Bayesian Hierarchical Non-Linear Modeling with LOESS Smoothing

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The Bayesian hierarchical approach offers many advantages over traditional frequentist methods when dealing with complex data structures and when prior knowledge is available:

- Incorporation of Prior Knowledge: This method allows for the integration of prior knowledge or expert opinion through the use of priors, which can improve estimates, especially when data is scarce or noisy.

- Probabilistic Interpretation: Bayesian inference provides a probabilistic interpretation of model parameters, giving a full distribution of possible values rather than a single point estimate, which adds richness to the understanding of parameter uncertainties.

- Flexibility in Model Specification: Hierarchical models can easily accommodate varying levels of randomness and complex data structures, which is often more cumbersome in frequentist frameworks.

- Direct Probability Statements: Bayesian statistics let you make direct probability statements about parameters (e.g., "The probability that a parameter is greater than zero"), whereas frequentist methods only provide probabilities of data given a hypothesis.

- Model Comparison and Averaging: Bayesian methods facilitate model comparison and averaging without the strict requirements of nested models or concerns over multiple testing that you encounter in the frequentist approach.

- Inference from the Posterior: In Bayesian analysis, inference is drawn from the posterior distribution, which is updated as new data arrives, enabling a natural form of sequential analysis.

- Robustness to Overfitting: Hierarchical Bayesian models can be more robust to overfitting, especially when informed priors are used, due to their ability to shrink estimates toward group-level averages.

- Handling of Missing Data: Bayesian hierarchical models can handle missing data more naturally by treating them as additional parameters to be estimated, under a coherent probabilistic framework.

- Endorsement of Uncertainty: Bayesian methods inherently account for uncertainty in all levels of the modeling process, making them inherently more cautious and, in some sense, more honest about the reliability of inferences.

- Continuous Learning: The Bayesian framework is conducive to continuous learning, as the posterior distribution can be updated with new data, reflecting an evolving state of knowledge without starting the analysis anew.

Setting a prior is a fundamental aspect of Bayesian analysis, and it's both an advantage and a point of criticism. Properly chosen priors can regularize estimates, inject substantive knowledge into the analysis, and improve inference, especially in small samples or complex models. However, the choice of prior must be made carefully to avoid undue influence on the results, and this process should be transparent and justified within the context of the analysis. Bayesian hierarchical models are especially powerful in situations where parameters are expected to vary by group or category but also share commonalities, as they can borrow strength across these groups to make more informed estimates. This is particularly useful in medical, biological, or social sciences where individual-level variability is nested within larger group-level trends.

Load libraries

Get the theophylline dataset from the base `datasets` library

Plot raw data to get a sense of how the data looks like

```
ggplot(df, aes(x = Time, y = conc)) + geom_point() +
  scale_x_continuous(breaks = seq(0,20,2))
```
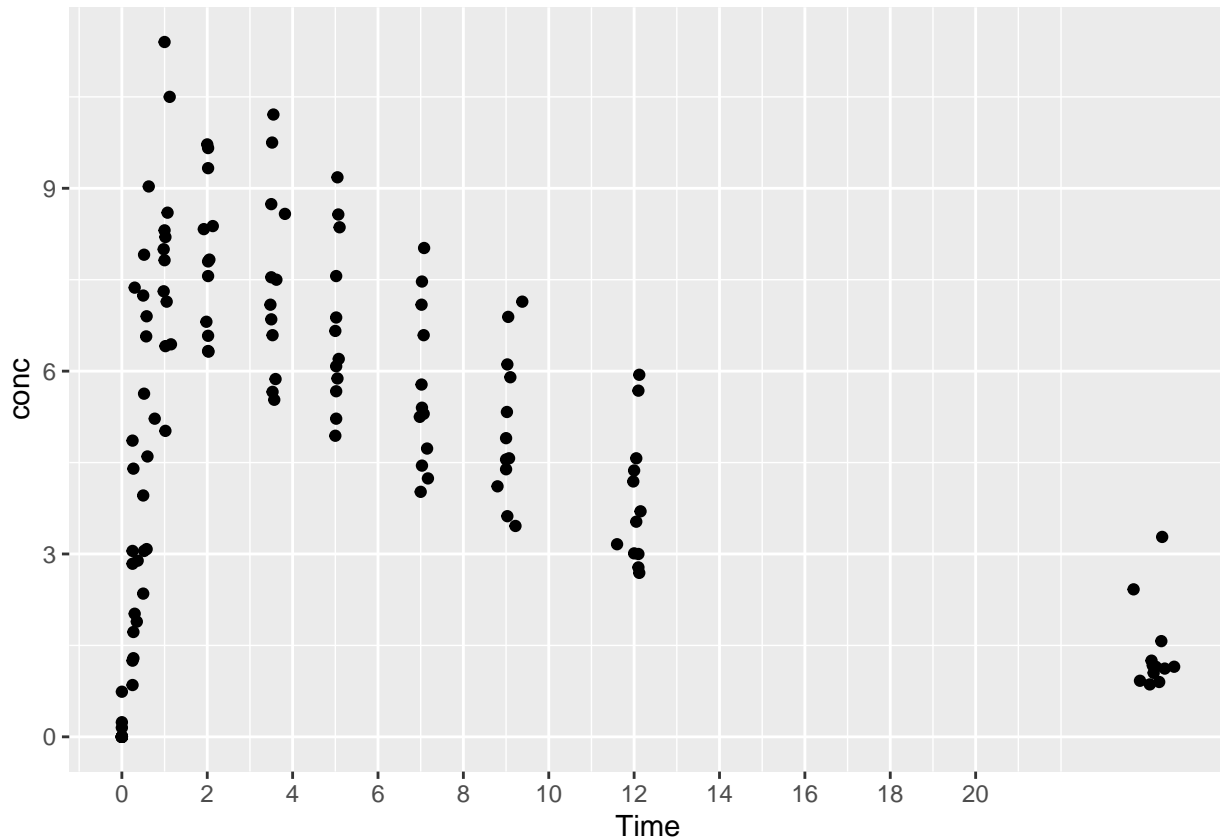

We can see in this plot that there is a very conspicuous change in the rate of concentration change at Time = 1s. A simple linear model won't cut it. Therefore, we need to build a non-linear model to account for this peculiarity.

