

Centre for Respiratory Infection









Epidemiology of TB and Diagnostic Delays





Onn Min Kon TB Clinics

R PA

St Mary's and Hammersmith Hospitals

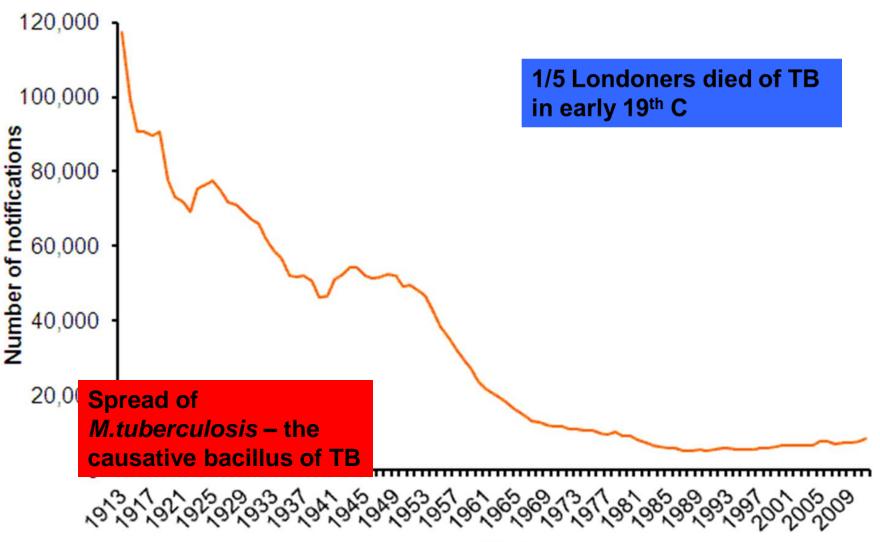
What happens after 'infection'

- May heal spontaneously
- May develop active disease
 - -over weeks and months
 - 'primary' TB
- Contained and dormant

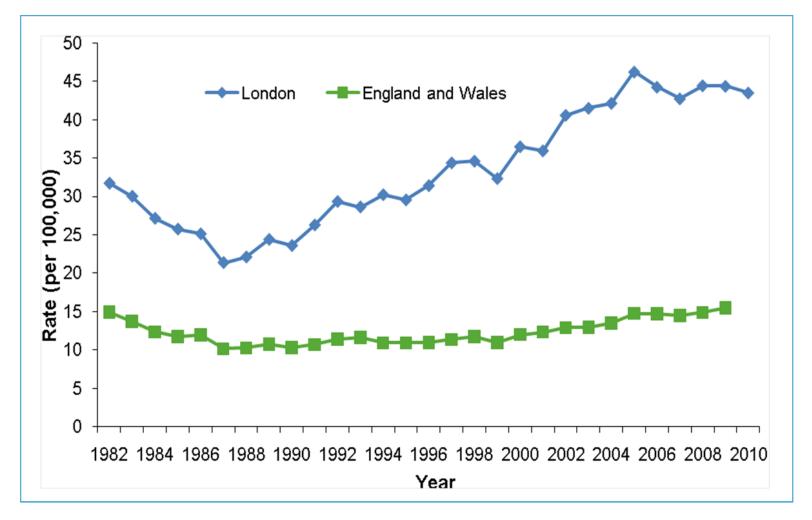
-may reactivate later if debilitated or immunocompromised

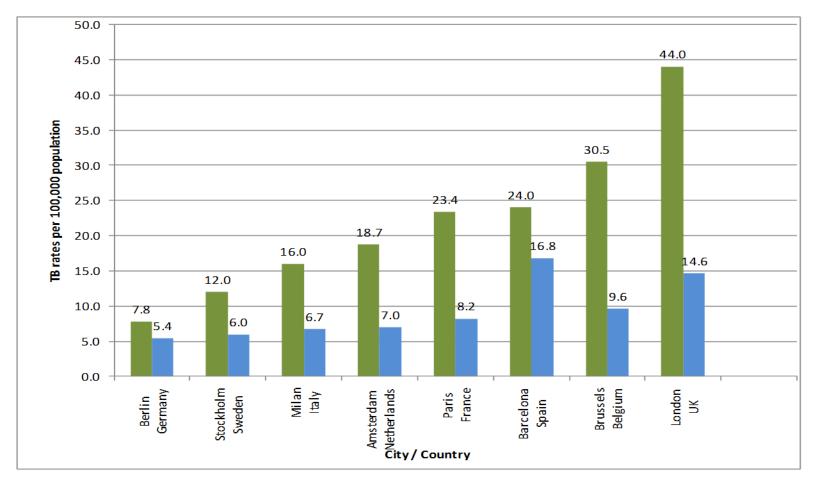
- 'post-primary' TB

lifetime cumulative risk after 'infection' - 10-15% (1.5% first year/ 5-10% five years)



