Stellingen behorende bij het proefschrift:

Pharmacokinetic profiling of fosfomycin and nitrofurantoin to optimize the treatment of uncomplicated urinary tract infections

- 1. Treatment failure with fosfomycin is more likely to be PD related (susceptibility of the uropathogen to fosfomycin) rather than PK related. (*This thesis*)
- 2. Adjusting the dose or formulation of nitrofurantoin is insufficient to reduce the risk of achieving sub-therapeutic urinary concentrations. (*This thesis*)
- 3. Clinical effectiveness of antibiotics in the treatment of urinary tract infections may not correlate well with *in vitro* activity, and therefore the clinical validation of PK/PD based dosing regimens should be a continuous process. (*This thesis*)
- 4. For the treatment of uncomplicated urinary tract infections, PK research in healthy volunteers is a correct approach as this population is representative of the real-life patients. (*This thesis*)
- 5. The influence of renal function on the effectiveness of the antibiotic treatment of urinary tract infections is overestimated in guidelines for drug dosage adjustment in clinical practice. (*This thesis*)
- 6. Infections caused by *Klebsiella pneumoniae* should never be treated with fosfomycin. (*Journal of Antimicrobial Chemotherapy 2018; 73: 709-719*)
- 7. To prevent treatment failure and emergence of resistance among uropathogens, fast and affordable diagnostic tests are needed to detect resistant subpopulations before starting fosfomycin treatment. (*Journal of Antimicrobial Chemotherapy 2018; 73: 2380-2387*)
- 8. Dosing regimens of antimicrobial agents should be designed based on their PK/PD characteristics instead of investigating the PK/PD after selecting the clinical dose, as is current practice in the process of drug development. (*Clinical Microbiology and Infection 2015; 21: 881–885*)
- 9. For therapy optimization, it is essential to set urine-specific susceptibility breakpoints for drugs indicated for the treatment of urinary tract infections. (*Clinical Microbiology and Infection 2012; 18: E37-E45*)
- 10. People make better decisions about some kinds of things but worse decisions about other kinds of things, when they have a strong urge to urinate. (*Neurourology and Urodynamics 2011; 30: 183-187*)
- 11. Everything is impossible until it has been done. (Johan Mouton)