

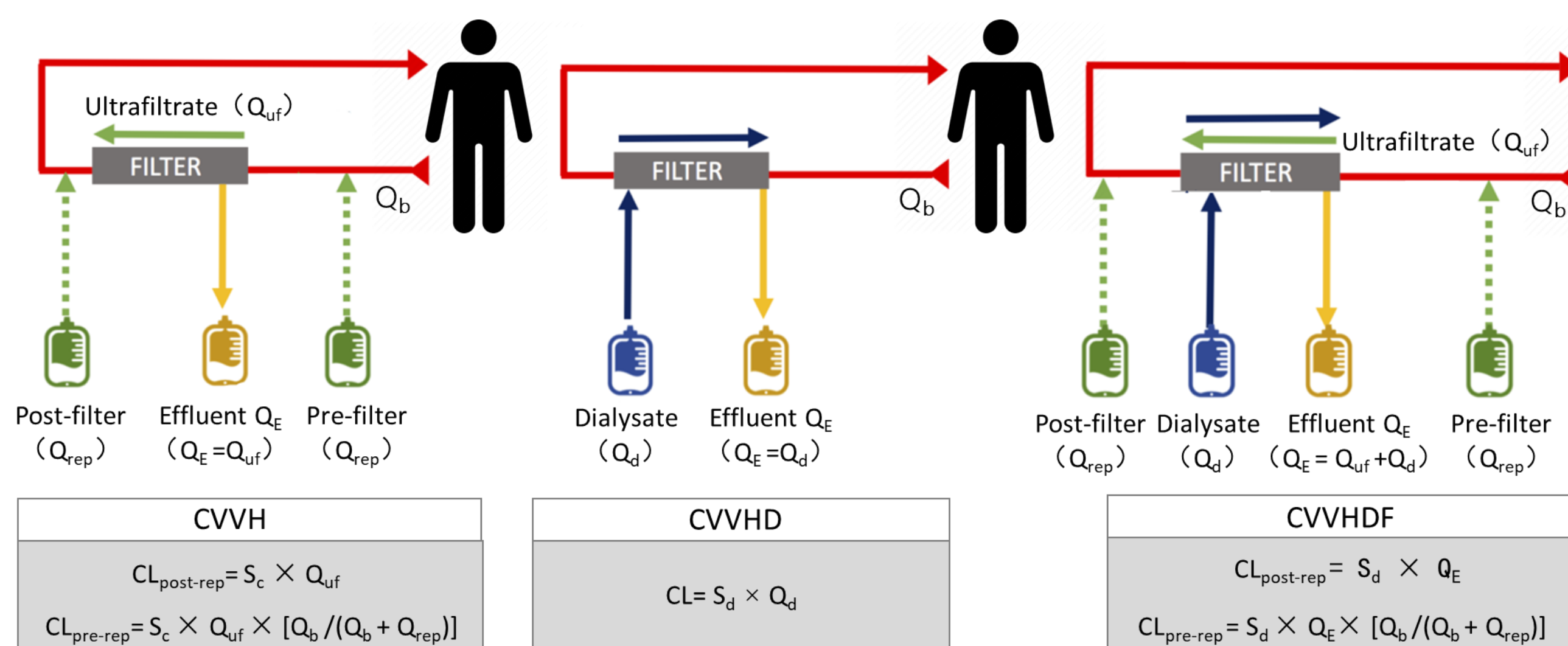
Population pharmacokinetic meta-analysis and dosing recommendation of meropenem in critically ill patients receiving continuous renal replacement therapy