TB rates in London, 1982-2010

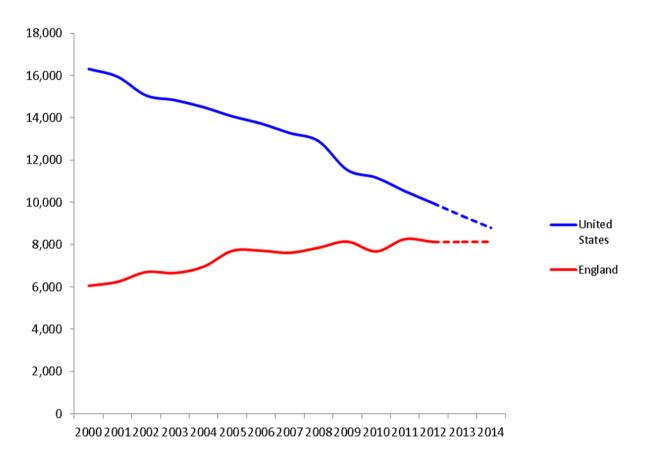




TB incidence in Western Europe, 2009

Collaborative TB Strategy - Consultation





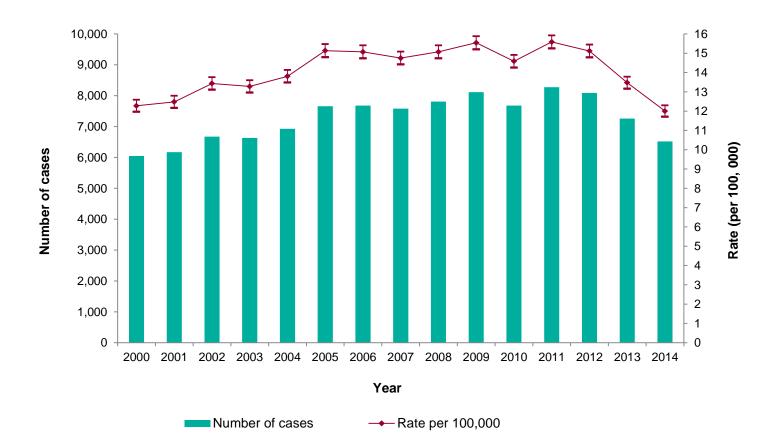
Trends in the annual number of cases of tuberculosis in England compared with the USA. Dotted lines show projected numbers, assuming present annual percentage change continues for 2 more years. (Based on data from Public Health England (UK) and the Centre for Disease Control and Prevention (USA). (Zenner et al 2013)

Collaborative TB Strategy - Consultation



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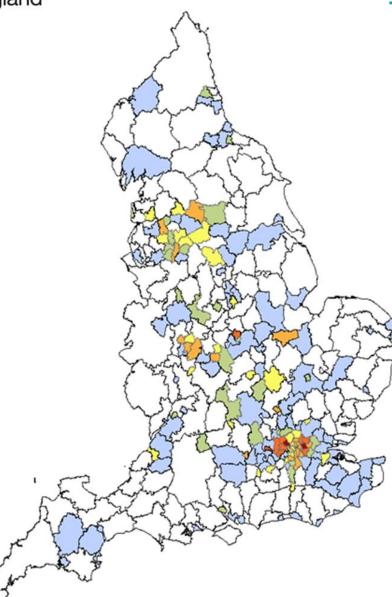
TB case notifications and rates, England, 2000-2014



Source: Enhanced Tuberculosis Surveillance (ETS), Office for National Statistics (ONS) Data extracted: March 2015. Prepared by: TB Section, National Infection Service, Public Health England



Tuberculosis in England: 2015 report



Three-year average TB rates by local authority district, England, 2012-2014



Tuberculosis rate (per 100,000)

| 0.0-4.9 |
|-----------|
| 5.0-9.9 |
| 10.0-14.9 |
| 15.0-24.9 |
| 25.0-39.9 |
| 40.0-69.9 |
| >70.0 |

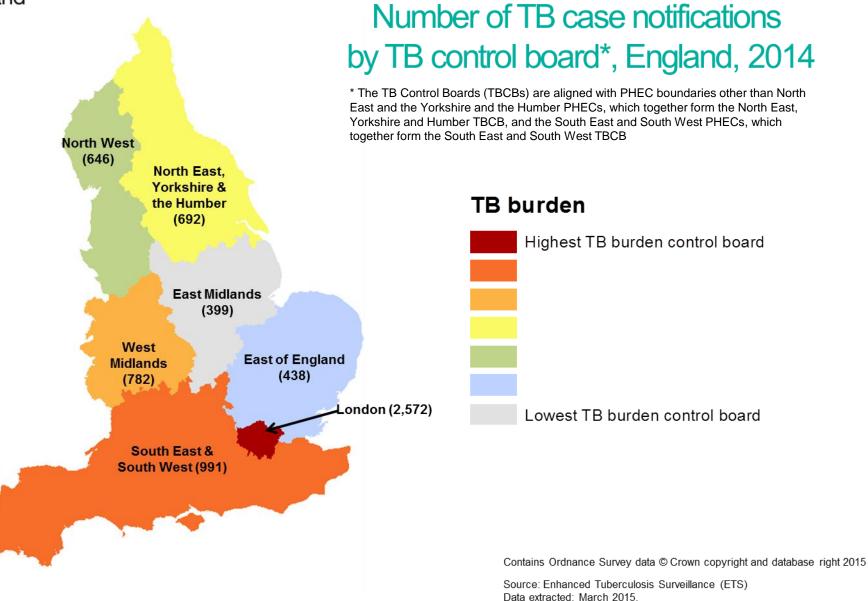
Contains Ordnance Survey data © Crown copyright and database right 2015. Contains National Statistics data © Crown copyright and database right 2015.

Source: Enhanced Tuberculosis Surveillance (ETS), Office for National Statistics (ONS) Data extracted: March 2015.

