/no). T-test and analysis of variance were used to compare the average MR between and across country groups, respectively. Logistic regression analysis was conducted to assess if and to what extent mesothelioma identification and MR explained asbestos ban. We calculated the variance inflation

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 Table 1
 Country status of asbestos ban and mesothelioma identification in 198 countries\*

		Asbestos ban		
		Banned	Not- banned	Total
Mesothelioma identification	Identified	48	54	102
	Not-identified	6	90	96
Total		54	144	198

P<0.0001 based on  $\chi^2$  test for independence of asbestos ban and mesothelioma identification.

See Methods for full definition of asbestos ban and mesothelioma identification.

\*Six 'early-banned' countries that banned asbestos in or before 1990, that is, Austria, Denmark, Iceland, Norway, Sweden and Switzerland, are precluded from this analysis and are analysed separately in table 4. See Methods for full explanation.

factor (VIF) using SAS and the VIF function in the car package<sup>20 21</sup> of R to investigate collinearity between the explanatory variables. Microsoft Excel (Microsoft Corporation, Redmond, Washington, USA), SAS V.9.4 and  $R^{21}$  were used for data analysis. Stata software V.17.0 (Stata

Corporation LLC, College Station, Texas, USA) was used to generate the box plot.

#### RESULTS

Table 1 shows the status of 198 countries in terms of asbestos ban and mesothelioma identification. Regarding asbestos ban, 54 (27%) of 198 countries banned asbestos and 144 (73%) did not. Regarding mesothelioma identification, 102 (52%) of 198 countries identified mesothelioma and 96 (48%) did not. When cross-stratified, 89% (48) of the 54 banned countries and 38% (54) of the 144 not-banned countries identified mesothelioma, and 48 (47%) of the 102 mesothelioma-identified countries and 6 (6%) of the 96 not-identified countries banned asbestos. Thus, asbestos ban and mesothelioma identification were closely associated (p<0.0001).

Table 2 shows mesothelioma MR by the status of asbestos ban and mesothelioma identification. The average MR was approximately two times as high in the asbestos-banned versus not-banned countries (5.14 vs 2.55 deaths per million per year, respectively; p=0.0004). The average MR was approximately 40% higher in the

 Table 2
 Global Burden of Disease Study (GBD)-estimated mortality rate (MR) of mesothelioma by country status and period in 198 countries\*

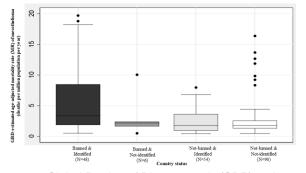
		GBD-estimated MR (average)	GBD-estimated MR (median)		
Country status	Status (countries with data, N)	(Deaths per million pe	r year)	P value†	
Asbestos ban	Banned (54)	5.14	3.13	0.0004	
	Not-banned (144)	2.55	1.79		
Mesothelioma identification	Identified (102)	3.80	2.26	0.0231	
	Not-identified (96)	2.69	1.85		
Ban and identification	Banned and identified (48)	5.39	3.34	<0.0001	
	Banned and not-identified (6)	3.15	2.18		
	Not-banned and identified (54)	2.39	1.79		
	Not-banned and not-identified (90)	2.65	1.79		
Period before year of ban or 2019‡					
1–5 years	Banned (54)	5.17	2.86	0.0002	
	Not-banned (144)	2.46	1.80		
6–10 years	Banned (48)	4.72	2.86	0.0018	
	Not-banned (144)	2.49	1.82		
>10 years	Banned (39)	4.23	2.60	0.0208	
	Not-banned (144)	2.59	1.82		
All years	Banned (54)	5.14	3.13	0.0004	
	Not-banned (144)	2.55	1.79		

See Methods for full definition of asbestos ban and mesothelioma identification.

\* Six 'early-banned' countries that banned asbestos in or before 1990, that is, Austria, Denmark, Iceland, Norway, Sweden and Switzerland, are precluded from this analysis and are analysed separately in table 4. See Methods for full explanation.