Build a Bayesian generalized non-linear multivariate multilevel (Hierarchical with each subject as a random intercept) model adjusting for participant weight and dose of theophylline; we can create a basic spline model and place a knot at T = 1s to account for the change in slope.

```
model <- brm(
  conc ~ bs(Time, knots = c(1)) + (1 | Subject) + Wt + Dose, # adjusting for weight and dose of theophy
  data = df,
  family = gaussian(), # for continuous outcomes
  prior = c(
    set_prior("normal(0,5)", class = "b"), # priors for fixed effects
    set_prior("normal(0,5)", class = "sd")  # priors for group-level effects
  ),
  chains = 2, iter = 5000, warmup = 2000,
  control = list(adapt_delta = 0.95)
```

```
)
```

```
## Compiling Stan program...

## Start sampling

##
## SAMPLING FOR MODEL 'anon_model' NOW (CHAIN 1).
## Chain 1:
## Chain 1: Gradient evaluation took 3.7e-05 seconds
## Chain 1: 1000 transitions using 10 leapfrog steps per transition would take 0.37 seconds.
## Chain 1: Adjust your expectations accordingly!
## Chain 1:
## Chain 1:
## Chain 1: Iteration:    1 / 5000 [  0%]  (Warmup)
## Chain 1: Iteration:  500 / 5000 [ 10%]  (Warmup)
## Chain 1: Iteration: 1000 / 5000 [ 20%]  (Warmup)
## Chain 1: Iteration: 1500 / 5000 [ 30%]  (Warmup)
## Chain 1: Iteration: 2000 / 5000 [ 40%]  (Warmup)
## Chain 1: Iteration: 2001 / 5000 [ 40%]  (Sampling)
## Chain 1: Iteration: 2500 / 5000 [ 50%]  (Sampling)
## Chain 1: Iteration: 3000 / 5000 [ 60%]  (Sampling)
## Chain 1: Iteration: 3500 / 5000 [ 70%]  (Sampling)
## Chain 1: Iteration: 4000 / 5000 [ 80%]  (Sampling)
## Chain 1: Iteration: 4500 / 5000 [ 90%]  (Sampling)
## Chain 1: Iteration: 5000 / 5000 [100%]  (Sampling)
## Chain 1:
## Chain 1:  Elapsed Time: 1.021 seconds (Warm-up)
## Chain 1:                1.195 seconds (Sampling)
## Chain 1:                2.216 seconds (Total)
## Chain 1:
##
## SAMPLING FOR MODEL 'anon_model' NOW (CHAIN 2).
## Chain 2:
## Chain 2: Gradient evaluation took 1.1e-05 seconds
## Chain 2: 1000 transitions using 10 leapfrog steps per transition would take 0.11 seconds.
## Chain 2: Adjust your expectations accordingly!
## Chain 2:
## Chain 2:
## Chain 2: Iteration:    1 / 5000 [  0%]  (Warmup)
## Chain 2: Iteration:  500 / 5000 [ 10%]  (Warmup)
## Chain 2: Iteration: 1000 / 5000 [ 20%]  (Warmup)
## Chain 2: Iteration: 1500 / 5000 [ 30%]  (Warmup)
## Chain 2: Iteration: 2000 / 5000 [ 40%]  (Warmup)
## Chain 2: Iteration: 2001 / 5000 [ 40%]  (Sampling)
## Chain 2: Iteration: 2500 / 5000 [ 50%]  (Sampling)
## Chain 2: Iteration: 3000 / 5000 [ 60%]  (Sampling)
## Chain 2: Iteration: 3500 / 5000 [ 70%]  (Sampling)
## Chain 2: Iteration: 4000 / 5000 [ 80%]  (Sampling)
## Chain 2: Iteration: 4500 / 5000 [ 90%]  (Sampling)
## Chain 2: Iteration: 5000 / 5000 [100%]  (Sampling)
## Chain 2:
## Chain 2:  Elapsed Time: 1.04 seconds (Warm-up)
## Chain 2:                1.485 seconds (Sampling)
## Chain 2:                2.525 seconds (Total)
```

```
## Chain 2:
```

Generate a model summary

```
summary(model)
```

```
##  Family: gaussian
##   Links: mu = identity; sigma = identity
## Formula: conc ~ bs(Time, knots = c(1)) + (1 | Subject) + Wt + Dose
##    Data: df (Number of observations: 132)
##   Draws: 2 chains, each with iter = 5000; warmup = 2000; thin = 1;
##          total post-warmup draws = 6000
##
## Multilevel Hyperparameters:
## ~Subject (Number of levels: 12)
##               Estimate Est.Error l-95% CI u-95% CI Rhat Bulk_ESS Tail_ESS
## sd(Intercept)     0.87      0.29     0.44     1.58 1.00     1820     3142
##
## Regression Coefficients:
##                 Estimate Est.Error l-95% CI u-95% CI Rhat Bulk_ESS Tail_ESS
## Intercept          -8.50     24.27   -55.72    40.28 1.00     2576     3110
## bsTimeknotsEQc11    7.10      0.51     6.12     8.11 1.00     4369     4068
## bsTimeknotsEQc12    9.59      1.09     7.50    11.71 1.00     4124     4305
## bsTimeknotsEQc13   -2.68      1.78    -6.18     0.81 1.00     3904     3900
## bsTimeknotsEQc14    1.95      0.57     0.82     3.04 1.00     4803     3909
## Wt                  0.05      0.19    -0.32     0.43 1.00     2554     3149
## Dose                0.98      2.40    -3.94     5.69 1.00     2605     3169
##
## Further Distributional Parameters:
##       Estimate Est.Error l-95% CI u-95% CI Rhat Bulk_ESS Tail_ESS
## sigma     1.36      0.09     1.20     1.55 1.00     4804     4070
##
## Draws were sampled using sampling(NUTS). For each parameter, Bulk_ESS
## and Tail_ESS are effective sample size measures, and Rhat is the potential
## scale reduction factor on split chains (at convergence, Rhat = 1).
```

