313. Tuberculosis: epidemiological and public health features

P2858

Agreement between Quantiferon-TB Gold In Tube, T-SPOT.TB and tuberculin skin testing for diagnosing latent tuberculosis infection in a contact tracing study

Neus Altet1, Irene Latorre1, Malú De Souza2, M. Angeles Jimenez2, Celia Mila3, J. Solsona2, E. Lara2, Marta Dolores Ferrer1, Joan Caylà1, Jose Antonio Dominguez Benitez1 1Servei de Microbiologia, Institut d’Investigació en Ciències de l’Àlter, Badalona Barcelona, Spain; 2Unitat de Prevenció i Control de la Tuberculosi de Barcelona, Institut Català de la Salut, Barcelona, Spain; 3Agència de Salut Pública de Barcelona, Agència de Salut Pública de Barcelona, Spain

Aim: To determine the agreement between Quantiferon-TB Gold In Tube (QFN), T-SPOT.TB and TST in diagnosing LTBI in a contact study.

Methods: 753 individuals from contact tracing studies were included in the study. In all cases QFN and TST were performed, and in 141 patients the T-SPOT was also performed. TST was negative when the induration was less than 5 mm.

Results: The QFN and TST obtained concordant result in 478 cases from the 753 patients (the overall agreement was 63%), being both tests negative in 145 cases, and positive in 333 cases. From the 275 discordant results, in only one case the TST was negative and the QFN positive (corresponds with a high degree of exposure to the index case), and in 274 cases the TST was positive and the QFN negative, corresponding in 239 cases to BCG-vaccinated patients, and without significant difference between time of exposure to the index case.

With regards to the 141 patients tested with T-SPOT both in vitro tests were concordant in 120 cases (85.1%), being in 61 cases both tests negative, and in 59 cases positive. From the 27 discordant results, in 5 cases the QFN was positive and the T-SPOT negative, and in 16 cases the QFN was negative but the T-SPOT was positive, being in 15 of them the time of exposure significantly higher.

Conclusion: QFN and T-SPOT.TB have a high concordance in the diagnosis of LTBI. T-SPOT.TB shows a higher number of positive results than QFN. The main discordant results between TST and QFN should be attributed to the BCG-vaccination. Both tests seem useful for the diagnosis of LTBI in the contact studies.

The study was supported by a grant from FIS (08/1738).

P2859

Tuberculosis in health care staff in Romania, 2006-2010

Nicoleta Cioran, Horia Cocoș, Elmina Breaim. Central Coordination Unit of Romanian National Tuberculosis Program, “Marius Nasta” Institute of Pulmonology, Bucharest, Romania

Introduction: Health care staff represents a well recognized high risk group for TB.

Objective: To analyze characteristics of TB cases out of health care staff in Romania during 2006-2010 by demographic, clinical, bacteriological parameters and treatment outcomes.

Methods: Retrospective descriptive study of TB cases notified among staff of health care facilities in Romania from 2006 to 2010. Data and information used are from the electronic National TB Register.

Results: Total number of TB cases reported in health care workers in Romania from 2006 to 2010 was 843, declining from 224 in 2006 to 139 in 2009 and slightly increasing (to 150) in 2010. Most of them were aged from 25 to 39 years. Conversely to the general population, female gender was predominant (over 70%).

Abstract printing supported by Chiesi. Visit Chiesi at Stand B2.10
as the residence in urban area (over 70% as well). Even the staff in TB network varied between 2006 and 2008 from 12.9% to 7.2% of all medical staff, TB incidence rate in this group was 114.5/1000, versus 51.3/1000 in other medical staff in 2010. Pulmonary cases were from 74.5% in 2007 to 85.9% in 2008. New cases and relapses represented over 96% of all cases (96.8% in 2006 and 99.3% in 2010). In the five years have been reported 18 MDR-TB cases – from 6 in 2008 (6.9%) to 3 in 2010 (respectively 3.3%) and none in 2007. Overall success rate was 98.3% in 2007 and 89.3% in 2006.

Conclusions: TB incidence rate in health care staff in TB facilities is 2.2 folds higher than that in the personnel in other health care facilities. New pulmonary cases were predominant, with a few MDR-TB cases and a therapeutic success rate over 89%.

