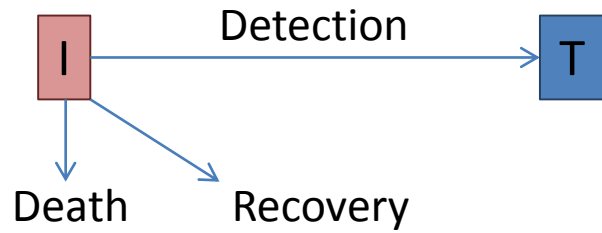


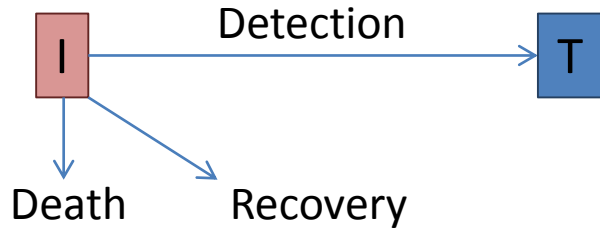
# Simulating ACF interventions in high-burden countries - how can we deal with uncertain case detection rates?

James Trauer





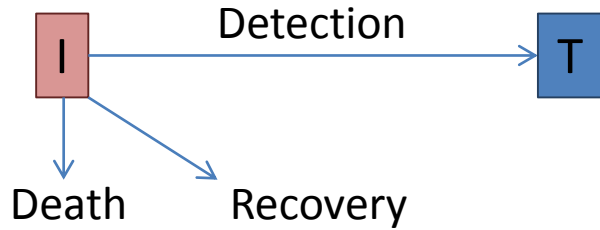
$$CDR = \frac{\textit{detection}}{\textit{detection} + \textit{death} + \textit{recovery}}$$



$$CDR = \frac{\textit{detection}}{\textit{detection} + \textit{death} + \textit{recovery}}$$

Known

The diagram shows the word 'Known' at the bottom with three arrows pointing upwards to the terms in the equation: 'CDR', 'detection', and 'death + recovery'.



$$CDR = \frac{\textit{detection}}{\textit{detection} + \textit{death} + \textit{recovery}}$$

Known

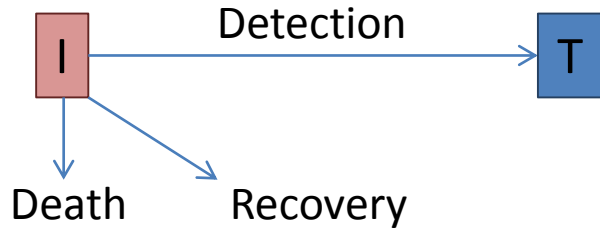
OPEN ACCESS Freely available online

PLoS one

### Natural History of Tuberculosis: Duration and Fatality of Untreated Pulmonary Tuberculosis in HIV Negative Patients: A Systematic Review

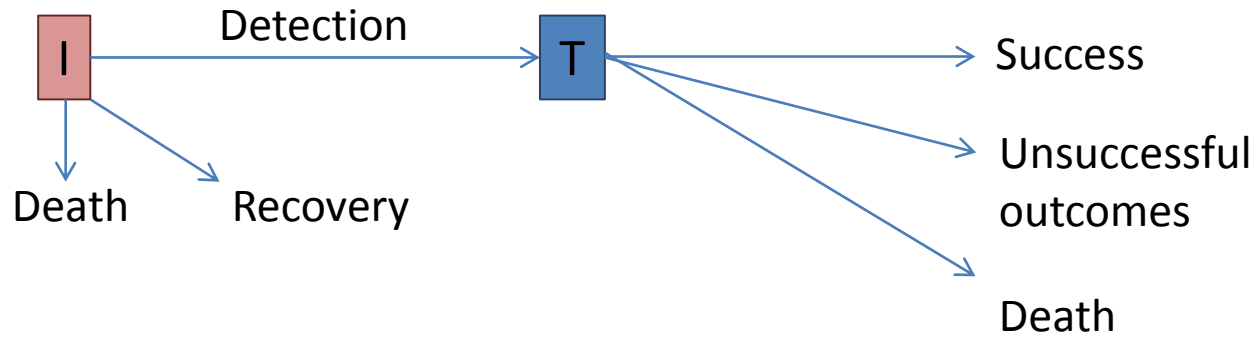
Edine W. Tiemersma<sup>1,2\*</sup>, Marieke J. van der Werf<sup>1,2</sup>, Martien W. Borgdorff<sup>3</sup>, Brian G. Williams<sup>4</sup>, Nico J. D. Nagelkerke<sup>5</sup>

