

# Anatomy of MERS epidemic

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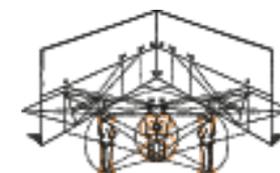
ISI Foundation  
Turin, Italy

**Inserm**



Institut national  
de la santé et de la recherche médicale

**UPMC**  
PARISUNIVERSITAS

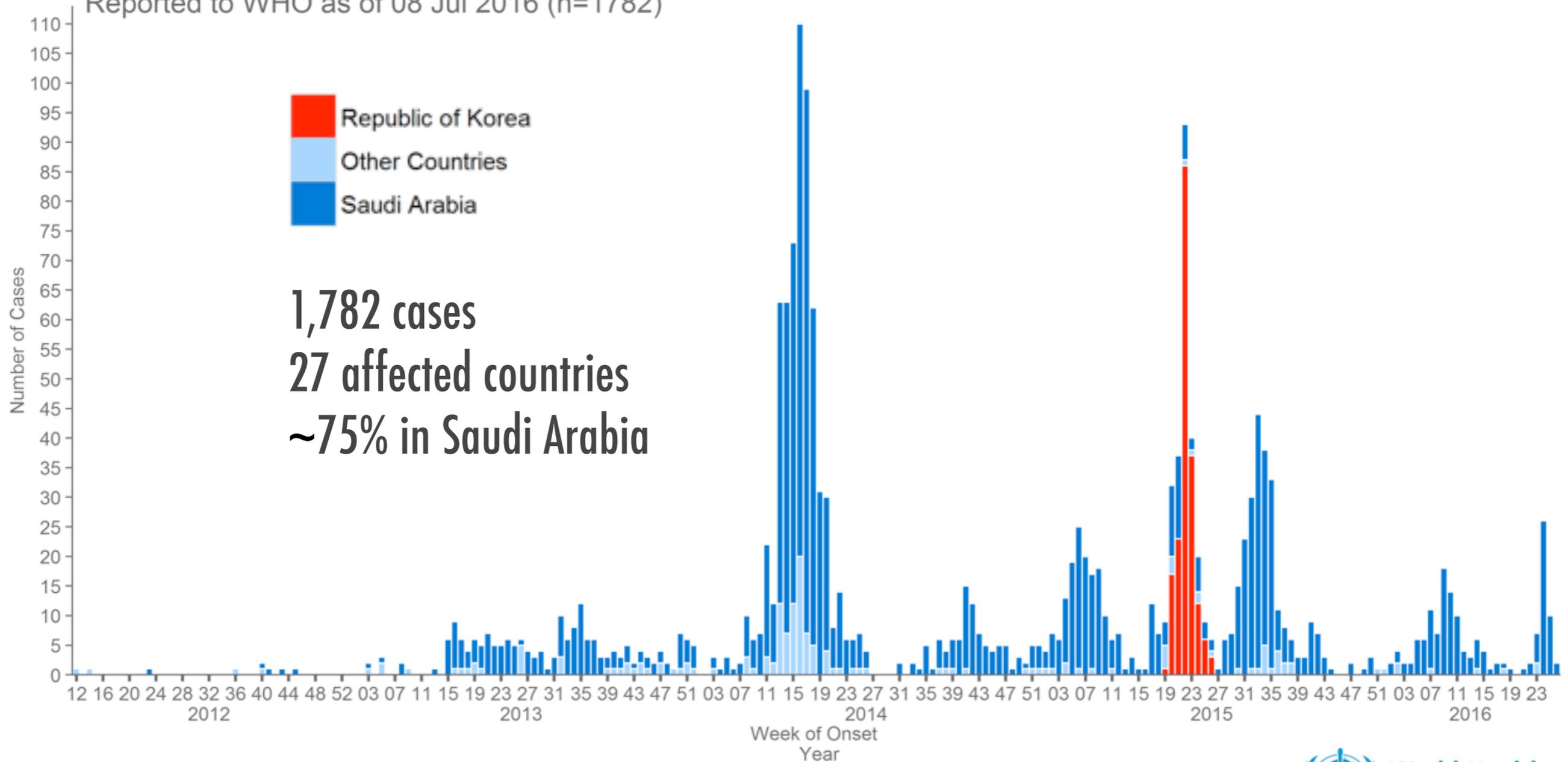


**ISI Foundation**

# MERS epidemic

## Confirmed global cases of MERS-CoV

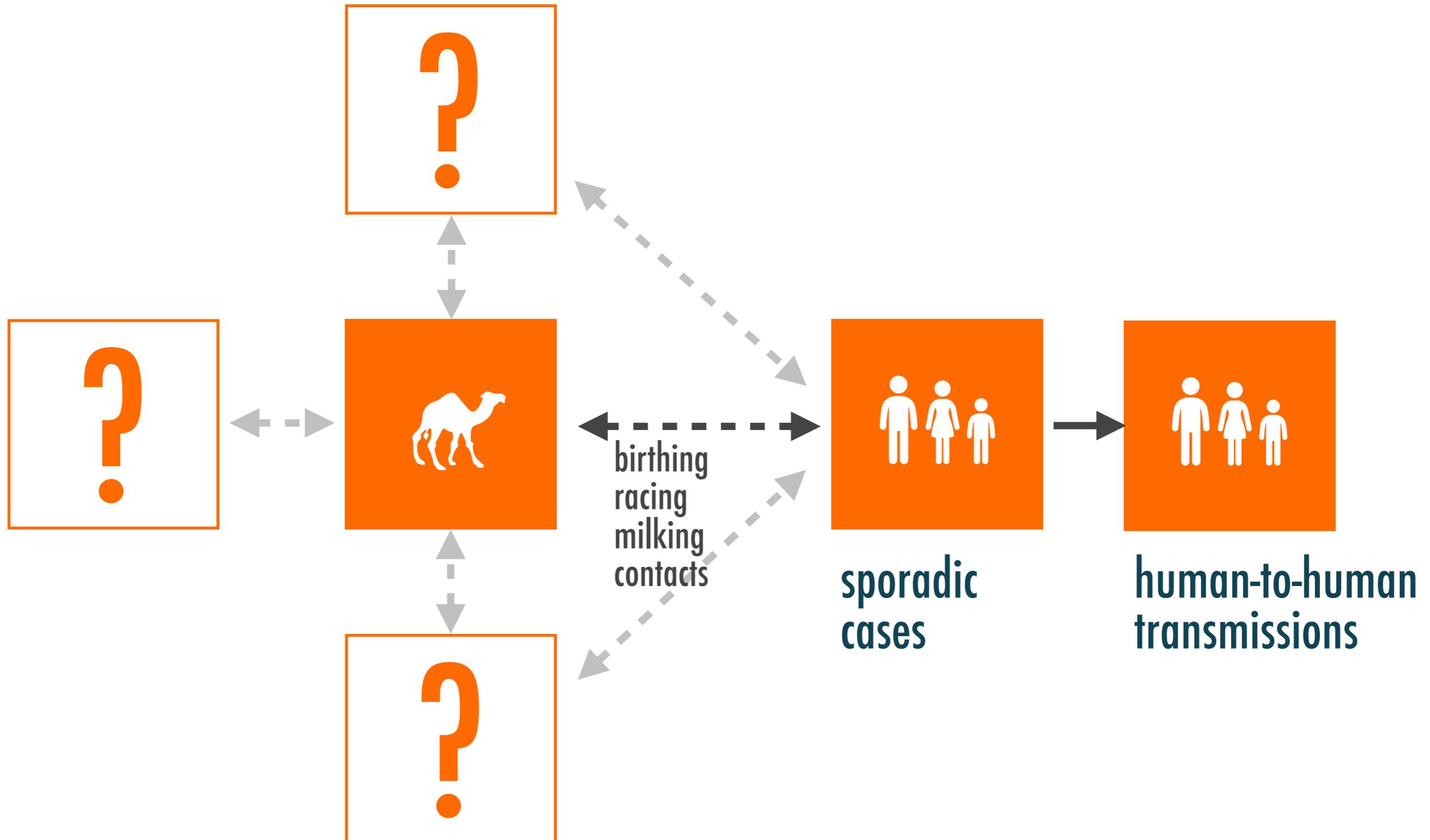
Reported to WHO as of 08 Jul 2016 (n=1782)



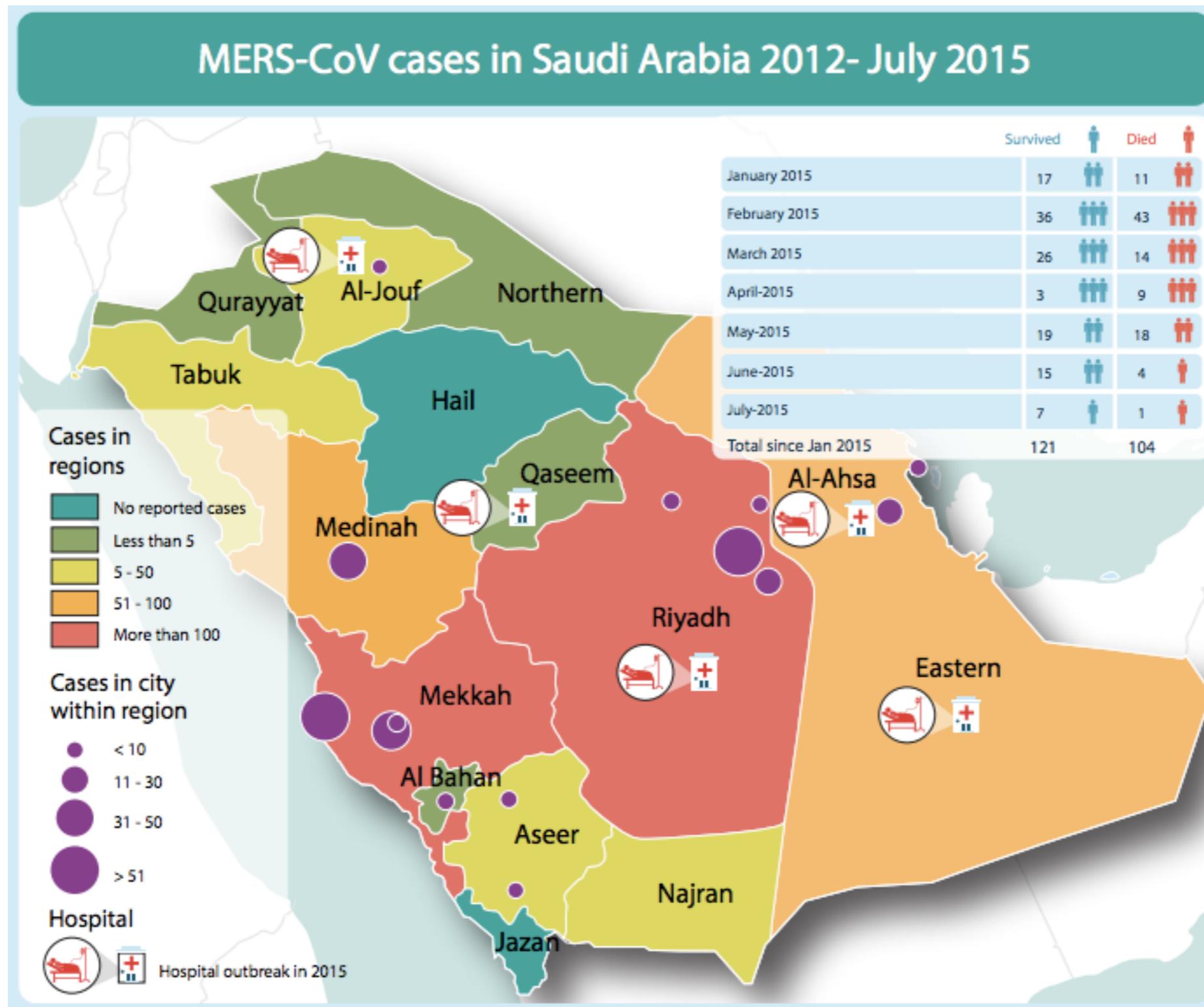
Other countries: Algeria, Austria, Bahrain, China, Egypt, France, Germany, Greece, Iran, Italy, Jordan, Kuwait, Lebanon, Malaysia, Netherlands, Oman, Philippines, Qatar, Thailand, Tunisia, Turkey, United Arab Emirates, United Kingdom, United States of America, Yemen  
Please note that the underlying data is subject to change as the investigations around cases are ongoing. Onset date estimated if not available.



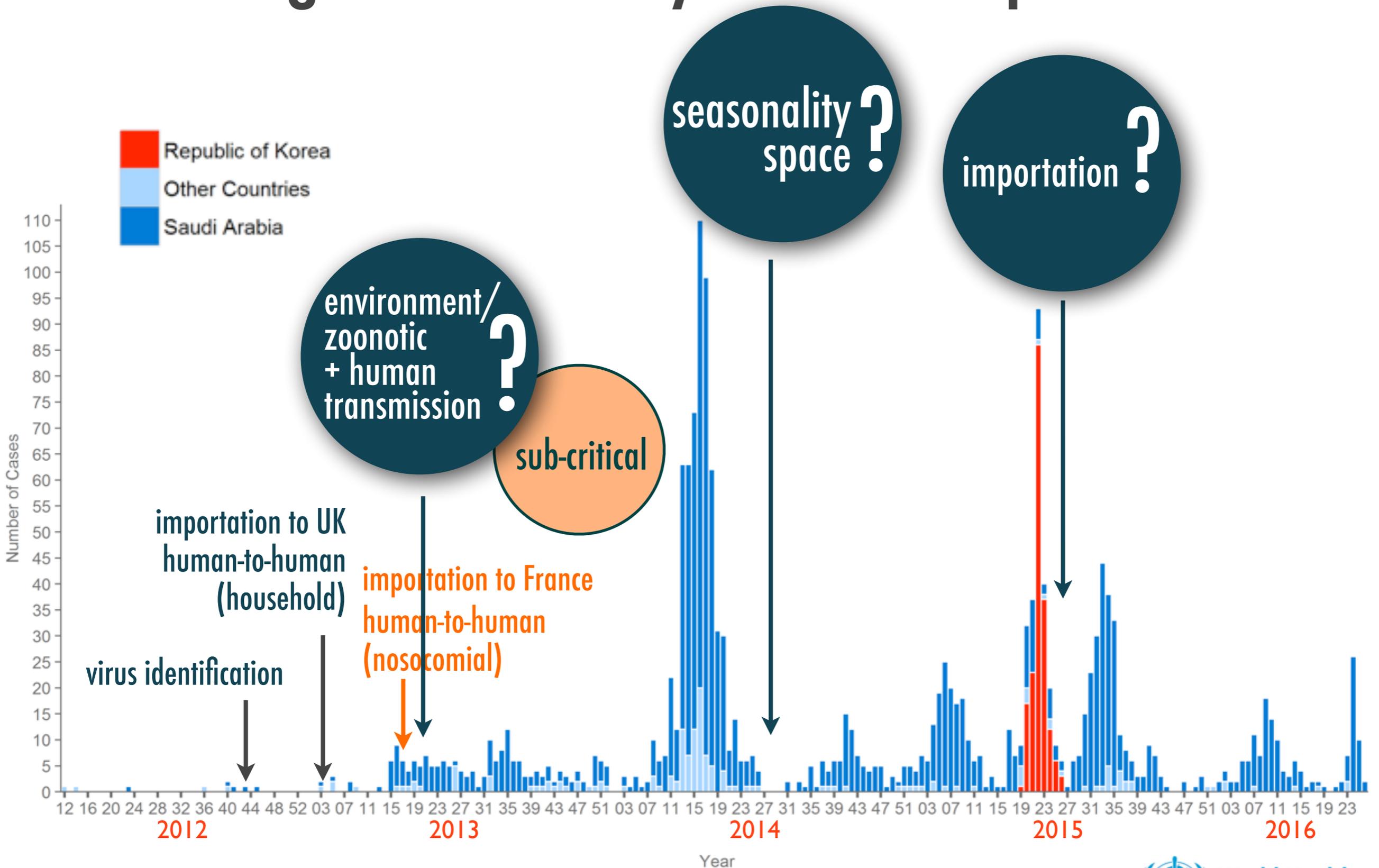
# MERS ecology



# MERS geography



# uncovering the anatomy of MERS epidemic

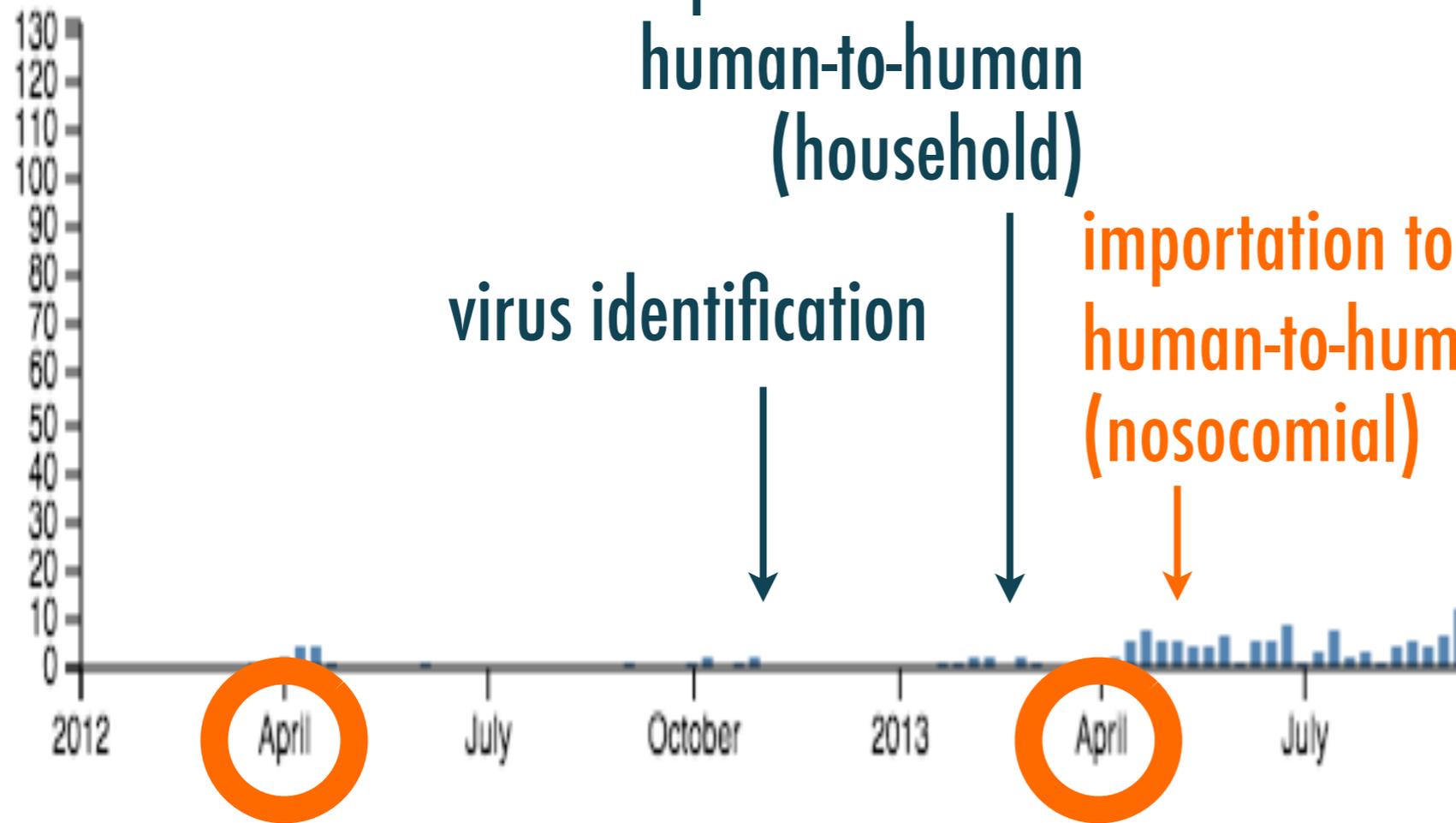


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# Spring 2014

seasonality  
space ?