†P value is for difference of average GBD-estimated MR between two groups or across groups.

‡Period before the year of ban in asbestos-banned countries or period before 2019 in not-banned countries.



Global Burden of Disease study (GDB)-estimated Figure 1 age-adjusted mortality rates (MR) of mesothelioma in countries grouped by the status of asbestos ban and mesothelioma identification in 198 countries. (1) Six 'earlybanned' countries that banned asbestos in or before 1990, that is, Austria, Denmark, Iceland, Norway, Sweden and Switzerland, are precluded from this analysis and are analysed separately in table 4. See Methods for full explanation. The outlier countries in the respective country groups indicated by points are as follows: 'Banned and identified': the UK (19.7), Australia (18.8); 'banned and notidentifed': Monaco (10.1), Algeria (0.5); "not-banned and identified': Amenia (8.0); 'not-banned and not-identified'): Andorra (16.4), Lesotho (13.7), Namibia (12.7), Eswatini (11.9), Botswana (9.9), Greenland (9.2), San Marinho (8.3). Number within brackets indicate values of mesothelioma MR. See Methods for full definition of asbestos ban and mesothelioma identification. This box plot shows the 25th percentile (Q1; bottom of box), median (Q2; horizontal line in box) and 75th percentile (Q3; top of box). Outliers outside the interquartile range are displayed as 'whisker' lines of points.

mesothelioma-identified versus not-identified countries (3.80 vs 2.69 deaths per million per year, respectively; p=0.0231). When cross-stratified, average MR was highest in the 'banned and identified' group, followed by the 'banned and not-identified', 'not-banned and not-identified' and 'not-banned and identified' groups at 5.39, 3.15, 2.65 and 2.39 deaths per million per year, respectively (p<0.0001 for difference in the average across groups). There was no difference in the median MR of the two 'not-banned' groups. By grouping the period before the ban year, the average MR of banned countries increased and the difference in the average MR widened between banned and not-banned countries, incrementally towards the ban year.

Figure 1 shows the data distribution of mesothelioma MR as a box plot in the four country groups. Several outlier countries (indicated as points) of the 'not-banned and not-identified' and 'banned and not-identified' groups had mesothelioma MR exceeding the 75th percentile (third quartile or Q3) of the 'banned and identified' group: Andorra (16.4), Lesotho (13.7), Namibia (12.7), Eswatini (11.9) and Botswana (9.9) in the 'not-banned and not-identified' group and Monaco (10.1) in the 'banned and not-identified' group (MR values shown in brackets; unit: deaths per million per year).

Table 3 shows findings from the logistic regression analyses explaining asbestos bans in 198 countries. In the single-variate models, the odds of asbestos ban: (1) increased 13.3-fold (95% CI 5.4 to 33.2) when the country identified mesothelioma; (2) increased incrementally towards the ban year by 16% (4% to 29%), 21% (9% to 34%) and 25% (13% to 39%) per unit increase of MR (ie, one death per million per year) during>10 years (p=0.0057), 6–10 years (p=0.0003) and 1–5 years (p<0.0001), before ban, respectively; and (3) increased by 22% (11% to 35%) per unit increase of MR during all years before ban (p<0.0001).

In the two-variate models, asbestos ban was regressed on a combination of explanatory variables that showed the highest statistical significance in the single-variate model: the common explanatory variable was mesothelioma identification, and the other explanatory variable was mesothelioma MR during 1-5 years before ban or that during all years before ban. Both models explained asbestos ban with high statistical significance (p<0.0001); the former model showed that the odds of asbestos ban increased 14.1-fold (5.3 to 37.9) for mesothelioma identification combined with a 26% (12% to 42%) increase per unit increase of MR during 1-5 years before ban. Collinearity of the two explanatory variables, mesothelioma MR and identification, on the outcome variable of asbestos ban was assessed by VIF, which yielded 1.026 and 1.025, respectively. Given that these values are effectively 1, collinearity was not an issue for the modelling.

