

Supplementary Materials: The Combination of Predictive Factors of Pharmacokinetic Origin Associates with Enhanced Disease Control During Treatment of Pediatric Crohn's Disease with Infliximab

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Table S1: Predictive factors of PK origin and time to CRP based clinical remission in pediatric CD

			Time (days)	HR	p value
Infusion 2					
Clearance above 0.294 L/day	0.294	110±27		ref.	
Clearance below 0.294 L/day		71±14	1.3 (0.9 to 2.0)	0.188	
IFX below 20µg/mL		95±18	ref.		
IFX above 20µg/mL		69±13	1.4 (0.9-2.1)	0.140	
Infusion 3					
Clearance above 0.294 L/day	0.294	138±28		ref.	
Clearance below 0.294 L/day		76±13	1.5 (1.0 to 2.3)	0.047	
IFX below 15µg/mL		139±20	ref.		
IFX above 15µg/mL		46±12	2.3 (1.5-3.6)	<0.001	
Infusion 4					
Clearance above 0.294 L/day	0.294	188±35		ref.	
Clearance below 0.294 L/day		68±11	2.1 (1.3 to 3.2)	0.001	
IFX below 10µg/mL		142±21			
IFX above 10µg/mL		53±12	2.2 (1.4-3.3)	<0.001	

Table S2: PF of PK origin either alone or combined at the end of induction and during maintenance and disease control over maintenance period.

	parameter*	estimates
Time and IFX >5µg/mL	θ_{pop} θ_{time} $\theta_{\text{concentration}}$ -2LL	-0.38±0.53 (p=0.473) 0.004±0.002 (p=0.046) 1.53±0.43 (p<0.001) 361.6
Time and Clearance <0.294 L/day	θ_{pop} θ_{time} θ_{Cl} -2LL	1.53±0.58 (p=0.008) 0.004±0.002 (p=0.046) 2.57±0.50 (p<0.001) 353.7 (Δ =-7.9; p=0.005)
Time and PF of PK origin (IFX >5µg/mL with Clearance <0.294 L/day)	θ_{pop} θ_{pop} $\theta_{\text{both PF of PK}}$ -2LL	1.06 ±0.48 (p<0.001) 0.004±0.001 (p<0.001) 2.97±0.81 (p<0.001) 396 (Δ =-12.6; ; p<0.001)
Time and both PF of PK origin at infusion 4 (IFX >10µg/mL with Clearance <0.294 L/day) and during maintenance	θ_{pop} $\theta_{\text{both PF of PK 4th inf}}$ θ_{time} $\theta_{\text{both PF of PK}}$ -2LL	0.35±0.50 (p=0.012) 2.44±0.64 (p<0.001) 0.002±0.001 (p=0.046) 2.51±0.73 (p<0.001) 377.8 (Δ =-23.2; p<0.001)

*Model: $\text{logit}(\text{Probability of CRP based remission}) = \theta_{\text{pop}} + \theta_{\text{covi}} * \text{covi} + \dots$

Figure S1: IFX concentration and Clearance at each of the Induction time point and maintenance.
Panel A: infusion 2; panel B: infusion 3; panel C: infusion 4; panel D: maintenance