Generate a model report

```
report(model)
```

```
## Response residuals not available to calculate mean square error. (R)MSE
##    is probably not reliable.
```

```
## Start sampling
```

```
##
## SAMPLING FOR MODEL 'anon_model' NOW (CHAIN 1).
## Chain 1:
## Chain 1: Gradient evaluation took 1.1e-05 seconds
## Chain 1: 1000 transitions using 10 leapfrog steps per transition would take 0.11 seconds.
## Chain 1: Adjust your expectations accordingly!
## Chain 1:
## Chain 1:
## Chain 1: Iteration:    1 / 5000 [  0%]  (Warmup)
```

```
## Chain 1: Iteration:  500 / 5000 [ 10%]  (Warmup)
## Chain 1: Iteration: 1000 / 5000 [ 20%]  (Warmup)
## Chain 1: Iteration: 1500 / 5000 [ 30%]  (Warmup)
## Chain 1: Iteration: 2000 / 5000 [ 40%]  (Warmup)
## Chain 1: Iteration: 2001 / 5000 [ 40%]  (Sampling)
## Chain 1: Iteration: 2500 / 5000 [ 50%]  (Sampling)
## Chain 1: Iteration: 3000 / 5000 [ 60%]  (Sampling)
## Chain 1: Iteration: 3500 / 5000 [ 70%]  (Sampling)
## Chain 1: Iteration: 4000 / 5000 [ 80%]  (Sampling)
## Chain 1: Iteration: 4500 / 5000 [ 90%]  (Sampling)
## Chain 1: Iteration: 5000 / 5000 [100%]  (Sampling)
## Chain 1:
## Chain 1:  Elapsed Time: 0.822 seconds (Warm-up)
## Chain 1:                1.277 seconds (Sampling)
## Chain 1:                2.099 seconds (Total)
## Chain 1:
##
## SAMPLING FOR MODEL 'anon_model' NOW (CHAIN 2).
## Chain 2:
## Chain 2: Gradient evaluation took 6e-06 seconds
## Chain 2: 1000 transitions using 10 leapfrog steps per transition would take 0.06 seconds.
## Chain 2: Adjust your expectations accordingly!
## Chain 2:
## Chain 2:
## Chain 2: Iteration:    1 / 5000 [  0%]  (Warmup)
## Chain 2: Iteration:  500 / 5000 [ 10%]  (Warmup)
## Chain 2: Iteration: 1000 / 5000 [ 20%]  (Warmup)
## Chain 2: Iteration: 1500 / 5000 [ 30%]  (Warmup)
## Chain 2: Iteration: 2000 / 5000 [ 40%]  (Warmup)
## Chain 2: Iteration: 2001 / 5000 [ 40%]  (Sampling)
## Chain 2: Iteration: 2500 / 5000 [ 50%]  (Sampling)
## Chain 2: Iteration: 3000 / 5000 [ 60%]  (Sampling)
## Chain 2: Iteration: 3500 / 5000 [ 70%]  (Sampling)
## Chain 2: Iteration: 4000 / 5000 [ 80%]  (Sampling)
## Chain 2: Iteration: 4500 / 5000 [ 90%]  (Sampling)
## Chain 2: Iteration: 5000 / 5000 [100%]  (Sampling)
## Chain 2:
## Chain 2:  Elapsed Time: 0.866 seconds (Warm-up)
## Chain 2:                1.088 seconds (Sampling)
## Chain 2:                1.954 seconds (Total)
## Chain 2:

## Response residuals not available to calculate mean square error. (R)MSE
##   is probably not reliable.

## We fitted a Bayesian linear mixed model (estimated using MCMC sampling with 2
## chains of 5000 iterations and a warmup of 2000) to predict conc with Time, Wt
## and Dose (formula: conc ~ bs(Time, knots = c(1)) + Wt + Dose). The model
## included Subject as random effect (formula: ~1 | Subject). Priors over
## parameters were set as student_t (location = 5.30, scale = 3.30) distributions.
## The model's explanatory power is substantial (R2 = 0.78, 95% CI [0.74, 0.81],
## adj. R2 = 0.76) and the part related to the fixed effects alone (marginal R2)
## is of 0.72 (95% CI [0.67, 0.76]).  Within this model:
##
##   - The effect of b Intercept (Median = -8.85, 95% CI [-55.72, 40.28]) has a
```

```
## 64.50% probability of being negative (< 0), 64.35% of being significant (<
## -0.14), and 63.10% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4384)
##   - The effect of b bsTimeknotsEQc11 (Median = 7.10, 95% CI [6.12, 8.11]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4096)
##   - The effect of b bsTimeknotsEQc12 (Median = 9.57, 95% CI [7.50, 11.71]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 3887)
##   - The effect of b bsTimeknotsEQc13 (Median = -2.70, 95% CI [-6.18, 0.81]) has a
## 93.15% probability of being negative (< 0), 92.03% of being significant (<
## -0.14), and 84.63% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4833)
##   - The effect of b bsTimeknotsEQc14 (Median = 1.96, 95% CI [0.82, 3.04]) has a
## 99.97% probability of being positive (> 0), 99.90% of being significant (>
## 0.14), and 96.95% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 2623)
##   - The effect of b Wt (Median = 0.05, 95% CI [-0.32, 0.43]) has a 61.55%
## probability of being positive (> 0), 31.50% of being significant (> 0.14), and
## 0.00% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2604)
##   - The effect of b Dose (Median = 1.02, 95% CI [-3.94, 5.69]) has a 66.50%
## probability of being positive (> 0), 64.08% of being significant (> 0.14), and
## 52.77% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2585)
##
## Following the Sequential Effect eXistence and sIgnificance Testing (SEXIT)
## framework, we report the median of the posterior distribution and its 95% CI
## (Highest Density Interval), along the probability of direction (pd), the
## probability of significance and the probability of being large. The thresholds
## beyond which the effect is considered as significant (i.e., non-negligible) and
## large are |0.14| and |0.86| (corresponding respectively to 0.05 and 0.30 of the
## outcome's SD). Convergence and stability of the Bayesian sampling has been
## assessed using R-hat, which should be below 1.01 (Vehtari et al., 2019), and
## Effective Sample Size (ESS), which should be greater than 1000 (Burkner,
## 2017)., We fitted a Bayesian linear mixed model (estimated using MCMC sampling
## with 2 chains of 5000 iterations and a warmup of 2000) to predict conc with
## Time, Wt and Dose (formula: conc ~ bs(Time, knots = c(1)) + Wt + Dose). The
## model included Subject as random effect (formula: ~1 | Subject). Priors over
## parameters were set as normal (mean = 0.00, SD = 5.00) distributions. The
## model's explanatory power is substantial (R2 = 0.78, 95% CI [0.74, 0.81], adj.
## R2 = 0.76) and the part related to the fixed effects alone (marginal R2) is of
## 0.72 (95% CI [0.67, 0.76]).  Within this model:
##
##   - The effect of b Intercept (Median = -8.85, 95% CI [-55.72, 40.28]) has a
## 64.50% probability of being negative (< 0), 64.35% of being significant (<
## -0.14), and 63.10% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4384)
##   - The effect of b bsTimeknotsEQc11 (Median = 7.10, 95% CI [6.12, 8.11]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4096)
```

```
##   - The effect of b bsTimeknotsEQc12 (Median = 9.57, 95% CI [7.50, 11.71]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 3887)
##   - The effect of b bsTimeknotsEQc13 (Median = -2.70, 95% CI [-6.18, 0.81]) has a
## 93.15% probability of being negative (< 0), 92.03% of being significant (<
## -0.14), and 84.63% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4833)
##   - The effect of b bsTimeknotsEQc14 (Median = 1.96, 95% CI [0.82, 3.04]) has a
## 99.97% probability of being positive (> 0), 99.90% of being significant (>
## 0.14), and 96.95% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 2623)
##   - The effect of b Wt (Median = 0.05, 95% CI [-0.32, 0.43]) has a 61.55%
## probability of being positive (> 0), 31.50% of being significant (> 0.14), and
## 0.00% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2604)
##   - The effect of b Dose (Median = 1.02, 95% CI [-3.94, 5.69]) has a 66.50%
## probability of being positive (> 0), 64.08% of being significant (> 0.14), and
## 52.77% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2585)
##
## Following the Sequential Effect eXistence and sIgnificance Testing (SEXIT)
## framework, we report the median of the posterior distribution and its 95% CI
## (Highest Density Interval), along the probability of direction (pd), the
## probability of significance and the probability of being large. The thresholds
## beyond which the effect is considered as significant (i.e., non-negligible) and
## large are |0.14| and |0.86| (corresponding respectively to 0.05 and 0.30 of the
## outcome's SD). Convergence and stability of the Bayesian sampling has been
## assessed using R-hat, which should be below 1.01 (Vehtari et al., 2019), and
## Effective Sample Size (ESS), which should be greater than 1000 (Burkner,
## 2017)., We fitted a Bayesian linear mixed model (estimated using MCMC sampling
## with 2 chains of 5000 iterations and a warmup of 2000) to predict conc with
## Time, Wt and Dose (formula: conc ~ bs(Time, knots = c(1)) + Wt + Dose). The
## model included Subject as random effect (formula: ~1 | Subject). Priors over
## parameters were set as normal (mean = 0.00, SD = 5.00) distributions. The
## model's explanatory power is substantial (R2 = 0.78, 95% CI [0.74, 0.81], adj.
## R2 = 0.76) and the part related to the fixed effects alone (marginal R2) is of
## 0.72 (95% CI [0.67, 0.76]).  Within this model:
##
##   - The effect of b Intercept (Median = -8.85, 95% CI [-55.72, 40.28]) has a
## 64.50% probability of being negative (< 0), 64.35% of being significant (<
## -0.14), and 63.10% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4384)
##   - The effect of b bsTimeknotsEQc11 (Median = 7.10, 95% CI [6.12, 8.11]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4096)
##   - The effect of b bsTimeknotsEQc12 (Median = 9.57, 95% CI [7.50, 11.71]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 3887)
##   - The effect of b bsTimeknotsEQc13 (Median = -2.70, 95% CI [-6.18, 0.81]) has a
## 93.15% probability of being negative (< 0), 92.03% of being significant (<
## -0.14), and 84.63% of being large (< -0.86). The estimation successfully
```