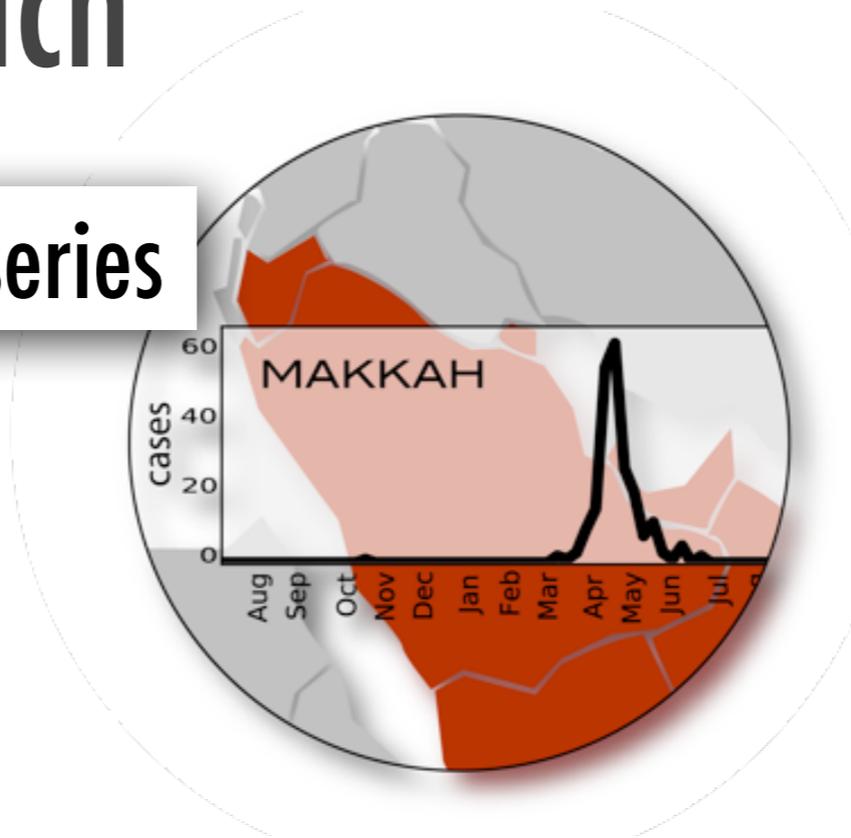
cases by date



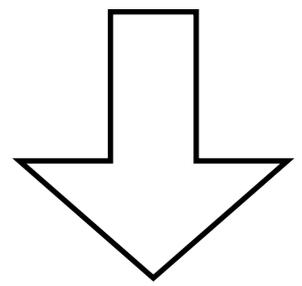
# integrated approach

incidence time series

# 1

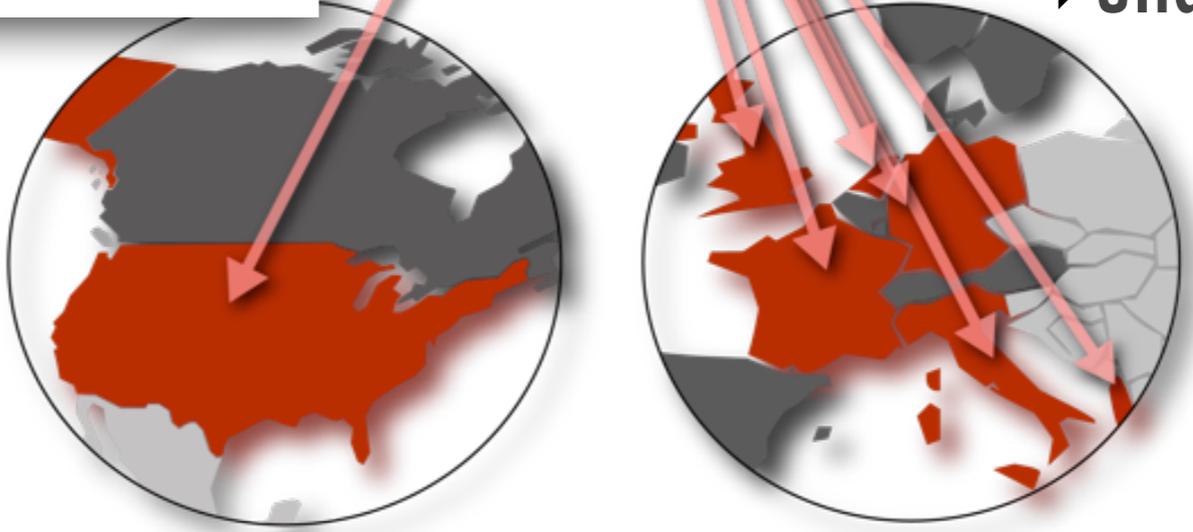


- ▶ generation sporadic cases  $\rho_{sp}^r(t)$
- ▶ human transmissions  $R^r(t)$



seasonality space

case importation



▶ underascertainment  $\rho$

■ countries with  $n_j=0$   
■ countries with  $n_j>0$

# geo-temporal variation

$$p_{sp}^r = \alpha_r p_{zoon}$$

$$R^r = \beta_r R$$

spatial  
dependence

temporal  
dependence

$p_{sp}$

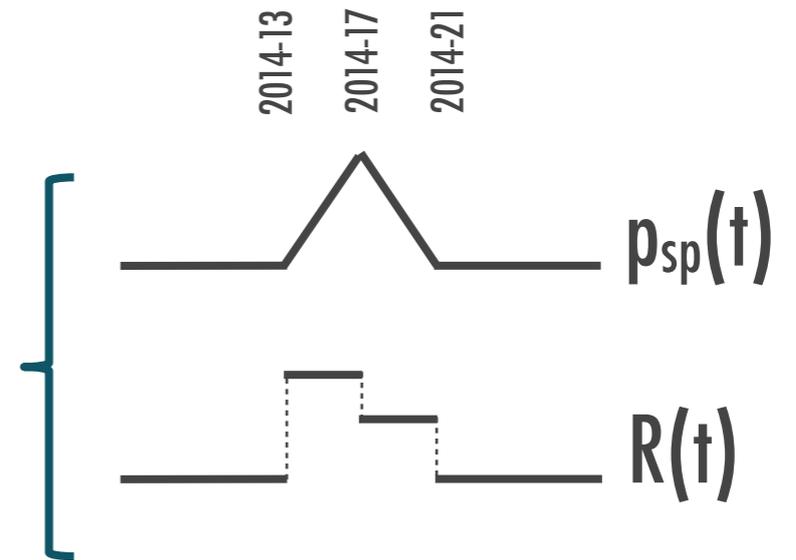
?

?

$R$

?

?

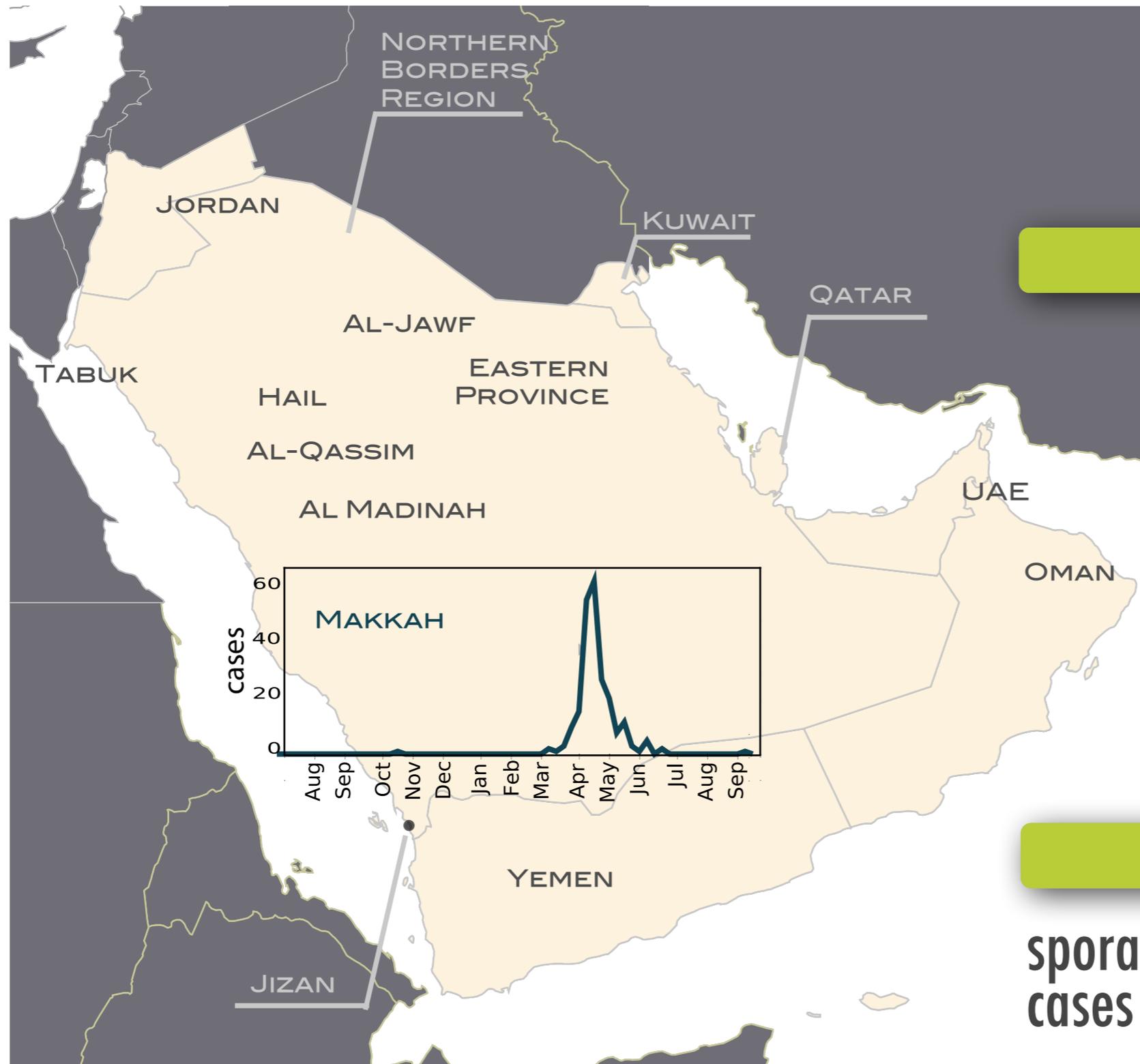


# 17 regions



11 provinces Saudi Arabia  
Qatar  
Oman  
Kuwait  
Jordan  
UAE  
Yemen

# onset time series of cases incidence



complete info scenario

66%

34%

sporadic cases

transmissions

[WHO]