P2860 Pattern of tuberculosis (TB) among healthcare workers (HCW) attending a revised national TB control programme (RNTCP) unit in Kottayam Medical College (Kerala India)

Paul Davis, Anesh Cherukulam. Pulmonary Medicine, Medical College, Kottayam, Kerala, India

Background: There is paucity of data with regard to pattern of TB in HCW

Aim: To study the pattern of TB among HCW.

Study setting: RNTCP unit of a Medical College in Kerala, India from October 2009-March 2011.

Methods: Clinical profile of patients referred to RNTCP unit with a proven diagnosis or with a clinical and radiologic diagnosis made by a specialist medical teacher were gathered in a preset oral questionnaire and clinical examination.

Result: Out of 1222 TB patients 5.72% (n=70) were HCW-90% of HCW (n=63) were nurses or nursing students, 10% (n=7) were paramedical staff. No doctors or medical students registered in control programme for treatment. 68.6% (n=48) had extra pulmonary TB & 31.4% (n=22) pulmonary TB (OD 2.72: 63.6% (14/22) of pulmonary TB was smear positive. Occurrence of TB lymphadenitis (n=24) and TB meningitis (n=5) were more common among HCW compared to general category patients (p value of <0.05). 92.8% were newly diagnosed (n 65), 4.2% (n 3) retreatment & 2.8% (n 2) MDR. None had brain or menigcal TB. 92.8% were BCG vaccinated.

Conclusion: TB lymphadenitis was more common among HCW compared to general category patients. Majority of HCW coming for treatment in RNTCP were nurses or nursing students. No doctors registered for treatment in control programme.

References:

P2861 Tuberculosis (TB) notifications in healthcare workers (HCW) in Liverpool, UK

Nicholas Coombes1, Syed M.H. Kazmi2, Sean Mackin1, Clifford Bissace2, Peter Davies1, School of Medicine, University of Liverpool, Merseyside, United Kingdom; 2York Centre, Liverpool Community Health, Liverpool, Merseyside, United Kingdom

Background: The number of overseas HCW in Liverpool has grown sharply in recent years. In 2005, an audit demonstrated a rise in the number of TB notifications in the Liverpool TB service. We compared this data with results from the previous audits.

Methods:

1. TB notification data for the calendar year 2011 was collected from the Liverpool TB service. We compared this data with results from the previous audits.

2. A second QFT was performed in 118 HCWs. Reversion rate was 42% (n=50 of 118 tested IGRA-positive) in the first follow-up test upon completion of the training there was again one conversion, one reversion and one constantly positive test result over the three years (own TB anamnesis). No case of active tuberculosis was diagnosed over the three-year observation period.

Conclusion: Prevalence and infection rates are low among trainees. Negative IGRA test results proved very constant. Therefore IGRA testing in this low risk group is feasible. However, screening should focus on trainees with personal risk factors for TB. All others should only be tested after they have been in close contact with a TB index patient.

P2864 The effect of the introduction of IGRA in screening French healthcare workers for tuberculosis

Dominique Tripodi1, Benedicte Courtois 1, Albert Nienhaus2, 1Occupational Health and Safety, University Medical Centre Hamburg-Eppendorf, Hamburg, Germany; 2Occupational Safety and Health, Vivantes, Berlin, Germany; 3Pneumology, University Medical Center Hannover, Germany

Introduction: In France pre-employment screening for tuberculosis (TB) is performed for healthcare workers (HCW). Screening is repeated, when exposure to TB patients or infectious material occurs. The results of the TB screenings were analysed in a retrospective analysis.

Method: TB screenings were performed with Tuberculin Skin Test (TST) and Interferon-Gamma Release Assays (Quantiferon Gold in tube QFT). If TST was >10mm or TST increased by >10 mm, X-ray and pneumology consultation regarding preventive treatment of latent TB infection (LTBI) was performed. The screening results of 637 HCWs on whom QFT was performed were extracted from the files of the University Hospital of Nantes.

Results: In 3 (0.5%) HCWs the QFT was indeterminate. In 22.2% the QFT was positive. A second QFT was performed in 118 HCWs. Reversion rate was 42% (5 out of 12). Conversion rate was 6% (6 out of 98). TST was performed in 466 (73.5%) of the HCWs. TST >10mm results were 77.4%. In those with TST <10mm, QFT was positive in 14% and in those with TST >10mm, QFT was positive in 26.7%. When based on QFT results, X-ray and pneumology consultation should have been reduced to 28.6% of those selected by TST.