```
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4833)
##   - The effect of b bsTimeknotsEQc14 (Median = 1.96, 95% CI [0.82, 3.04]) has a
## 99.97% probability of being positive (> 0), 99.90% of being significant (>
## 0.14), and 96.95% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 2623)
##   - The effect of b Wt (Median = 0.05, 95% CI [-0.32, 0.43]) has a 61.55%
## probability of being positive (> 0), 31.50% of being significant (> 0.14), and
## 0.00% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2604)
##   - The effect of b Dose (Median = 1.02, 95% CI [-3.94, 5.69]) has a 66.50%
## probability of being positive (> 0), 64.08% of being significant (> 0.14), and
## 52.77% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2585)
##
## Following the Sequential Effect eXistence and sIgnificance Testing (SEXIT)
## framework, we report the median of the posterior distribution and its 95% CI
## (Highest Density Interval), along the probability of direction (pd), the
## probability of significance and the probability of being large. The thresholds
## beyond which the effect is considered as significant (i.e., non-negligible) and
## large are |0.14| and |0.86| (corresponding respectively to 0.05 and 0.30 of the
## outcome's SD). Convergence and stability of the Bayesian sampling has been
## assessed using R-hat, which should be below 1.01 (Vehtari et al., 2019), and
## Effective Sample Size (ESS), which should be greater than 1000 (Burkner,
## 2017)., We fitted a Bayesian linear mixed model (estimated using MCMC sampling
## with 2 chains of 5000 iterations and a warmup of 2000) to predict conc with
## Time, Wt and Dose (formula: conc ~ bs(Time, knots = c(1)) + Wt + Dose). The
## model included Subject as random effect (formula: ~1 | Subject). Priors over
## parameters were set as normal (mean = 0.00, SD = 5.00) distributions. The
## model's explanatory power is substantial (R2 = 0.78, 95% CI [0.74, 0.81], adj.
## R2 = 0.76) and the part related to the fixed effects alone (marginal R2) is of
## 0.72 (95% CI [0.67, 0.76]).  Within this model:
##
##   - The effect of b Intercept (Median = -8.85, 95% CI [-55.72, 40.28]) has a
## 64.50% probability of being negative (< 0), 64.35% of being significant (<
## -0.14), and 63.10% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4384)
##   - The effect of b bsTimeknotsEQc11 (Median = 7.10, 95% CI [6.12, 8.11]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4096)
##   - The effect of b bsTimeknotsEQc12 (Median = 9.57, 95% CI [7.50, 11.71]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 3887)
##   - The effect of b bsTimeknotsEQc13 (Median = -2.70, 95% CI [-6.18, 0.81]) has a
## 93.15% probability of being negative (< 0), 92.03% of being significant (<
## -0.14), and 84.63% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4833)
##   - The effect of b bsTimeknotsEQc14 (Median = 1.96, 95% CI [0.82, 3.04]) has a
## 99.97% probability of being positive (> 0), 99.90% of being significant (>
## 0.14), and 96.95% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 2623)
##   - The effect of b Wt (Median = 0.05, 95% CI [-0.32, 0.43]) has a 61.55%
## probability of being positive (> 0), 31.50% of being significant (> 0.14), and
```

```
## 0.00% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2604)
##   - The effect of b Dose (Median = 1.02, 95% CI [-3.94, 5.69]) has a 66.50%
## probability of being positive (> 0), 64.08% of being significant (> 0.14), and
## 52.77% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2585)
##
## Following the Sequential Effect eXistence and sIgnificance Testing (SEXIT)
## framework, we report the median of the posterior distribution and its 95% CI
## (Highest Density Interval), along the probability of direction (pd), the
## probability of significance and the probability of being large. The thresholds
## beyond which the effect is considered as significant (i.e., non-negligible) and
## large are |0.14| and |0.86| (corresponding respectively to 0.05 and 0.30 of the
## outcome's SD). Convergence and stability of the Bayesian sampling has been
## assessed using R-hat, which should be below 1.01 (Vehtari et al., 2019), and
## Effective Sample Size (ESS), which should be greater than 1000 (Burkner,
## 2017)., We fitted a Bayesian linear mixed model (estimated using MCMC sampling
## with 2 chains of 5000 iterations and a warmup of 2000) to predict conc with
## Time, Wt and Dose (formula: conc ~ bs(Time, knots = c(1)) + Wt + Dose). The
## model included Subject as random effect (formula: ~1 | Subject). Priors over
## parameters were set as normal (mean = 0.00, SD = 5.00) distributions. The
## model's explanatory power is substantial (R2 = 0.78, 95% CI [0.74, 0.81], adj.
## R2 = 0.76) and the part related to the fixed effects alone (marginal R2) is of
## 0.72 (95% CI [0.67, 0.76]).  Within this model:
##
##   - The effect of b Intercept (Median = -8.85, 95% CI [-55.72, 40.28]) has a
## 64.50% probability of being negative (< 0), 64.35% of being significant (<
## -0.14), and 63.10% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4384)
##   - The effect of b bsTimeknotsEQc11 (Median = 7.10, 95% CI [6.12, 8.11]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4096)
##   - The effect of b bsTimeknotsEQc12 (Median = 9.57, 95% CI [7.50, 11.71]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 3887)
##   - The effect of b bsTimeknotsEQc13 (Median = -2.70, 95% CI [-6.18, 0.81]) has a
## 93.15% probability of being negative (< 0), 92.03% of being significant (<
## -0.14), and 84.63% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4833)
##   - The effect of b bsTimeknotsEQc14 (Median = 1.96, 95% CI [0.82, 3.04]) has a
## 99.97% probability of being positive (> 0), 99.90% of being significant (>
## 0.14), and 96.95% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 2623)
##   - The effect of b Wt (Median = 0.05, 95% CI [-0.32, 0.43]) has a 61.55%
## probability of being positive (> 0), 31.50% of being significant (> 0.14), and
## 0.00% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2604)
##   - The effect of b Dose (Median = 1.02, 95% CI [-3.94, 5.69]) has a 66.50%
## probability of being positive (> 0), 64.08% of being significant (> 0.14), and
## 52.77% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2585)
##
```

```
## Following the Sequential Effect eXistence and sIgnificance Testing (SEXIT)
## framework, we report the median of the posterior distribution and its 95% CI
## (Highest Density Interval), along the probability of direction (pd), the
## probability of significance and the probability of being large. The thresholds
## beyond which the effect is considered as significant (i.e., non-negligible) and
## large are |0.14| and |0.86| (corresponding respectively to 0.05 and 0.30 of the
## outcome's SD). Convergence and stability of the Bayesian sampling has been
## assessed using R-hat, which should be below 1.01 (Vehtari et al., 2019), and
## Effective Sample Size (ESS), which should be greater than 1000 (Burkner,
## 2017)., We fitted a Bayesian linear mixed model (estimated using MCMC sampling
## with 2 chains of 5000 iterations and a warmup of 2000) to predict conc with
## Time, Wt and Dose (formula: conc ~ bs(Time, knots = c(1)) + Wt + Dose). The
## model included Subject as random effect (formula: ~1 | Subject). Priors over
## parameters were set as normal (mean = 0.00, SD = 5.00) distributions. The
## model's explanatory power is substantial (R2 = 0.78, 95% CI [0.74, 0.81], adj.
## R2 = 0.76) and the part related to the fixed effects alone (marginal R2) is of
## 0.72 (95% CI [0.67, 0.76]).  Within this model:
##
##   - The effect of b Intercept (Median = -8.85, 95% CI [-55.72, 40.28]) has a
## 64.50% probability of being negative (< 0), 64.35% of being significant (<
## -0.14), and 63.10% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4384)
##   - The effect of b bsTimeknotsEQc11 (Median = 7.10, 95% CI [6.12, 8.11]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4096)
##   - The effect of b bsTimeknotsEQc12 (Median = 9.57, 95% CI [7.50, 11.71]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 3887)
##   - The effect of b bsTimeknotsEQc13 (Median = -2.70, 95% CI [-6.18, 0.81]) has a
## 93.15% probability of being negative (< 0), 92.03% of being significant (<
## -0.14), and 84.63% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4833)
##   - The effect of b bsTimeknotsEQc14 (Median = 1.96, 95% CI [0.82, 3.04]) has a
## 99.97% probability of being positive (> 0), 99.90% of being significant (>
## 0.14), and 96.95% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 2623)
##   - The effect of b Wt (Median = 0.05, 95% CI [-0.32, 0.43]) has a 61.55%
## probability of being positive (> 0), 31.50% of being significant (> 0.14), and
## 0.00% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2604)
##   - The effect of b Dose (Median = 1.02, 95% CI [-3.94, 5.69]) has a 66.50%
## probability of being positive (> 0), 64.08% of being significant (> 0.14), and
## 52.77% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2585)
##
## Following the Sequential Effect eXistence and sIgnificance Testing (SEXIT)
## framework, we report the median of the posterior distribution and its 95% CI
## (Highest Density Interval), along the probability of direction (pd), the
## probability of significance and the probability of being large. The thresholds
## beyond which the effect is considered as significant (i.e., non-negligible) and
## large are |0.14| and |0.86| (corresponding respectively to 0.05 and 0.30 of the
## outcome's SD). Convergence and stability of the Bayesian sampling has been
```

