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DEPARTMENT OF MEDICINE & ENVIRONMENTAL HEALTH

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March 23, 1981

Dr. David P. Martin, Director
LABORATORY ANIMAL SCIENCE & MEDICINE
LITTON BIONETICS, INC.
5516 Nicholson Lane
Kensington, Maryland 20795

RE: BO-77-2
LBI #20771

Dear David:

Based upon the recent conversations with either George Levinskas or myself, it is our understanding that the following modifications to the report entitled, "Effects of PCB on the Reproductive Performance of the Rhesus Monkey", have now been agreed upon.

PAGE 3, SECTION 2a, LAST PARAGRAPH

Replace the first sentence with: "Test animals were dosed for a period of six months before they were mated."

PAGE 6, SECTION 5b

The first five lines should be replaced with: "The study design called for measurement of various physiological parameters as well as physical ones, such as body weight, at fixed intervals. During the conduct of the study , however, female animals became pregnant at various times. This situation resulted in a potentially confounding variable, since pregnancy could effect many of the measurements. To minimize this effect, we identified hematology and biochemistry test results which....".

PAGE 12, SECTION IVA, 1a

The entire first paragraph should read: "Each male had been expected to produce at least two pregnancies from 16 mating periods, if there was no effect of test compound upon fertility and the animals performed according to colony history. None of the males was able to be scheduled for the predicted 16 meeting periods due to the lack of available females as a result of pregnancy. Extending the breeding period to May 31, 1978, to attempt to get more study females pregnant still did not allow 16

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mating periods to occur for the study males. Neverthless, the males exceeded the predicted conception rate and each male in each group produced at least two pregnancies.

PAGE 12, LAST LINE

...."1977 (Section IV D)."

PAGE 13, SECTION 4a

After (2) insert: "Statistical analysis of the data showed levels of significance in numerous categories. Examination of these instances revealed that the deviations fell into one of our categories:" Then replace (3), (4), (5) and (6) with (1), (2), (3), and (4) and insert: "Only those changes which fell into category (4) were considered further".

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Replace "test periods" or "testing periods" with "blood collection periods" in section (1), line 2 and section (2), lines 2 and 3.

In section (4) replace lines 3, 4, and 5 with: "Although no pattern of change was apparent, the frequency with which significant changes occurred suggests that a causal relationship to compound administration seems likely".

We prefer the word "suggests" to the phrase "was such that" in this sentence, but will accept the latter if you feel strongly about it.

PAGE 18, SECTION 5

What is now the first paragraph should read: "Births of the offspring from the study females began to occur in the fifth quarter of the study. Abnormal physical findings observed were as follows:

"In the high dose group there was only one birth. This was a live infant (B-8542) from female 312-1 which was observed to have swelling of the eyelids and lips at birth and at 30 days of age during the fifty quarter of the study."

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PAGE 27, PARAGRAPH 4, LINE 1

"compound-related" should be hyphenated and not spelled as one word.

It is our understanding that these changes in the text will not alter the meaning that you intend to convey but only provide clarification.

Unless there are still points of contention (in which case, please call me or George Levinskas), I suggest that the report now be issued in "final form" and twelve (12) copies sent along with the final invoice which we will forward for payment.

Sincerely yours,

Peter E. Berteau

cc: J. R. Craddock - MONSANTO COMPANY
Harvey Giss - LITTON BIONETICS, INC.
G. J. Levinskas- MONSANTO COMPANY