## References

1 Buelow E, Gonzalez TB, Versluis D et al. Effects of selective digestive decontamination (SDD) on the gut resistome. J Antimicrob Chemother 2014: 69: 2215-23.

2 Woodford N, Johnson AP, Morrison D et al. Vancomycin-dependent enterococci in the United Kingdom. J Antimicrob Chemother 1994; 33: 1066.

3 Tambyah PA, Marx JA, Maki DG. Nosocomial infection with vancomycindependent enterococci. Emerg Infect Dis 2004; 10: 1277-81.

4 Kirkpatrick BD, Harrington SM, Smith D et al. An outbreak of vancomycindependent Enterococcus faecium in a bone marrow transplant unit. Clin Infect Dis 1999; 29: 1268-73.

5 Millan AS, Depardieu F, Godreuil S et al. VanB-type Enterococcus faecium clinical isolate successively inducibly resistant to, dependent on, and constitutively resistant to vancomycin. Antimicrob Agents Chemother 2009: **53**: 1974-82.

6 de la Cal MA, Cerdá E, van Saene HK et al. Effectiveness and safety of enteral vancomycin to control endemicity of methicillin-resistant Staphylococcus aureus in a medical/surgical intensive care unit. J Hosp Infect 2004; 56: 175-83.

7 Daneman N, Sarwar S, Fowler RA et al. Effect of selective decontamination on antimicrobial resistance in intensive care units: a systematic review and meta-analysis. Lancet Infect Dis 2013; 13: 328-41.

8 Houben AJM, Oostdijk EAN, van der Voort PHJ et al. Selective decontamination of the oropharynx and the digestive tract, and antimicrobial resistance: a 4 year ecological study in 38 intensive care units in the Netherlands. J Antimicrob Chemother 2014; 69: 797-804.

J Antimicrob Chemother 2014 doi:10.1093/jac/dku292 Advance Access publication 8 September 2014

# Comment on: Measurement units for antibiotic consumption in outpatients

#### Samuel Coenen<sup>1,2\*</sup>, Robin Bruvndonckx<sup>3</sup>, Niel Hens<sup>3,4</sup>, Marc Aerts<sup>3</sup> and Herman Goossens<sup>1</sup>

<sup>1</sup>Laboratory of Medical Microbiology, Vaccine & Infectious Disease Institute (VAXINFECTIO), University of Antwerp, Antwerp, Belgium; <sup>2</sup>Centre for General Practice, Vaccine & Infectious Disease Institute (VAXINFECTIO), University of Antwerp, Antwerp, Belgium; <sup>3</sup>Interuniversity Institute for Biostatistics and statistical Bioinformatics (I-BIOSTAT), University of Hasselt, Diepenbeek, Belaium; <sup>4</sup>Centre for Health Economic Research and Modelling Infectious Diseases (CHERMID), Vaccine & Infectious Disease Institute (VAXINFECTIO), University of Antwerp, Antwerp, Belgium

\*Corresponding author. Tel: +32 3 265 25 25; Fax: +32 3 265 25 29; E-mail: samuel.coenen@uantwerpen.be

Keywords: antibiotic consumption, ambulatory care, dose change, measurement unit

Sir.

We are grateful for the interest of Čižman<sup>1</sup> in our work<sup>2</sup> on appropriate measurement units for outpatient antibiotic consumption. Based on data for Slovenia and referring to the correlation between antibiotic consumption and resistance, he argues that for an international benchmarking of antibiotic use in outpatients, the number of defined daily doses (DDDs) per 1000 inhabitants per day (DID) is a better measurement unit than the number of packages per 1000 inhabitants per day (PID), and that if substantial changes in the number of DDDs per package occur over time, additional measurement units should be used, such as PID and the number of prescriptions per 1000 inhabitants per year, to identify trends in national prescribing. The latter statement is completely in line with our recommendation to use a similar combination of measurement units or to exercise caution when interpreting trends based only on DDDs when such changes occur or are unknown.<sup>2</sup>

This recommendation was first based on the observation in Belgium that outpatient antibiotic consumption in terms of DID had not decreased since the start of the national public antibiotic awareness campaign, whereas we observed a substantial decrease in PID due to the less frequent treatment of fewer individuals. These contrasting trends coincided with a decrease in the proportion of pneumococci resistant to penicillins, tetracyclines and macrolides and are explained by a substantial increase in the number of DDDs per package for the most commonly used antibiotics.<sup>3</sup> In Slovenia, the latter is not the case based on the data provided by Čižman.<sup>1</sup> Meanwhile, we have shown that the number of DDDs per package increased for most commonly used antibiotics in Europe (31 countries), resulting in contrasting trends depending on whether DID or PID is used as the measurement unit and corroborating our recommendation to adopt PID to monitor outpatient antibiotic use in Europe.<sup>4</sup> Based on that study, Figure 1 shows the estimated linear trends in the number of DDDs per package for total outpatient antibiotic consumption in Europe, Belgium and Slovenia. These data explain why consumption trends expressed in DID or PID are contrasting for Belgium and Europe, and similar for Slovenia.





We are currently investigating whether the description of outpatient antibiotic consumption in DID and PID can be improved by including change points in the model,<sup>5</sup> e.g. to assess the impact of the European Antibiotic Awareness Day,<sup>6,7</sup> and what measurement unit correlates best with antimicrobial resistance in Europe. Expressing the consumption in PID and including a change point provided the best fitting model when assessing the correlation between macrolide, lincosamide, streptogramin B and tetracycline consumption and the prevalence of macrolide-resistant *Streptococcus pyogenes* in Belgium.<sup>8</sup>

### **Transparency declarations**

None to declare.

## References

**1** Čižman M. Measurement units for antibiotic consumption in outpatients. *J Antimicrob Chemother* 2014; **69**: 2877–8.

**2** Coenen S, Gielen B, Blommaert A *et al*. Appropriate international measures for outpatient antibiotic prescribing and consumption:

recommendations from a national data comparison of different measures. J Antimicrob Chemother 2014; **69**: 529–34.

**3** Goossens H, Coenen S, Costers M *et al*. Achievements of the Belgian Antibiotic Policy Coordination Committee (BAPCOC). *Euro Surveill* 2008; **13**: pii=19036.

**4** Bruyndonckx R, Hens N, Aerts M *et al.* Measuring trends of outpatient antibiotic use in Europe: jointly modelling longitudinal data in defined daily doses and packages. *J Antimicrob Chemother* 2014; **69**: 1981–9.

**5** Minalu G, Aerts M, Coenen S *et al.* Adaptive change-point mixed models applied to data on outpatient tetracycline use in Europe. *Stat Model* 2013; **13**: 253–74.

**6** Coenen S, Costers M, De Corte S *et al*. The first European Antibiotic Awareness Day after a decade of improving outpatient antibiotic use in Belgium. *Acta Clin Belg* 2008; **63**: 296–300.

**7** Earnshaw S, Monnet DL, Duncan B *et al*. European Antibiotic Awareness Day, 2008—the first Europe-wide public information campaign on prudent antibiotic use: methods and survey of activities in participating countries. *Euro Surveill* 2009; **14**: pii=19280.

**8** Van Heirstraeten L, Coenen S, Lammens C *et al.* Antimicrobial drug use and macrolide-resistant *Streptococcus pyogenes*, Belgium. *Emerg Infect Dis* 2012; **18**: 1515–8.