1 KNCV Tuberculosis Foundation, The Hague, The Netherlands, 2 Center for Infection and Immunity Amsterdam (CINIMA), Academic Medical Center, University of Amsterdam, Amsterdam, The Netherlands, 3 Cluster Infectious Disease Control, Municipal Health Service (GGD) Amsterdam, Amsterdam, The Netherlands, 4 South African Centre for Epidemiological Modelling and Analysis, Stellenbosch, South Africa, 5 Department of Community Medicine, Faculty of Medicine and Health Sciences, UAE University, Al Ain, United Arab Emirates



$$CDR = \frac{\textit{detection}}{\textit{detection} + \textit{death} + \textit{recovery}}$$

$$\therefore \textit{detection} = \frac{(\textit{death} + \textit{recovery}) \times CDR}{1 - CDR}$$



# WHO GLOBAL TB REPORT 2016

## ACCESS TO CARE

---

**6.1 million** people had  
**ACCESS TO QUALITY TB CARE**

**4.3 million** people  
**MISSED OUT**

A row of silhouettes representing a diverse group of people, including men and women of various ages and ethnicities, positioned behind the text for the missed out statistic.

# Time to diagnosis

**BMC Infectious Diseases**



Research article

**Open Access**

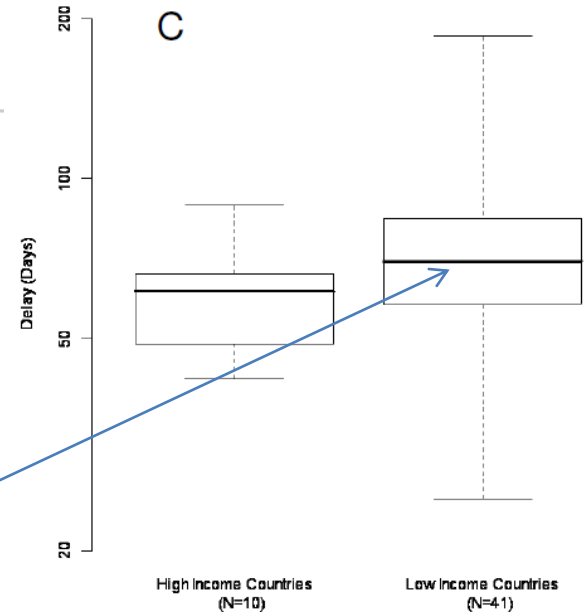
## Time delays in diagnosis of pulmonary tuberculosis: a systematic review of literature

Chandrashekhar T Sreeramareddy\*<sup>1,5</sup>, Kishore V Panduru<sup>2,6</sup>, Joris Menten<sup>3</sup> and J Van den Ende<sup>4</sup>

Address: <sup>1</sup>Department of Community Medicine, Manipal Teaching Hospital, Manipal College of Medical Sciences, Pokhara, Nepal, <sup>2</sup>Department of Internal Medicine, Manipal Teaching Hospital, Manipal College of Medical Sciences, Pokhara, Nepal, <sup>3</sup>Department of Public Health, Institute of Tropical Medicine, Antwerp, Belgium, <sup>4</sup>Department of Clinical Sciences, Institute of Tropical Medicine, Antwerp, Belgium, <sup>5</sup>Current address: Department of Community Medicine, Melaka-Manipal Medical College, Jalan Batu Hampar, Bukit Baru, Melaka, Malaysia and <sup>6</sup>Current address: Department of Respiratory Medicine, Ministry of Health, Brunei Darussalam

Email: Chandrashekhar T Sreeramareddy\* - chandrashekharats@yahoo.com; Kishore V Panduru - pandu\_vki@yahoo.com; Joris Menten - jmenten@itg.be; J Van den Ende - jvdende@itg.be

