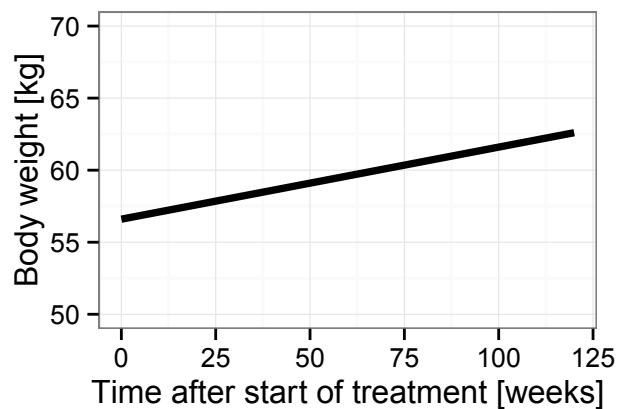
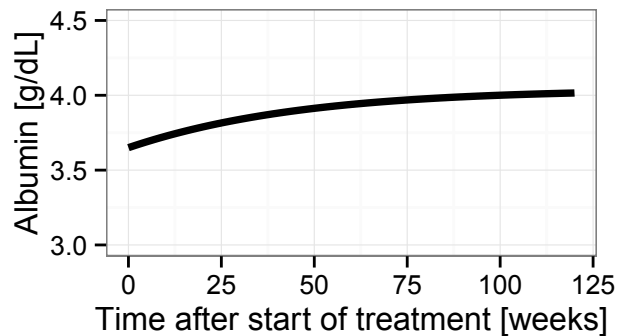
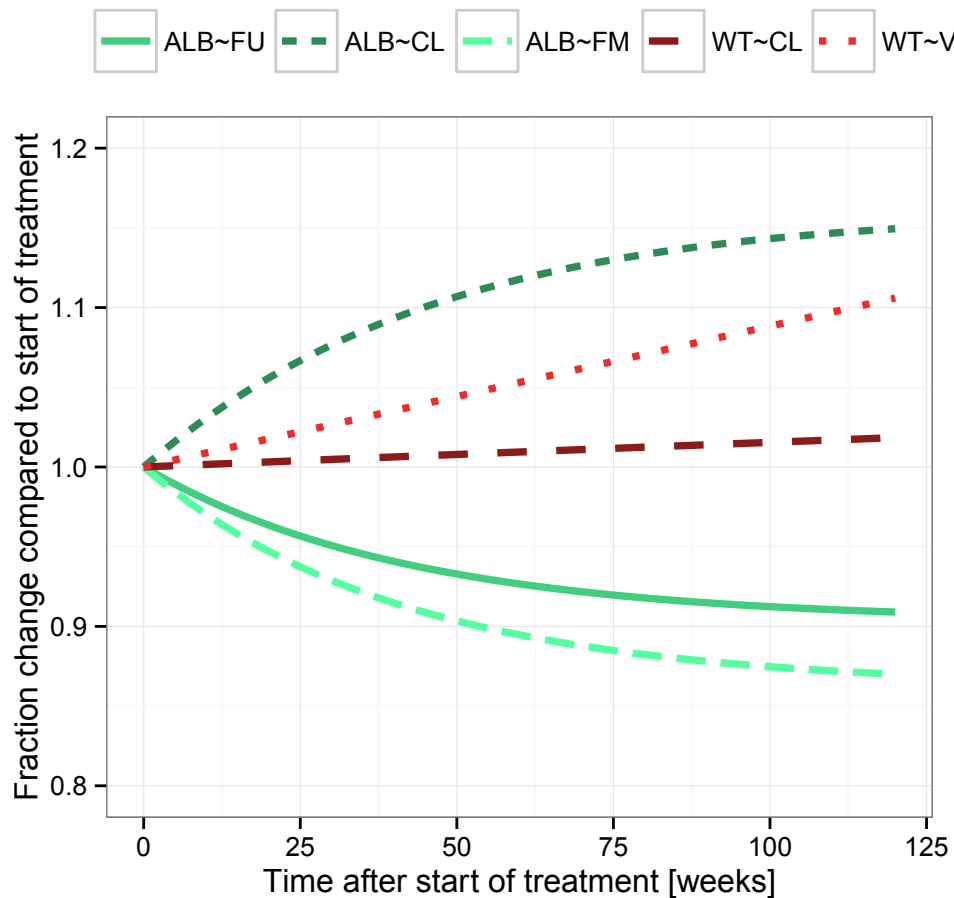