Prepared by: TB Section, National Infection Service, Public Health England

WW Public Health England

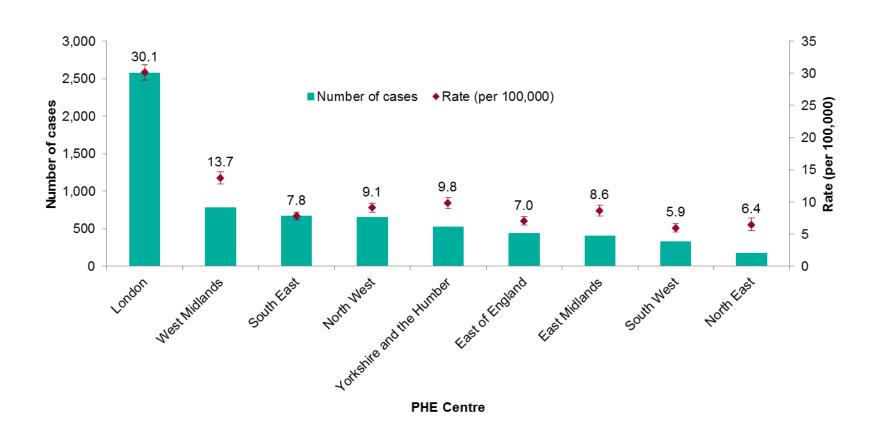
Prepared by: TB Section, National Infection Service, Public Health England





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TB case notifications and rates by PHE Centre, England, 2014



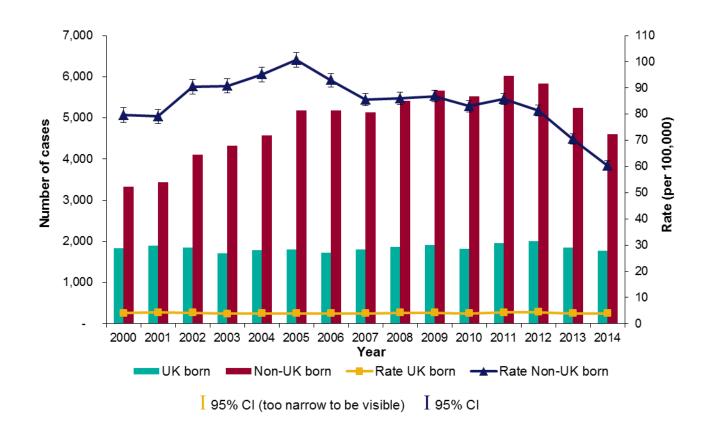
Source: Enhanced Tuberculosis Surveillance (ETS), Office for National Statistics (ONS) Data extracted: March 2015.

Prepared by: TB Section, National Infection Service, Public Health England

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TB case notifications and rates by place of birth, England, 2000-2014



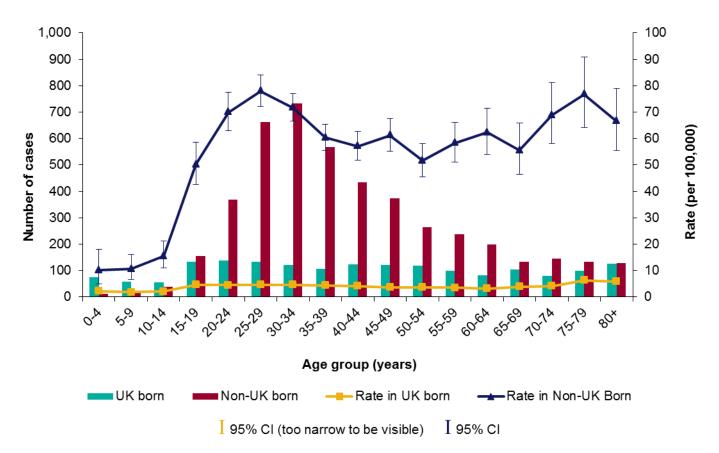
Source: Enhanced Tuberculosis Surveillance (ETS), Labour Force Survey (LFS) Data extracted: March 2015.

Tuberculosis in England: 2015 report

Prepared by: TB Section, National Infection Service, Public Health England

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TB case notifications and rates by age group and place of birth, England, 2014



Source: Enhanced Tuberculosis Surveillance (ETS), Labour Force Survey (LFS) Data extracted: March 2015. Prepared by: TB Section, National Infection Service, Public Health England