Yaru Peng, Zeneng Cheng, Feifan Xie*

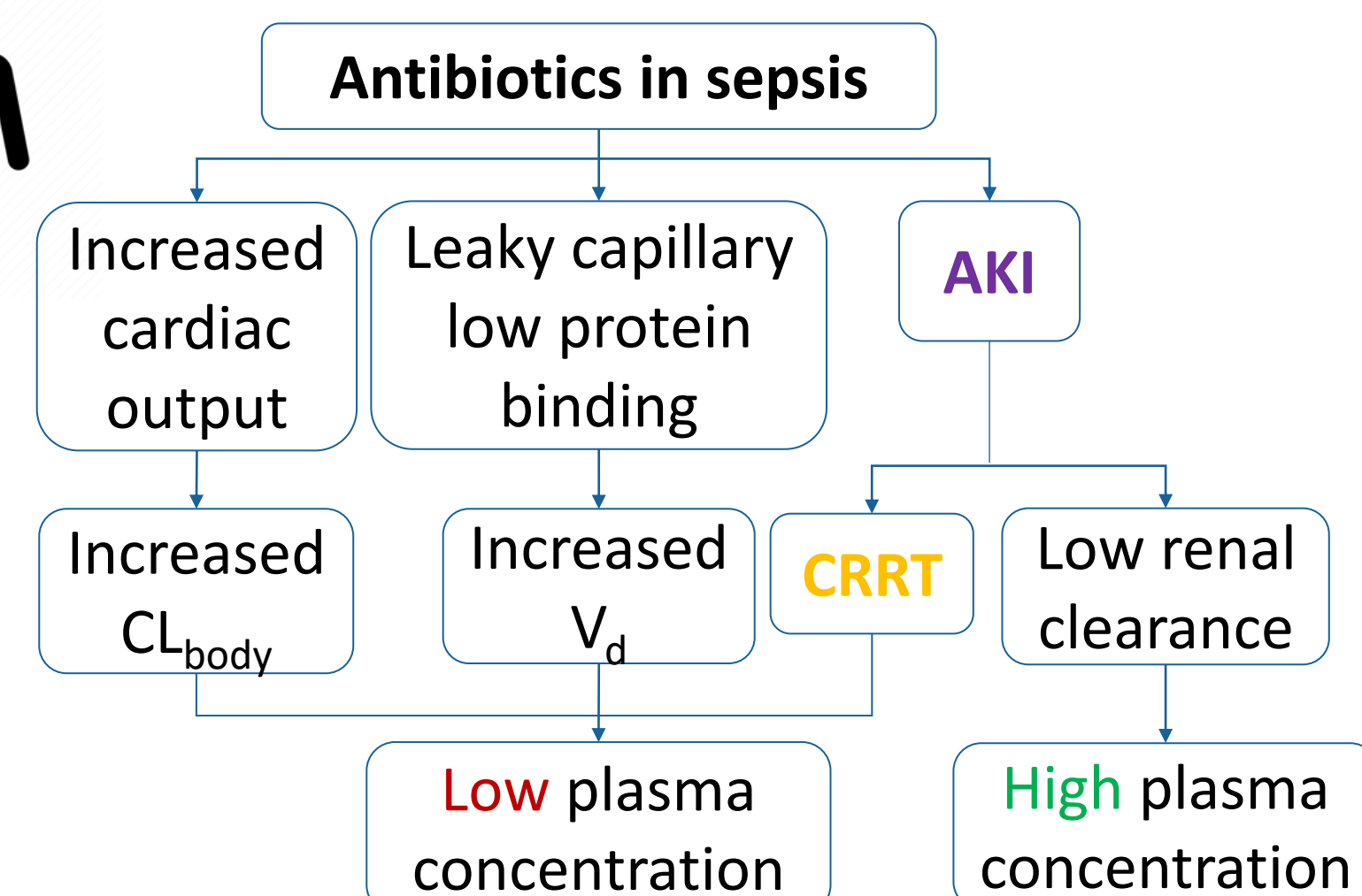
Division of Biopharmaceutics and Pharmacokinetics, Xiangya School of Pharmaceutical Sciences, Central South University, Changsha, China

Contact: feifan.xie@csu.edu.cn

Lack of drug dosing guidelines for critically ill patients receiving continuous renal replacement therapy (CRRT)

