

Questions and Comments Received From Webinar Attendees Directed to Specific Speakers: Jeanne Domoradzki

Q: Just to comment that subproportionality may also be due to changes in protein binding and elimination and not just changes to dissolution rate, transit time in the gastrointestinal tract, or ability to cross intestinal barriers. $\text{Area under the curve} = (\text{Bioavailability} \times \text{Dose}) / (\text{Total Clearance}) = (\text{Bioavailability} \times \text{Dose}) / (\text{fraction unbound} \times \text{intrinsic clearance})$.

A: That depends on whether one is considering absorption or bioavailability. Kinetically, these have different meanings and different determinants. For example, systemic elimination does not affect absorption.