\* Corresponding author



- Median total delay 68 days (IQR 25 to 185 days)
  - Patient delay 5 to 162 days
  - Health system delay 2 to 87 days



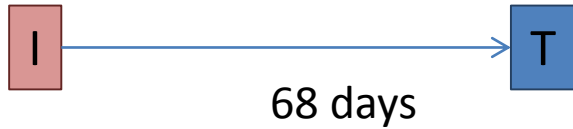
Detected



Never detected

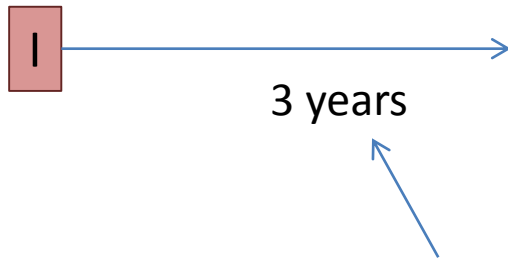


# Detected



Idealised

# Never detected



OPEN ACCESS Freely available online

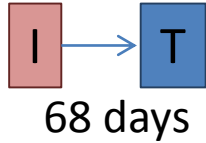
PLoS one

## Natural History of Tuberculosis: Duration and Fatality of Untreated Pulmonary Tuberculosis in HIV Negative Patients: A Systematic Review

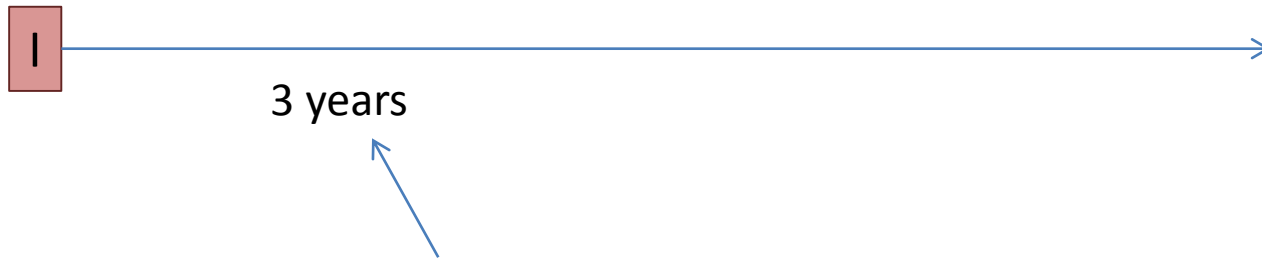
Edine W. Tiemersma<sup>1,2\*</sup>, Marieke J. van der Werf<sup>1,2</sup>, Martien W. Borgdorff<sup>3</sup>, Brian G. Williams<sup>4</sup>, Nico J. D. Nagelkerke<sup>5</sup>

1 KNCV Tuberculosis Foundation, The Hague, The Netherlands, 2 Center for Infection and Immunity Amsterdam (CIIMM) Academic Medical Center, University of Amsterdam, Amsterdam, The Netherlands, 3 Cluster Infectious Disease Control, Municipal Health Service (GGD) Amsterdam, Amsterdam, The Netherlands, 4 South African Centre for Epidemiological Modelling and Analysis, Stellenbosch, South Africa, 5 Department of Community Medicine, Faculty of Medicine and Health Sciences, UAE University, Al Ain, United Arab Emirates

# Detected



# Never detected



OPEN ACCESS Freely available online

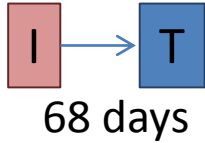


## Natural History of Tuberculosis: Duration and Fatality of Untreated Pulmonary Tuberculosis in HIV Negative Patients: A Systematic Review

Edine W. Tiemersma<sup>1,2\*</sup>, Marieke J. van der Werf<sup>1,2</sup>, Martien W. Borgdorff<sup>3</sup>, Brian G. Williams<sup>4</sup>, Nico J. D. Nagelkerke<sup>5</sup>