partial info scenario

>34%

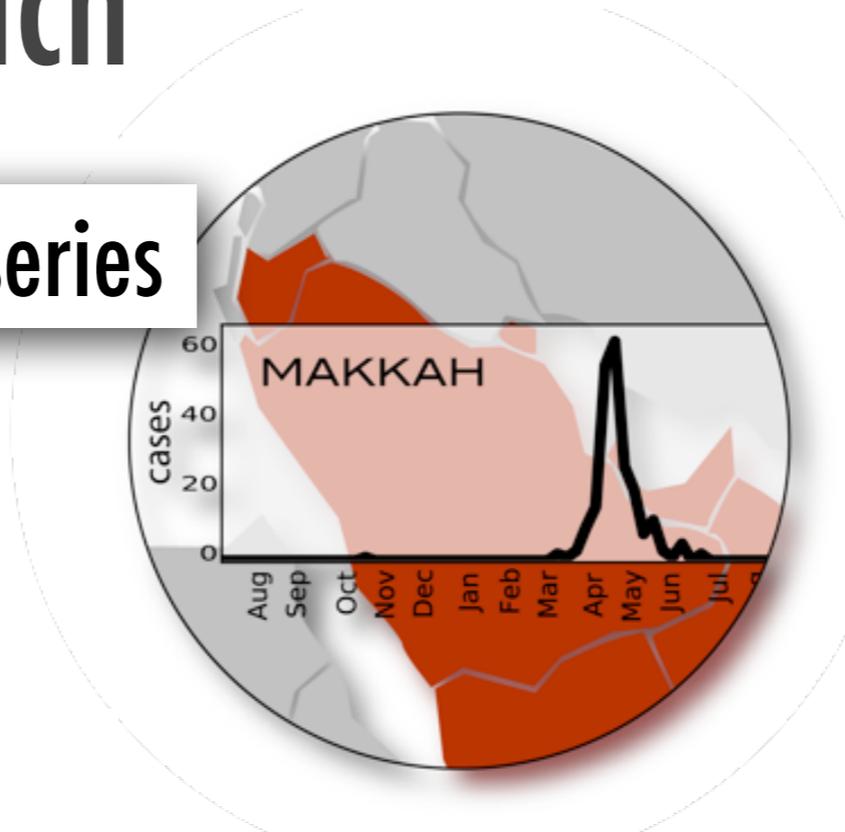
sporadic cases

transmissions

# integrated approach

incidence time series

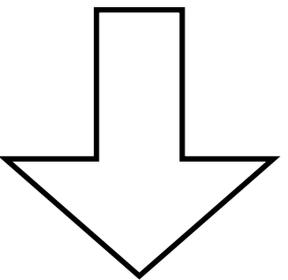
# 1



▶ generation sporadic cases  $\rho_{sp}^r(t)$

▶ human transmissions  $R^r(t)$

best fit values



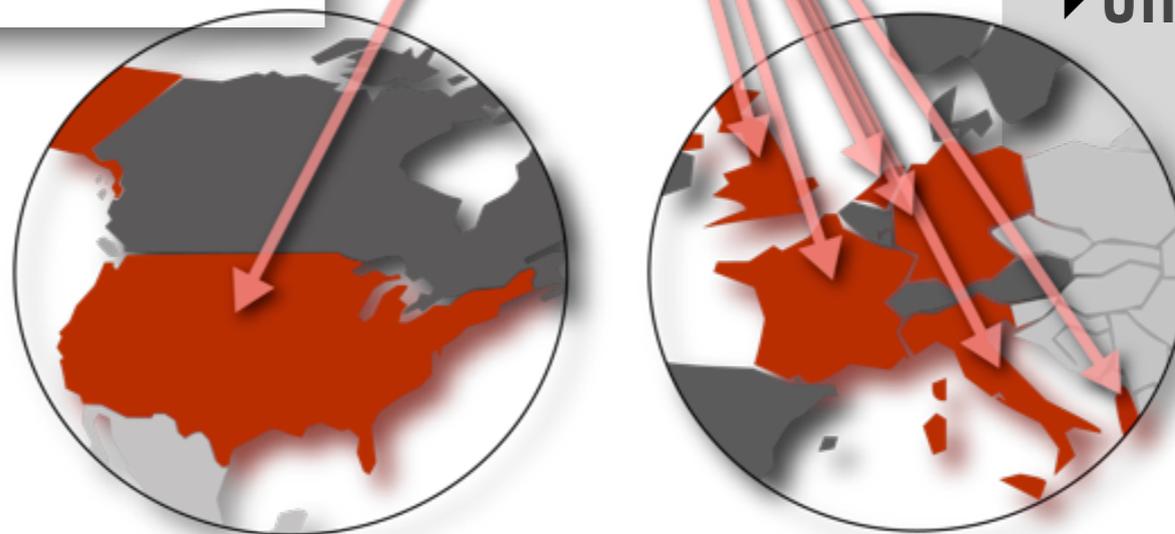
▶ underascertainment

$\rho$

seasonality space

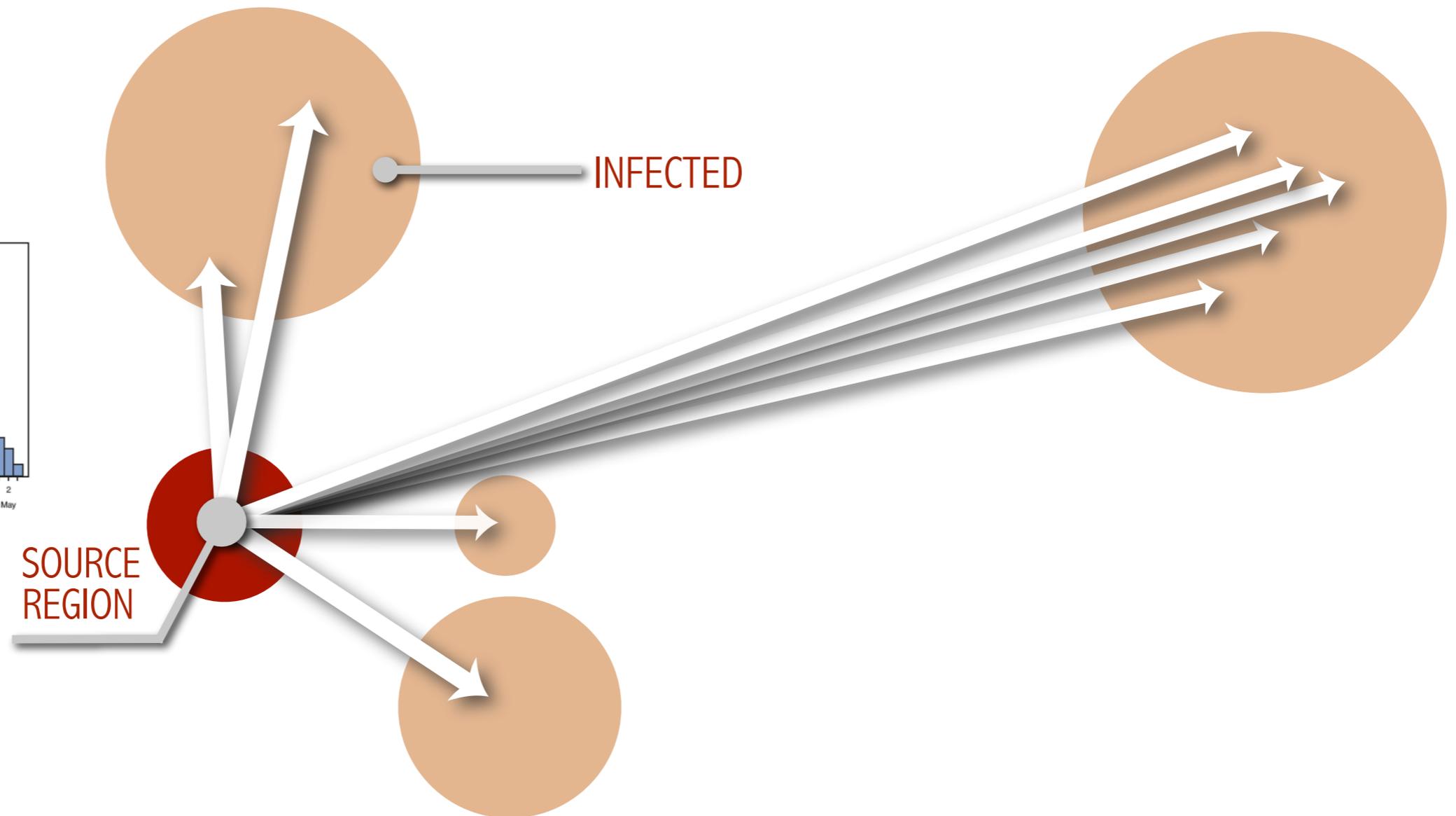
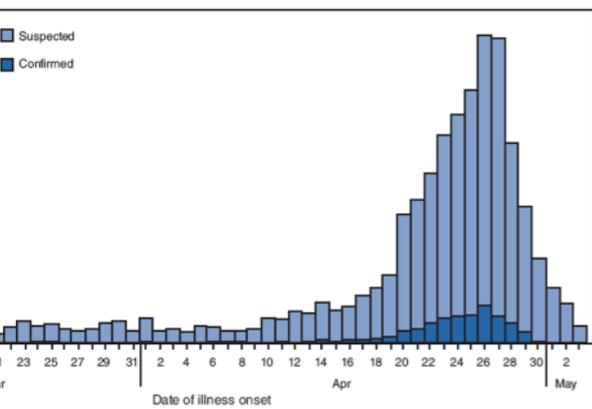
case importation

# 2



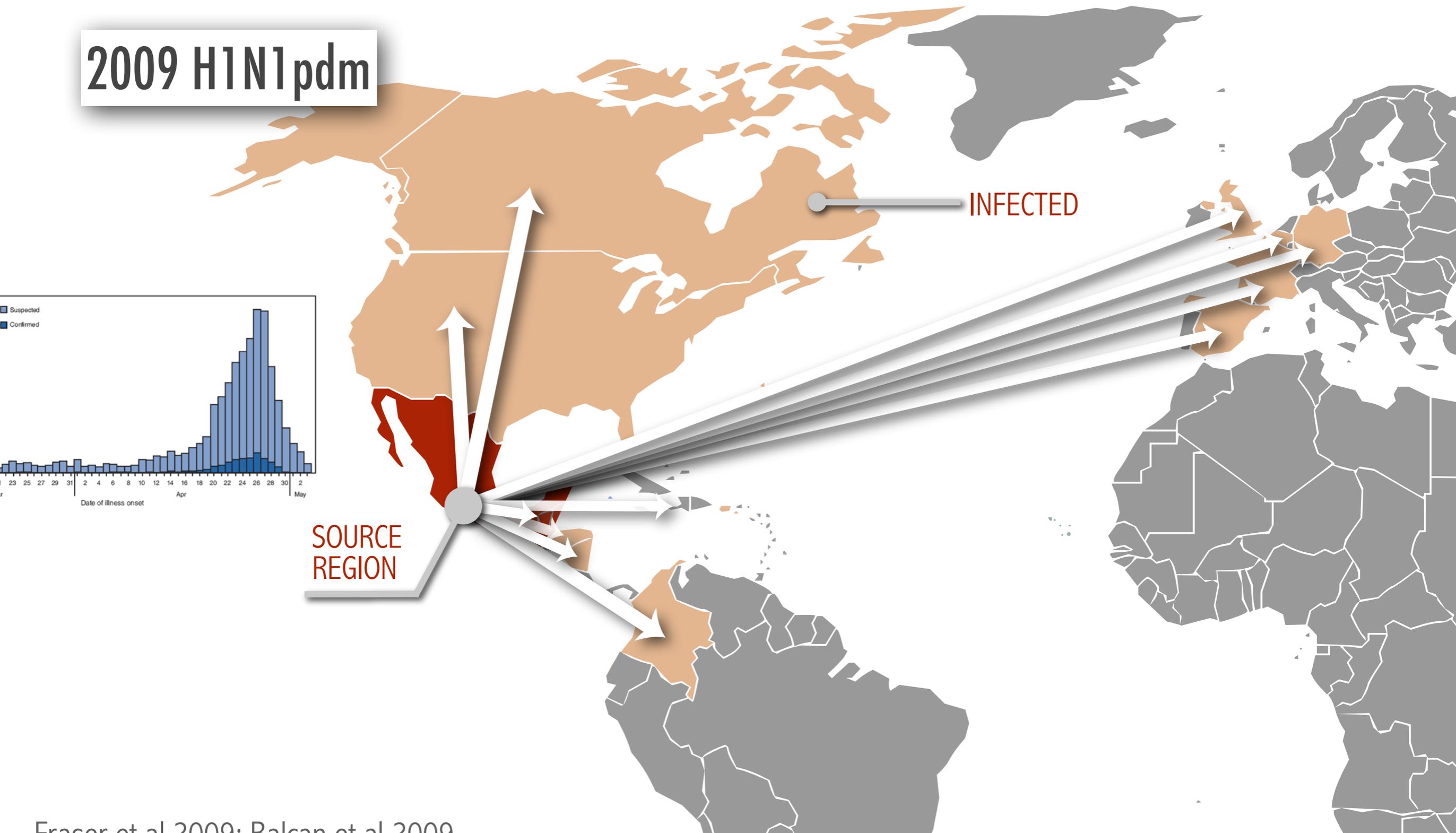
■ countries with  $n_j=0$   
■ countries with  $n_j>0$

# estimating $\rho$ from importations



# estimating $\rho$ from importations

2009 H1N1 pdm



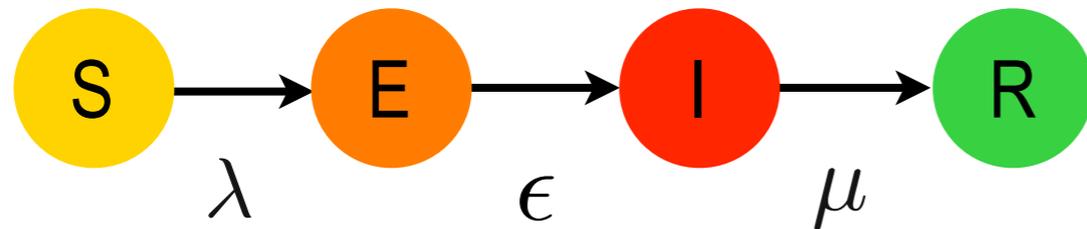


# CHARTING THE NEXT PANDEMIC



GLEAMViz.org

# sporadic + human-to-human transmission



5.2d: latency

7.6d: generation time

Assiri et al 2013

force of infection

$$\lambda = \lambda^{(SP)} + \lambda^{(H-H)}$$

↓

$$p_{sp}^r(t)$$

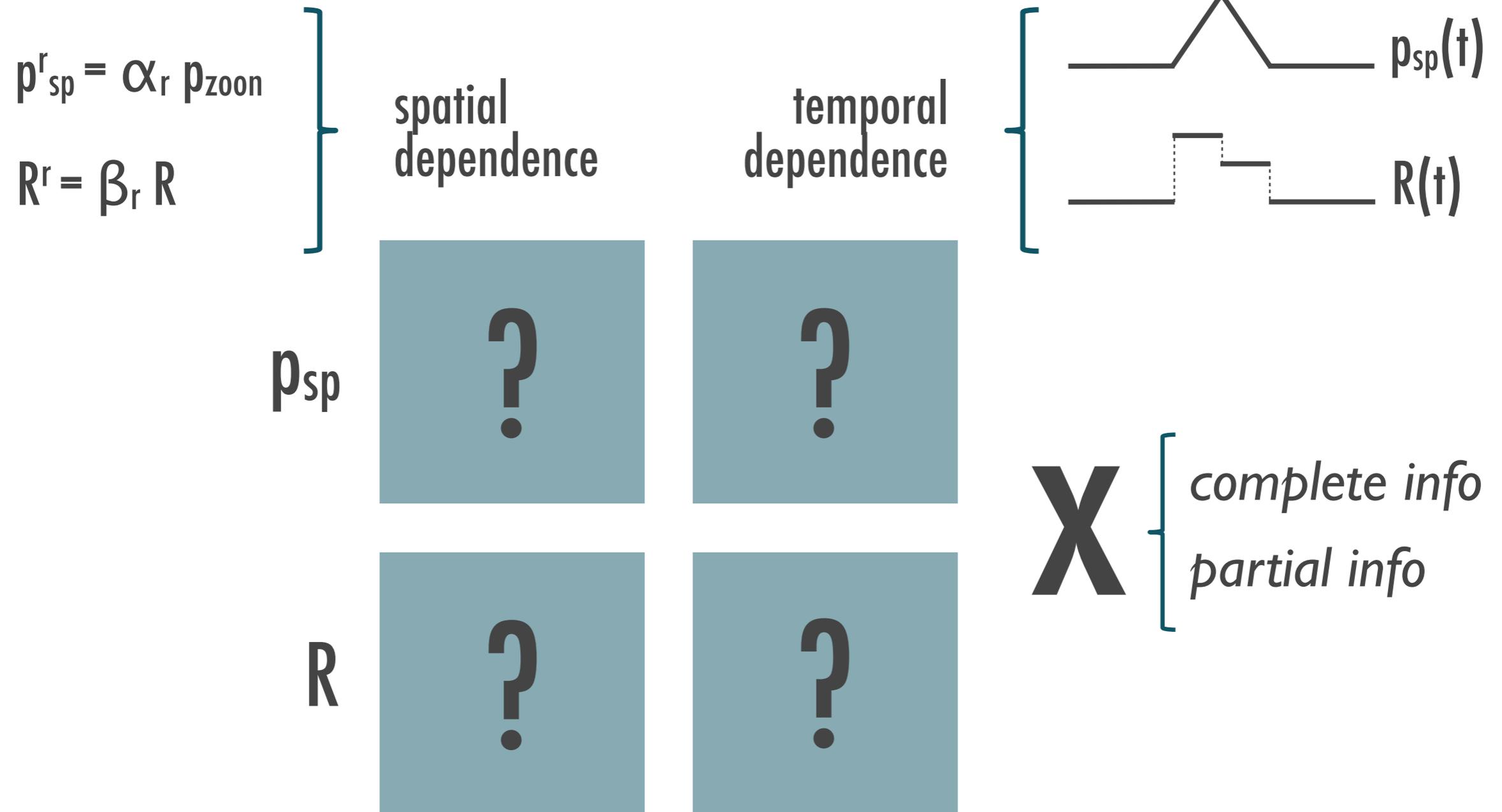
↓

$$R^r(t)$$

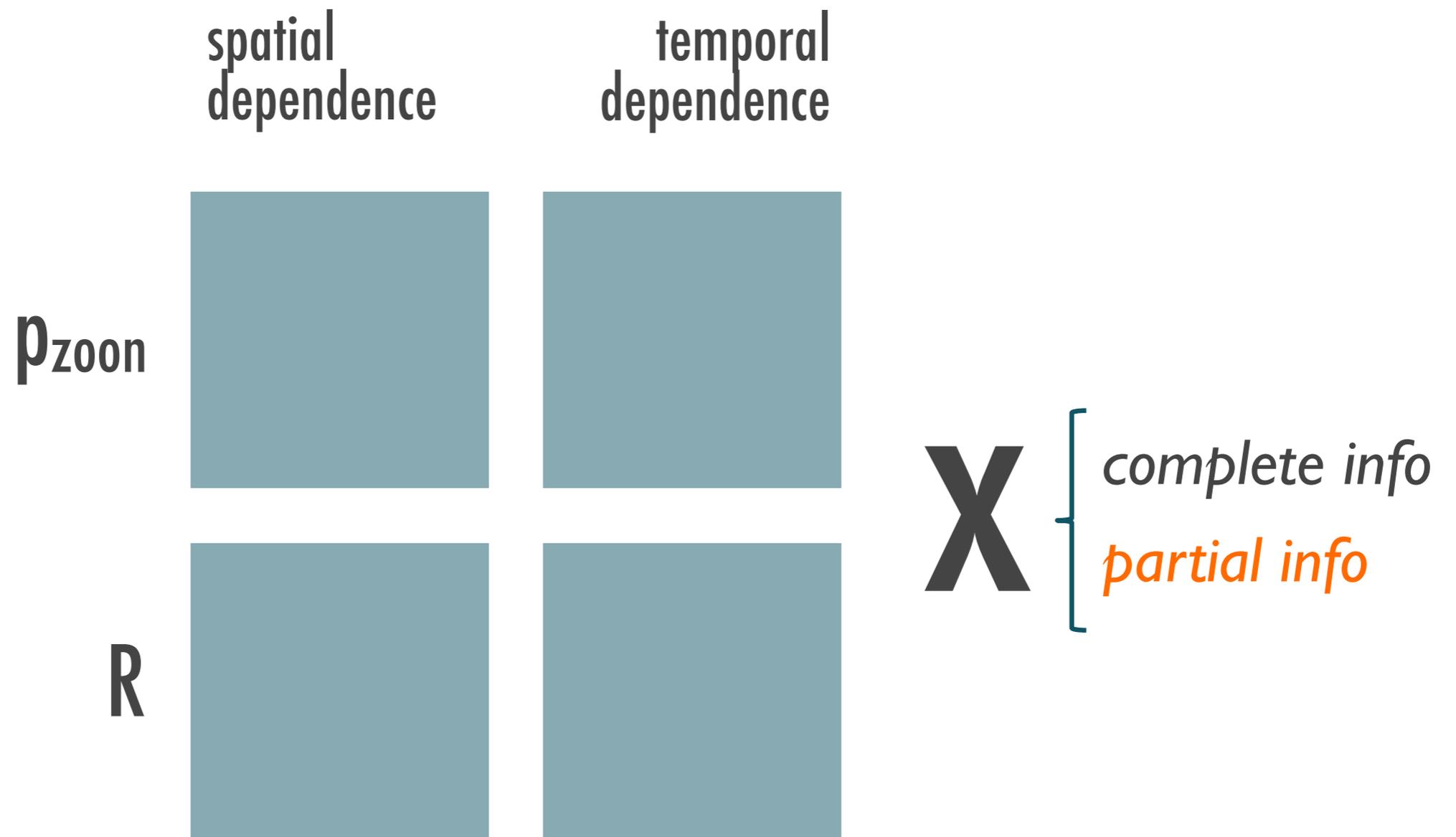
- 
- ▶ number of cases  $n^r(t)$
  - ▶ notifications

