The prevalence of HIV, HBV and HCV among Filipino blood donors and overseas work visa applicants

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Objective  Our aim was to estimate the prevalence of HIV, HBV and HCV among the general population of the Philippines using data sources outside of the limited existing active surveillance network.

Methods  We analysed aggregate HIV, HBV and HCV test results for hospital-based blood donors (BDs) and overseas Filipino worker candidates (OFWCs) that had been reported from licensed laboratories to the National STD/AIDS Cooperative Central Laboratory in Manila between 2002 and 2004.

Findings  From over 144,000 blood-screening results, the HIV prevalence was 0.006% in BDs and 0.001% in OFWCs; that of HBV was 4.2% in both groups; and that of HCV was 0.3% in BDs and 0.9% in OFWCs. Males were at increased risk of both HBV and HCV; among OFWCs, younger women were at increased risk. Laboratories that tested sequentially but stopped testing after the first positive result were far less likely to detect HCV, indicating that sequential testing protocols may underestimate HCV and HIV prevalence. OFWCs were at low risk of HIV, and the risk of testing positive for these viruses was not increased among OFWCs applying for a repeated work visa, compared with first-time applicants.

Conclusion  Based on these data, we conclude that HIV is rare in the Philippines. In contrast with prior reports, we found no evidence that OFWCs constitute a high-risk group for HIV. Further research is needed to understand why younger women are at increased risk of acquiring HBV.


Introduction

Limited surveillance data exist in the Philippines for the three bloodborne viruses (BBVs) of greatest importance: human immunodeficiency virus (HIV), hepatitis B virus (HBV) and hepatitis C virus (HCV). As of December 2005, only 2410 HIV-positive patients had been reported to the national HIV/AIDS registry. In addition to the HIV/AIDS registry, the Philippines conducts active surveillance on HIV and other sexually transmitted infections (STIs) among high-risk groups such as female sex workers (FSWs), men who have sex with men and injecting drug users, but has no surveillance system outside these high-risk groups by which to estimate BBV trends in the general population. The Philippines Department of Health was advised to conduct sentinel surveys for blood banks and overseas Filipino workers, which could contribute to the identification of new high-risk populations.

In this study, we investigated the prevalence of BBVs from two informal data sets, both collected for reasons other than disease surveillance. The first source was hospital-based blood banks. WHO/UNAIDS currently recommends testing of blood donors (BDs) as a reasonably low-cost approach for HIV surveillance in countries with low rates of HIV infection. The second source was persons applying for foreign work visa permits. These overseas Filipino worker candidates (OFWCs) are an important demographic because Filipino migrant workers contribute tremendously to the economy of the Philippines, and because it is believed that returning workers may be an important source of HIV infections in the country. Since 1999, more than 40% of HIV/AIDS registered cases have previously been overseas. Several overseas employees have responded to this by making HIV testing, and occasionally testing for other infections, a systematic requirement of their employer and/or host countries.

Our goals were to estimate the prevalence of HIV, HBV and HCV among these populations, to contrast the prevalence of these viruses between the two source groups, to explore epidemiologic risk factors for BBV positivity; and, because these sets represent convenience samples, to probe where possible for evidence of systematic biases that might impact our interpretation of the sets.

Methods

Data source

We analysed blood-screening data for HIV, HBV and HCV among BDs and OFWCs from 2002–04. These data are reported to the STD/AIDS Cooperative Central Laboratory (SACCL) in Manila. This network of laboratories submits
quartely aggregate results to SACCL on the number of tests performed and positive cases, categorized by sex and age. While the potential sources of data were quite large, actual reporting was far lower. Of the over 130 accredited laboratories for the OFWCs screening test, and over 150 licensed as blood banks, the number of laboratories with eligible data for this study that actually reported to SACCL was only 17, 13 and 11 in successive years for BDs; and 14, 16 and 9 for OFWCs for the years 2002, 2003 and 2004, respectively.