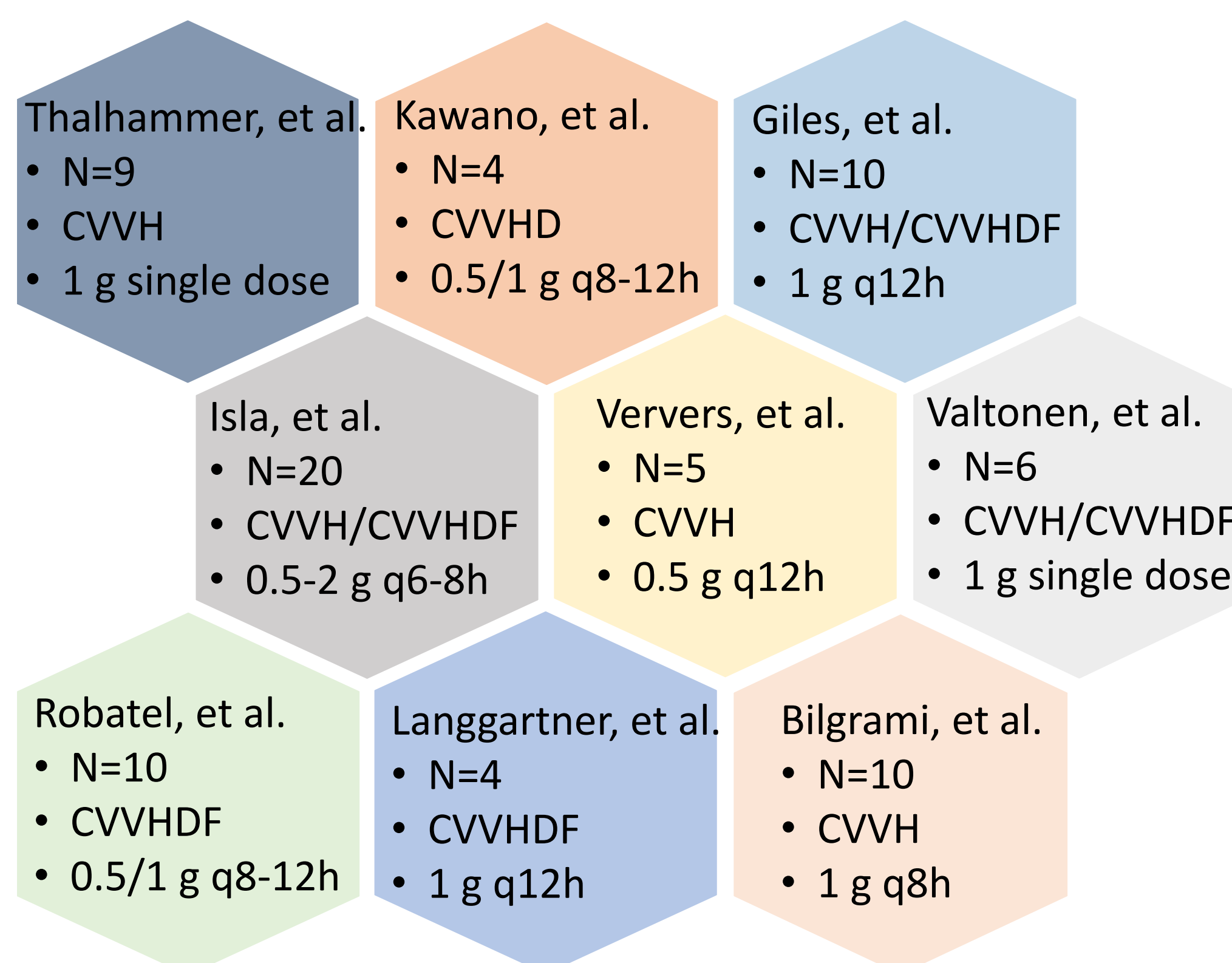


Extracorporeal clearance of different CRRT modalities (CL_{CRRT}) mainly depends on the effluent flow rate (Q_E , also termed as CRRT dose).