Conclusion: TST overestimated the prevalence of LTBI in this cohort. The de-
Introduction: Contact tracing is part of the tuberculosis (TB) elimination strategy. It is important to know which risk factors are associated with a positive screening.

Objectives: To identify risk factors associated with a positive screening.

Material and methods: During 2011, contacts of patients with pulmonary TB (sputum or broncho-alveolar lavage smear or culture positive), followed for screening in a TB reference centre, were questioned about their exposure to the index case. Positive screening was defined as active TB or latent infection. Contacts with incomplete characterization of exposure, unfinished screening or a past history of TB were excluded. A binary logistic model was used to analyze the variables.

Results: We observed 509 contacts of which 359 (153 men, median age: 32 years) were included in the analysis. 76 had a positive screening. Positive screening was associated with a positive spumum analysis of the index case (OR=2.62, 95%CI=1.33-5.14) and with coinfection (OR=3.42, 95%CI=1.66-7.07). Each additional year in age of the contact implied an increase in the odds for infection of 39%. OR(1-2-1-0) and each additional day of symptoms by the index case, previous to treatment, implied an increase in the odds for infection among his contacts of 1% (OR=1.01, 95%CI=1.00-1.02). No significant differences were found regarding size and ventilation of the exposure site.

Conclusions: This study shows that there is a significant increase in the risk of TB transmission to contacts for every day that the diagnosis of the index case is delayed. Increased risk was also shown for coinhabitants, contacts of older age and the presence of positive spumum smear or culture of the index case.

P2866 Approaching tuberculosis in a vulnerable group

Elmira Bregan 1, Nicloleta Cioran 1, Nicoleta Popescu 2, Florentina Liguia Purtunea 3, NTP, Marius Nasta Institute of Pulmonology, Bucharest, Romania; 2Medical Department, Samosocial, Bucharest, Romania; 3Public Health and Management, University of Medicine and Pharmacy Carol Davila, Bucharest, Romania

Introduction: The homeless represents a high risk group for TB, with poor access to health care services. TB prevalence among the around 5,000 homeless estimated to live in Bucharest is not known.

Aims and objectives: To detect active TB cases in homeless population in Bucharest.

Methods: A screening program was developed based on partnership between the National TB Programme and Samosocial Romania (NGO providing medical, psychological and social support to homeless people). Persons coming to the surgery service of Samosocial have been screened for TB in TB facilities. Any homeless giving a positive informed consent has been included and only exclusion criterion was having another chest X-ray in the previous 6 months.

Results: In the 248 cases registered from January to June 2011, 85.9% were males, mean age was 43.9 years (from 18 to 73 years old), 30.2% didn't have any ID document and half were at first medical consultation. TB has been suspected in 44.4% by clinical criteria, but from eligible persons only 48% were screened by chest X-ray, 14.1% refused and 35.5% didn't come back for screening. Active TB has been found in 8 cases (6.7% from the screened persons), 2 of them negative to clinical examination. None was previously examined even though they have free access to TB services. All cases were admitted in long-stay hospitals for treatment and counseling.

Conclusions: In this project the prevalence of active TB was found very high in the homeless population (6,700/1,000). Providing free access to TB services is not sufficient to detect TB cases in this high risk group and active screening programs are necessary.

P2867 Tuberculosis and migration: Predictors of epidemiological trends

Anna Goldberg 1, Viacheslav Zhamarlev 2, Natalya Skrynnik 1, 2Admiralty District, TB Dispensary, St. Petersburg, Russian Federation; 3The Molecular Genetics laboratory, The Research institute of Phthisiopulmonology, St. Petersburg, Russian Federation

Last years economic migration to the big cities of Russian Federation (RF) from other regions of RF and former Soviet republics has considerably increased. Since 2007 national free diagnosis of tuberculosis (TB) and free TB-treatment are not available for foreigners in St. Petersburg, RF.

Objective: To determine the impact of health care changes on the incidence of TB in migrants.

Methods: The study was conducted in the district of St. Petersburg. The population of this district was 178,000. From 01.01.2008 an office for migrants has opened at the TB dispensary. All persons including migrants who addressed the TB Dispensary during 2008-2011 were examined. The causes for examination were symptoms of tuberculosis or casual radiological findings. Examinations and treatment were free for all patients. Until 2008, there was no systematic recording of TB in migrants.