$$D^r(t) = \rho n^r(t)$$

# integrated approach: 32 models



# results



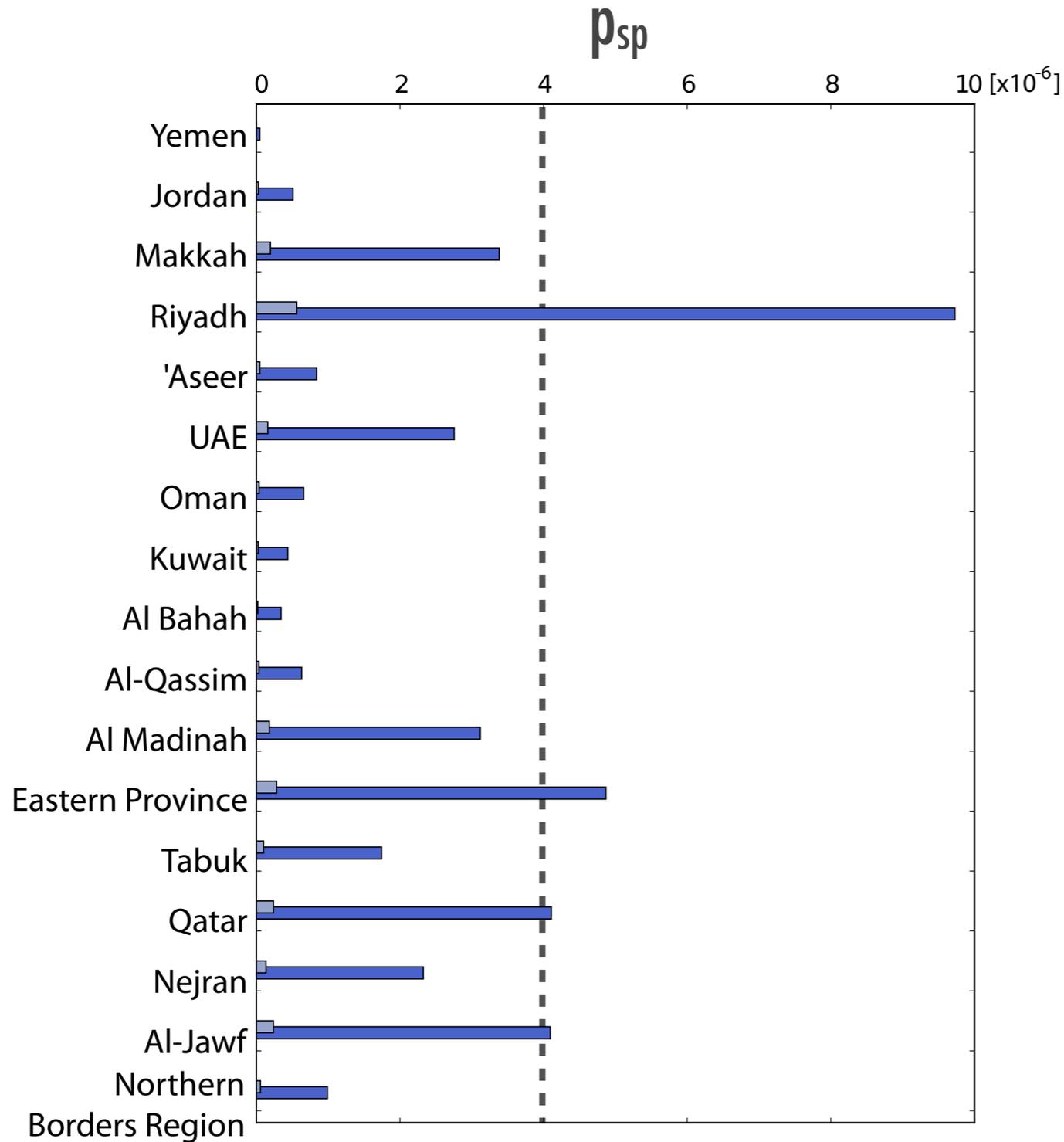
# results

	spatial dependence	temporal dependence	
$\rho_{\text{zoon}}$			<b>X</b> { <i>complete info</i> <i>partial info</i>
R			

# results

	spatial dependence	temporal dependence	
$\rho_{\text{zoon}}$			<b>X</b> { <i>complete info</i> <i>partial info</i>
R			

# geo-temporal variation

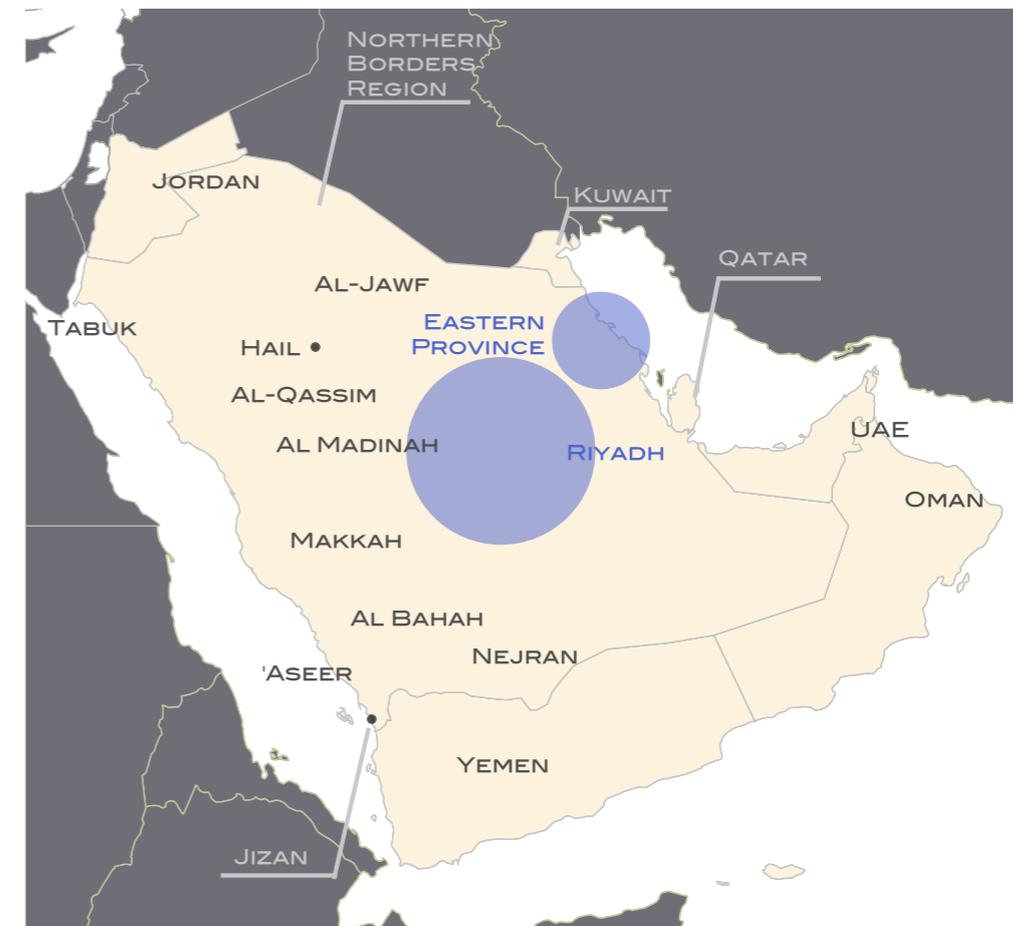


R

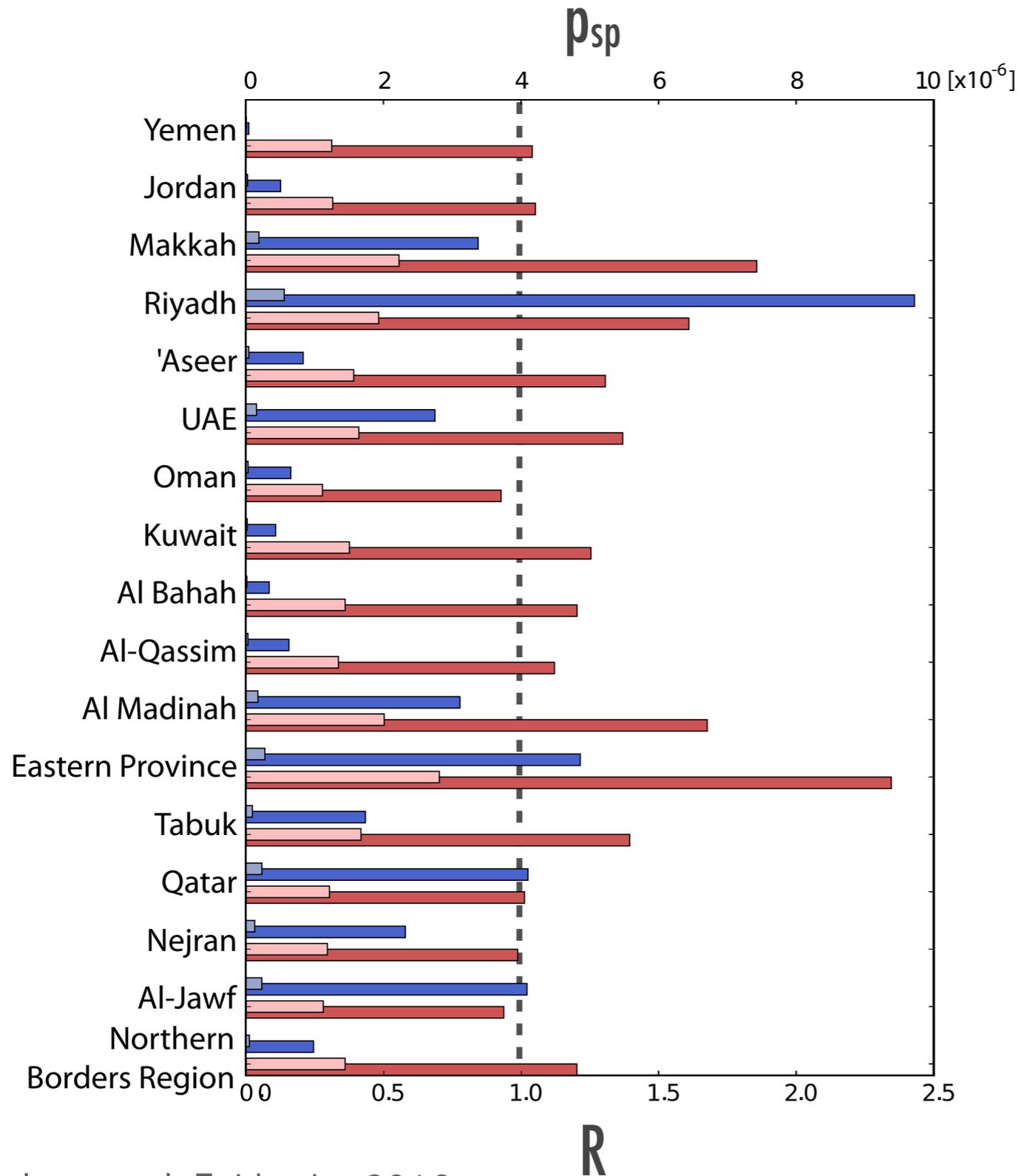
$p_{sp}$

baseline  $\rightarrow$  Spring 2014  
 6/week  $\rightarrow$  108/week  
 ~17-fold increase

acute epidemics in calves in Spring: Wernery et al 2015



# geo-temporal variation

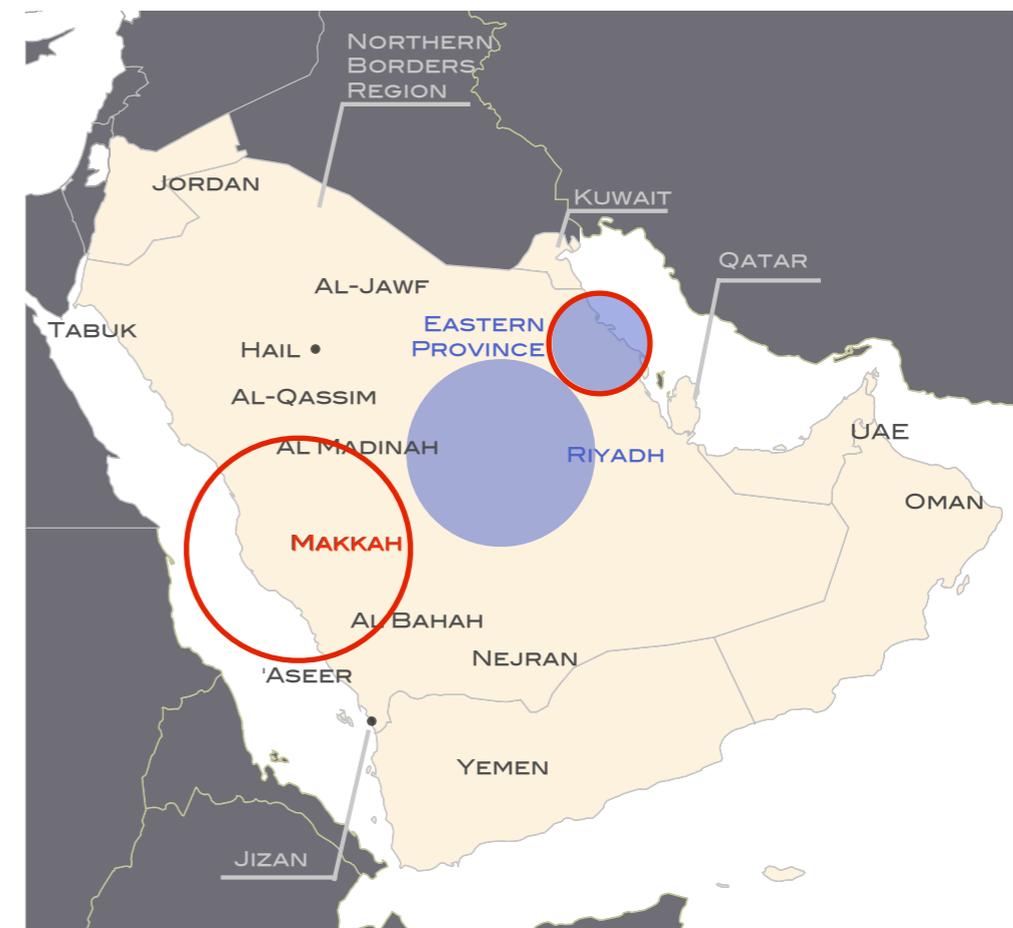


Poletto et al, Epidemics 2016

$p_{sp}$

baseline → Spring 2014  
 6/week → 108/week  
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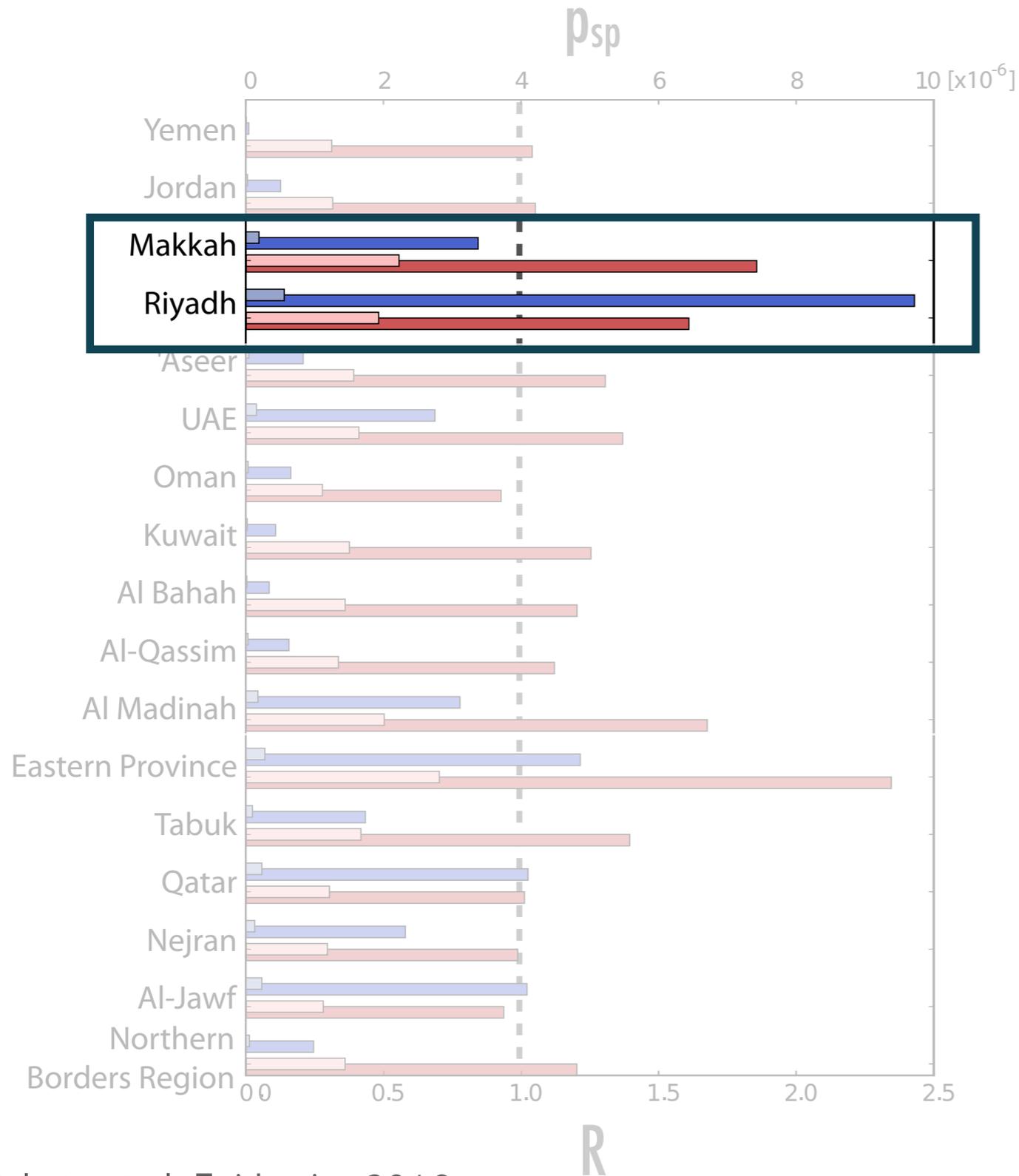
acute epidemics in calves in Spring: Wernery et al 2015