```
## assessed using R-hat, which should be below 1.01 (Vehtari et al., 2019), and
## Effective Sample Size (ESS), which should be greater than 1000 (Burkner,
## 2017)., We fitted a Bayesian linear mixed model (estimated using MCMC sampling
## with 2 chains of 5000 iterations and a warmup of 2000) to predict conc with
## Time, Wt and Dose (formula: conc ~ bs(Time, knots = c(1)) + Wt + Dose). The
## model included Subject as random effect (formula: ~1 | Subject). Priors over
## parameters were set as normal (mean = 0.00, SD = 5.00) distributions. The
## model's explanatory power is substantial (R2 = 0.78, 95% CI [0.74, 0.81], adj.
## R2 = 0.76) and the part related to the fixed effects alone (marginal R2) is of
## 0.72 (95% CI [0.67, 0.76]).  Within this model:
##
##   - The effect of b Intercept (Median = -8.85, 95% CI [-55.72, 40.28]) has a
## 64.50% probability of being negative (< 0), 64.35% of being significant (<
## -0.14), and 63.10% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4384)
##   - The effect of b bsTimeknotsEQc11 (Median = 7.10, 95% CI [6.12, 8.11]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4096)
##   - The effect of b bsTimeknotsEQc12 (Median = 9.57, 95% CI [7.50, 11.71]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 3887)
##   - The effect of b bsTimeknotsEQc13 (Median = -2.70, 95% CI [-6.18, 0.81]) has a
## 93.15% probability of being negative (< 0), 92.03% of being significant (<
## -0.14), and 84.63% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4833)
##   - The effect of b bsTimeknotsEQc14 (Median = 1.96, 95% CI [0.82, 3.04]) has a
## 99.97% probability of being positive (> 0), 99.90% of being significant (>
## 0.14), and 96.95% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 2623)
##   - The effect of b Wt (Median = 0.05, 95% CI [-0.32, 0.43]) has a 61.55%
## probability of being positive (> 0), 31.50% of being significant (> 0.14), and
## 0.00% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2604)
##   - The effect of b Dose (Median = 1.02, 95% CI [-3.94, 5.69]) has a 66.50%
## probability of being positive (> 0), 64.08% of being significant (> 0.14), and
## 52.77% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2585)
##
## Following the Sequential Effect eXistence and sIgnificance Testing (SEXIT)
## framework, we report the median of the posterior distribution and its 95% CI
## (Highest Density Interval), along the probability of direction (pd), the
## probability of significance and the probability of being large. The thresholds
## beyond which the effect is considered as significant (i.e., non-negligible) and
## large are |0.14| and |0.86| (corresponding respectively to 0.05 and 0.30 of the
## outcome's SD). Convergence and stability of the Bayesian sampling has been
## assessed using R-hat, which should be below 1.01 (Vehtari et al., 2019), and
## Effective Sample Size (ESS), which should be greater than 1000 (Burkner,
## 2017)., We fitted a Bayesian linear mixed model (estimated using MCMC sampling
## with 2 chains of 5000 iterations and a warmup of 2000) to predict conc with
## Time, Wt and Dose (formula: conc ~ bs(Time, knots = c(1)) + Wt + Dose). The
## model included Subject as random effect (formula: ~1 | Subject). Priors over
## parameters were set as normal (mean = 0.00, SD = 5.00) distributions. The
```

```
## model's explanatory power is substantial (R2 = 0.78, 95% CI [0.74, 0.81], adj.
## R2 = 0.76) and the part related to the fixed effects alone (marginal R2) is of
## 0.72 (95% CI [0.67, 0.76]).  Within this model:
##
##   - The effect of b Intercept (Median = -8.85, 95% CI [-55.72, 40.28]) has a
## 64.50% probability of being negative (< 0), 64.35% of being significant (<
## -0.14), and 63.10% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4384)
##   - The effect of b bsTimeknotsEQc11 (Median = 7.10, 95% CI [6.12, 8.11]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4096)
##   - The effect of b bsTimeknotsEQc12 (Median = 9.57, 95% CI [7.50, 11.71]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 3887)
##   - The effect of b bsTimeknotsEQc13 (Median = -2.70, 95% CI [-6.18, 0.81]) has a
## 93.15% probability of being negative (< 0), 92.03% of being significant (<
## -0.14), and 84.63% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4833)
##   - The effect of b bsTimeknotsEQc14 (Median = 1.96, 95% CI [0.82, 3.04]) has a
## 99.97% probability of being positive (> 0), 99.90% of being significant (>
## 0.14), and 96.95% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 2623)
##   - The effect of b Wt (Median = 0.05, 95% CI [-0.32, 0.43]) has a 61.55%
## probability of being positive (> 0), 31.50% of being significant (> 0.14), and
## 0.00% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2604)
##   - The effect of b Dose (Median = 1.02, 95% CI [-3.94, 5.69]) has a 66.50%
## probability of being positive (> 0), 64.08% of being significant (> 0.14), and
## 52.77% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2585)
##
## Following the Sequential Effect eXistence and sIgnificance Testing (SEXIT)
## framework, we report the median of the posterior distribution and its 95% CI
## (Highest Density Interval), along the probability of direction (pd), the
## probability of significance and the probability of being large. The thresholds
## beyond which the effect is considered as significant (i.e., non-negligible) and
## large are |0.14| and |0.86| (corresponding respectively to 0.05 and 0.30 of the
## outcome's SD). Convergence and stability of the Bayesian sampling has been
## assessed using R-hat, which should be below 1.01 (Vehtari et al., 2019), and
## Effective Sample Size (ESS), which should be greater than 1000 (Burkner, 2017).
## and We fitted a Bayesian linear mixed model (estimated using MCMC sampling with
## 2 chains of 5000 iterations and a warmup of 2000) to predict conc with Time, Wt
## and Dose (formula: conc ~ bs(Time, knots = c(1)) + Wt + Dose). The model
## included Subject as random effect (formula: ~1 | Subject). Priors over
## parameters were set as student_t (location = 0.00, scale = 3.30) distributions.
## The model's explanatory power is substantial (R2 = 0.78, 95% CI [0.74, 0.81],
## adj. R2 = 0.76) and the part related to the fixed effects alone (marginal R2)
## is of 0.72 (95% CI [0.67, 0.76]).  Within this model:
##
##   - The effect of b Intercept (Median = -8.85, 95% CI [-55.72, 40.28]) has a
## 64.50% probability of being negative (< 0), 64.35% of being significant (<
## -0.14), and 63.10% of being large (< -0.86). The estimation successfully
```

```
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4384)
##    - The effect of b bsTimeknotsEQc11 (Median = 7.10, 95% CI [6.12, 8.11]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4096)
##    - The effect of b bsTimeknotsEQc12 (Median = 9.57, 95% CI [7.50, 11.71]) has a
## 100.00% probability of being positive (> 0), 100.00% of being significant (>
## 0.14), and 100.00% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 3887)
##    - The effect of b bsTimeknotsEQc13 (Median = -2.70, 95% CI [-6.18, 0.81]) has a
## 93.15% probability of being negative (< 0), 92.03% of being significant (<
## -0.14), and 84.63% of being large (< -0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 4833)
##    - The effect of b bsTimeknotsEQc14 (Median = 1.96, 95% CI [0.82, 3.04]) has a
## 99.97% probability of being positive (> 0), 99.90% of being significant (>
## 0.14), and 96.95% of being large (> 0.86). The estimation successfully
## converged (Rhat = 1.000) and the indices are reliable (ESS = 2623)
##    - The effect of b Wt (Median = 0.05, 95% CI [-0.32, 0.43]) has a 61.55%
## probability of being positive (> 0), 31.50% of being significant (> 0.14), and
## 0.00% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2604)
##    - The effect of b Dose (Median = 1.02, 95% CI [-3.94, 5.69]) has a 66.50%
## probability of being positive (> 0), 64.08% of being significant (> 0.14), and
## 52.77% of being large (> 0.86). The estimation successfully converged (Rhat =
## 1.000) and the indices are reliable (ESS = 2585)
##
## Following the Sequential Effect eXistence and sIgnificance Testing (SEXIT)
## framework, we report the median of the posterior distribution and its 95% CI
## (Highest Density Interval), along the probability of direction (pd), the
## probability of significance and the probability of being large. The thresholds
## beyond which the effect is considered as significant (i.e., non-negligible) and
## large are |0.14| and |0.86| (corresponding respectively to 0.05 and 0.30 of the
## outcome's SD). Convergence and stability of the Bayesian sampling has been
## assessed using R-hat, which should be below 1.01 (Vehtari et al., 2019), and
## Effective Sample Size (ESS), which should be greater than 1000 (Burkner, 2017).
```
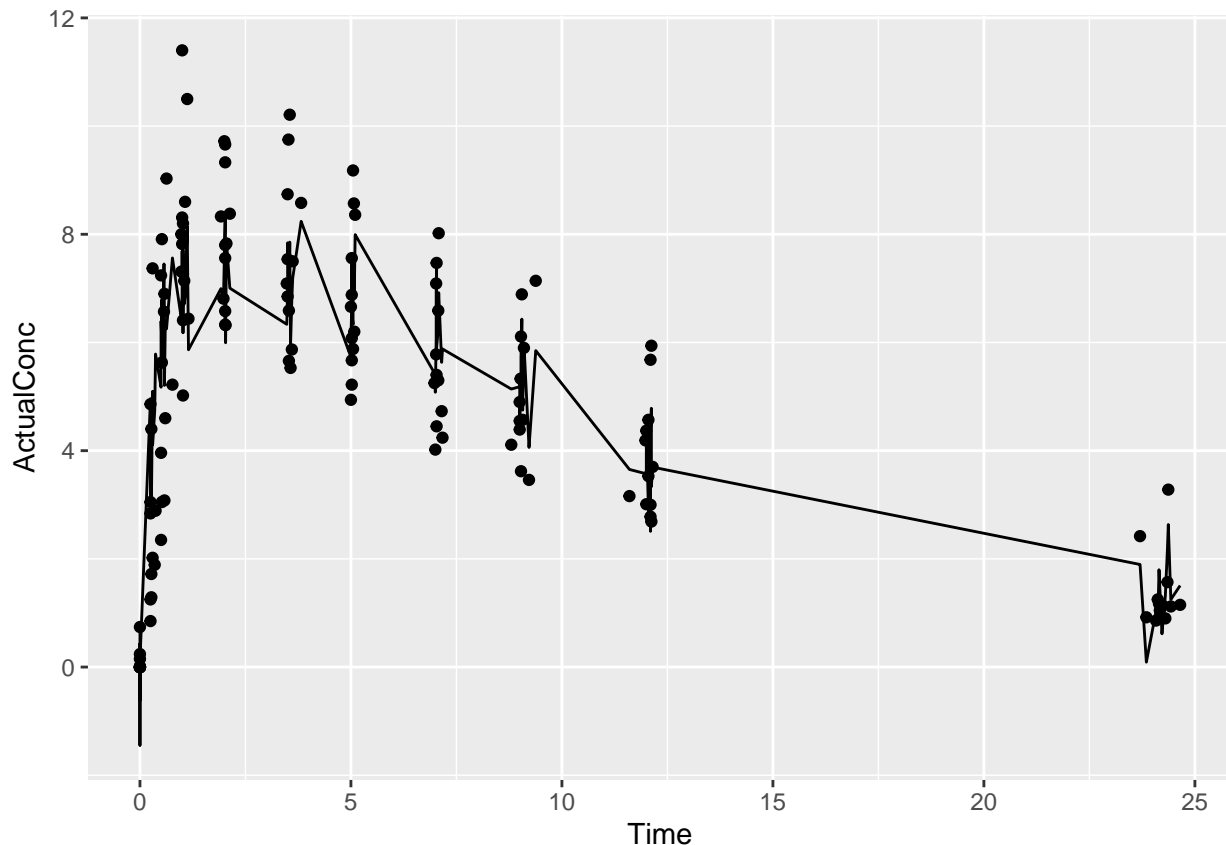
Plot the fitted model

```r
fitted_values <- fitted(model)
df_fitted <- data.frame(Time = df$Time, ActualConc = df$conc, FittedConc = fitted_values)

# plotting
ggplot(df_fitted, aes(x = Time)) +
  geom_point(aes(y = ActualConc)) + # actual data points
  geom_line(aes(y = FittedConc.Estimate)) # fitted spline curve
```