Results: During 2008-2010 the number of new TB cases in local population decreased, the number of infectious TB has decreased in local population and in migrants. The number of new cases of TB in children (local population and migrants) has decreased. In 2011, the trend reversed.

Number of TB cases

<table>
<thead>
<tr>
<th>Year</th>
<th>Total</th>
<th>Migrants</th>
<th>Total in Migrants</th>
</tr>
</thead>
<tbody>
<tr>
<td>2008</td>
<td>55</td>
<td>51</td>
<td>4</td>
</tr>
<tr>
<td>2009</td>
<td>37</td>
<td>31</td>
<td>6</td>
</tr>
<tr>
<td>2010</td>
<td>13</td>
<td>12</td>
<td>1</td>
</tr>
<tr>
<td>2011</td>
<td>30</td>
<td>22</td>
<td>8</td>
</tr>
</tbody>
</table>

Conclusions: To ensure effective control of tuberculosis free access to health care is required for all TB patients, including migrants.

P2868 New-entrant screening for tuberculosis at port of entry in the U.K.: Is it time to change policy?

Vikas Panamitry, Henke Kunst, Martin Dedidacut, Joy Troko, Department of Respiratory Medicine & Physiology, Heart of England NHS Foundation Trust, Birmingham, West Midlands, United Kingdom

Background: It is shown that 20% of tuberculosis (TB) cases are diagnosed in persons entering UK within 2 years & in 45% within 5 years of arrival. Therefore the first 2-5 years of arrival in UK presents a period of high risk of reactivation of new-entrants with LTBI. This clearly indicates the importance of early screening & reducing the risk of active TB by giving chemoprophylaxis to patients with LTBI.

UK policy at the moment advocates identification of active TB by chest x-ray for all new arrivals intending to stay for >6 months from countries with a TB incidence >40/100,000; however, screening is very arbitrary & not standardised. The results are forwarded to local NHS TB services where new entrants intend to settle for complete screening.

Methods: A retrospective review was conducted of all new-entrants who were referred to our institution during a 2 year period, from January 2010, with suppos- edly an abnormal chest x-ray or port of entry. All new-entrants were screened for active & latent TB by TST or IGRA

Results: 103 patients, 51 females & 52 males were referred, however only 42 patients (40.7%) (26 females & 14 males) attended the TB clinic,of which 31(73.8%) were < 35 years old. Of the 42, 13 patients (31%) were diagnosed to have LTBI while none were found to have active TB. Six patients with LTBI were from the sub-Saharan Africa, & from the Indian subcontinent. Of 13 patients with LTBI, 10 patients completed 3 month course of chemoprophylaxis.

Conclusion: Screening new-entrants at port of entry is inadequate. New-entrants often do not attend clinic appointments since they have moved elsewhere or do not understand the necessity of being screened for latent or active TB.

P2869 Impaired pulmonary function and the risk of tuberculos is – A population based cohort study

Malin Hjemberg 1, Claes-Geörn Loftdahl 2, Niclas Winquist 1

1Section for Infection Medicine, Department of Clinical Sciences, Lund University, Lund, Sweden; 2Respiratory Medicine & Allergology, Department of Clinical Sciences Lund, Lund University, Lund, Sweden; 3Cardiovascular Epidemiology Research Group, Department of Clinical Sciences Malmö, Lund University, Lund, Sweden

Background: Even though COPD is a frequent co-morbid condition in elderly with active tuberculosis relatively little is known to what extent impaired lung function may influence the progression of active pulmonary tuberculosis in population based studies.

Methods: A case control study was conducted with 180 patients with active pulmonary tuberculosis and 180 matched controls with no evidence of chronic lower respiratory disease.

Results: The patients with active tuberculosis had an impaired pulmonary function compared to the controls (mean FEV1 0.60 vs 1.20 L, p < 0.01). The patients with a FEV1 < 0.80 L had an increased risk of hospitalization for active tuberculosis (OR 1.9, 95% CI 1.1-3.3).

Conclusion: Impaired pulmonary function is a risk factor for tuberculosis in elderly and may influence the progression of active pulmonary tuberculosis.
function increases the incidence of active tuberculosis in excess of the direct effect of smoking.