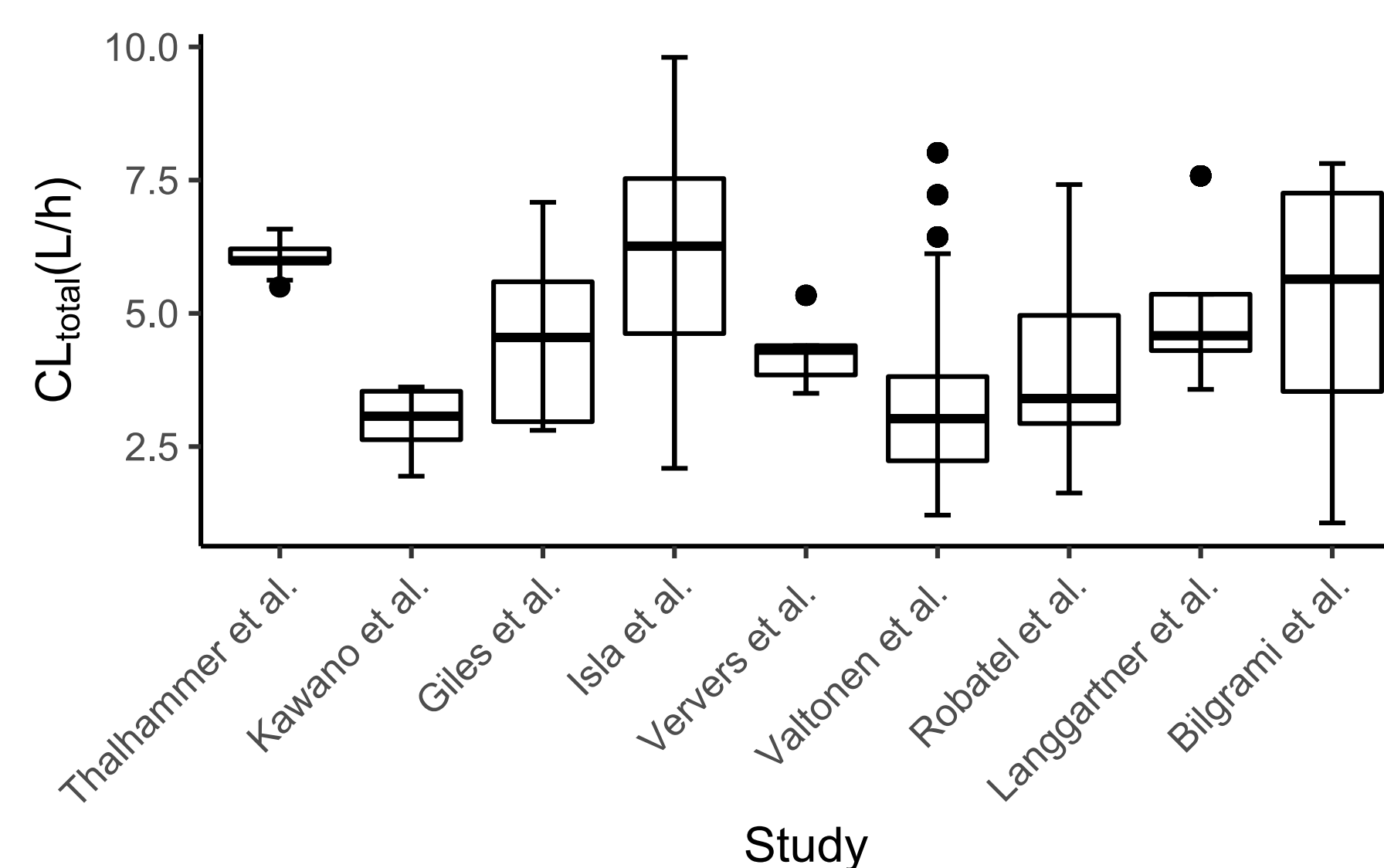


The pharmacokinetics (PK) of an antibiotic is altered by the pathophysiological state and CRRT.

Population pharmacokinetic meta-analysis of meropenem: data source and main findings



Included published PK studies for present population PK meta-analysis of meropenem in CRRT patients. (501 observations from 78 patients)

