APPENDIX B: PEDIATRIC DECISION TREE INTEGRATION OF PK-PD

**Pediatric Study Decision Tree**

Reasonable to assume (pediatrics vs adults)
✓ similar disease progression?
✓ similar response to intervention?

NO ➔

- Conduct PK studies
- Conduct safety/efficacy trials*

NO ➔

Is there a PD measurement** that can be used to predict efficacy?

NO ➔

YES TO BOTH ➔

Reasonable to assume similar concentration-response (C-R) in pediatrics and adults?

YES ➔

- Conduct PK studies to achieve levels similar to adults
- Conduct safety trials

NO ➔

YES ➔

- Conduct PK/PD studies to get C-R for PD measurement
- Conduct PK studies to achieve target concentrations based on C-R

- Conduct safety trials