This fitted model had a lot of noise at specific time points. However, the model works well for our purpose. We can play around with refining prior distributions, setting more degrees of freedom for knots, or using cubic splines. We can use a quick and dirty technique to smoothen the curve - the Locally Estimated Scatterplot Smoothing (LOESS) technique.

Smoothening the noisy curve

```
df_fitted <- data.frame(Time = df$Time, FittedConc = fitted_values)

# apply a LOESS smoother to the fitted values
loess_fit <- loess(FittedConc.Estimate ~ Time, data = df_fitted, span = 0.4)  # ajust 'span' for smooth

# generate smooth predictions from the LOESS model
df_fitted$SmoothConc <- predict(loess_fit)

# plotting
ggplot(df, aes(x = Time)) +
    geom_point(aes(y = conc)) +  # actual data points
    geom_line(data = df_fitted, aes(y = SmoothConc), size = 1) # smoother curve
```

```
## Warning: Using `size` aesthetic for lines was deprecated in ggplot2 3.4.0.
## i Please use `linewidth` instead.
## This warning is displayed once every 8 hours.
## Call `lifecycle::last_lifecycle_warnings()` to see where this warning was
## generated.
```
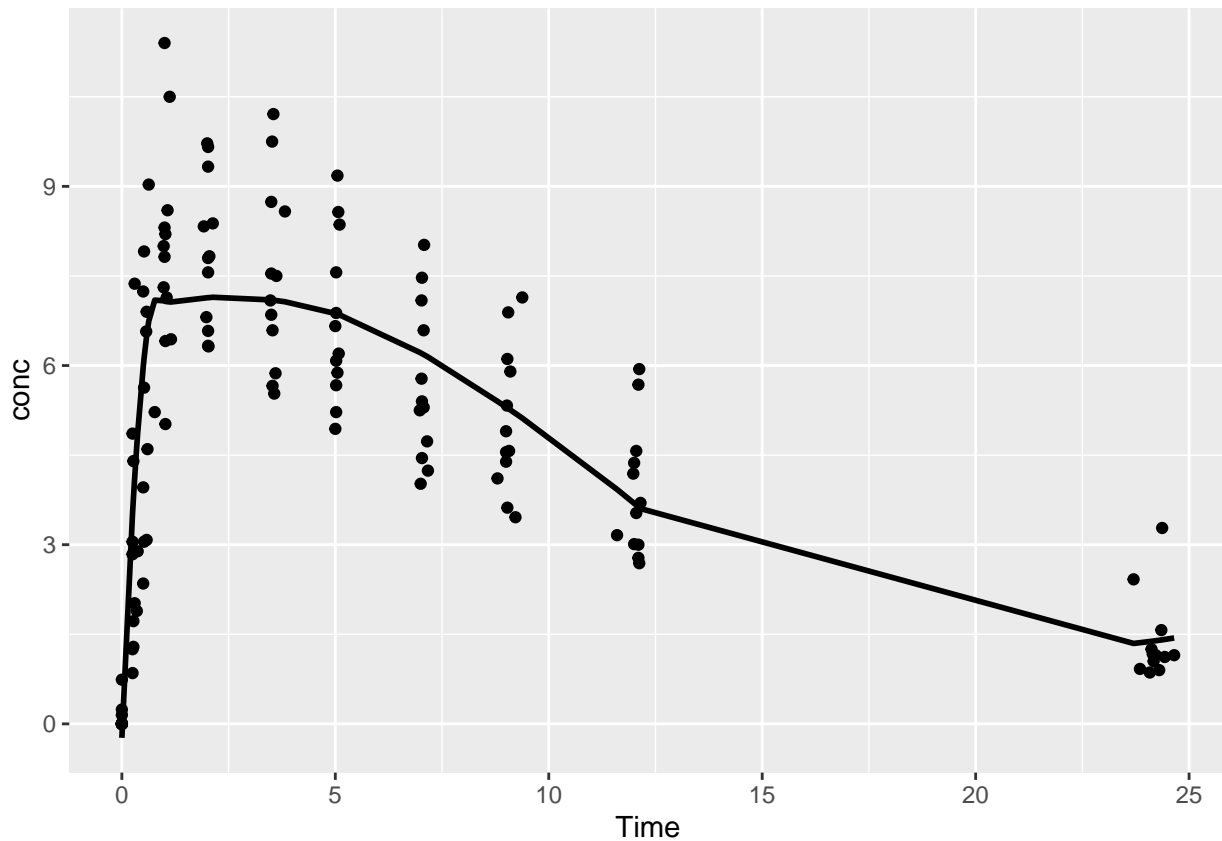
Visualizing prior and posterior distribution curves at various knotted time points

```r
posterior_samples <- as.data.frame(posterior_samples(model))
```
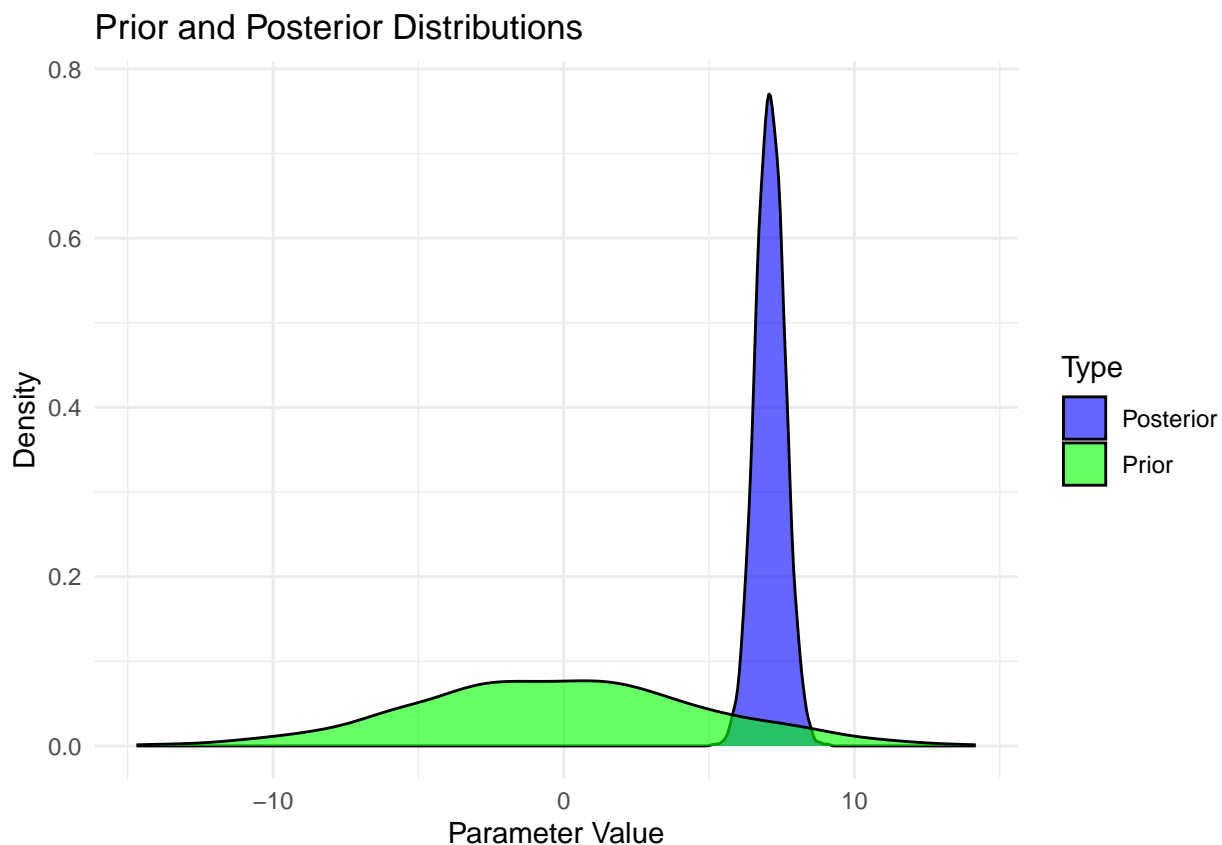
```
## Warning: Method 'posterior_samples' is deprecated. Please see ?as_draws for
## recommended alternatives.
```

```r
prior_distribution <- data.frame(Parameter = rnorm(1000, mean = 0, sd = 5))

posterior_distribution <- posterior_samples %>%
    select(b_bsTimeknotsEQc11) %>%
    rename(Parameter = b_bsTimeknotsEQc11)

combined_distribution <- rbind(
    data.frame(Parameter = prior_distribution$Parameter, Type = "Prior"),
    data.frame(Parameter = posterior_distribution$Parameter, Type = "Posterior")
)

ggplot(combined_distribution, aes(x = Parameter, fill = Type)) +
    geom_density(alpha = 0.6) +
    scale_fill_manual(values = c("blue", "green")) +
    labs(title = "Prior and Posterior Distributions", x = "Parameter Value", y = "Density") +
    theme_minimal()
```