BBV test results for BDs derived from hospital-based blood banks. Procedures for blood screening varied between laboratories, with some adopting either sequential or simultaneous testing protocols for the three BBVs. In the sequential testing system, a blood unit would be rejected after the first virus found positive, with no further testing done. Simultaneous testing laboratories generate results for a given BBV, irrespective of the results for the other two BBVs. Because we were usually unable to determine from the laboratories or from SACCL directly whether a given laboratory followed a sequential versus simultaneous testing protocol, we presumptively classified laboratories as either simultaneous or sequential testers by examining the total number of tests for each virus. Because HBV was the most common virus detected, if the number of HIV or HCV samples tested was approximately equal to the number of HBV tested minus the number of HBV positive results we considered them to be simultaneous testers. If the numbers of tests done on each virus were approximately equal, we considered them to be simultaneous testers. Fig. 1 shows a flowchart of the BBV tests for BDs and OFWCs.

For OFWCs, we collected data from accredited OFWC clinics/hospitals. In contrast with blood banks, where screening protocols are dictated by hospital policy, the decision as to which viruses to screen for in OFWCs — or the decision to screen at all — depended on the visa requirements for the host countries to which OFWCs were applying.

Laboratory methods
For HIV, the initial anti-HIV-1 and HIV-2 screening tests were conducted at the laboratories, with confirmatory tests done at SACCL for OFWCs and at the Research Institution for Tropical Medicine for BDs, using immunofluorescent antibody assays and Western blotting. Since HBV screening was through detection of hepatitis B surface antigen (HBsAg), the test only detects acute or chronic active viral hepatitis B. HCV was screened via anti-HCV antibody. The screening test kits varied between laboratories, although they were evaluated by the Department of Health.

Analytic methods
Because the data are reported to SACCL only in aggregate form, we were unable to perform statistical analyses at the level of individual patients. We calculated BD and OFWC positivity rates as the number of positive samples divided by the total number screened. For comparisons of HIV, HBV and HCV infections between groups of people, we calculated relative risks (RR) and 95% confidence intervals (95% CI).

Results
Our final data set consisted of approximately 144,000 blood samples. The precise number of patients could not be ascertained from the aggregate data, but based on the most common test performed (HBsAg) would have been at least 144,624. The majority of samples were collected from large cities. Among 63,249 HIV tests for BDs, 17,326 (27%) were reported from the National Capital Region, 10,589 (17%), 10,345 (16%), and 9,714 (15%) from Southern Mindanao, Southern Tagalog and the Central Visayas, respectively. Of the 69,123 HIV tests for OFWCs, 64,764 (94%) were from the National Capital Region.

Table 1 shows prevalence ratios among BDs and OFWCs. The sex ratio differed substantially between the two groups, with 92% of BDs being male versus ca. 42% for OFWCs. Because HIV cases were few and the prevalence of HCV and HBV were similar across the three years of data collection, we combined the data from 2002–2004 in our analyses.

BBV = bloodborne viruses; HBV = hepatitis B virus; HCV = hepatitis C virus; HIV = human immunodeficiency virus; neg. = negative; pos. = positive.
HIV rates were extremely low in both BDs and OFWCs. HIV was detected in only 4 of 63,249 BDs screened (prevalence 0.006%) and only 1 of 69,123 OFWCs (prevalence 0.001%). All 5 positive cases were men.

HBsAg was detected in 6,030 out of 144,624 screened (2704 of 64,954 BDs [prevalence, 4.2%] and 3,326 of 79,670 OFWCs [prevalence, 4.2%]). The HBV rates were similar between BDs and OFWCs (RR, 1.00 [95% CI: 0.95–1.05]). The risk was higher in males for both BDs (male prevalence, 4.3% and female prevalence, 2.9%; RR 1.14 [95% CI: 1.24–1.71]), and OFWCs (4.5% versus 4.0%; RR 1.14 [95% CI: 1.06–1.22]). Among the groups tested, age proved to be an important risk factor only in the case of hepatitis B among female OFWCs. In this group, HBV prevalence was inversely related to age, with a consistent gradient of risk when comparing each age group to that above it (Table 2). Women under 19 years were far more likely to test positive for HBV when compared with any of the higher age categories. Similarly, women aged 19–29 years and women aged 30–39 years were also more likely to test positive for HBV compared with the higher age categories.