Table 4 shows the characteristics of the six 'earlybanned' countries that were analysed separately due to the non-availability of pre-ban mesothelioma MR. All six 'early-banned countries' identified mesothelioma, and their mesothelioma MR for 1990–1994 (ie, the earliest available period) were high relative to those of the 'banned and identified' group: four exceeded the 75th percentile (Q3) and two were below Q3 but exceeded the average and median values.

Online supplemental figure 1 highlights the relationship between the ban year and mesothelioma MR in asbestos-banned countries. In the 'banned and identified' group, higher MR generally corresponded to earlier years of ban (eg, the Netherlands, Italy and the UK), although with notable exceptions: Canada (and Turkey and South Africa, to a lesser extent) was late to ban asbestos despite having a relatively high mesothelioma MR whereas Brunei and Kuwait were early to ban asbestos despite having a relatively low MR. Online supplemental table 1 presents the values of all country variables for each country.

### DISCUSSION

Asbestos-banned countries comprised less than a third, and mesothelioma-identified countries approximately half of the 198 analysed countries. The country status of asbestos ban and mesothelioma identification (each a binary variable) were closely associated: the majority of banned countries identified mesothelioma but less than 40% of not-banned countries did, and close to half of

Table 3         Logistic regression analyses explaining asbestos ban in 198 countries*				
	Parameter estimate	OR	95% CI	P value
One variate model				
Mesothelioma identification	2.59	13.33	5.35 to 33.23	< 0.0001
Global Burden of Disease Study-estimated mortality rate (MR) of mesothelioma during:				
All years before (year of ban or 2019)	0.20	1.22	1.11 to 1.35	< 0.0001
1–5 years before (year of ban or 2019)	0.22	1.25	1.13 to 1.39	< 0.0001
6-10 years before (year of ban or 2019)	0.19	1.21	1.09 to 1.34	0.0003
>10 years before (year of ban or 2019)	0.15	1.16	1.04 to 1.29	0.0057
Two variate model				
I. Mesothelioma identification and MR of mesothelioma during† ‡	Model p<0.000	1		
Mesothelioma identification	2.60	13.53	5.151 to 35.52	< 0.0001
All years before (year of ban or 2019)	0.20	1.22	1.10 to 1.37	0.0004
II. Mesothelioma identification and MR of mesothelioma during†‡	Model p<0.000	1		
Mesothelioma identification	2.65	14.13	5.27 to 37.86	<0.0001
1–5 years before (year of ban 2019)	0.23	1.26	1.12 to 1.42	0.0001

See Methods for full definition of asbestos ban and mesothelioma identification.

\*Six 'early-banned' countries that banned asbestos in or before 1990, that is, Austria, Denmark, Iceland, Norway, Sweden and Switzerland, are precluded from this analysis and are analysed separately in table 4. See Methods for full explanation.

†) Period before the year of ban in asbestos-banned countries and period before 2019 in not-banned countries.

<sup>‡</sup>) Variance inflation factor (VIF) was used to assess collinearity between explanatory variable; VIF were 1.026 for mesothelioma identification and 1.025 for mesothelioma MR during 1–5 years before year of ban or 2019.

Table 4Characteristics of six 'early-banned' countriesseparately analysed by year of ban and GBD-estimatedmortality rate (MR) of mesothelioma during 1990–1994,contrasted with the MR of 48 'banned and identified'countries

Country	Year of ban	GBD-estimated mesothelioma MR during 1990–1994 (deaths per million per year)	
'Early-banned' country			
Austria	1990	5.66	
Denmark	1980	9.41	
Iceland	1983	7.05	
Norway	1984	15.55	
Sweden	1986	8.37	
Switzerland	1989	10.23	
'Banned and identified'	1991–2019	Average: 5.32, median 3.14	
countries (n=48)		(First quartile 1.82, third quartile 8.07)	
See Methods for full definition of asbestos ban.			