Prior and Posterior Distributions


```r
posterior_samples <- as.data.frame(posterior_samples(model))
```

```
## Warning: Method 'posterior_samples' is deprecated. Please see ?as_draws for
## recommended alternatives.
```
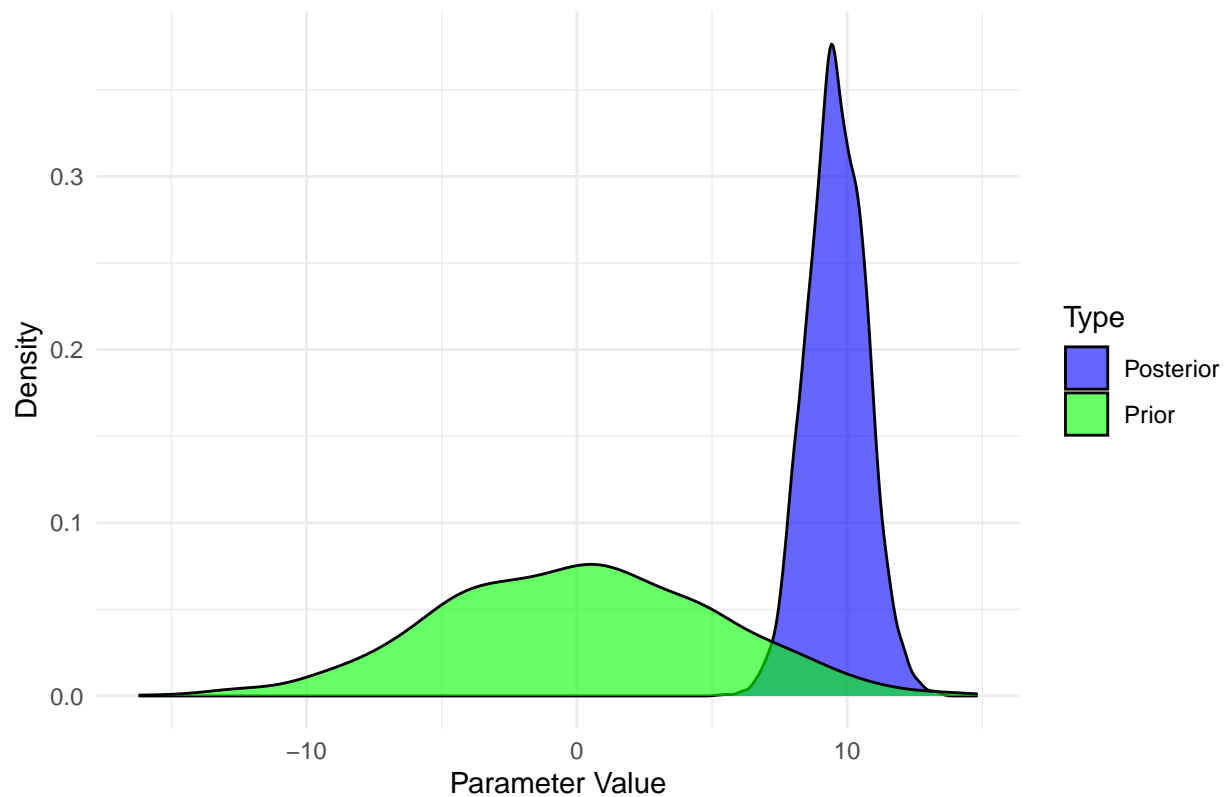
```r
prior_distribution <- data.frame(Parameter = rnorm(1000, mean = 0, sd = 5))

posterior_distribution <- posterior_samples %>%
    select(b_bsTimeknotsEQc12) %>%
    rename(Parameter = b_bsTimeknotsEQc12)

combined_distribution <- rbind(
    data.frame(Parameter = prior_distribution$Parameter, Type = "Prior"),
    data.frame(Parameter = posterior_distribution$Parameter, Type = "Posterior")
)

ggplot(combined_distribution, aes(x = Parameter, fill = Type)) +
    geom_density(alpha = 0.6) +
    scale_fill_manual(values = c("blue", "green")) +
    labs(title = "Prior and Posterior Distributions", x = "Parameter Value", y = "Density") +
    theme_minimal()
```