1 KNMIV Tuberculosis Foundation, The Hague, The Netherlands, 2 Center for Infection and Immunity Amsterdam (CIIMM) Academic Medical Center, University of Amsterdam, Amsterdam, The Netherlands, 3 Cluster Infectious Disease Control, Municipal Health Service (GGD) Amsterdam, Amsterdam, The Netherlands, 4 South African Centre for Epidemiological Modelling and Analysis, Stellenbosch, South Africa, 5 Department of Community Medicine, Faculty of Medicine and Health Sciences, UAE University, Al Ain, United Arab Emirates

# Detected

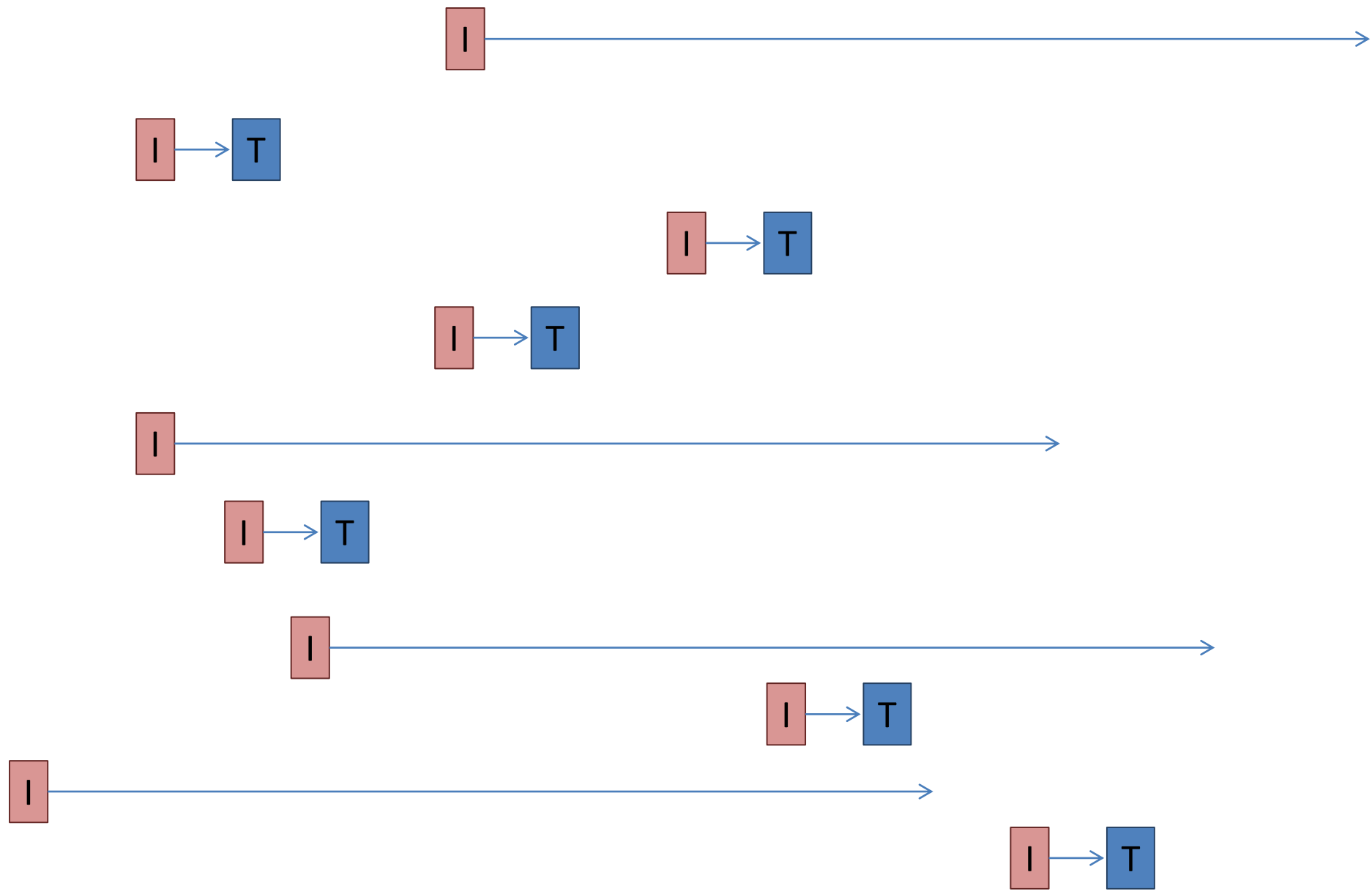


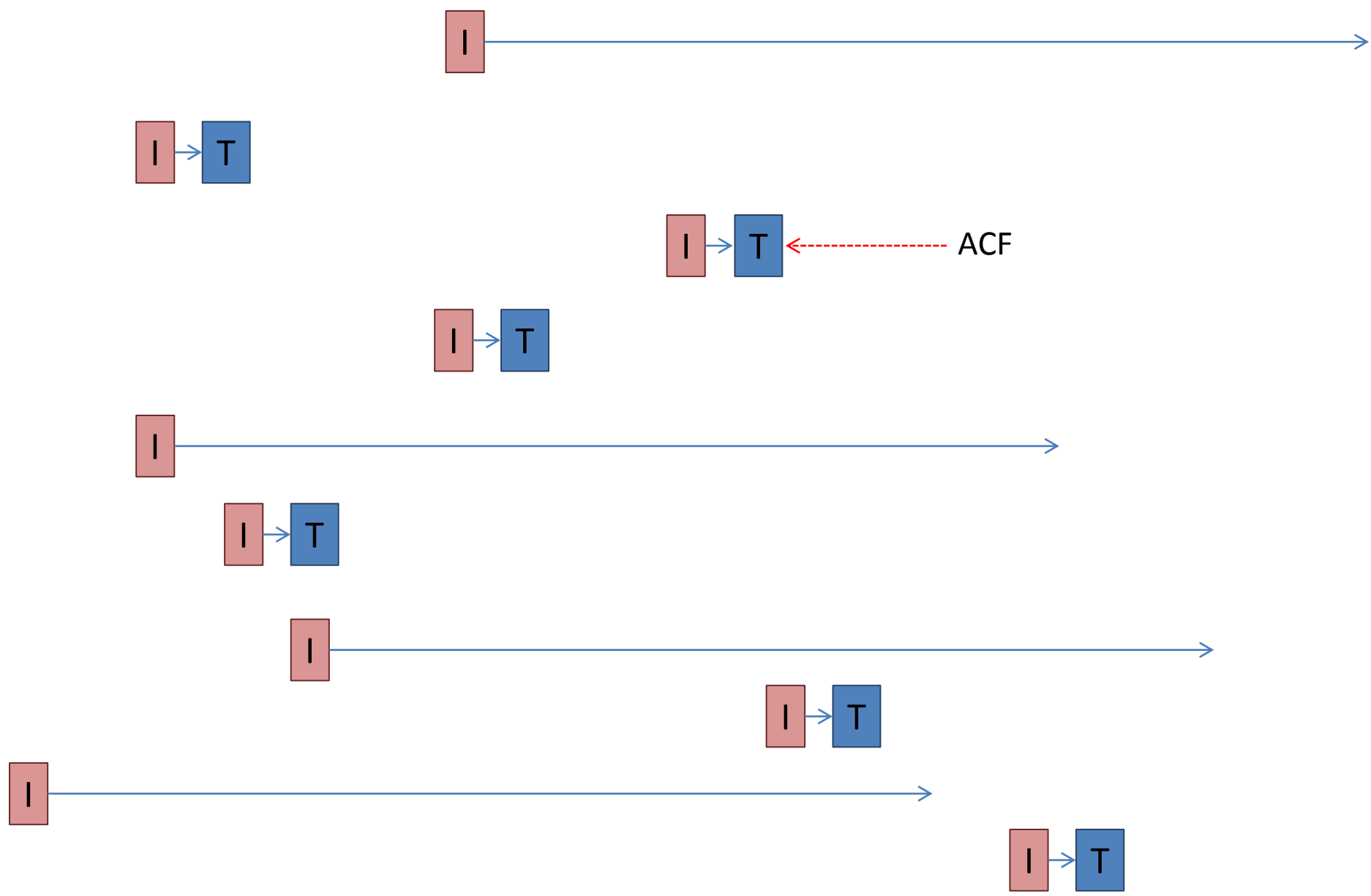
x 60% → <10% of transmission