HCV was detected in 31 of 74,180 screened (208 of 63,166 BDs [prevalence, 0.3%] and 103 of 11,014 OFWCs [prevalence, 0.9%]). HCV prevalence in BDs was almost one-third in OFWCs (RR 0.35 [95% CI: 0.28–0.45]). The HCV prevalence was significantly higher among male BDs than female BDs (RR 3.60 [95% CI: 1.48–8.74]), but did not differ significantly between male OFWCs and female OFWCs (RR 0.97 [95% CI: 0.66–1.45]).

versus simultaneous testing among BDs

Given that the HCV rates among BDs were far lower than among OFWCs, whereas the HBV rates were similar for each group, we considered whether some of this difference could be an artefact of the sequential testing approach used at many blood banks, a bias that should not apply to the OFWCs. Therefore, we performed a sensitivity analysis in which we contrasted the rates of HBV and HCV depending on whether the laboratory used a sequential versus a simultaneous testing protocol for the BDs. Because HBV is the first virus screened for (due to the prevalence of the disease and the low cost of the assay relative to HIV or HCV), we hypothesized that this would have the effect of systematically under reporting the prevalence of HCV and HIV. As shown in Table 3, 8 blood banks reported or appeared to be performing sequential testing, whereas 18 blood banks conducted sequential testing. Approximately 5% fewer tests were submitted for HIV and HCV than for HBV among the sequential testing laboratories. Compared with simultaneous testing laboratories, sequential testers (Table 4) were far less likely to detect HCV infections (RR, 0.33 [95% CI: 0.22–0.41]), whereas no such effect was seen for HBV infections (RR, 0.99 [95% CI: 0.92–1.06]). The HIV prevalence was also lower among the sequential testing laboratories, though this difference was not statistically significant (RR, 0.30 [95% CI: 0.03–2.85]).

The effect of sequential testing

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should have an increased risk of BBVs compared with those applying for their first trip abroad.

We analysed data from the 17 OFWC testing clinics/hospitals that reported data separately for first-time and repeat OFWCs. Contrary to our expectations, the prevalence of HBV did not vary between the two groups as a function of whether the OFWC was a first-time or a repeat applicant (Table 4, RR, 1.05 [95% CI: 0.97–1.14]). In the case of HCV, first-time OFWCs were actually more likely to test positive than repeated OFWCs (RR, 1.53 [95% CI: 1.03–2.28]).

Discussion
In the published literature, our study was the first to examine the effect of first-time versus repeat visa application as a risk factor for BBVs among OFWCs, and the first to contrast directly the BBV prevalence between BDs and OFWCs. Our analysis showed that HIV was extremely uncommon among this population of BDs and OFWCs, whereas HBV was very common and the prevalence of HCV was between the two. Significantly, our data suggested a far lower prevalence of HIV among OFWCs than previous estimates of 0.01%. In contrast with previously identified high-risk groups, such as injecting drug users, men who have sex with men and FSWs, our analysis showed a negligible risk for HIV among those applying for foreign work visa permits — only a single case of HIV among about 70,000 OFWC samples tested. Moreover, the risk of all three viruses among OFWCs applying for their second or subsequent permits to work abroad, among whom the prevalence of BBVs should logically have been higher if prior overseas work was a risk factor for infection, was no different for HIV and HBV. For HCV infection, the risk was lower among repeat OFWCs.

How can we reconcile our findings with previous reports on the prevalence of BBVs among workers returning from overseas? One possible explanation is the manner in which HIV cases are registered in the Philippines. To date, overseas Filipino workers have been the largest single demographic group in the HIV/AIDS registry. However, this could largely be due to ascertainment bias, given that the visa testing requirements imposed on OFWCs do not apply to the general population. Because the OFWCs are oversampled in the national registry, they may appear more important than they really are, principally because these totals do not consider the denominator that represents the population from which they were drawn.

Likewise, HBV seroprevalence was comparable to the 2–5% among the general population in previous studies, but far lower than a WHO report of 8–10% overall, 10% among males, and 10% in injecting drug users. The HCV prevalence in the study was slightly lower than previous data of 2–4% for the general population, but far lower than the 70% reported for injecting drug users.

We observed higher rates for HBV and HCV infection among males than females. Why this might be the case cannot be explained using the aggregate

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<th>Table 3. Effect of age on HBsAg positivity among female OFWCs</th>
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HBsAg, hepatitis B surface antigen; OFWCs, overseas Filipino worker candidates.