Country of birth and time from UK entry to TB notification, England, 2014

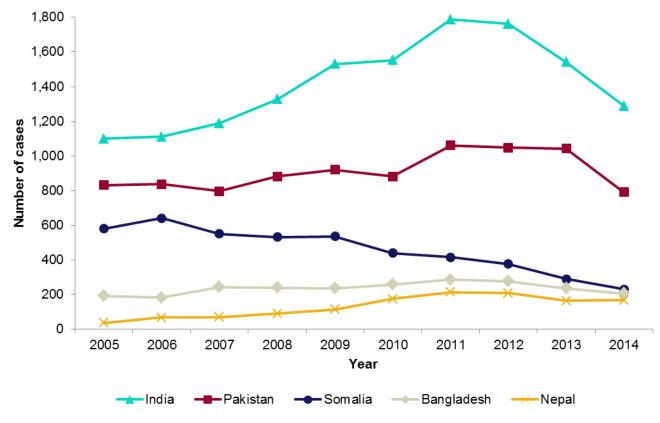
| Country of birth | Number of cases | Percentage of cases (%)* | Median time in years since entry to UK (IQR) | |
|-------------------|--------------------|--------------------------------|---|--|
| UK | 1,774 | 28.2 | - | |
| India | 1,288 | 20.5 | 7 (3 - 14) | |
| Pakistan | 791 | 12.6 | 10 (3 - 25) | |
| Somalia | 230 | 3.7 | 10 (3 - 14) | |
| Bangladesh | 207 | 3.3 | 8 (4 - 19) | |
| Nepal | 168 | 2.7 | 4 (3 - 9) | |
| Nigeria | 118 | 1.9 | 7 (2 - 17) | |
| Philippines | 111 | 1.8 | 9 (4 - 13) | |
| Zimbabwe | 107 | 1.7 | 11 (9 - 12) | |
| Afghanistan | 96 | 1.5 | 8 (4 - 13) | |
| Romania | 88 | 1.4 | 1 (0 - 6) | |
| Eritrea | 83 | 1.3 | 3 (0 - 8) | |
| Kenya | 81 | 1.3 | 19 (8 - 41) | |
| Sri Lanka | 78 | 1.2 | 11 (4 - 15) | |
| Poland | 70 | 1.1 | 6 (2 - 8) | |
| Others (each <1%) | 1,007 | 15.8 | 8 (3 -16) | |
| Total* | 6,297 | 100.0 | 9 (3 - 20) | |

* Where country of birth was known



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Trend in TB case notifications for top five countries of birth* of non-UK born cases, England, 2005-2014



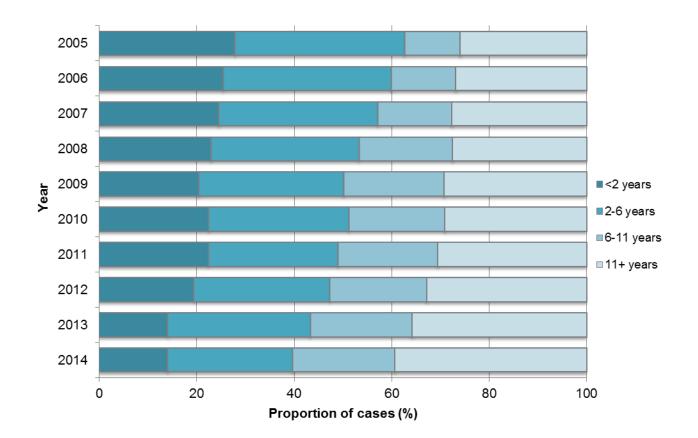
^{*} Five most frequent countries of birth in 2014

Tuberculosis in England: 2015 report

Source: Enhanced Tuberculosis Surveillance (ETS) Data extracted: March 2015. Prepared by: TB Section, National Infection Service, Public Health England



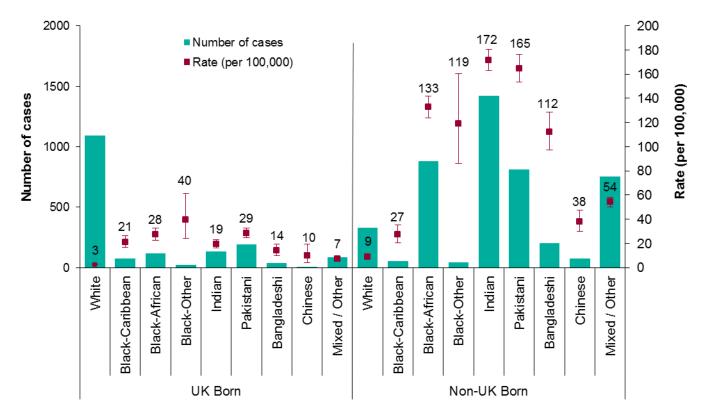
Time between entry to the UK & TB notification for non-UK born cases by year, England, 2005-2014





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TB case notifications and rates by place of birth and ethnic group, England, 2014

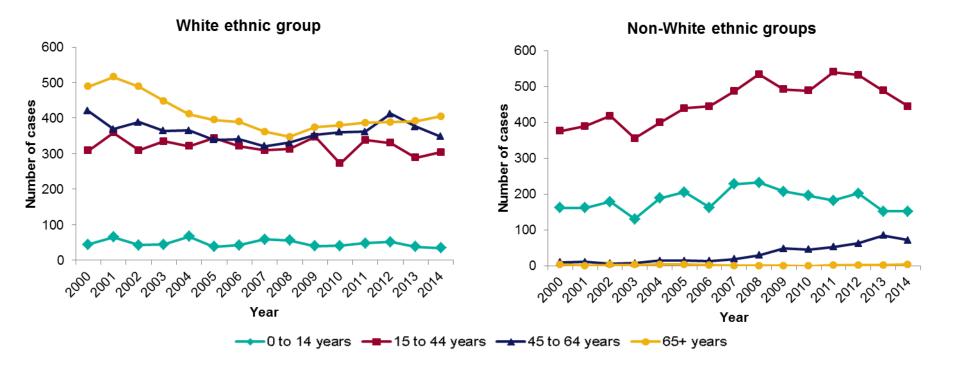


Place of birth/Ethnic group

Source: Enhanced Tuberculosis Surveillance (ETS) Data extracted: March 2015. Prepared by: TB Section, National Infection Service, Public Health England



