May 23, 1950.

Mr. Carlton Fredericks,
Institute of Nutrition Research,
62 West 45 Street,
New York 19, N.Y.

Dear Mr. Fredericks,

Thank you for your reply and extended reply to my hasty query concerning the character of the evidence for your statement that "the father's diet before conception also affects the child." I must certainly agree with you that this evidence is quite diffuse when it rests on a) primitive practices b) an irrelevant issue - the fertility of the male as influenced by nutrition, which can scarcely be questioned - and c) unpublished and undocumented experiments.

It may help a little to go over your letter paragraph by paragraph in numerical order.

P2. I am not sure what you mean by "Murphy's dictum". If you mean that genetic mutations can occur at all times, I agree. But then you must show that nutrition has a direct effect on mutation rates, a conclusion which few American geneticists will accept.

P3. Our medical library has a copy of a book entitled "Nutrition and Physical Degeneration" by Weston Price, published in 1932-40 by Heuber rather than by the American Academy of Applied Nutrition. Is this the book to which you refer? On page 342 there is a passing mention (not "details of an experiment") of a series of four litters by the same dachshund sire, in each of which there occurred pups with deformities. [There is a slight inconsistency: Fig. 120 refers to five bitches, but this may have been a typographical error.] There is no mention here of any evidence that the hereditary abnormalities transmitted by the propositus sire were induced by his nutritional status, nor does your letter refer to any sort of control, which might raise this observation to the status of an experiment. Prase's conclusion that the genotype of the male parent may influence congenital abnormalities is not objectionable.

P4. I would be very much interested to see the details of the experiments on cats — would you favor me with a reprint when they are published? This type of experiment requires a genetic purity of parental lines that I doubt has been attained in cats.

P5. While I agree that due consideration must be given to primitive practices in social organization and hygienic behavior, for civilized peoples have
a great deal to learn about how to live, you can hardly cite this as evi-
dence. After all, some primitive peoples have not connected intercourse
with conception, and others placate the gods of the elements, which does
not imply that our biology and our meteorology are false. But even if this
type of argument were acceptable, the examples cited by Price might be
directed towards the maintenance of fertility, nutritional effects on which
I could not controvert.

P.6. Agreed, or rather agreeable.

P.7. You are begging the question, not citing evidence.

P.8. This might be interesting, if there could be adequate controls. Was
the mother's nutrition independent of the father's? Are you ignoring the
discordant examples, of which there must be many, and selecting only those
chance sequences which fit the concept? This type of reasoning should be
backed up by a careful statistical analysis. Otherwise, anyone with a broad
experience can select, consciously or otherwise, those sequences which fit
his a priori concepts.

P.9. Perhaps they agree that the evidence is grossly inadequate.

P.10. I think you have misinterpreted the Drosophila experiments. I don't know
of anyone who has claimed to restore the normal genotype by nutritional means.
Some mutants may be modified in their immediate phenotypic expression by
nutrition, which is a very different thing. I am certainly unacquainted with
any suggestion of an influence of paternal nutrition.

P.11. Heredity is not immutable — at least with respect to spontaneous mutations,
and genetic changes induced by X-rays and certain very toxic chemicals.
I do not deny that the possibility exists of nutritional effects in the male,
but we are not discussing this. We are discussing whether there is any evidence
for it, and of this I am left entirely unconvincing. My colleagues in this de-
partment, many of whom are actively engaged both in nutritional and genetic
factors in animal reproduction, are of a like mind,

This criticism is perhaps close to carping, directed as it is against a
very narrow aspect of your highly commendable work. Certainly we do not wish
to neglect the nutritional status of fathers, for many reasons besides the
likelihood of fertility effects and the remote possibility of effects on con-
genital malformations. The lay public is however often poorly informed on
these matters, in particular with the current Lysenkist controversy. It is
important on that basis to be extremely cautious in propagating well-established
facts as distinguished from speculations. For my part, I would be extremely
hesitant to propound uncertain data in another specialty than my own, for example
nutrition.

Sincerely yours,

Joshua Lederberg
Assistant Professor
of Genetics