the identified countries banned asbestos but only a small fraction (6%) of not-identified countries did. Mesothelioma MR was higher in the banned (vs not-banned) and in the identified (vs not-identified) countries. When cross-stratified, MR was highest in the 'banned and identified' group and lowest in the two 'not-banned' groups. Over time, MR increased incrementally and was consistently higher in the banned than not-banned countries as the ban year approached. Logistic regression analyses showed that asbestos ban is explained by the mesothelioma MR during the 5-year period before a ban, independently, and in combination with mesothelioma identification. The present study thus demonstrated that the pre-ban mesothelioma burden is an important driver to ban asbestos.

Asbestos-banned countries were more likely to identify mesothelioma, and mesothelioma-identified countries were more likely to ban asbestos. Although cause and effect cannot be discerned because mesothelioma identification did not necessarily precede ban, we believe that an association between asbestos ban and mesothelioma identification is plausible, because a high burden of disease is likely to prompt practitioners and authorities to identify and notify the disease. However, this may depend on other factors that we did not examine, such as the level of knowledge of stakeholders regarding the relationship between asbestos and mesothelioma. Banned countries incurred higher pre-ban mesothelioma MR than not-banned countries. Notably, the incremental increases of MR in banned countries and of the difference in MR between banned and not-banned countries towards the ban year strengthen the notion that the preceding mesothelioma burden becomes increasingly important over time to ban asbestos. We speculate that mesothelioma burdens reinforce knowledge and raise concern of various stakeholders, culminating in a societal imperative to ban asbestos.

Logistic regression analyses showed that the odds of a country banning asbestos increased incrementally in response to pre-ban MR towards the ban year. Mesothelioma identification also explained asbestos ban in both the single-variate and two-variate models. The two-variate models were not impacted by collinearity between the explanatory variables of mesothelioma MR and mesothelioma identification. The ARD burden at a certain point in time includes a *latent* component (due to the latency between exposure and disease) which is not detectable until the latency period saturates. In our study, the preban mesothelioma MR in asbestos-banned countries represented only the *detectable* component. In contrast, mesothelioma identification (defined for the period until 2019) reflected the detectable and latent components. Thus, it is reasonable that pre-ban mesothelioma MR and mesothelioma identification emerged as independent explanatory factors for countries to ban asbestos.

The fundamental strategy to eliminate ARD is to cease asbestos use.<sup>6</sup> International cooperation for ARD emphasises the banning of asbestos<sup>22</sup> to effectively prevent future disease burdens. Although the importance of asbestos bans cannot be understated, asbestos bans per se have no impact in alleviating the level of current ARD burden which result from historical asbestos use. Asbestos-banned countries likely learnt lessons from their historical policies of using asbestos via the mesothelioma burden and identification. International cooperation should thus integrate the theme of mesothelioma identification, including the sharing of relevant technologies, in the roadmap to ban asbestos. The experience, expertise and technology acquired by the 'banned and identified' countries are valuable assets that can be used to support the global elimination of ARD. From a clinical standpoint, novel diagnostic tools are needed, such as easy-to-use kits for immunohistochemical staining, highutility tissue and blood biomarkers and telepathology.

Many studies have featured themes of asbestos ban status<sup>24–26</sup> or processes.<sup>15</sup> Several studies reported higher mesothelioma burdens in asbestos-banned countries, relative to not-banned countries, based on straightforward, cross-sectional comparisons.<sup>24–26</sup> Only a few previous studies have undertaken global assessment to identify factors that may subsequently promote asbestos bans. Seeking to explain 'why some, but not all, countries banned asbestos', Bahk *et al*<sup>27</sup> analysed the crude MR of mesothelioma as a factor that may lead to an asbestos ban. However, the Bahk *et al* study did not yield a definitive