Trend of TB case notifications by age group and ethnic group* in UK born cases, England, 2000-2014



* Cases with Black-Caribbean, Black-African, Black-Other, Indian, Pakistani, Bangladeshi, Chinese and Mixed/Other ethnic groups were grouped as 'non-White'

Source: Enhanced Tuberculosis Surveillance (ETS) Data extracted: March 2015. Prepared by: TB Section, National Infection Service, Public Health England



TB case notifications by site of disease, England, 2014

| Site of disease* | Number of cases | Percentage** | |
|----------------------------|-----------------|--------------|--|
| Pulmonary | 3,434 | 52.9 | |
| Extra-thoracic lymph nodes | 1,445 | 22.3 | |
| Intra-thoracic lymph nodes | 863 | 13.3 | |
| Unknown extra-pulmonary | 638 | 9.8 | |
| Pleural | 566 | 8.7 | |
| Other extra-pulmonary | 504 | 7.8 | |
| Gastrointestinal | 368 | 5.7 | |
| Bone – spine | 310 | 4.8 | |
| Miliary | 179 | 2.8 | |
| Bone – not spine | 168 | 2.6 | |
| CNS – meningitis | 148 | 2.3 | |
| Genitourinary | 129 | 2.0 | |
| CNS – other | 99 | 1.5 | |
| Cryptic disseminated | 39 | 0.6 | |
| Laryngeal | 16 | 0.2 | |

* With or without disease at another site

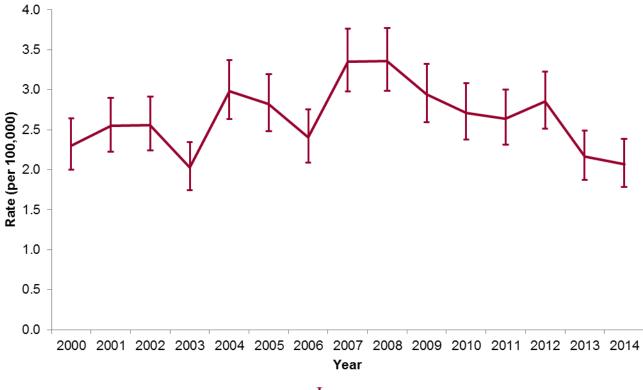
** Total percentage exceeds 100% due to disease at more than one site

CNS - Central Nervous System



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Rate of TB in UK born children (0-14 years old), England, 2000-2014



I 95% CI



Number and proportion of cases with unique and clustered MIRU-VNTR strain types, England, 2010-2014

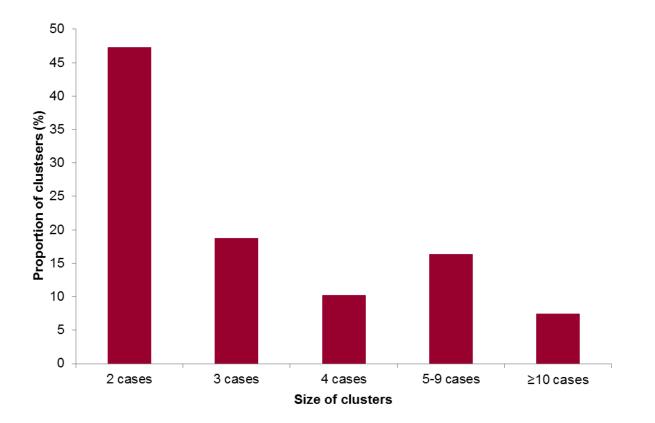
| Year | Notified cases | Culture c cas | | Strain type (≥23 le | | Unique | cases | Cluster | ed cases * | New clusters (per year) ** |
|-------|-------------------|------------------|------|------------------------|------|--------|-------|---------|------------|-------------------------------|
| | n | n | % | n | % | n | % | n | % | n |
| 2010 | 7,677 | 4,609 | 60.0 | 3,229 | 70.1 | 1,391 | 43.1 | 1,838 | 56.9 | 366 |
| 2011 | 8,276 | 5,029 | 60.8 | 4,267 | 84.8 | 1,846 | 43.3 | 2,421 | 56.7 | 536 |
| 2012 | 8,086 | 4,895 | 60.5 | 4,301 | 87.9 | 1,754 | 40.8 | 2,547 | 59.2 | 534 |
| 2013 | 7,257 | 4,390 | 60.5 | 3,656 | 83.3 | 1,526 | 41.7 | 2,130 | 58.3 | 401 |
| 2014 | 6,520 | 3,914 | 60.0 | 3,319 | 84.8 | 1,512 | 45.6 | 1,807 | 54.4 | 408 |
| Total | 37,816 | 22,837 | 60.4 | 18,772 | 82.2 | 8,029 | 42.8 | 10,743 | 57.2 | 2,245 |

* Clustered in time period (2010-2014), clustered cases notified in year

** A new cluster forms at the point when a second case is notified with the same MIRU-VNTR strain type as an existing case



Proportion of MIRU-VNTR clusters by size, England, 2010-2014





Number and proportion of pulmonary TB cases by delay from symptom onset to treatment start, England, 2011-2014