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<th>Table 4. Effect of sequential versus simultaneous testing protocol on risk of HBV/HIV/HCV positivity</th>
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<td>Testing protocol</td>
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RR = 0.99* 95% CI: 0.92–1.06
RR = 0.30* 95% CI: 0.03–2.85
RR = 0.33* 95% CI: 0.22–0.41

Ab, antibody; CI, confidence interval; HBsAg, hepatitis B surface antigen; HBV, hepatitis B virus; HCV, hepatitis C virus; HIV, human immunodeficiency virus; RR, relative risk.

* Risk is for sequential versus simultaneous.
data that was available to us. However, we hypothesize that it could represent different patterns of sexual activity or higher rates of injecting drug use among males. The finding that among OFWCs younger women were at higher risk of HBV might also represent high-risk sexual behaviour in this group. In partial support of this theory, a recent study showed that the Chlamydia infection rate of antenatal women was significantly higher among younger women.\(^7\)

Our report sheds light on the limitations of sequential testing among BDs. Among sequential testing groups, HIV and HCV tests were done only for blood units that tested negative for HBsAg. Because it is unlikely that co-infection rates for HBV and HCV and/or HIV occur in a random fashion, this would lead to a proportionally greater underestimation of these infections. Previous studies have shown that 7–18% of those with HBV infection were co-infected with HCV.\(^8\) Also, a significant percentage of HIV-infected patients were co-infected with HBV and/or HCV.\(^9\)–\(^23\) To provide safe blood within limited budgets, many blood banks have adopted the sequential testing approach to contain costs associated with HIV and HCV tests. This ongoing testing system for donated blood misses these co-infection cases. Given the residual uncertainty regarding whether laboratories were truly using simultaneous versus sequential testing, our analysis is vulnerable to misclassification. However, such misclassification would actually underrepresent the true bias from sequential testing.

Our analysis has several limitations. First, there is a sampling bias. Our data were not randomly collected from all licensed laboratories but from a comparatively small number of laboratories. Because of aggregate data reporting, we were unable to assess co-infection rates, or to do patient-level analysis to control for the effect of repeat donors. Also, the reliability of the data could not be confirmed, and was frequently suspect. For example, some reports lacked information on age, sex and the purpose for testing, and had unreadable handwriting and obvious errors (e.g. the number of positive cases was greater than total tested), and as such, they were rejected from our analysis. Similarly, there was no mechanism in this analysis for quality control of the test results themselves. The effect of this would be to underestimate the prevalence of the BBVs to some degree. Perhaps most importantly, while our data set was extremely large, it nevertheless represents a ‘convenience sample’ in that it was collected for non-research purposes and was not collected systematically. Consequently, there could easily be undetectable biases in the data that reflect the non-random manner in which patients chose to become (or were rejected from becoming) blood donors or to seek overseas visas. We also identify several points at which the prevalence of the BBVs could be underestimated. The first is the bias introduced by sequential testing protocols, for which we provide empirical evidence. In addition, HBV prevalence in BDs may be underestimated due to the frequent practice of excluding those considered to be at high risk, such as those with tattoos or a remote history of IDU.\(^11\) Similarly, the practice believed to be used at some laboratories of pretesting for HBV with a rapid test before doing a formal screen would also underrepresent the total number of patients identified with BBVs, though such patients would not be included in the denominator of those tested in this report.

Nevertheless, we feel confident in concluding that the prevalence of HIV in the Philippines is extremely low, since for the aggregate effect of these biases to increase the prevalence of HIV from our observed level of 0.0038% (5 positives of 132,372 samples tested) to a range more comparable with that found in neighbouring south-east Asian countries (ca. 1%)\(^12\) would require the biases to underestimate the prevalence by 2–3 orders of magnitude, which is very unlikely. This conclusion is further strengthened by a significant proportion of the donors in the Philippines often being not anonymous volunteers but replacement donors, meaning donors recruited from among a patient’s family or acquaintances, who may paradoxically be at increased risk of HIV or other diseases.\(^26\)\(^,\)\(^29\)

