

To Dr. H. Gell

From Dr. H. C. Knowlton, Jr.

Date May 17, 1967

This concerns the research proposal of Drs. Ben Friedman and Eugene Sawyer entitled "The Therapeutic Effect of Total Body Irradiation Followed by Infusion of Stored Autologous Marrow in Humans". After study of the proposal and listening to testimony this morning from Drs. Ben Friedman and Harry Horwitz (serving as consultant) I give provisional approval to the proposal. I have two suggestions, however.

First, I believe a statement should be inserted to the effect that the radiation therapy to be used is an accepted form of treatment and that beneficial therapy is not being withheld.

Second, I believe a statement might be made that means of measurement of dependent variables in question are judged satisfactory to determine a difference between groups II and III. In addition, some statement might be made to the effect that sampling of the population will be at random and that all efforts will be made to reduce the number of independent variables and keep heterogeneity at a minimum. I assume all this is present in the investigators' minds. It would help to have it on paper.

I realize that my second request deals with experimental design, a matter which is really not the responsibility of the Committee. Nevertheless, we are balancing the good to come against hazards encountered, and the good to come is related to experimental design.

I believe that merely a few sentences can be added to the protocol to clarify these points. I am leaving all of this to your discretion, however.

H. C. Knowlton