
 Poster

[P26-4] P26-4: Central nervous system drugs (3)

Chair: Christoph Hiemke, Germany

Tue. Sep 26, 2017 12:30 PM - 1:30 PM Annex Hall (1F)

(Tue. Sep 26, 2017 12:30 PM - 1:30 PM Annex Hall)

[P26-4-3] The estimation of pharmacokinetic parameters of lithium in children by pharmacokinetic modeling: a pilot study at Yuwaprasart Waithayopathum Child Psychiatric Hospital

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Keywords: Lithium, Pharmacokinetics, Children, Pharmacokinetic Modeling

Background

Since the dose adjustment of Lithium in pediatric patients still have problematic. In our hospital, more than half of the patients had serum Lithium outside therapeutic range (0.6-1.2 mg/L). The estimation of pharmacokinetic parameters of Lithium in children by pharmacokinetic modeling to produce the dose adjustment may be useful. Therefore, the purposed of the study was to estimate the pharmacokinetic parameters and calculate the precision (%RMSE) of the predicted Lithium concentration equation.

Method:

This research was study from blood Lithium levels of Children who had visit Yuwaprasart Waithayopathum Child Psychiartric Hospital during 1 January 2011-31 December 2009 by the following criteria; Age lower than 18 years, Receiving Litium continuously not less than 1 week, the laboratory of renal and liver function were available, good compliance. These blood levels information were used as a part of pharmacokinetic modeling process to establish the equation for prediction of Lithium concentrations in blood. The accuracy and precision of the equation were evaluated by the mathematical principle. The assist package software (WIN-NONLIN) was used to create pharmacokinetic parameters.

Results Twenty nine patients were recruited. The characteristics presented as the following; age (mean \pm SD) was 15.79 ± 2.64 years, weight (mean \pm SD) was 69.75 ± 22.28 kg and the average daily dose of Lithium (mean+ SD) was 858.62 ± 274.53 mg. The trough Lithium concentrations (mean \pm SD) was 0.56 ± 19 mg/L. The kinetic parameters (Mean \pm SD) presented as the following $t_{1/2}$ (7.23 ± 3.38 hr), K_e (0.12 ± 0.07 hr⁻¹), V_d (48.83 ± 15.60 L), AUC_{0-12} (11.01 ± 5.61 mg*hr/L). The population pharmacokinetic parameters; K_e , and $AUC(0-\infty)$ presented as 0.0151 hr⁻¹ and 169.8113 , mg*hr/L respectively. The modified equation from Yukawa equation for Lithium clearance calculation and prediction Lithium concentrations were CL (mL/min) = $[36.5 + (0.242 \times BW$ (kg) $- 7.79)] / Scr$ (mg/dL) and Lithium Concentration (mg/L) = $4 \times Dose$ (mg) / CL (mL/min) $\times 73.89$, respectively with 6.39 %RMSE (test for 20 patients data).

Conclusions

There were variation of Lithium pharmacokinetic parameters in children. The modified Yukawa equation may be able use for predict Lithium concentration with 6.39% RMSE. However, prospective use of this equation in large population is needed.