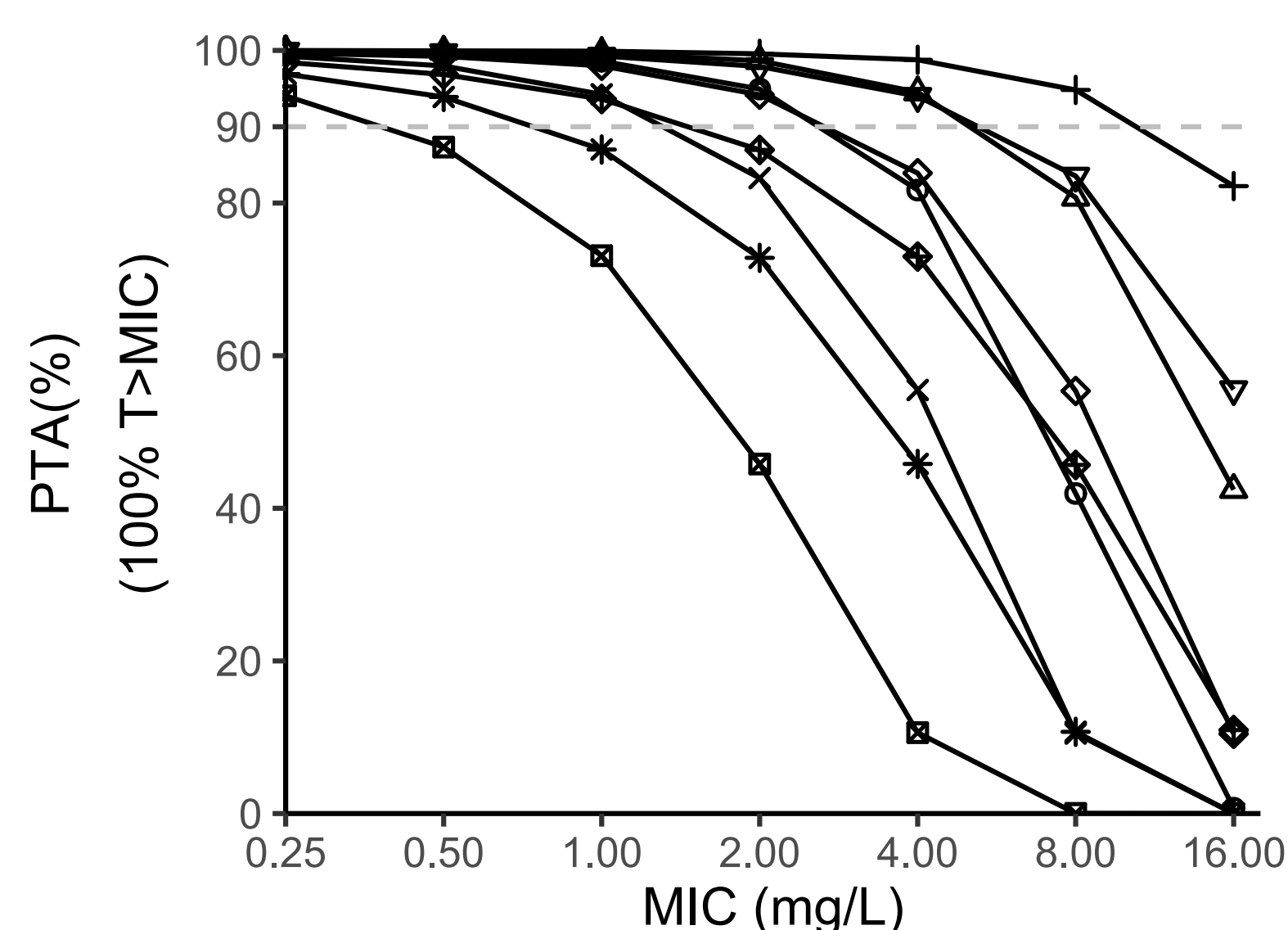


$$CL_{\text{total}}(L/h) = \left(\theta_{CL_{\text{body}}(\text{non-trauma})} \text{ or } \theta_{CL_{\text{body}}(\text{trauma})} \right) \times \exp^{\eta_{CL_{\text{body}}}} + CL_{\text{CRRT}}$$