In conclusion, we suggest that the prevalence of the three BBVs among BDs, and possibly OFWCs as well, probably offers a reasonable if imperfect source for monitoring prevalence. Our analyses raise several additional questions, such as why males overall had higher rates of HBV and HCV, and
why younger female OFWCs appeared to be at higher risk of HBV. Our surprising result was that OFWCs were not at increased risk of HIV or other viruses, a finding that contrasts with previous reports and could be explained by ascertainment bias in the national registry statistics. Our analysis suggests the possibility that the national statistics oversampled OFWCs, leading to an overestimation of the importance of this contribution of this group to overall numbers of HIV infections. However, our analysis cannot estimate the magnitude of the effect of any such sampling bias. We provide empirical evidence that the sequential testing procedures introduce under reporting bias, and suggest that this practice be abandoned in favour of simultaneous testing. This issue will likely grow in importance as the HIV epidemic increases in the future. We provide additional evidence that HIV remains extremely uncommon in the Philippines — a fascinating and somewhat surprising observation given the rising rates of HIV in neighbouring south-east Asian nations. While our data set cannot explain the reason for this, this evidence at least prompts inquiry into the very relevant issue of what cultural, biologic, geographic or political factors have so far left the Philippines comparatively unscathed by the HIV pandemic.

Competing interests: None declared.

Resumen

Prevalencia de las infecciones por el VIH, el VHB y el VHC entre los filipinos donantes de sangre y los solicitantes de un visado de trabajo para el extranjero

Objetivo Decidimos estimar la prevalencia del VIH, el VHB y el VHC en la población general de Filipinas empleando fuentes de datos distintas de la limitada red de vigilancia activa existente.

Métodos Analizamos los resultados globales de las pruebas del VIH, el VHB y el VHC a que se sometió a donantes de sangre (DS) en hospitales y a filipinos candidatos a trabajar en el extranjero (FCTE) sobre los que laboratorios autorizados informaron al Laboratorio Central Nacional de Cooperación en materia de ETS/SIDA en Manila entre 2002 y 2004.

Resultados Los resultados de las más de 144 000 pruebas de cribado de sangre realizadas revelaron que la prevalencia de infección por VIH era del 0,006% entre los DS y del 0,001% entre los FCTE; la prevalencia del VHB era del 4,2% en los dos grupos; y la del VHC era del 0,3% entre los DS y del 0,9% entre los FCTE. Los hombres presentaban un mayor riesgo de infección tanto por VHB como por VHC; entre los FCTE, el riesgo era mayor entre las mujeres más jóvenes. Los laboratorios que realizaban las pruebas de manera secuencial pero las interrumpían tras obtener el primer resultado positivo tenían mucha menos probabilidad de detectar el VHC, lo que lleva a pensar que posiblemente los protocolos de pruebas secuenciales subestiman la prevalencia del VIH y del VHB. Los FCTE presentaban un bajo riesgo de infección por el VIH, y la probabilidad de dar positivo para esos virus no aumentaba entre los FCTE que repetían su solicitud de un visado de trabajo, en comparación con quienes presentaban la solicitud por vez primera.

Conclusión Se desprende de estos datos que la infección por VIH es infrecuente en Filipinas. A diferencia de otros trabajos anteriores, éste no aporta indicio alguno de que los FCTE constituyan un grupo de alto riesgo de infección por VIH. Habrá que realizar nuevas investigaciones para averiguar por qué las mujeres jóvenes presentan un mayor riesgo de adquisición del VHB.
معيد انتشار فيروس الإيدز و فيروس التهاب الكبد بين الفلبينيين المترأكون بالدم وطالي تأشيرات العمل بالخارج

الهدف: إن إجراء هذه الدراسة هو تقييم معدل انتشار فيروس الإيدز و فيروس التهاب الكبد بين الفلبينيين المتبرعين بالدم و طالي تأشيرات العمل بالخارج.

الطريقة: فما تتيح تحليل الاختبارات للمجموعة المتنوعة من الفلبينيين، جزئيا غير متماثلة، ومن ثم أكبر عدد من فحوصات فيروس التهاب الكبد، C. نظرية إيجابية، لأنها تكون قد أقرت، من قبل علماء في هذه المجال.

الاستنتاج: أظهرت هذه الاختبارات تقريرات أثر إيجابية، بما في ذلك انتشار فيروس التهاب الكبد، مع ذكر نتائج بقيمة 0.3% بين الفلبينيين المتبرعين بالدم و 0.9% بين الفلبينيين طالبي تأشيرات العمل بالخارج. برنامج隔离 في هذا المجال، من خلال التعليم المتعلق بالخطر المتعلق.

المراجعات