# Never detected

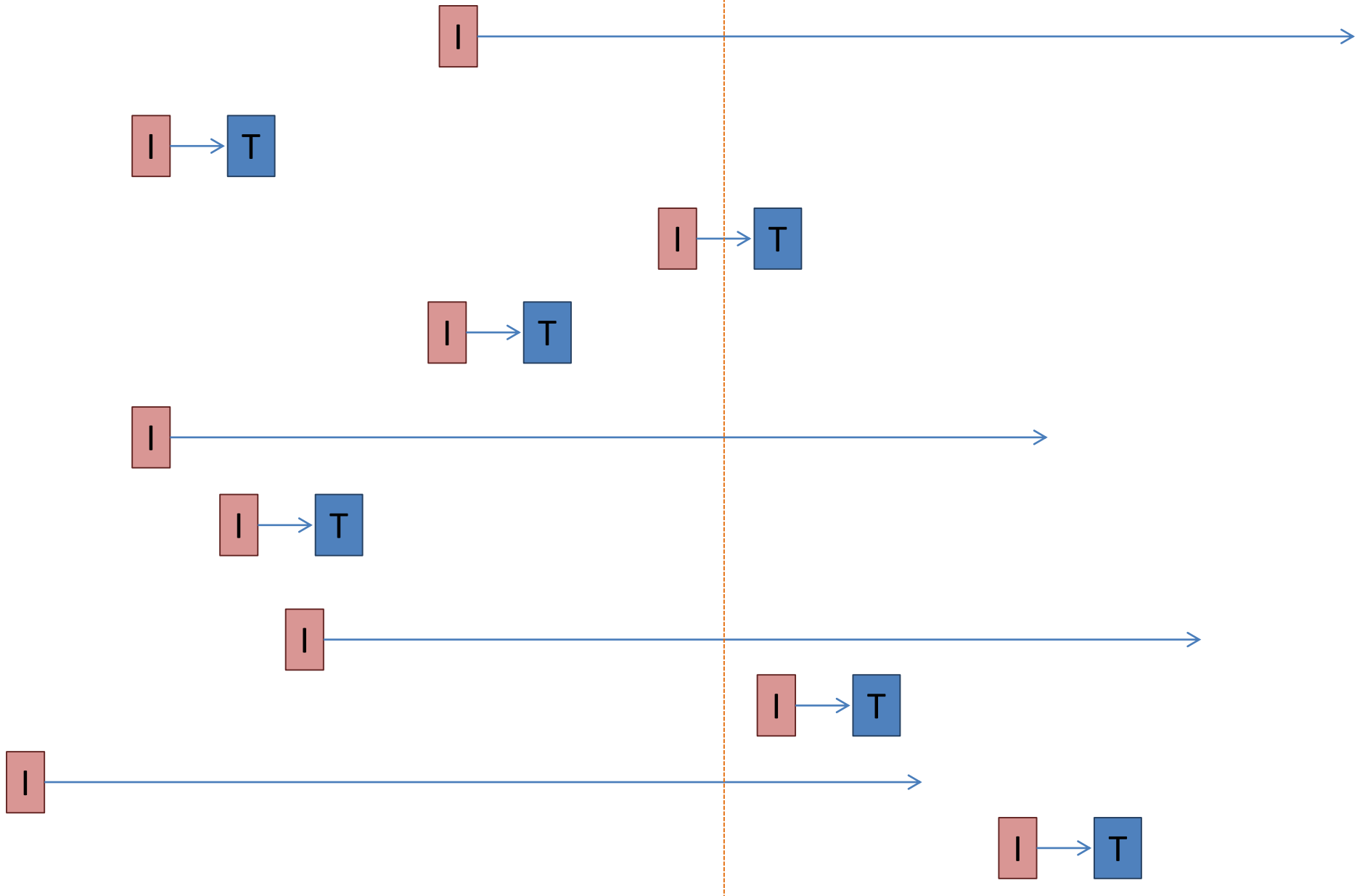


x 40% → >90% of transmission





ACF



# Idealised

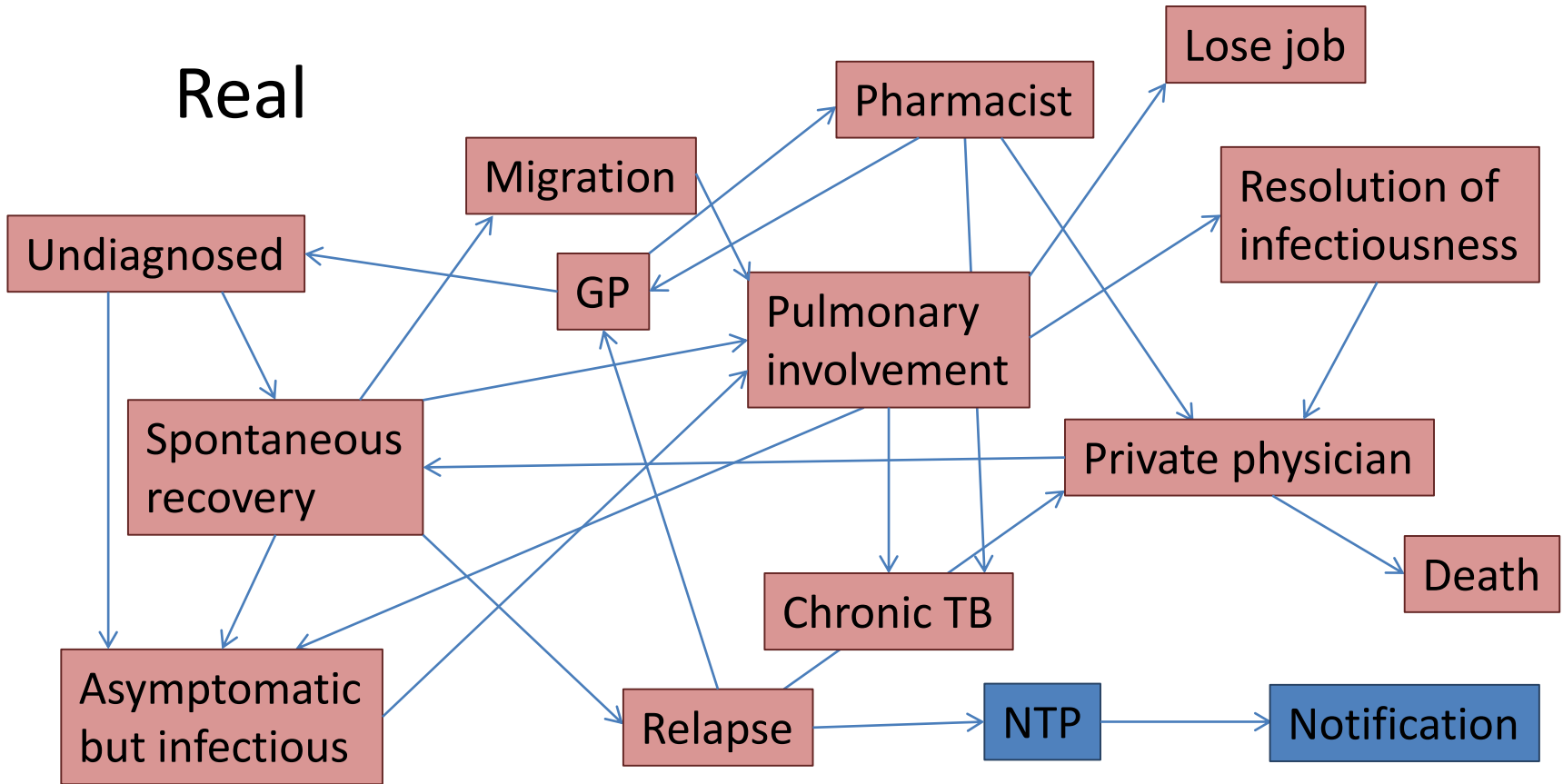




# Idealised



# Real



# Heterogeneities in pre-treatment pathways

- Biological
  - Symptoms
  - Severity
  - Infectiousness
  - Resolution/recovery
- Patient-related
  - Priorities
  - Disability and loss of income
- Health system-related
  - Provider awareness
  - Multiple health systems
  - Access

# Conclusions

- Case detection parameter important
- Unknown cases critical
- Mechanism of ACF important
- Models simplify realities of case detection