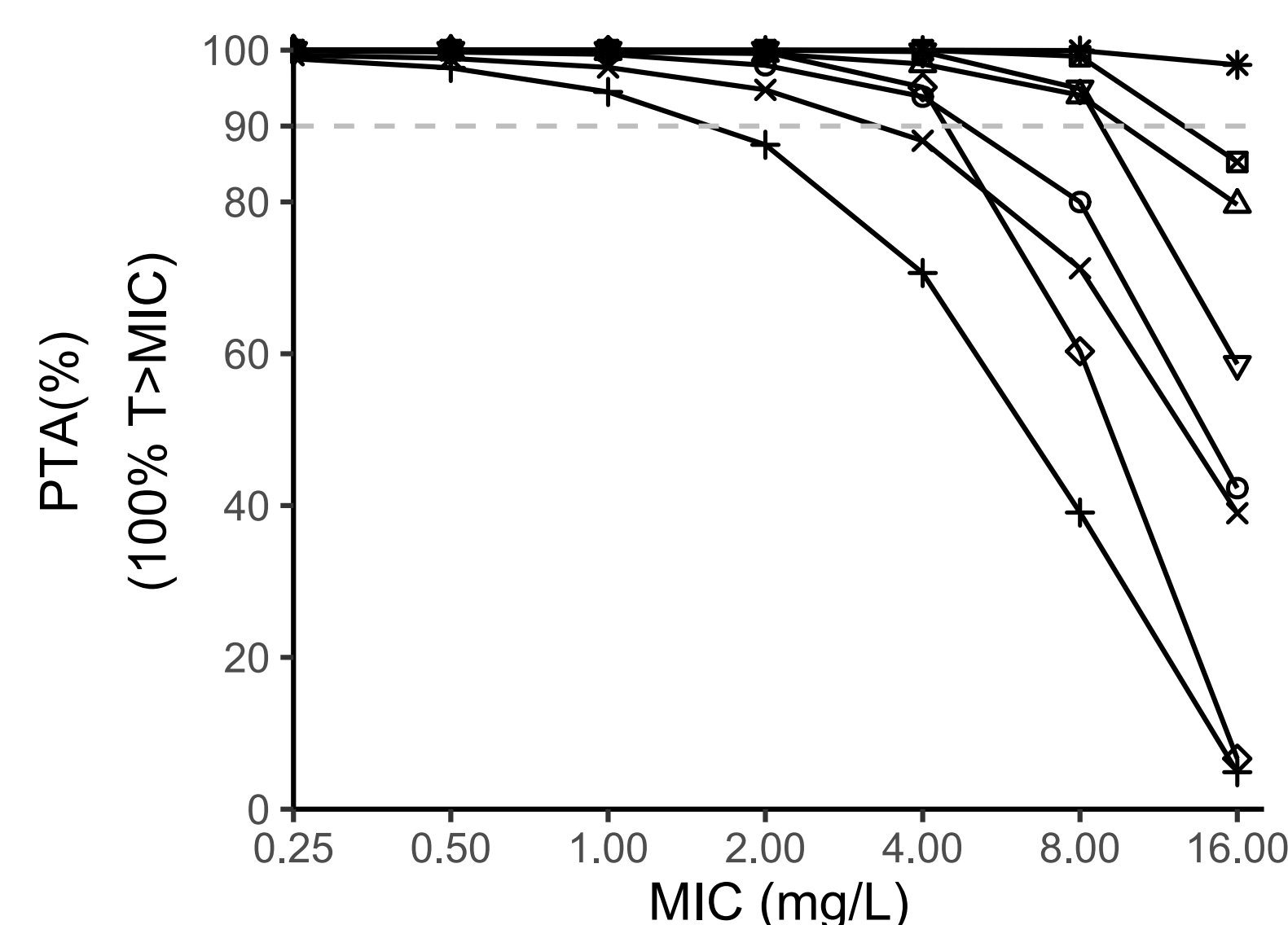
- Non-trauma patients: considerable discrepancy in total clearance (CL_{total}) was found among the studies (median CL_{total} ranged from 3.06 to 6.07 L/h);
- Trauma patients: typical CL_{total} (62.7 L/h) in trauma patients was increased by 20-fold compared to non-trauma patients.

Meropenem dosing recommendation

Non-trauma CRRT patients: 1 g q6h by short-term infusion, 1 g q8h by prolonged infusion, or 2-4 g q24h by continuous infusion; Trauma CRRT patients: 3-4 g q24h by continuous infusion



- 0.5 g q6h infused over 30mins
- △ 1 g q6h infused over 30mins
- + 2 g q6h infused over 30mins
- × 0.5 g q8h infused over 30mins
- ◇ 1 g q8h infused over 30mins
- ▽ 2 g q8h infused over 30mins
- 0.5 g q12h infused over 30mins
- * 1 g q12h infused over 30mins
- ◆ 2 g q12h infused over 30mins



- 1 g q8h infused over 3h
- △ 2 g q8h infused over 3h
- + 1 g q12h infused over 3h
- × 2 g q12h infused over 3h
- ◇ 2 g q24h infused over 24h
- ▽ 3 g q24h infused over 24h
- 4 g q24h infused over 24h
- * 6 g q24h infused over 24h