R

baseline → Spring 2014  
 ~3-fold increase

# geo-temporal variation

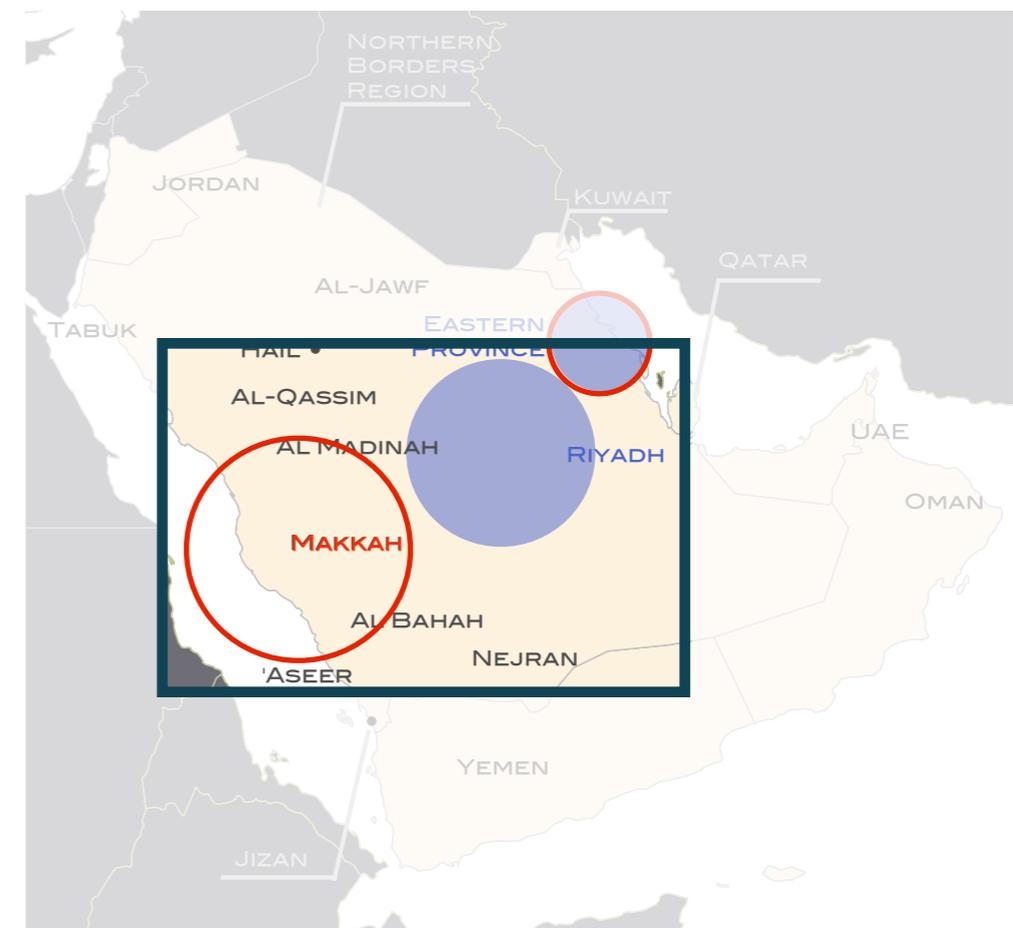


Poletto et al, Epidemics 2016

$p_{sp}$

baseline → Spring 2014  
 ~6/week → ~108/week  
 ~17-fold increase

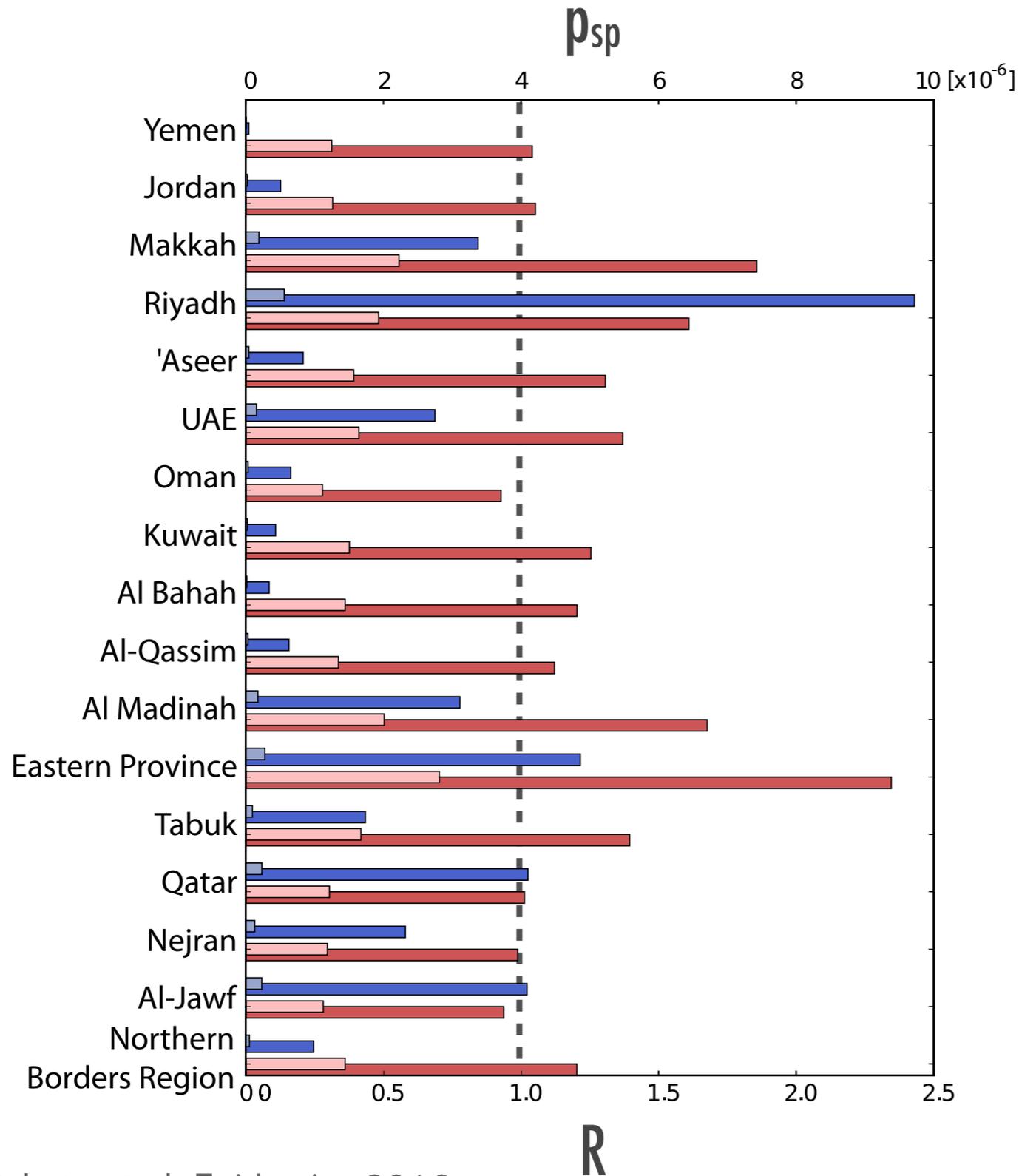
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$R$

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# geographical variation



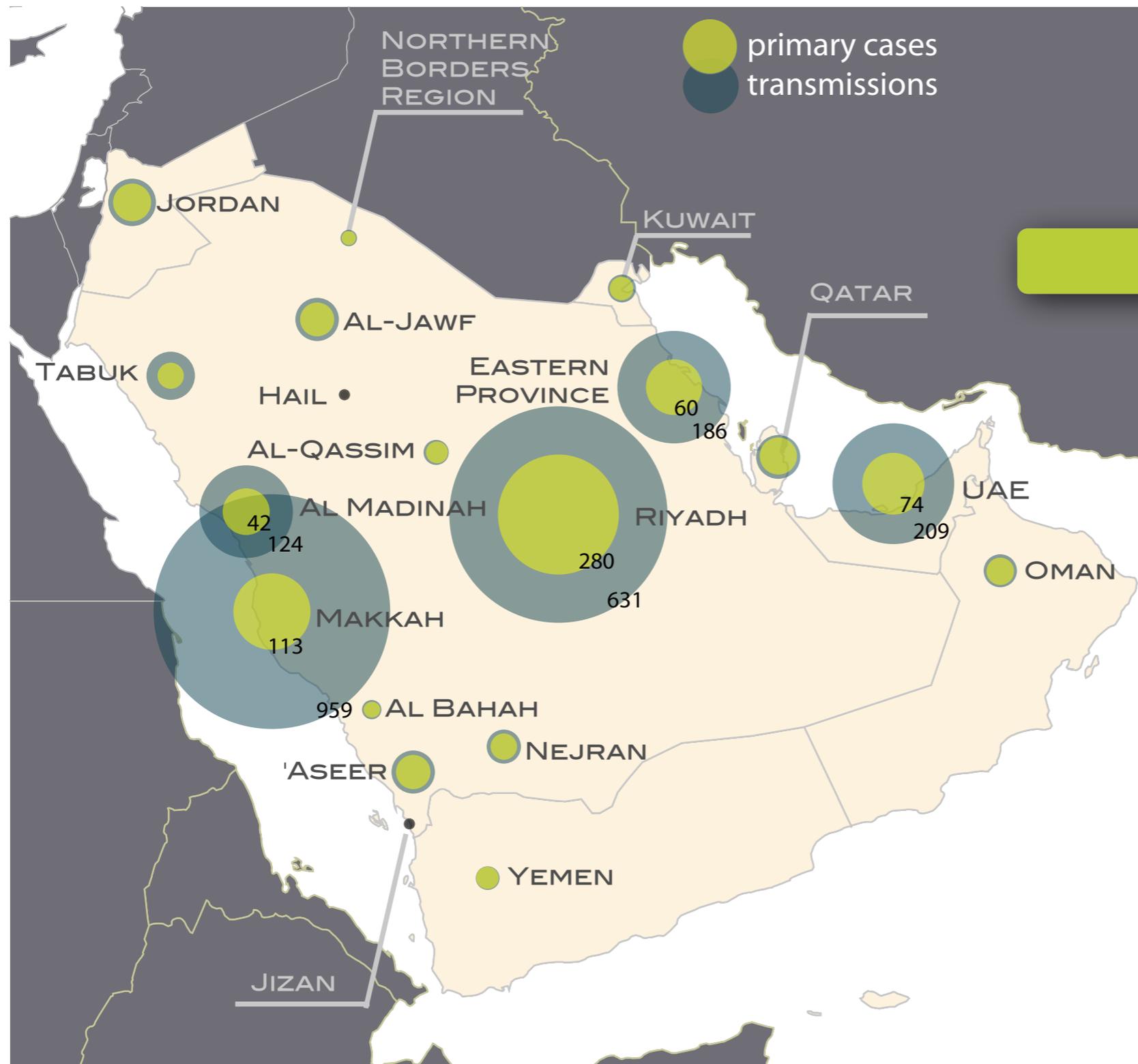
$p_{sp}$

baseline → Spring 2014  
coefficient of variation: **0.99**

$R$

baseline → Spring 2014  
coefficient of variation: **0.20**

# sporadic vs. human-transmission cases



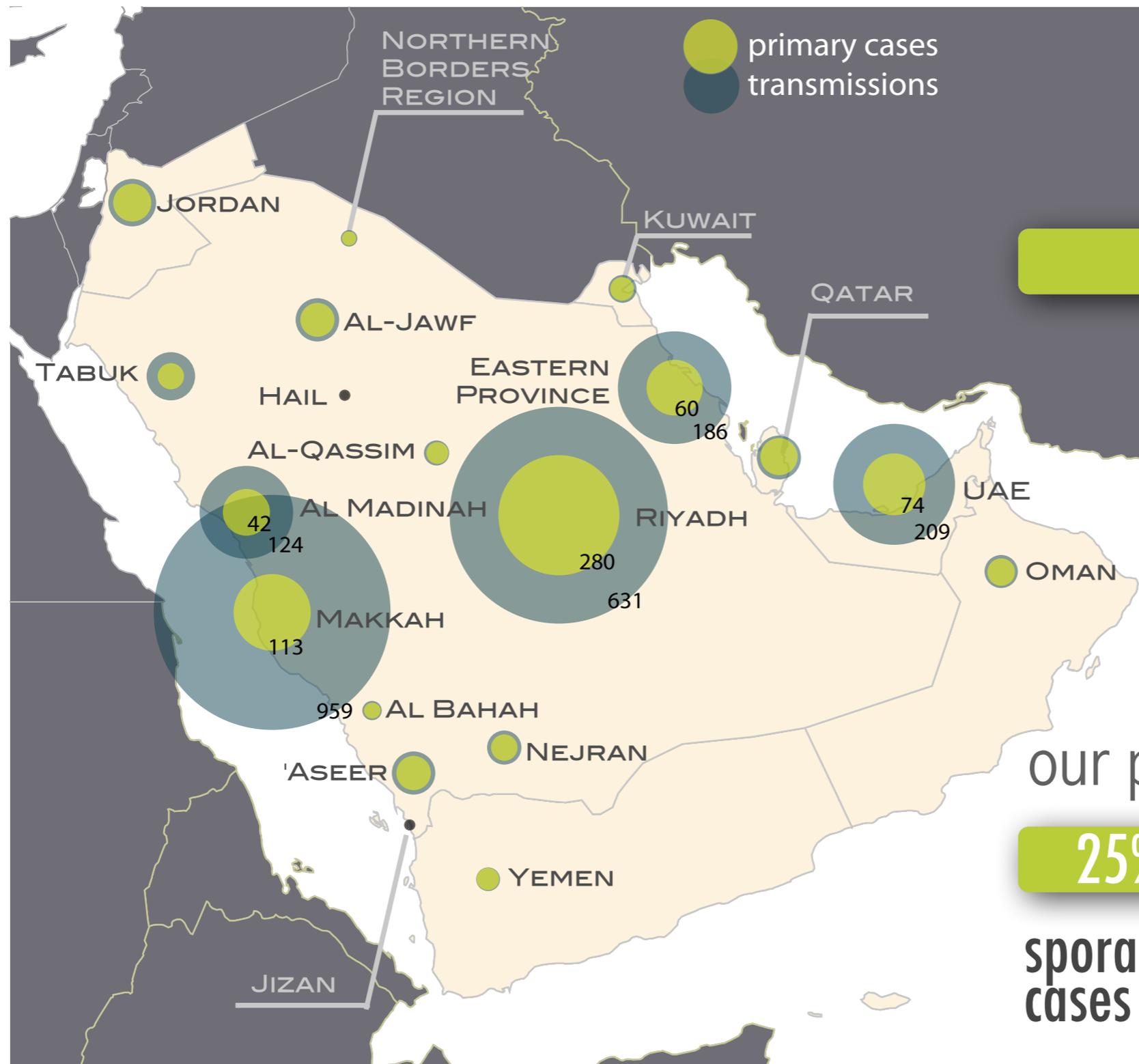
WHO:



sporadic cases

transmissions

# sporadic vs. human-transmission cases



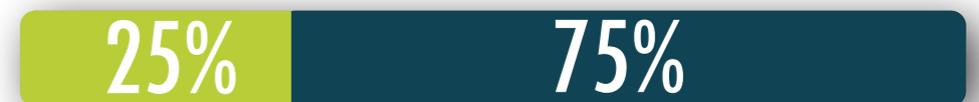
WHO:



sporadic cases

transmissions

our predictions:

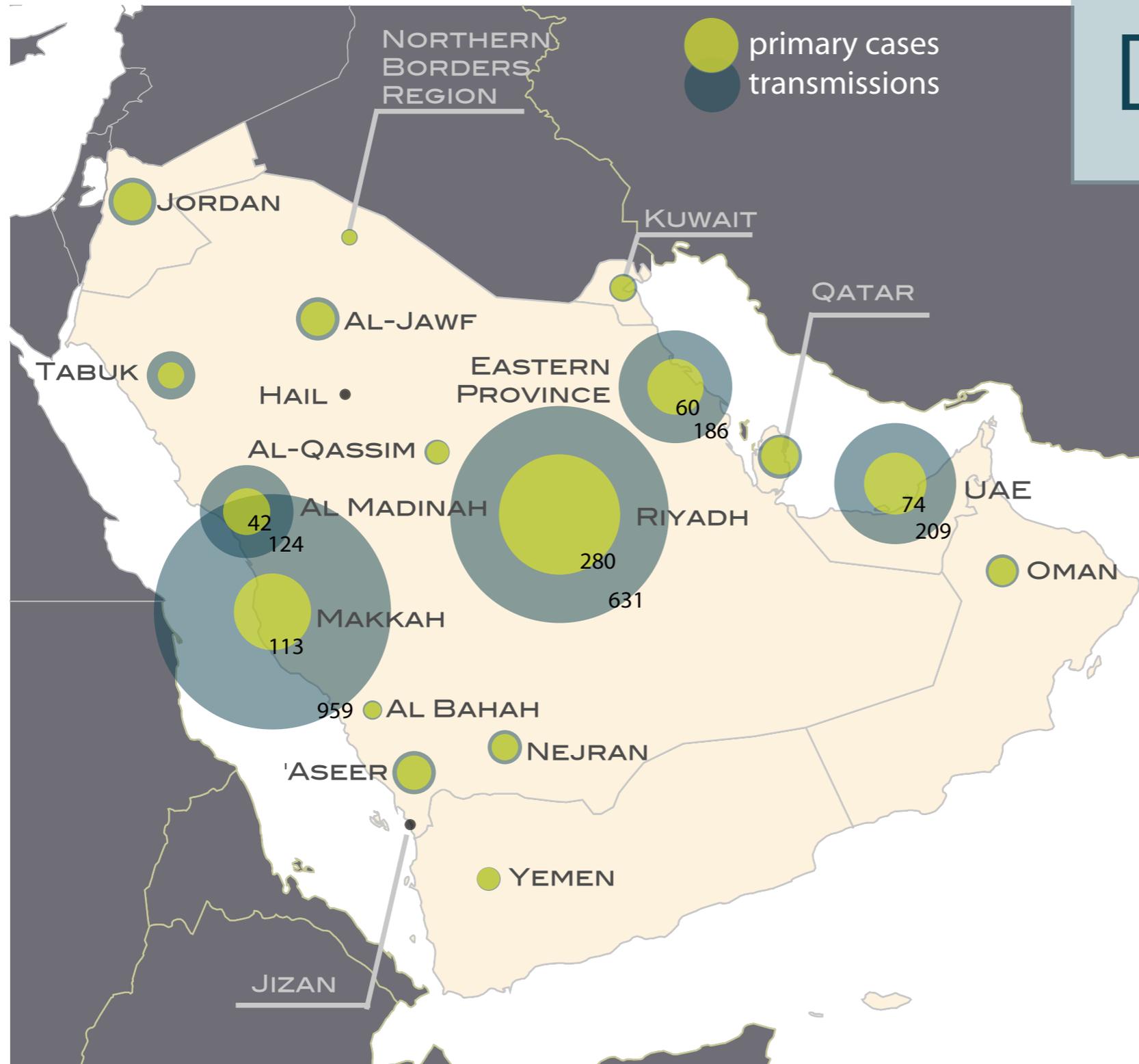


sporadic cases

transmissions

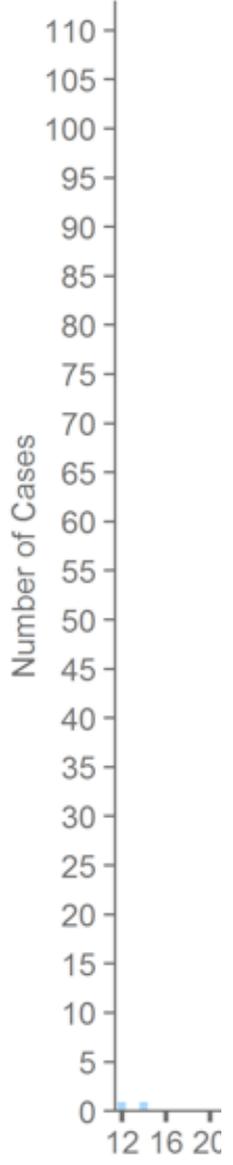
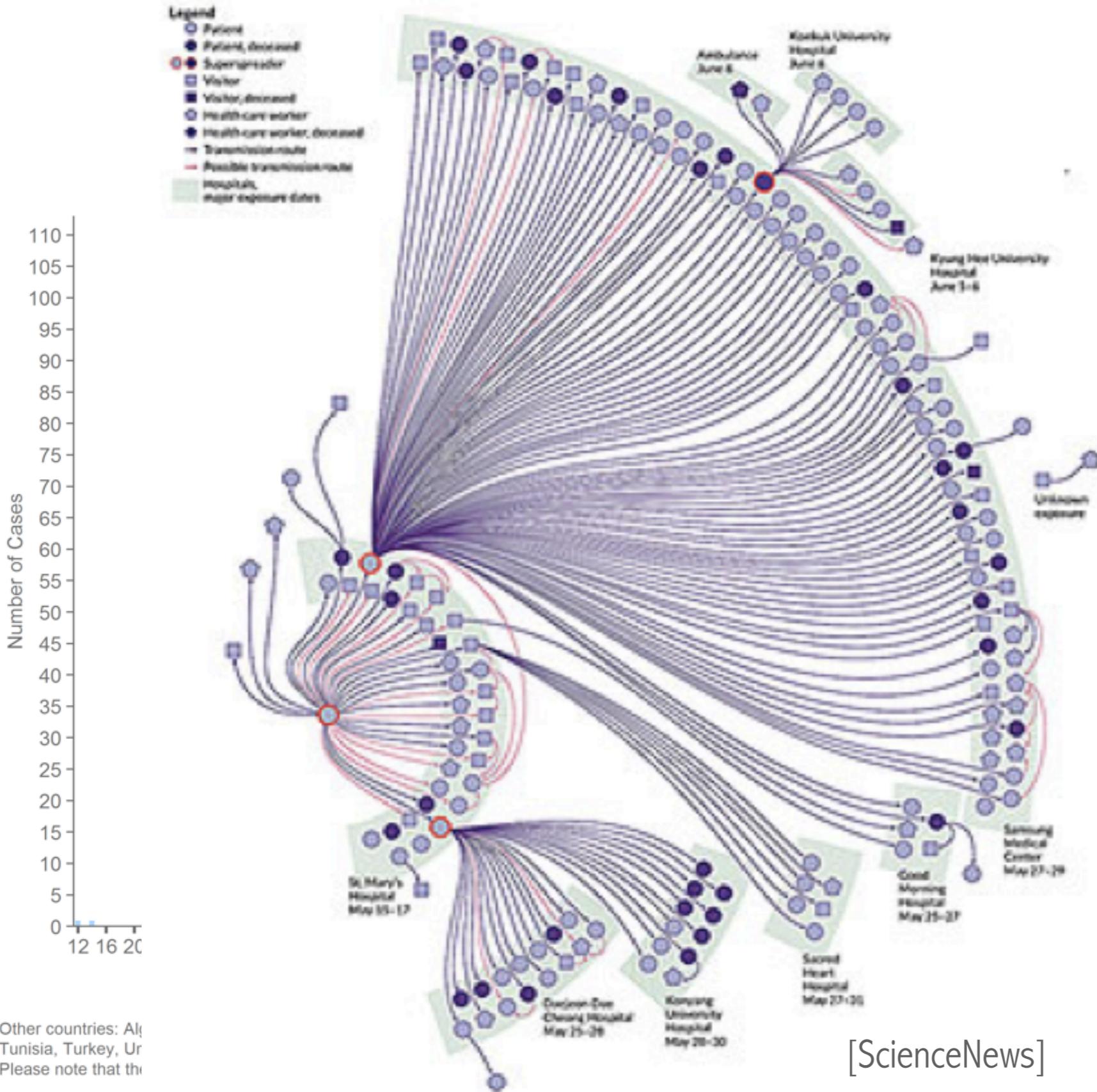
# underascertainment

4-fold underascertainment  
[2-6]

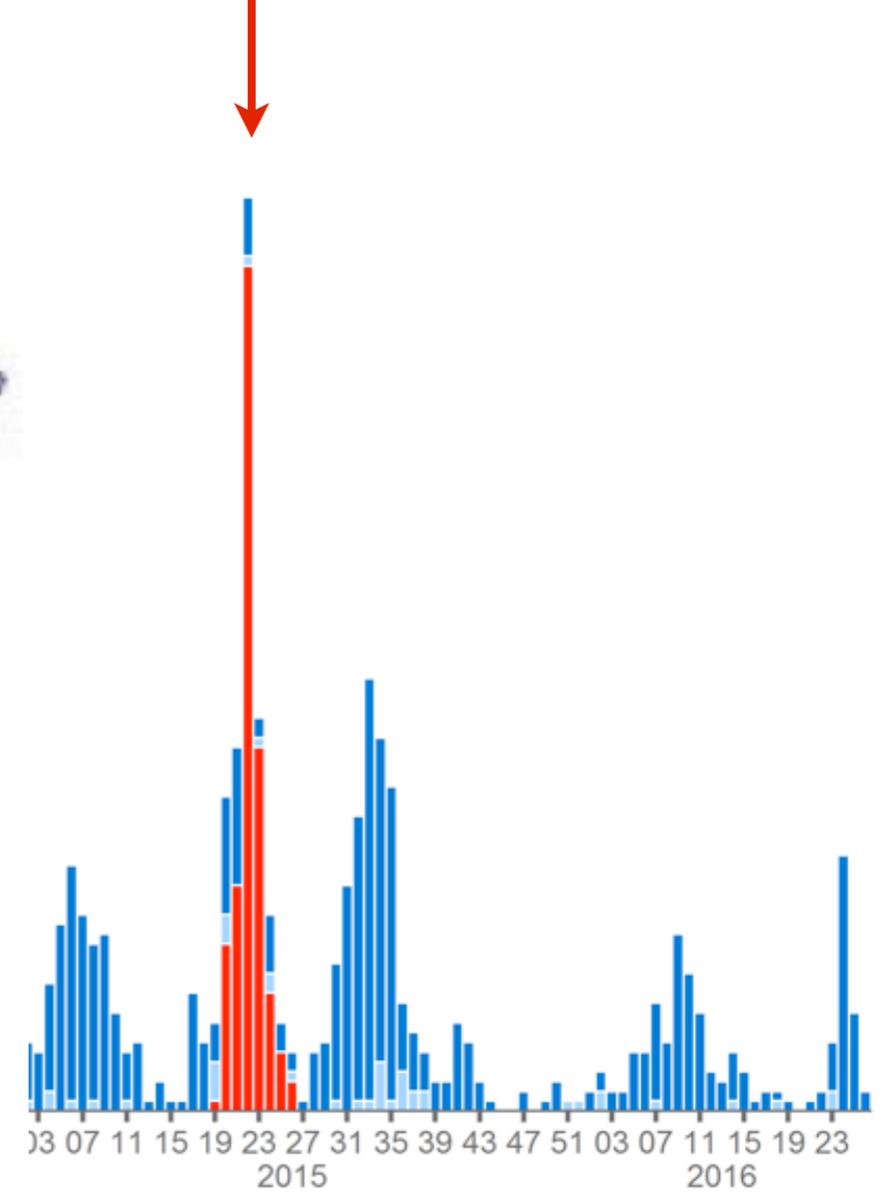


see nationwide seroprevalence investigation in SA: Muller et al 2015

# Summer 2015



South Korea



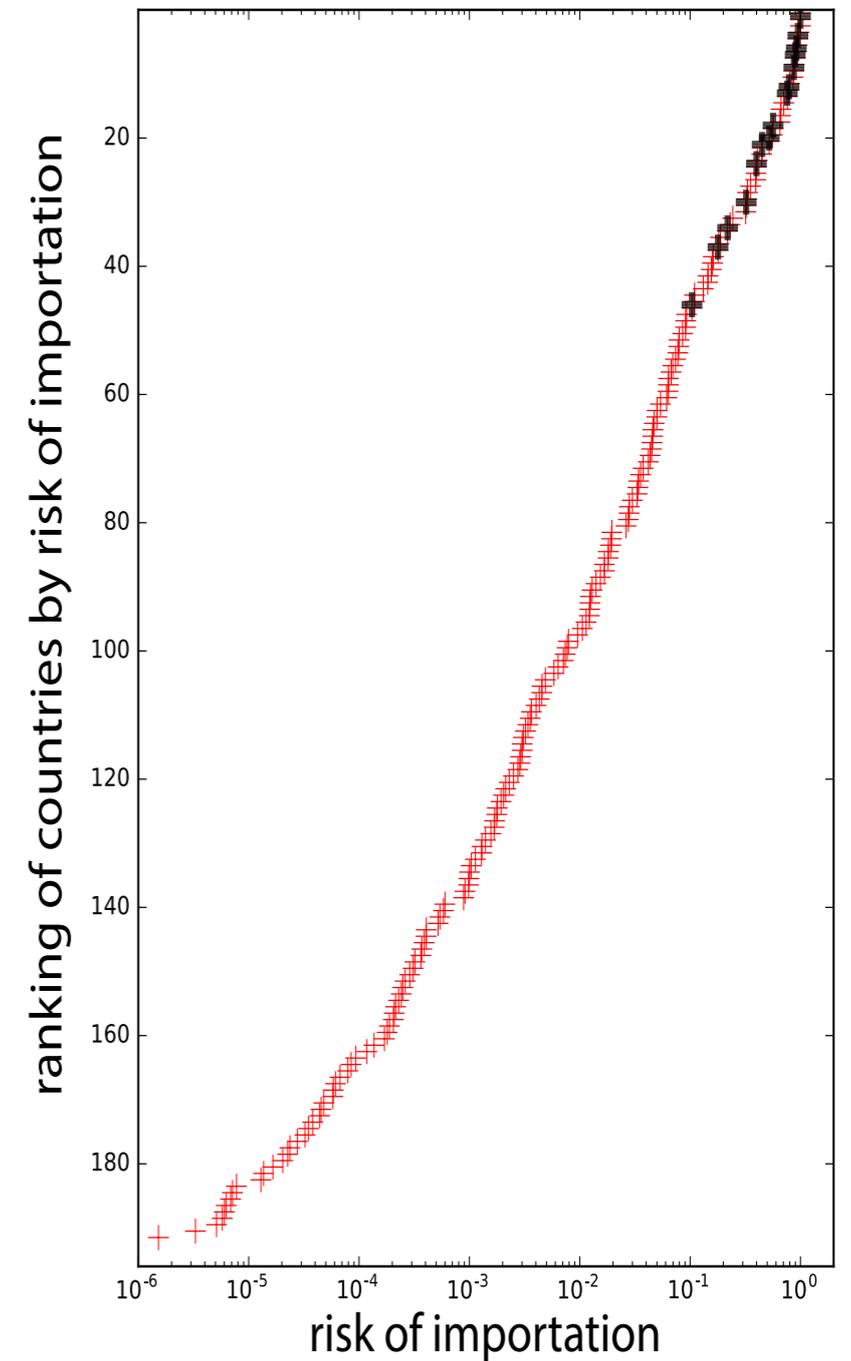
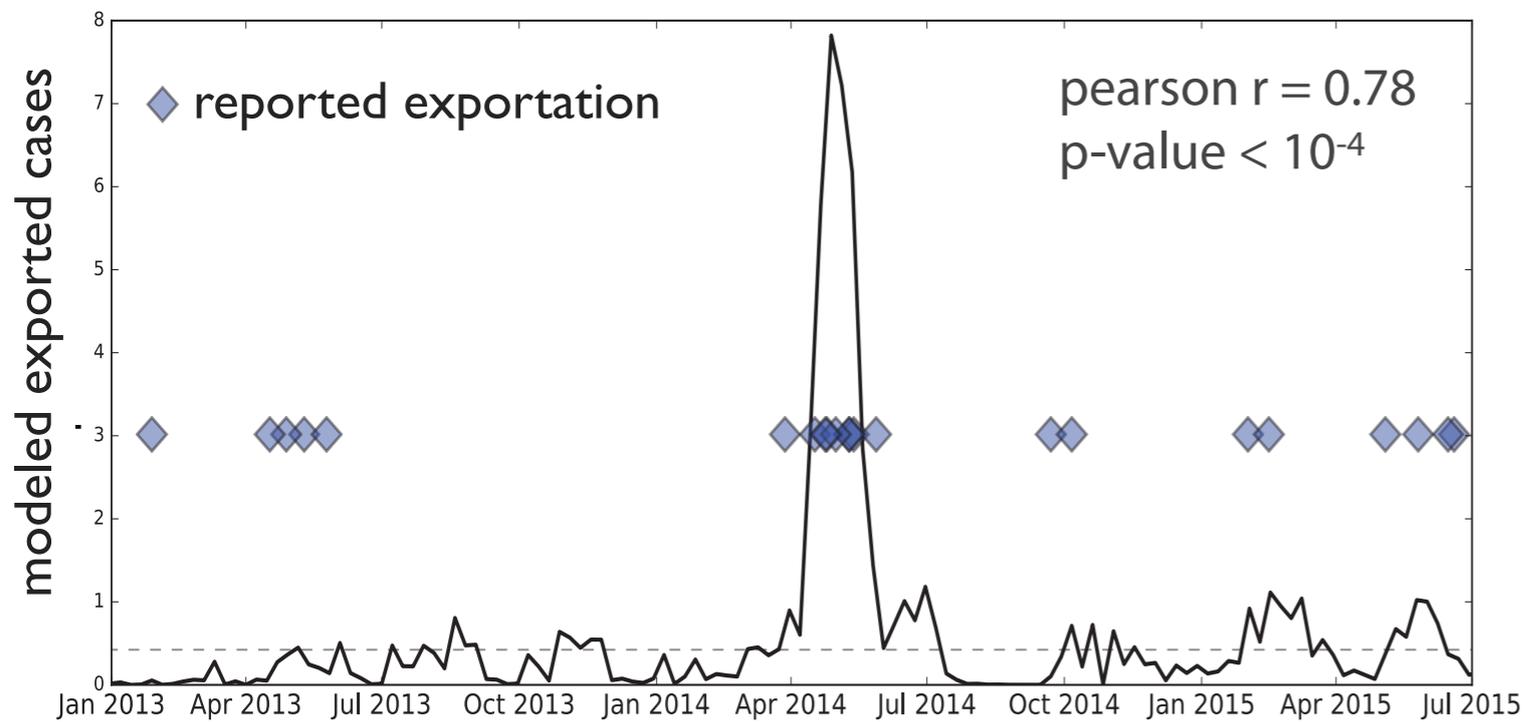
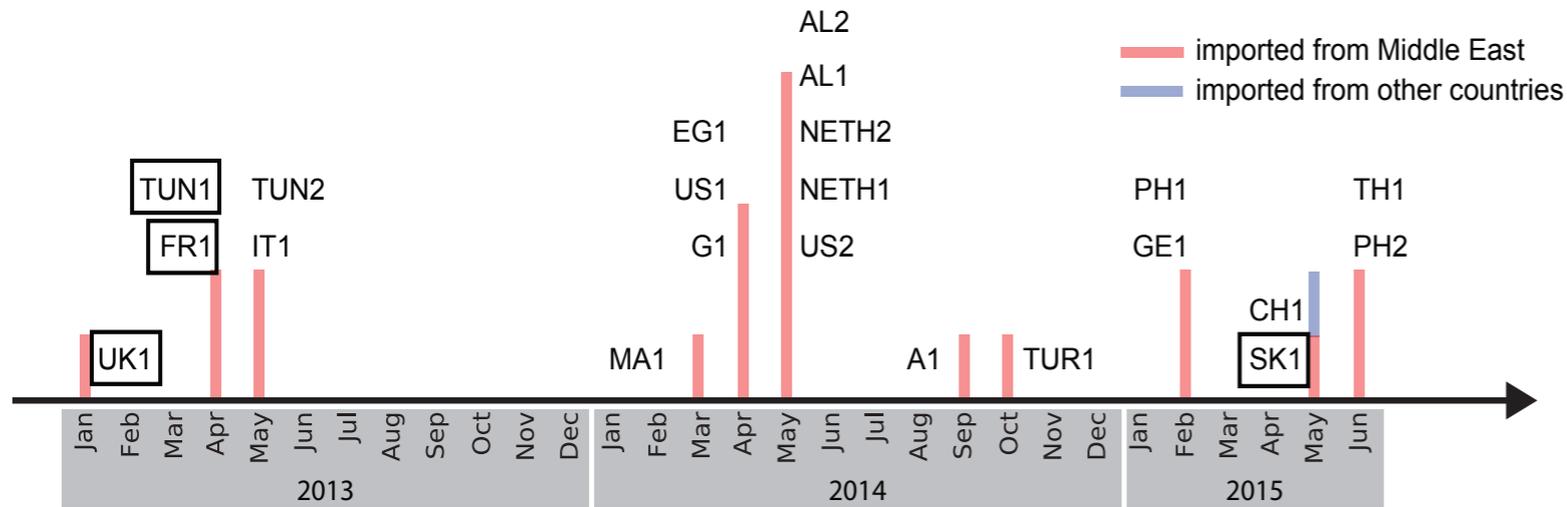
Other countries: Al  
Tunisia, Turkey, Ur  
Please note that th