Prior and Posterior Distributions

```r
posterior_samples <- as.data.frame(posterior_samples(model))
```
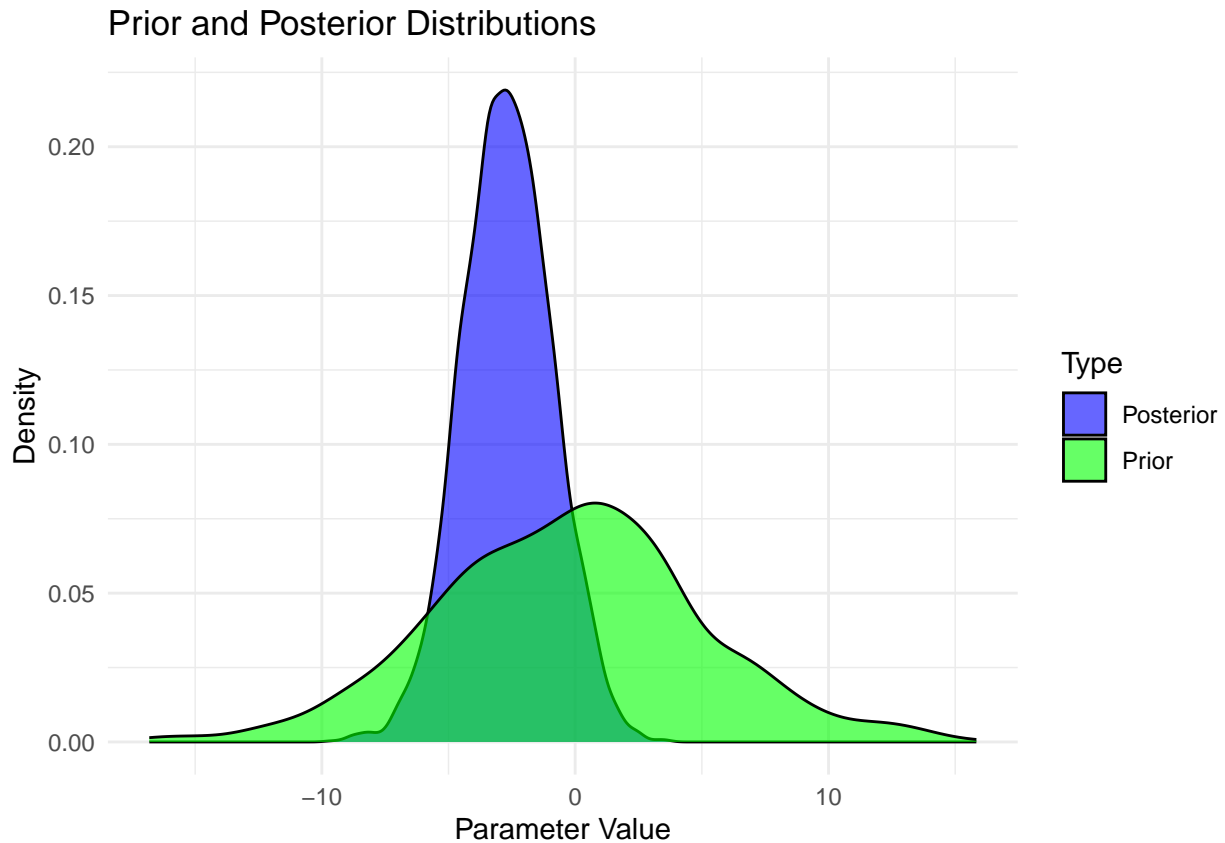
```
## Warning: Method 'posterior_samples' is deprecated. Please see ?as_draws for
## recommended alternatives.
```

```r
prior_distribution <- data.frame(Parameter = rnorm(1000, mean = 0, sd = 5))

posterior_distribution <- posterior_samples %>%
    select(b_bsTimeknotsEQc13) %>%
    rename(Parameter = b_bsTimeknotsEQc13)

combined_distribution <- rbind(
    data.frame(Parameter = prior_distribution$Parameter, Type = "Prior"),
    data.frame(Parameter = posterior_distribution$Parameter, Type = "Posterior")
)

ggplot(combined_distribution, aes(x = Parameter, fill = Type)) +
    geom_density(alpha = 0.6) +
    scale_fill_manual(values = c("blue", "green")) +
    labs(title = "Prior and Posterior Distributions", x = "Parameter Value", y = "Density") +
    theme_minimal()
```

Prior and Posterior Distributions

```
posterior_samples <- as.data.frame(posterior_samples(model))
```

```
## Warning: Method 'posterior_samples' is deprecated. Please see ?as_draws for
## recommended alternatives.
```

```
prior_distribution <- data.frame(Parameter = rnorm(1000, mean = 0, sd = 5))

posterior_distribution <- posterior_samples %>%
    select(b_bsTimeknotsEQc14) %>%
    rename(Parameter = b_bsTimeknotsEQc14)

combined_distribution <- rbind(
    data.frame(Parameter = prior_distribution$Parameter, Type = "Prior"),
    data.frame(Parameter = posterior_distribution$Parameter, Type = "Posterior")
)

ggplot(combined_distribution, aes(x = Parameter, fill = Type)) +
    geom_density(alpha = 0.6) +
    scale_fill_manual(values = c("blue", "green")) +
    labs(title = "Prior and Posterior Distributions", x = "Parameter Value", y = "Density") +
    theme_minimal()
```

Prior and Posterior Distributions