Methods: 28,907 participants of the Malmo Preventive Project performed a spirometry at baseline examination and were followed for a mean of 25 years. Pulmonary function was measured as FVC and FEV1, % of predicted (standardized for age, height and gender) and classified according to the GOLD criteria. Cases of incident tuberculosis, notified in the local tuberculosis register 1989-2008 were identified. Hazard ratios (HR) for subsequent tuberculosis according to FVC and FEV1 were estimated using Cox regression.

Main results: A total of 26 cases of incident tuberculosis were identified corresponding to an overall incidence of 5.2 (95% confidence interval [95% CI] 3.5-7.6) per 100,000 person-years. The incidence rate was inversely correlated with FEV1 (% of predicted). HR per 10% unit increase 0.71 (95% CI 0.59-0.86). The results persisted after adjustment for smoking and age at screening. HR per 10% unit increase in FEV1 (% of predicted) 0.75 (95% CI 0.61-0.91). The incidence of TB increased with GOLD-stage, stage I, crude HR 1.9 (95% CI 0.5-6.7), stage II, HR 5.64 (95% CI 2.2-14.7), stage III+IV, HR 6.9 (95% CI 0.9-52.6), p<0.001 for linear trend, although only one case of TB occurred in GOLD-stage III+IV.

Conclusions: Impaired pulmonary function or COPD is associated with an increased incidence of active tuberculosis.

P2870 Incidence of tuberculosis: The application of capture-recapture method to compare two sources of information

Francisco Muñiz1, Maríta López2, Noelia Curarréco2, Sara Raposo2, Piedad Rivera3, Silvia García1, José Guerra2 *Respiratory Department, University Hospital, León, Spain; "Internal Medicine, University Hospital, León, Spain

Introduction: The true incidence of tuberculosis is higher than that in national and international records. Underreporting is estimated to vary between 7% and 27% according to studies.

Objective: Estimate the rate of tuberculosis in an area of Leon for the years 2008 and 2009 using the capture-recapture method to compare two sources of information: tuberculostatic drugs prescribed (rilampcin-isomiazid association) and the register of the regional epidemiological surveillance system (SIVE).

Method: Retrospective descriptive study in an area of 351,086 inhabitants of the cases of tuberculosis using as source (i), information on tuberculostatic drugs prescribed, and (ii), the SIVE register. We calculated incidence rates for each source by the capture-recapture method. We analyzed epidemiological and demographic data, symptoms, diagnosis, treatment and monitoring.

Results: The incidence obtained in 2008 using the SIVE data was 18.80/100000 and using the pharmacy register was 26.77. In 2009, the SIVE data gave an estimate of 18.23 and the pharmacy register 22.50. When applied the capture-recapture method, the annual incidence for 2008 was 44.14 (95%CI 37.85 - 50.41) and for 2009 of 34.17 (95% CI 30.19-38.17). In each of the years studied the number of cases obtained from the pharmacy register was higher.

Conclusions: The SIVE data on the incidence of tuberculosis in the study area understates the true incidence rate. The source of information that involves recording cases of tuberculosis in the community is underestimated. The capture-recapture method is a good alternative to measure the incidence of tuberculosis and to check the surveillance systems.

P2872 Efficiency of molecular methods for epidemiological investigation in tuberculosis (TB)

Katarzyna Kruczał1, Ewa Augustynowicz-Kopeć2, Monika Kozinska2, Grazyna Fassak-Standa1, Mariusz Duplaga1, Zofia Zwołka1, Ewa Nizankowska-Mogilnicka1 *Department of Pulmonology, Jagiellonian University Medical College, Krakow, Poland; 2Department of Microbiology, The Institute of Tuberculosis and Lung Diseases, Warsaw, Poland; 3Referential Tuberculosis Laboratory, John Paul Hospital, Cracow, Poland; 4Institute of Public Health, Jagiellonian University Medical College, Krakow, Poland

Introduction: High prevalence of tuberculosis in Poland may relate to active transmission.

Objectives: The assessment of the transmission of TB in the city of Krakow with genetic methods and with standardized epidemiological interview and the comparison of these methods.

Methods: In years 2007-2011 genomic DNA samples isolated from Mycobacterium tuberculosis complex strains coming from 274 patients were analysed by spoligotyping and IS6110-Mtb1-Mtb2 PCR method. The strains were assessed as identical if their DNA patterns were the same in both methods. In IS6110-Mtb1-Mtb2 PCR method DNA profile obtained in both PCR reactions should be identical.