[ScienceNews]

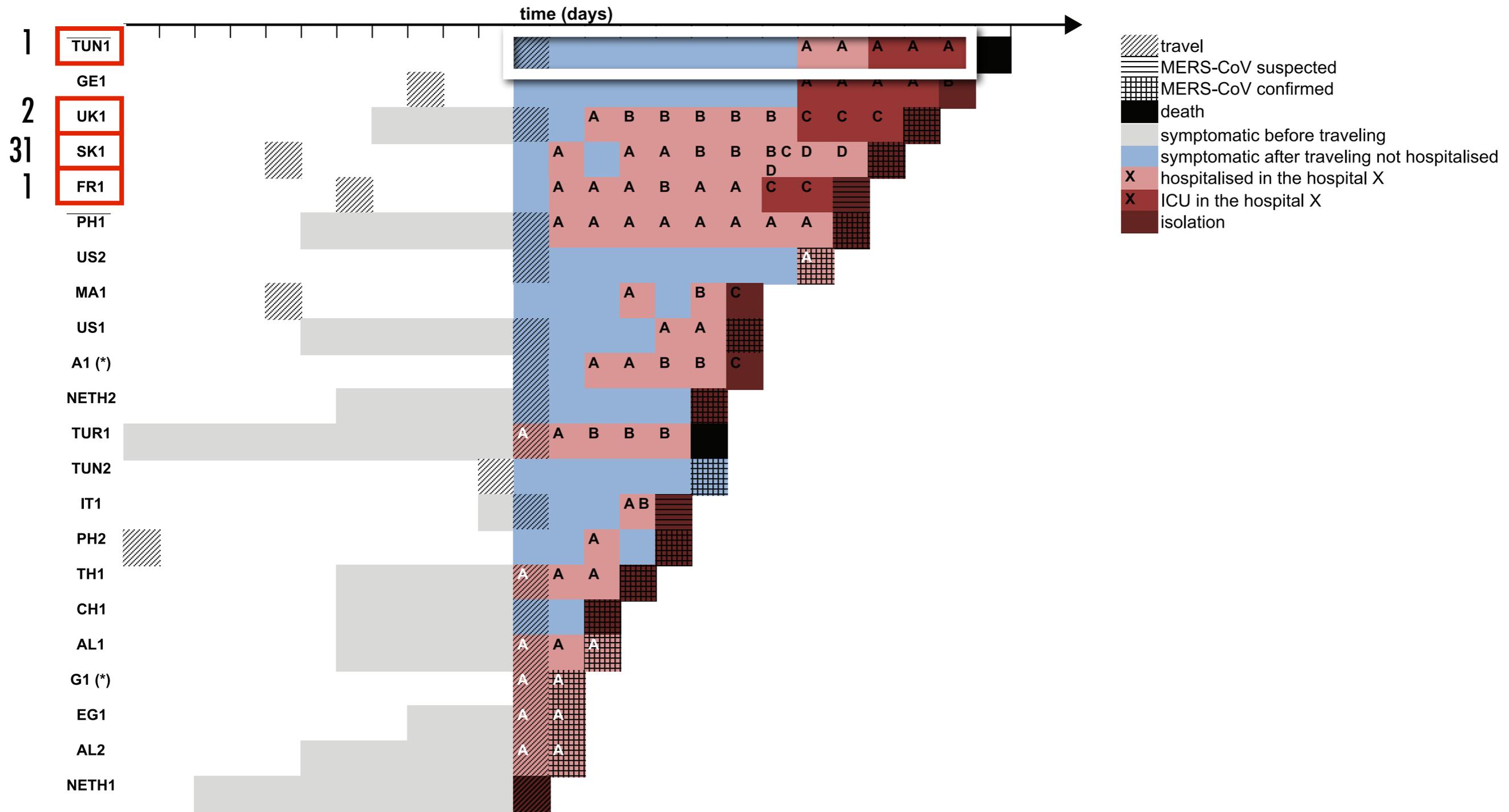
n, Philippines, Qatar, Thailand,



# 22 importations (2012-2015)

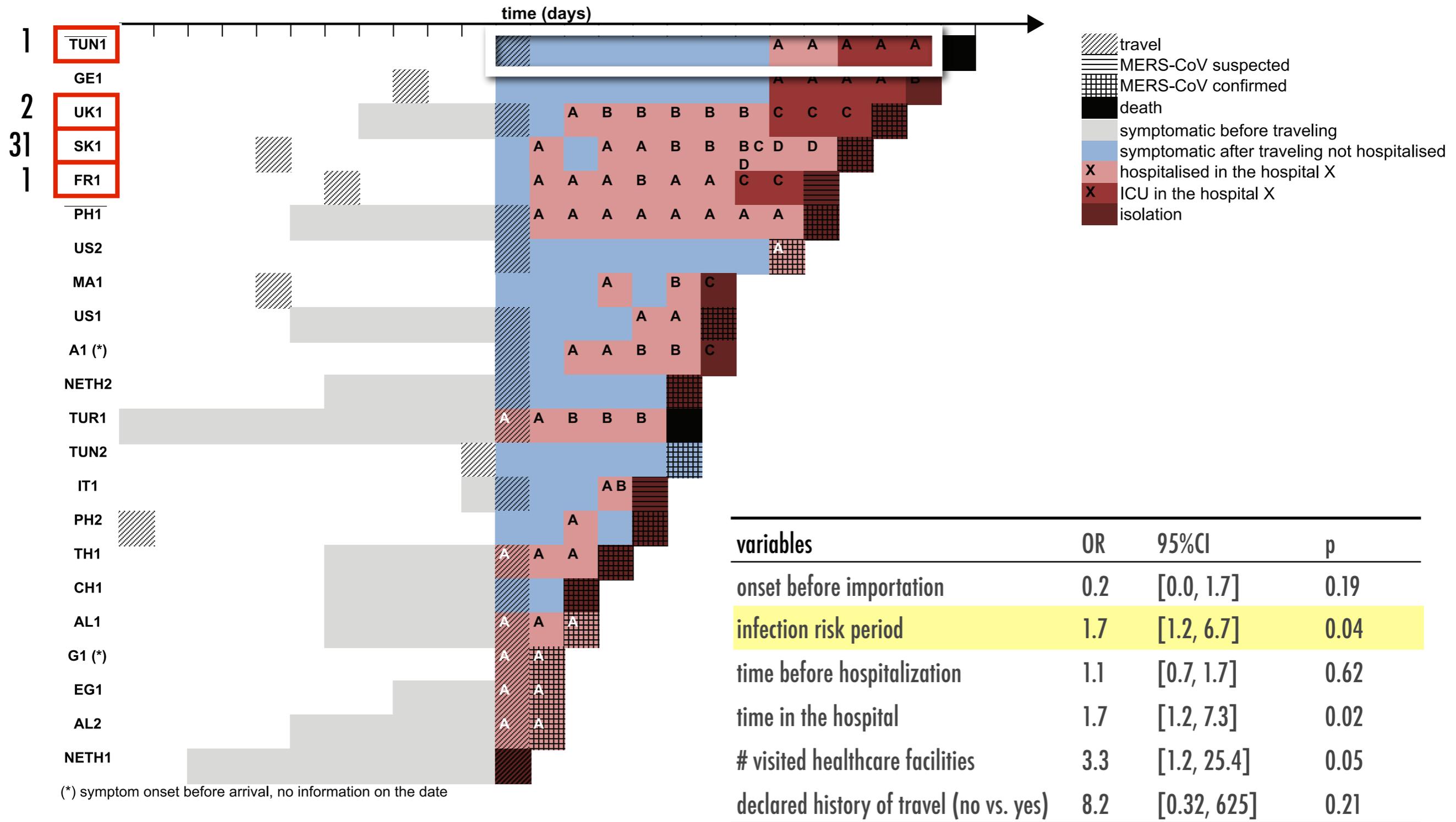


# imported case history

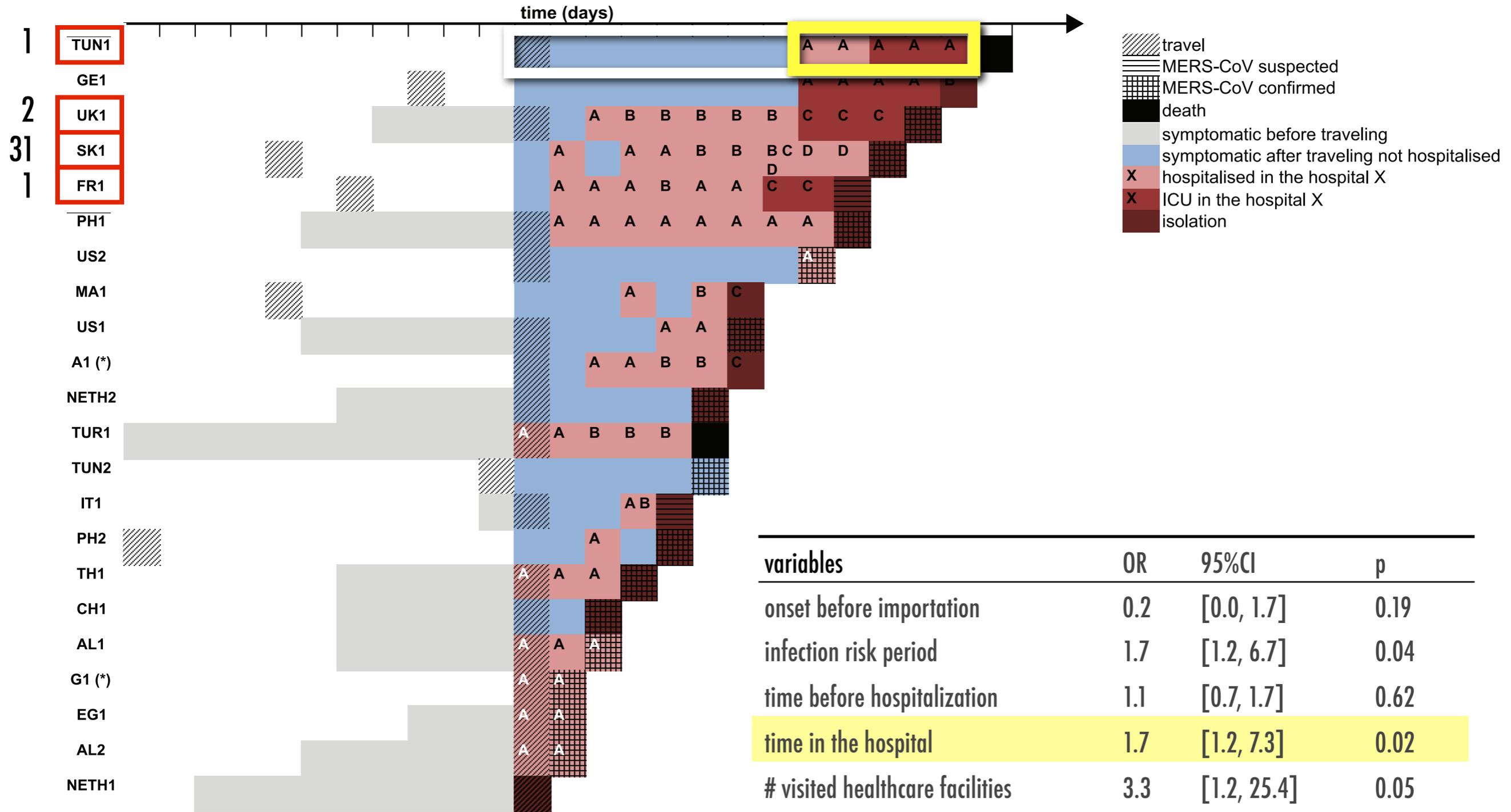


(\*) symptom onset before arrival, no information on the date

# risk of onward transmission



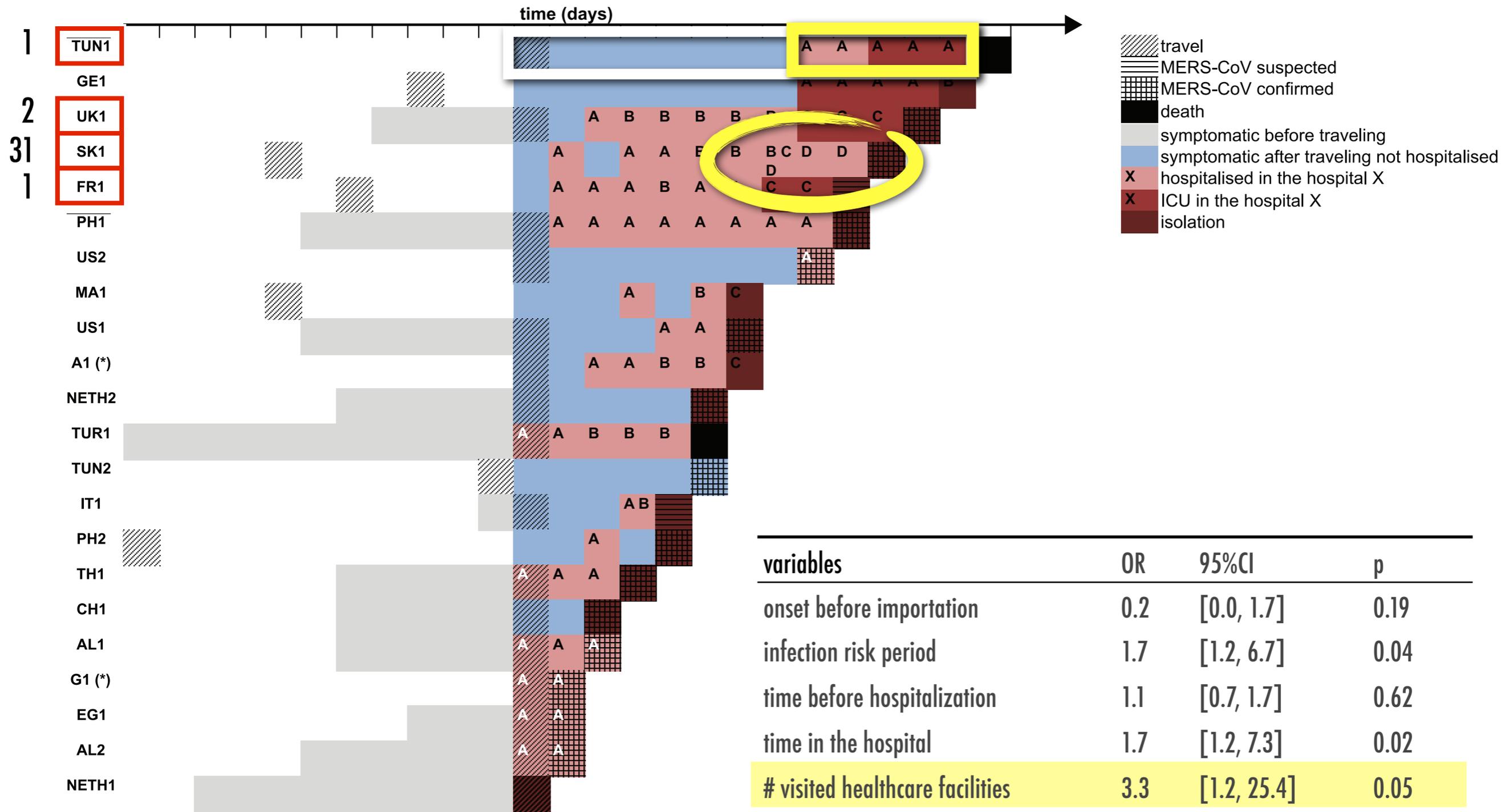
# hospitalization period



(\*) symptom onset before arrival, no information on the date

variables	OR	95%CI	p
onset before importation	0.2	[0.0, 1.7]	0.19
infection risk period	1.7	[1.2, 6.7]	0.04
time before hospitalization	1.1	[0.7, 1.7]	0.62
time in the hospital	1.7	[1.2, 7.3]	0.02
# visited healthcare facilities	3.3	[1.2, 25.4]	0.05
declared history of travel (no vs. yes)	8.2	[0.32, 625]	0.21

