

ng·h/mL, and $t_{1/2}$ of 10.03 and 13.12 hours.

C_{max} and AUC values increased with dose level, but less than dose proportionally. No changes were noted in any parameters after repeated dosing.

In summary, C_{max} and \overline{AUC} values after single dosing increased with dose level; a dose proportionality relationship was observed between 3 and 10 mg/kg; however, both C_{max} and AUC values were almost leveled off at 30 and 60 mg/kg. At the dose level ranged 3 to 60 mg/kg, there were no sex differences in any parameters and exposure of TAK-070 M-II seemed not to differ after repeated dosing. Additionally, TAK-070 M-II showed biphasic elimination from plasma, although it was the evaluation was of data from a small number of sampling points in the terminal elimination phase.

10. References

- 1) Validation report, Study number P12-31801,
- 2) Control background data, Vol. 59, , 2011.

11. Personnel Engaged in the Study

Study Director:

MS, DJST

, Japan

Test Article Supervisor:

Analytical Supervisors

Analysis of Dosing Formulations:

Plasma Cortisol Measurement:

Animal Experiment Supervisor (dosing, clinical sign, body weight, food consumption, blood pressure, and respiration rate):

Electrocardiography:

Clinical Pathology Supervisor:

Necropsy Supervisor:

Pathological Specimen Preparation Supervisor:

Study Pathologist:

DVM, PhD, DJSTP, JCVP

Statistical Analysis Supervisor:

12. Storage Locations for Records, Data, and Specimens

All records and data, and the archive specimens will be stored at the locations in (, Japan) listed below for 10 years after preparation of this Final Report.

Archives of SNBL DSR

Protocol and amendments to the Protocol

Study schedule

Records and data on the test and control articles

Records and data on the test system

Records and data on maintenance conditions

Acclimation records

Dosing records

Clinical signs observation records

Body weight measurement records

Food consumption measurement records

Electrocardiography records
Blood pressure records
Respiration rate records
Hematology records
Blood chemistry records
Plasma cortisol measurement record
Necropsy records
Organ weight records
Histopathology records
Slide specimen preparation records
Records and data related to toxicokinetics
Statistical analysis records
Draft Final Report
Final Report
All other data related to the study

Organ Storage Room of ...
Formalin-fixed specimens (vacuum packed)
Paraffin-embedded specimens
Slide specimens