Covariate value over time for the typical individual



Change per relationship over time



Change per parameter over time

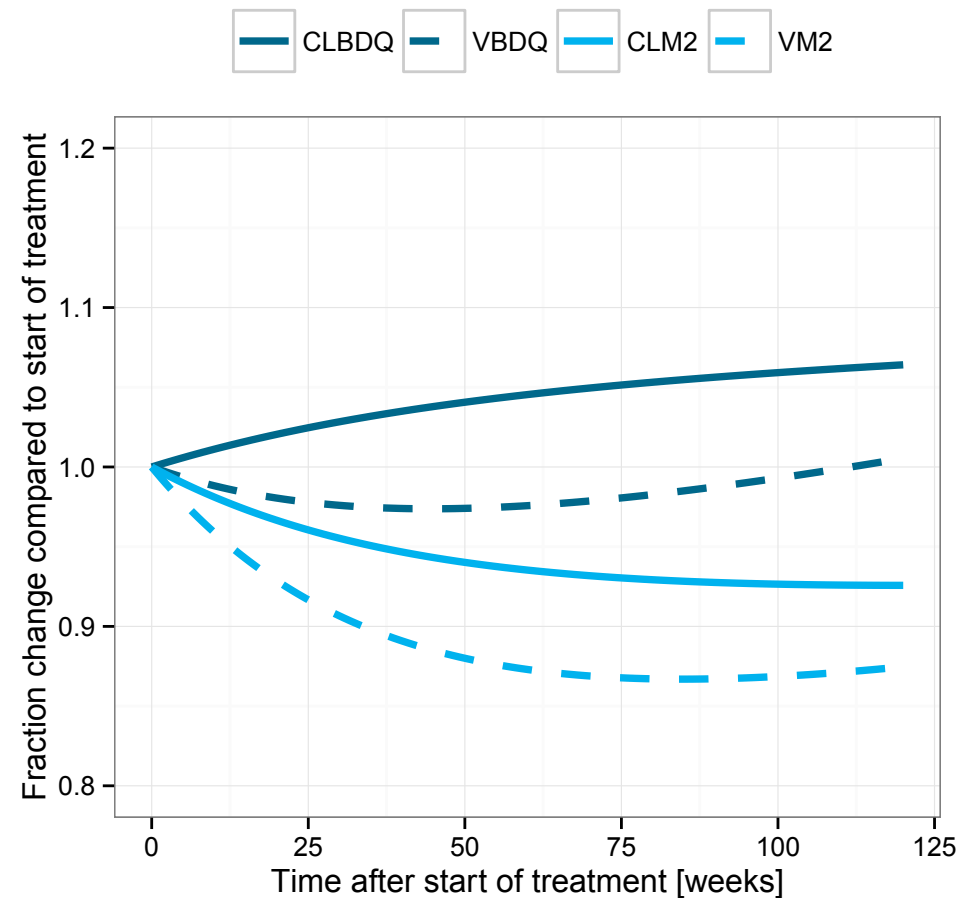


Figure S2. Overview of the effects of the relationships between selected PK parameters and time-varying covariate body weight and albumin concentration showing the typical change in the covariates over time on treatment (left panel), the fractional change per relationship over time after start of treatment (middle panel) and total fractional change in selected PK parameters (right panel). Specific individual weight and albumin dynamics yields different total effects on the PK parameters, which can be notably stronger than the typical or even go in the opposite direction.