answer because the question was set in a larger framework of societal context interwoven with several hypotheses. Lin *et al*<sup>28</sup> identified two international conventions and government effectiveness as facilitators of asbestos bans. To our knowledge, our paper is the first to directly investigate if and to what extent mesothelioma burden has an impact on countries deciding to ban asbestos. The association among pre-ban mesothelioma MR, mesothelioma identification and asbestos ban did not apply to certain countries. Relative to the MR of the 'banned and identified' countries, we found high MR in the 'not-banned and not-identified' countries of Andorra Lesotho Namibia Eswatini and Botswana and

in the 'not-banned and not-identified' countries of Andorra, Lesotho, Namibia, Eswatini and Botswana and in the 'banned and not-identified' country of Monaco (figure 1). These countries can learn from the policies of the 'banned and identified' countries, as such policies may 'diffuse across countries of similar background'.27 Although the history of asbestos and ARD is unique to each country,<sup>15</sup> several common patterns emerged regarding the relationship between the timing of a ban and the mesothelioma MR: (1) countries with high MR that banned asbestos early (eg, Netherlands, Italy and the UK and the six 'early-banned' countries-see below for the latter); (2) some countries without high MR that banned asbestos early (eg, Brunei and Kuwait); and (3) some countries with high MR that banned asbestos late (eg, Canada, South Africa and Turkey-historically asbestos-producing and asbestos-exporting countries) (online supplemental figure 1). Separately, political factors play important roles as typified by the EU Directive which effectually mandated all EU member states that had not yet banned asbestos to do so by January 2005.<sup>8</sup>

We did not include the six 'early-banned' countries in our main analysis because data on their pre-ban mesothelioma MR were not available in GBD. We thus substituted the pre-ban MR with the 1990-1994 MR in a separate analysis and found that the 1990-1994 MR of the 'early-banned countries' was compatible with the upper range of that of the 'banned and identified' countries (table 4). We speculate that the pre-ban MR of the 'early-banned countries' was also in the upper range of that of the 'banned & identified' countries. This speculation is supported by our observation that the pre-ban standardised incidence rates of male pleural mesothelioma in the early-banned countries of Denmark, Norway and Sweden were higher than or similar to that of Finland,<sup>29</sup> which had an upper range pre-ban MR of the 'banned and identified' countries. Thus, the associations of asbestos bans with mesothelioma MR and identification found in this study would likely be strengthened if we could include the 'early-banned' countries in the main analysis.

A major strength of our study was that the national experiences of a wide range of countries were incorporated based on comparable data from international sources,<sup>71619</sup> enabling a global assessment. Another advantage was that distinct aspects of key variables were taken into consideration, that is, the level and identification of

the mesothelioma burden and the status and timing of ban. Also, the time sequence of mesothelioma burden and asbestos ban were adequately considered (the former variable was defined as preceding the latter). A major limitation of our study was our total dependence on GBD estimates which are prone to error particularly for developing countries due to uncertainties in the original information source as well as our narrow focus on the relationship between mesothelioma and bans. The dynamics leading to an asbestos ban are complex with many factors interacting in myriad ways.<sup>15</sup> Thus, our approach may be oversimplified and may have overlooked important factors such as the level of stakeholders' knowledge (as mentioned earlier), national economic status (eg, GDP) and political motives (eg, camouflage of mesothelioma status). We were unable to directly analyse data of the six 'early-banned' countries, which required a separate analysis. Furthermore, our study did not account for variations in completeness of asbestos ban. Finally, mesothelioma identification was not limited to the pre-ban period because the disease category was only first introduced into the WHO mortality database in 1994.

In conclusion, we herein report that mesothelioma burden had an impact on, and together with its identification, explained the banning of asbestos in many countries. These countries likely banned asbestos by taking lessons from their historical policies of using asbestos because the mesothelioma burden and its identification followed historical asbestos use. The identification of mesothelioma as a 'signal tumour' of the ARD burden is important to enhance the pace of asbestos bans. The world should adopt a comprehensive preventative strategy for ARD elimination that combines the banning of asbestos with the identification of mesothelioma.

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