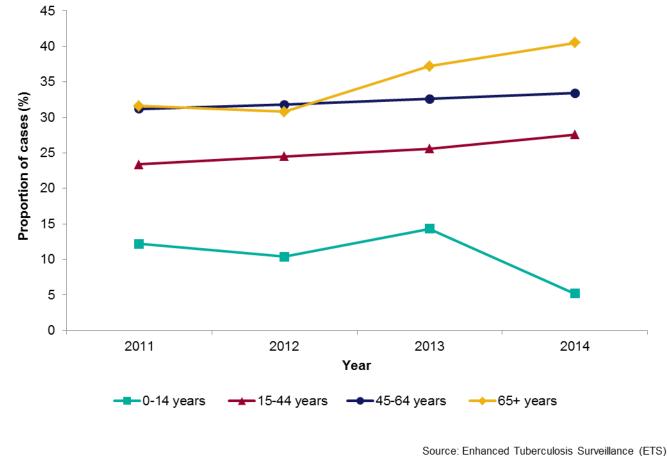
| | 0-2 months | | 2-4 months | | >4 m | Total* | |
|------|------------|------|------------|------|------|--------|-------|
| Year | n | % | n | % | n | % | n |
| 2011 | 1,317 | 45.0 | 856 | 29.3 | 753 | 25.7 | 2,926 |
| 2012 | 1,369 | 44.0 | 925 | 29.8 | 814 | 26.2 | 3,108 |
| 2013 | 1,222 | 41.2 | 900 | 30.4 | 841 | 28.4 | 2,963 |
| 2014 | 1,137 | 39.5 | 868 | 30.2 | 870 | 30.3 | 2,875 |

* The number of pulmonary cases with time between onset of symptoms to start of TB treatment available, excluding those diagnosed post-mortem and those that were known to not start treatment

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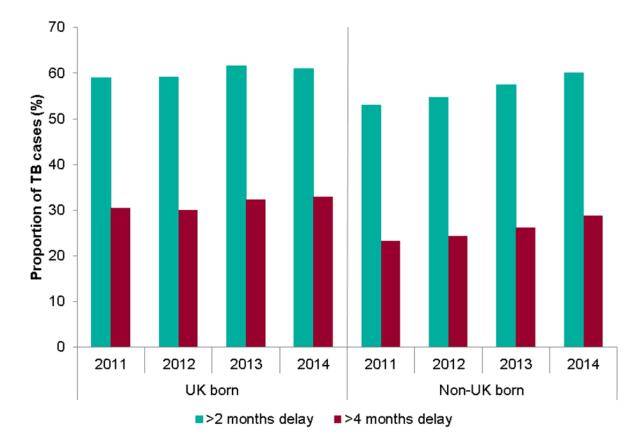
Proportion of pulmonary cases with a delay from symptom onset to treatment start of >4 months by age group, England, 2011-2014



Tuberculosis in England: 2015 report

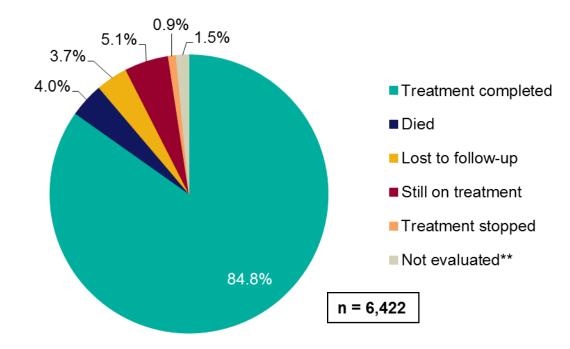
Source: Enhanced Tuberculosis Surveillance (ETS) Data extracted: March 2015. Prepared by: TB Section, National Infection Service, Public Health England WW Public Health England

Proportion of cases with a delay from symptom onset to treatment start by place of birth, England, 2011-2014



Public Health England

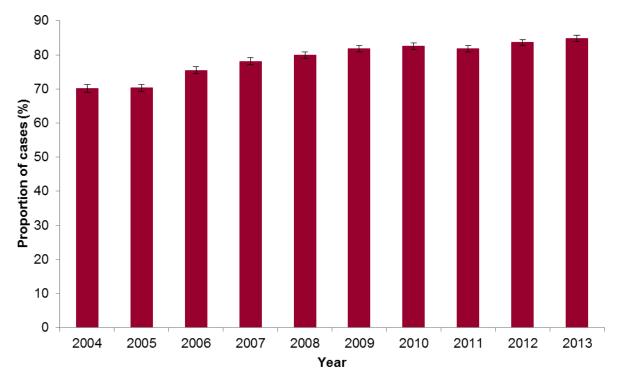
TB outcome at 12 months for drug sensitive cases with expected treatment duration <12 months*, England, 2013



* Excludes initial and amplified to rifampicin resistant TB and MDR-TB cases and MDR-TB treated cases and those with CNS, spinal, miliary or cryptic disseminated TB ** Not evaluated includes missing, unknown and transferred out

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Treatment completion at 12 months for drug sensitive TB cases with expected treatment duration <12 months*, England, 2004-2013



I 95% CI

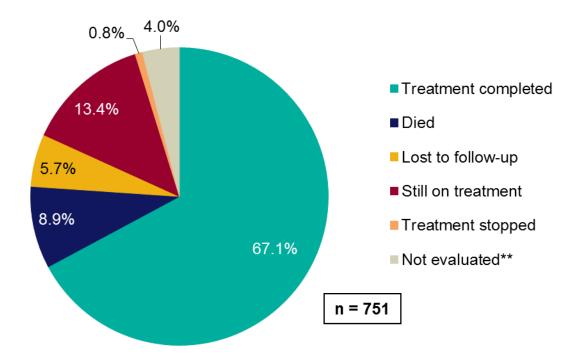
* Excludes initial and amplified to rifampicin resistant TB and MDR-TB cases and MDR-TB treated cases and those with CNS, spinal, miliary or cryptic disseminated TB