Results: Among 274 strains, 122 genetic patterns (spoligotypes) were identified. Unique spoligotypes occurred in 91 strains; remaining 183 strains belonged to 31 clusters stemming from the same spoligotypes. The application of the IS6110-Mtb1-Mtb2 PCR in next stage of the analysis, allowed reducing the number of clusters to 18 with 91 strains. The clusters included 2-14 patients and covered one third of analysed samples. The clusters were dominated by men below 50 years, AFB+, with strains sensitive to first-choice antituberculous agents. The standard epidemiological interviews did not reveal neither direct close nor periodic contacts between the patients included in clusters (only three clusters comprised prisoners from the same penitentiary and three homeless persons with periodic contact).

Conclusion: The assessment of transmission of TB should be based on scrupulous epidemiological interview and on molecular genetic methods. Active transmission in Krakow in years 2007-2011 was responsible for 30% of diagnosed TB cases.

P2874 Administration of BCG vaccination: Survey of practice in the Mersey region, UK

Sean Mackin1, Syed M.H. Kazmi2, Nick Coombes1, Mohammed Hussain2, Clifford Bissacet3, Peter Davies2 *School of Medicine, University of Liverpool, United Kingdom; 3Respiratory Medicine, Liverpool Heart and Chest Hospital, Liverpool, United Kingdom; 4TB Centre, Liverpool Community Health, Liverpool, United Kingdom

Background: A UK wide survey in 2005 showed wide variation in BCG practice and that the providers were uncertain in various aspects of the administration. The National Institute of Clinical Excellence (NICE) published guidelines in 2006, which along with Department of Health guidance aimed to standardise practice. We intended to assess regional BCG service 6 years later.

Method: 4 TB centres providing BCG vaccination across Mersey region were contacted and data regarding current practice of BCG administration was collected.

Results: The adherence to the guidelines was recorded as compliance. See table.

Questions
Administration of BCG at sites other than upper arm
Re-vaccination in the absence of BCG scar in children
Prior assessment of HIV status
Prior evaluation of anaphylaxis risk
Availability of rehydration equipment
Formal training of staff in paediatric resuscitation

Guidelines Compliance (%)
50%*
100%
100%
100%
75%
100%

* The other sites used were right upper arm or upper thigh. Reasons included patient preference or confusing scar.

There were no recorded episodes of anaphylaxis in the past 12 months. All centres referred patients with severe adverse local reactions appropriately.

Conclusions: Our survey demonstrates that NICE recommendations and their implementation were essential in elucidating areas of uncertainty in the administration of BCG vaccination and subsequently the practice is now uniform across the Mersey region. We intend to extend this survey to national level.

P2875 Enhancing patient safety: New WHO guidance on pharmacovigilance in tuberculosis care

Shanthi N. Pal1, Christian Lienhardt2, Sten Olsson3, Dennis Falzon3 *STOP TB Department, World Health Organization, Geneva, Switzerland; 2Department of Essential Medicines and Pharmaceutical Policies, World Health Organization, Geneva, Switzerland; 3The Uppsala Monitoring Centre, (UMC), Uppsala, Sweden

Adverse drug reactions (ADRs) can lead to a patient interrupting tuberculosis (TB) treatment before completion, and contribute to avoidable morbidity, treatment failure, loss in quality of life, or death. While many national TB control programmes have a long tradition of monitoring patient care, the surveillance of drug-related problems, or pharmacovigilance, has not been systematic. The increasing worldwide use of more extensive regimens for drug-resistant TB, the concomitant use of antiretroviral therapy in patients with HIV-associated TB, and the imminent release on the market of new classes of medicines to treat TB make the case for pharmacovigilance even stronger.

WHO produced guidance this year on pharmacovigilance for TB through the financial support of the European Commission Seventh Framework Programme. The manual discusses how pharmacovigilance can be effectively implemented in a programme through key stakeholders, and provides a step-by-step approach on how to identify signals, assess relationships between an event and a drug, determine causality, and communicate findings. It presents three methodologies of pharmacovigilance which can be applied for the detection, assessment, understanding and prevention of adverse events or any other drug-related problem under field situations. The first two - spontaneous and targeted spontaneous reporting - can be built into national programmes of routine pharmacovigilance and/or tuberculosis control. The third type, cohort event monitoring (CEM), is an active form of surveillance, similar in design and management to an epidemiological cohort study. CEM would be particularly well suited to the post-marketing surveillance of new drugs.