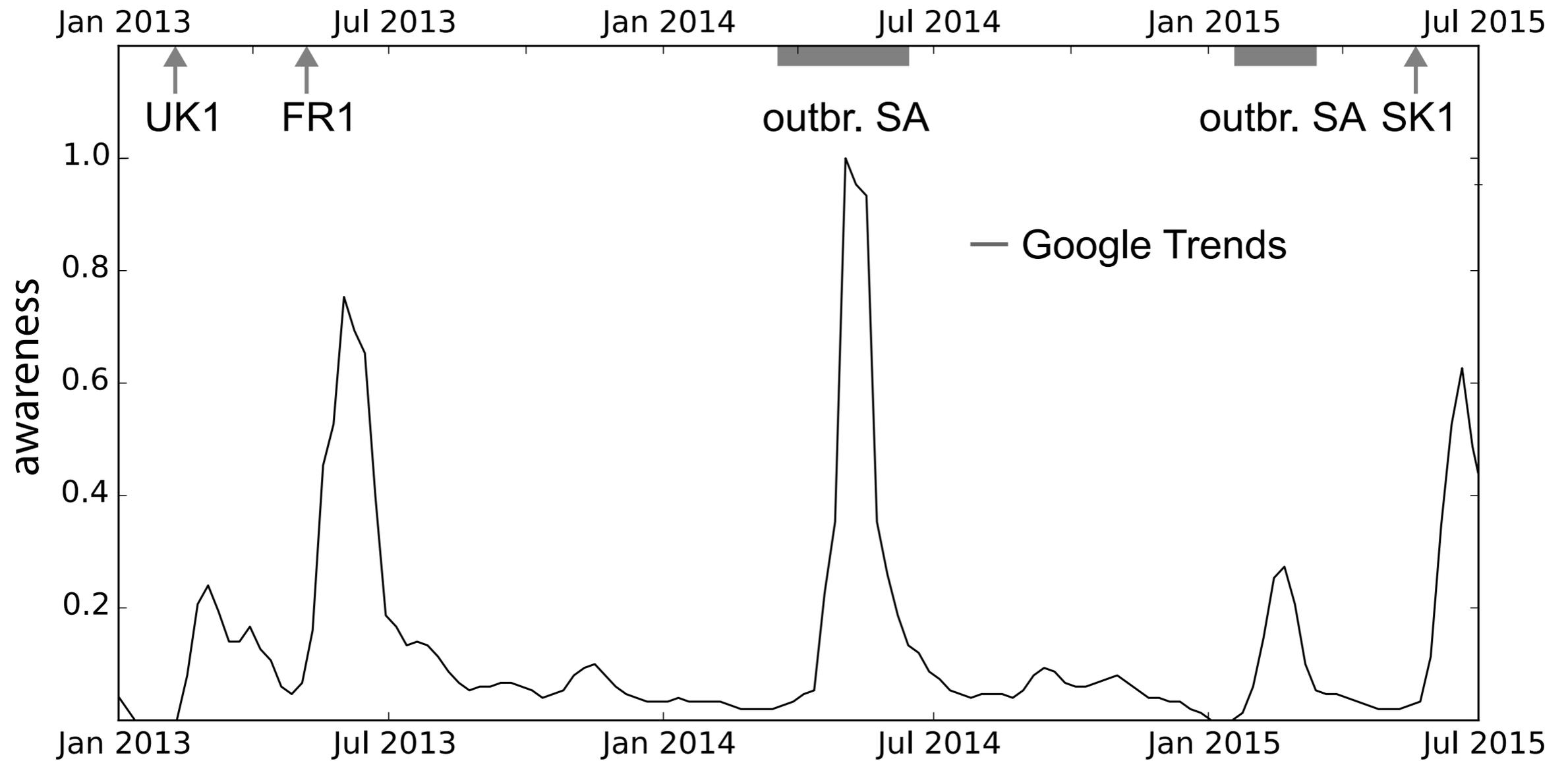
# # healthcare centers



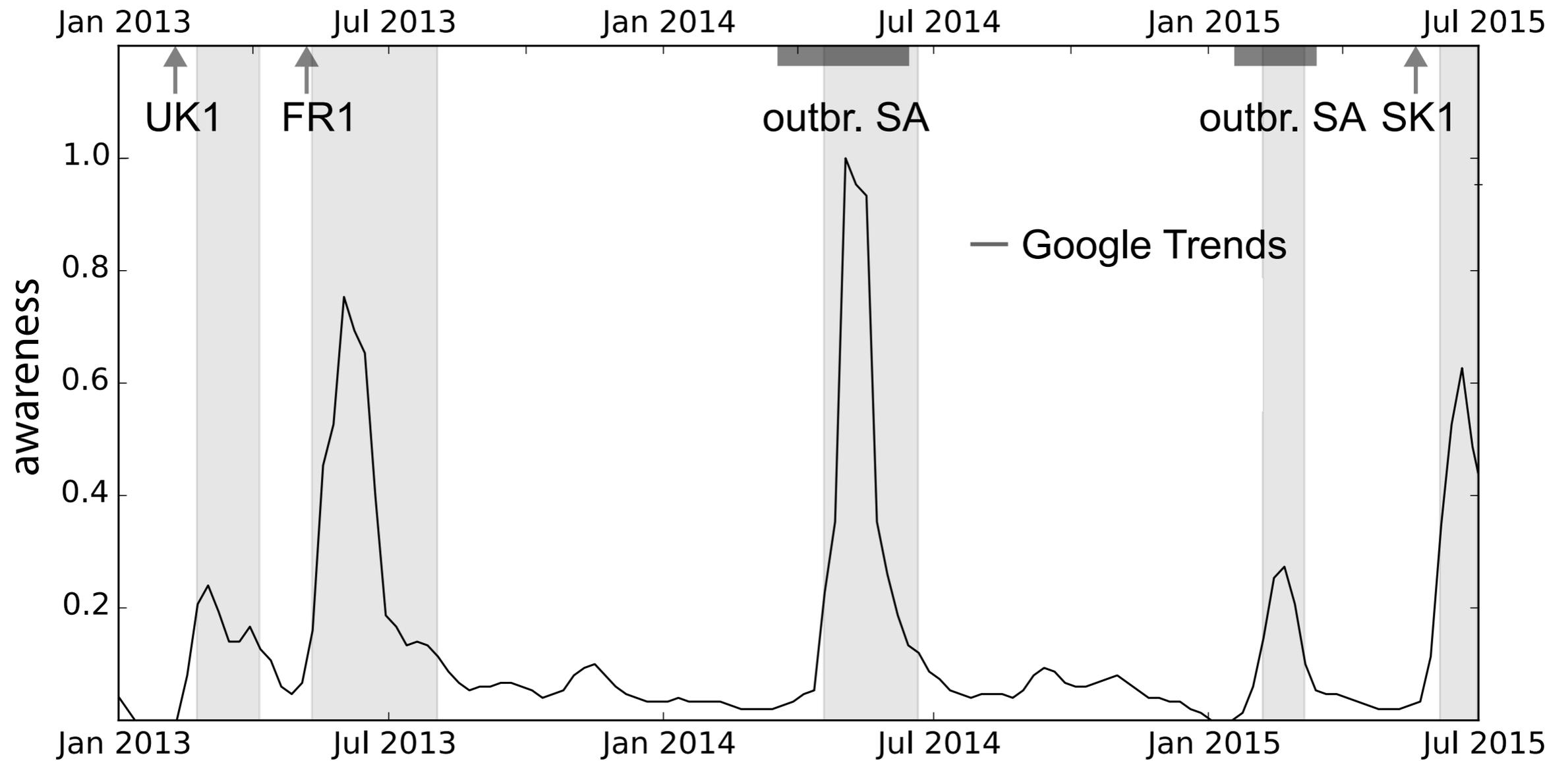
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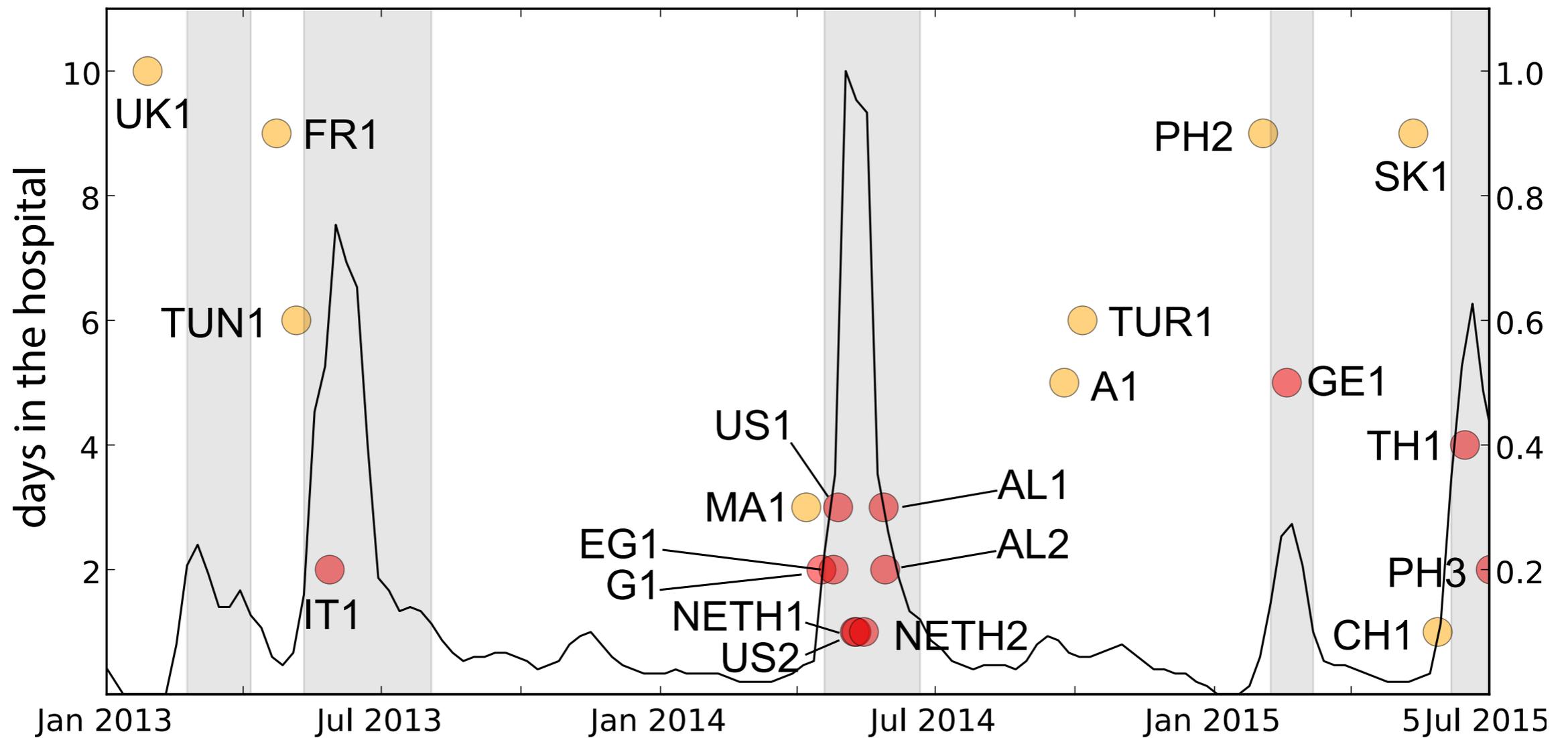
# awareness



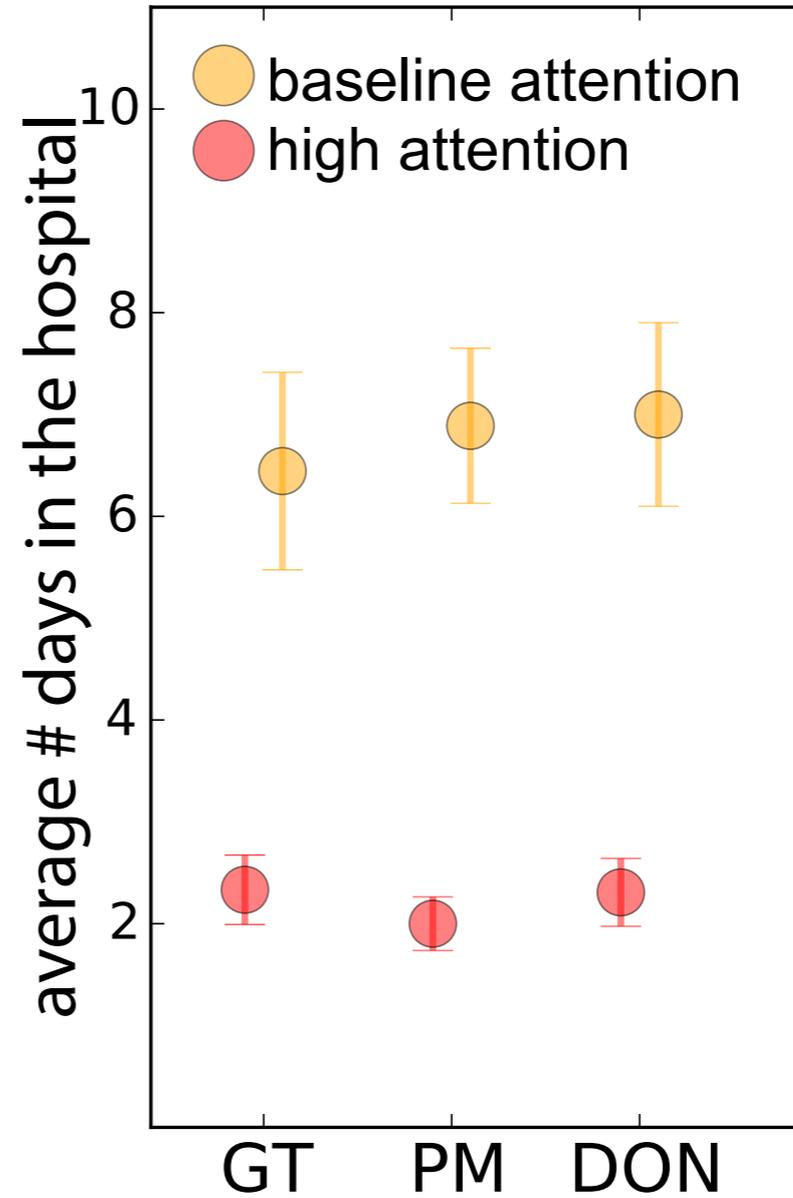
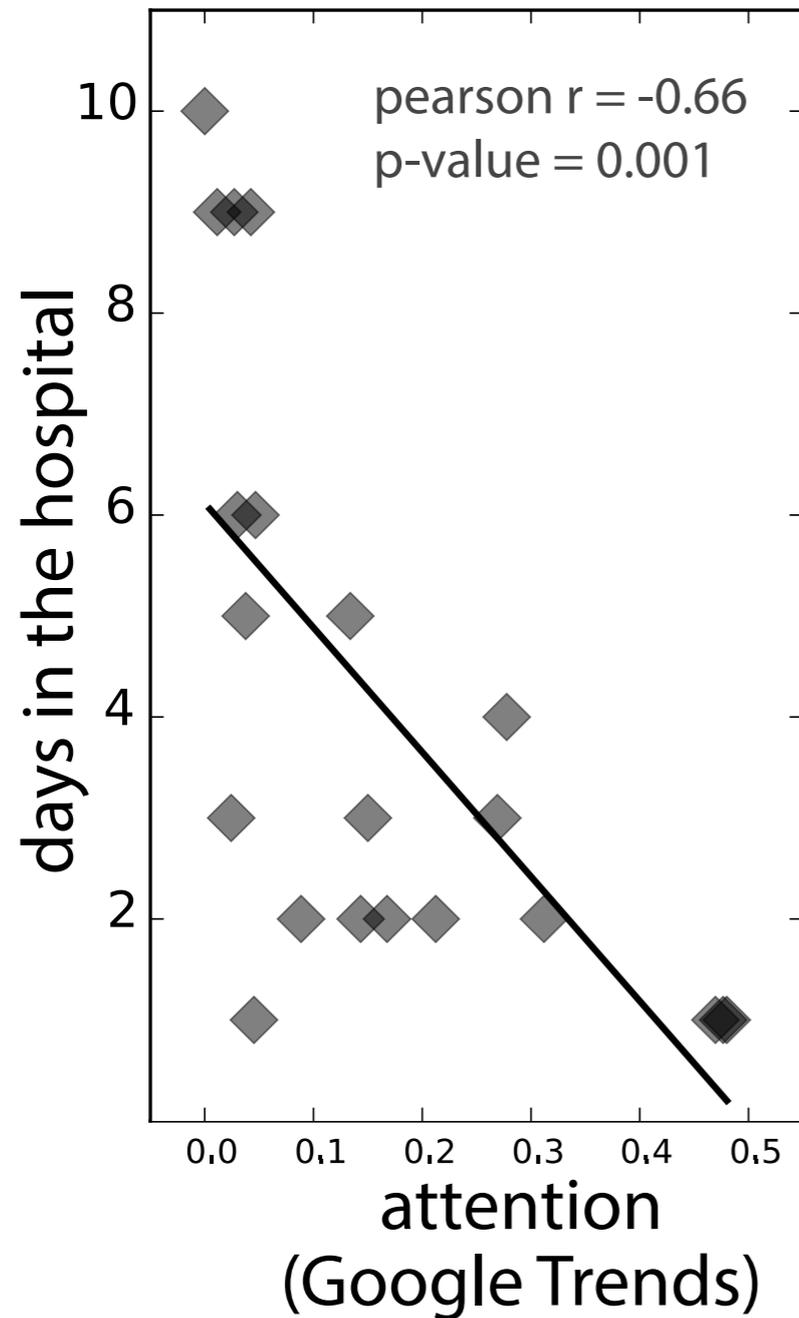
# awareness



# awareness vs response



# awareness vs response



days in the community vs. GT:  
 $r=0.22$ ,  $p=0.32$

# acknowledgments



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Davide Colombi

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Poletto et al. Eurosurveillance (2014)

Poletto et al. Epidemics (2016)

doi:10.1016/j.epidem.2015.12.001

Poletto et al. under revision



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