Tuberculosis in England: 2015 report

Source: Enhanced Tuberculosis Surveillance (ETS) Data extracted: March 2015. Prepared by: TB Section, National Infection Service, Public Health England



Last recorded TB outcome for drug sensitive cohort with CNS, spinal, miliary or cryptic disseminated* TB, England, 2013



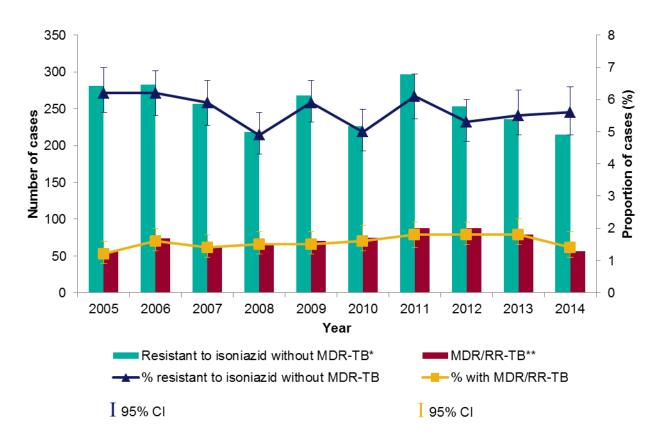
* Excludes initial and amplified to rifampicin resistant TB and MDR-TB cases and MDR-TB treatment cases and only includes drug sensitive TB cases with CNS, spinal, military or cryptic disseminated TB

** Not evaluated includes missing, unknown and transferred out

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Number and proportion of TB cases with initial drug resistance, England, 2005-2014



* Culture confirmed cases with DST results for at least isoniazid and rifampicin resistant to isoniazid without MDR-TB

** Culture confirmed cases with DST results for at least isoniazid and rifampicin resistant to rifampicin, including those with MDR-TB

Tuberculosis in England: 2015 report

Source: Enhanced Tuberculosis Surveillance (ETS) Data extracted: March 2015. Prepared by: TB Section, National Infection Service, Public Health England



Number and proportion of TB cases with drug resistance by age group, England, 2014

| Age group (years) | Resistant to isoniazid without MDR-TB cases* | | MDR/RR- | Total cases [#] | |
|----------------------|--|-----|------------------|-----------------------------|-------|
| | n | % | n | % | n |
| 0-4 | 0 | 0.0 | 0 | 0.0 | 12 |
| 5-14 | 2 | 4.3 | 0 | 0.0 | 46 |
| 15-24 | 31 | 5.6 | 9 | 1.6 | 558 |
| 25-34 | 62 | 5.7 | 24\$ | 2.2 | 1,088 |
| 35-44 | 51 | 6.8 | 11 ^{\$} | 1.5 | 746 |
| 45-54 | 33 | 6.3 | 7 | 1.3 | 525 |
| 55-64 | 20 | 5.9 | 4 | 1.2 | 339 |
| 65+ | 16 | 2.8 | 1 | 0.2 | 575 |
| Total | 215 | 5.5 | 56 | 1.4 | 3,889 |

* Culture confirmed cases with DST results for at least isoniazid and rifampicin who are resistant to isoniazid without MDR-TB

** Culture confirmed cases with DST results for at least isoniazid and rifampicin who are resistant to rifampicin, including those with MDR-TB

All culture confirmed cases with DST results for at least isoniazid and rifampicin

^{\$} Three cases aged 25-34 and one case aged 35-44 were resistant to rifampicin without MDR-TB

Most frequent countries of birth of TB cases with drug resistance, England, 2014

| Country of birth* | Total cases** | isoniazid without | | | MDR/RR-TB cases ^{\$} | | |
|-------------------|------------------|-------------------|------|-----|----------------------------------|--|--|
| | n | n | % | n | % | | |
| UK | 967 | 52 | 5.4 | 6 | 0.6 | | |
| India | 780 | 40 | 5.1 | 10^ | 1.3 | | |
| Pakistan | 469 | 27 | 5.8 | 3 | 0.6 | | |
| Somalia | 129 | 10 | 7.8 | 0 | 0.0 | | |
| Philippines | 75 | 7 | 9.3 | 2 | 2.7 | | |
| Afghanistan | 59 | 7 | 11.9 | 1 | 1.7 | | |
| Bangladesh | 107 | 7 | 6.5 | 0 | 0.0 | | |
| Nepal | 113 | 6 | 5.3 | 2 | 1.8 | | |
| Lithuania | 46 | 5 | 10.9 | 11 | 23.9 | | |
| Eritrea | 53 | 5 | 9.4 | 0 | 0.0 | | |
| Nigeria | 81 | 4 | 4.9 | 3^ | 3.7 | | |
| Zimbabwe | 69 | 2 | 2.9 | 3 | 4.3 | | |
| Ethiopia | 27 | 1 | 3.7 | 3 | 11.1 | | |
| Latvia | 15 | 1 | 6.7 | 3 | 20.0 | | |

* Top 14 countries of birth for cases resistant to isoniazid without MDR and MDR/RR-TB cases in 2014.

** Culture confirmed cases with DST results for at least isoniazid and rifampicin

Culture confirmed cases with DST results for at least isoniazid and rifampicin who are resistant to isoniazid without MDR-TB

\$ Culture confirmed cases with DST results for at least isoniazid and rifampicin who are resistant to rifampicin, including those with MDR-TB

^ Two cases from Nigeria and one case from India were resistant to rifampicin without MDR-TB



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Last recorded TB outcome for entire

drug resistant cohort, England, 2012*

TB outcome at 24 months for drug resistant cohort, England, 2012*

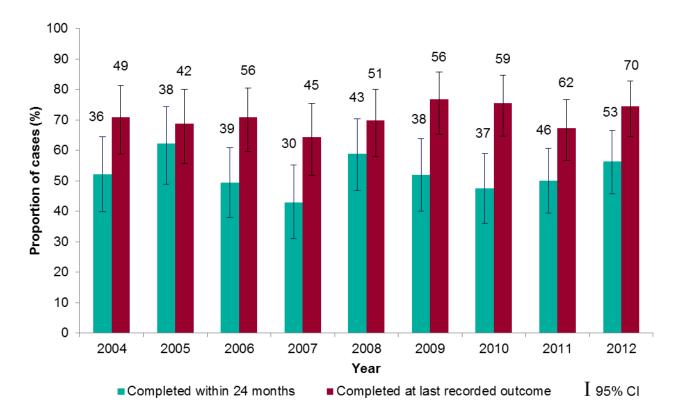
1.1% 5.3% 5.3% 6.4% 9.6% 24.5% 4.3% 56.4% 9.6% 74.5% n = 94 n = 94 3.2% Treatment completed Died Lost to follow up Still on treatment Treatment stopped Not evaluated**

* Includes initial and amplified MDR/RR-TB and MDR-TB treatment cases only

** Not evaluated includes missing, unknown and transferred out

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Treatment completion for drug resistant TB cases, England, 2004-2012*



* Includes initial and amplified MDR/RR-TB and MDR-TB treatment cases only Data labels display the number of cases completing treatment

> Source: Enhanced Tuberculosis Surveillance (ETS) Data extracted: March 2015. Prepared by: TB Section, National Infection Service, Public Health England

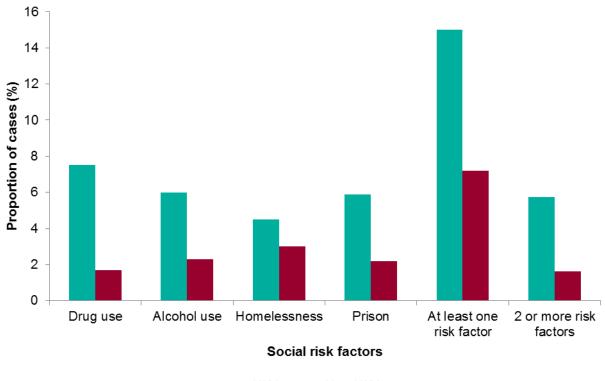
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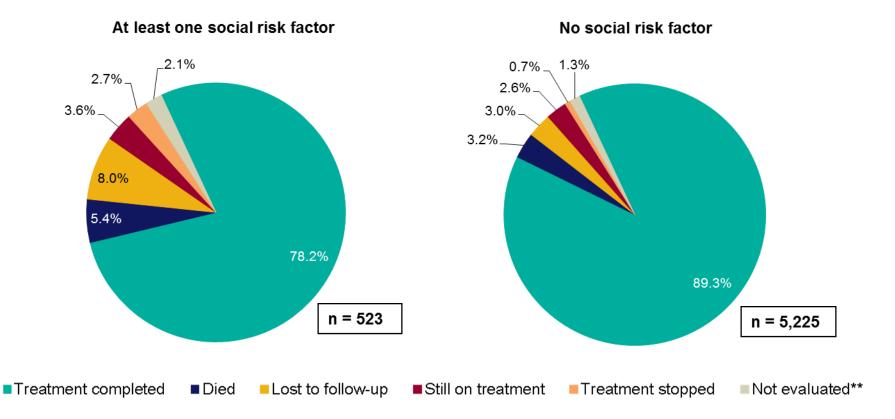
Proportion of TB cases with a social risk factor by place of birth, England, 2014



■ UK born ■ Non-UK born



Last recorded TB outcome for entire drug sensitive cohort by social risk factor*, England, 2013



* Excludes initial and amplified to rifampicin resistant TB and MDR-TB cases and MDR-TB treated cases and those with CNS, spinal, miliary or cryptic disseminated TB ** Not evaluated includes missing, unknown and transferred out

Tuberculosis in England: 2015 report

Source: Enhanced Tuberculosis Surveillance (ETS) Data extracted: March 2015. Prepared by: TB Section, National Infection Service, Public Health England

Summary

- Clinical suspicion key
 - 'great mimicker'
 - epidemiology more sensitive than symptoms (or plain CXR)
- Epidemiology young non UK born versus caucasian elderly UK born
- Pulmonary/ extrapulmonary presentations chest radiology
- Radiology 'normal' symptomatic cases need assessing
- Refer to local TB clinic
- Latent TB screening
 - Contact screening
 - New Entrant Screening