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## IMMUNE AGENTS MAY PLAY Ca ROLE

Researchers find that antigens and antibodies can cause mitosis in cultures of human blood cells

Antigens and antibodies may play a role in leukocytosis and neoplastic disease.

This suggestion comes from two widely separated teams of researchers. A Finnish research group bases its hypothesis on the finding that antibodies can induce mitosis in human leukocytes. New Zealand investigators reached the same conclusion after getting similar results with the antigen tuberculin.

While the two groups have come to much the same general conclusions, both their experimental studies and their conclusions lie along different paths.

The Finnish workers— Drs. Ralph Gräsbeck, Clas Nordman and Albert de la Chapelle of Helsingfors' Minerva Foundation Institute for Medical Research and the Folkhälsan Institute for Genetics—immunized rabbits with a series of five intravenous injections of a preparation of whole leukocytes obtained from five healthy human volunteers. The resulting antisera, harvested

ten days after the last injection, were added to cultures of human peripheral leukocytes.

For a control series, serum samples were taken from some of the rabbits before immunization; one rabbit was "immunized" with saline; and serum samples from other nonimmunized rabbits were used.

Dr. Gräsbeck's group found that four out of five antisera induced mitoses in the leukocyte cultures. The fifth antiserum had a toxic effect only.

"The results demonstrate a clear mitogenic action," they report in the *Lancet*, adding that "since the injected leukocyte preparations contained other blood cells, one cannot say against which cell type the antibody responsible for the mitogenic activity was directed."

It is possible that leukocyte antigens are liberated in rabbits and are contained in the resulting antiserum, but at present the Finnish investigators believe that an antileukocytic immune globulin is responsible for the effect. "Our view is that some kind of biochemical chain reaction is involved when a cell is stimulated to divide. Apparently, numerous stimuli can trigger this reaction, and antigens acting on presensitized cells are one such group."

Such stimuli, they add, could include both phytohemagglutinin (a well-known inducer of cell division) and antileukocytic immune serum. "Adhering to the cell surface, they cause the cell to divide, no presensitization apparently being required."

## Findings from New Zealand

The New Zealand investigators—Drs. R. R. Lycette, G. E. Pearmain and P. H. Fitzgerald—based their studies on the hypothesis that the mitogenic action of phytohemagglutinin (PHA) might have an immunological basis.

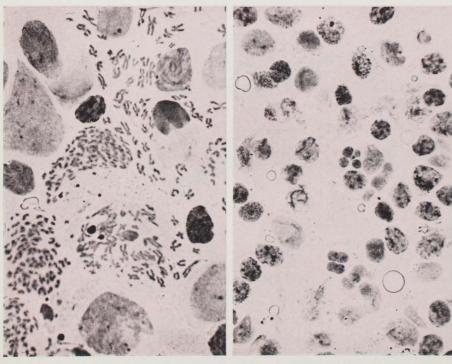
Substituting tuberculin for PHA in peripheral blood leukocyte cultures made from individuals sensitized to tuberculin, they discovered mitoses similar to those obtained with PHA.

They then took lymphocytes from five patients with severe hay fever, and cultured them with grass pollens. Again, they found that blast transformation cells and mitoses occurred consistently after four to six days of incubation at 37° C. Lymphocytes cultured from normal persons as controls showed no such effect.

Further studies on lymphocyte cultures with Sabin polio vaccine also produced blast transformation cells and mitoses in four to five days.

These findings, they suggest, may be a clue to a possible explanation of the abnormal blood pictures in such conditions as sarcoidosis, berylliosis, rheumatoid arthritis and Hashimoto's disease, where the lymphocyte is histologically predominant.

They also point out that the abnormal leukocytes encountered in routine differential counts in young children, particularly those suffering from infections, may be the equivalent of blast transformation cells present in lymphocyte-antigen cultures.



**LEUKOCYTES** cultured in antileukocyte serum show numerous mitoses with blast formations (left). Effect is absent in cells that are grown with nonimmunized serum (right).

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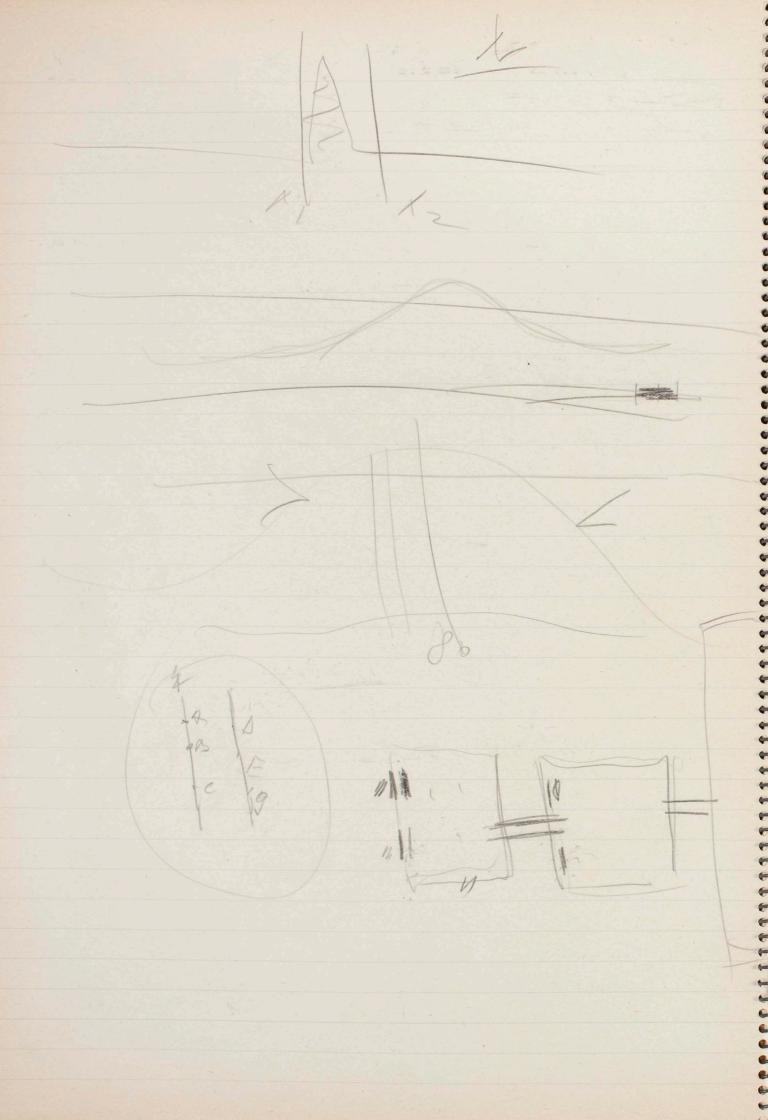
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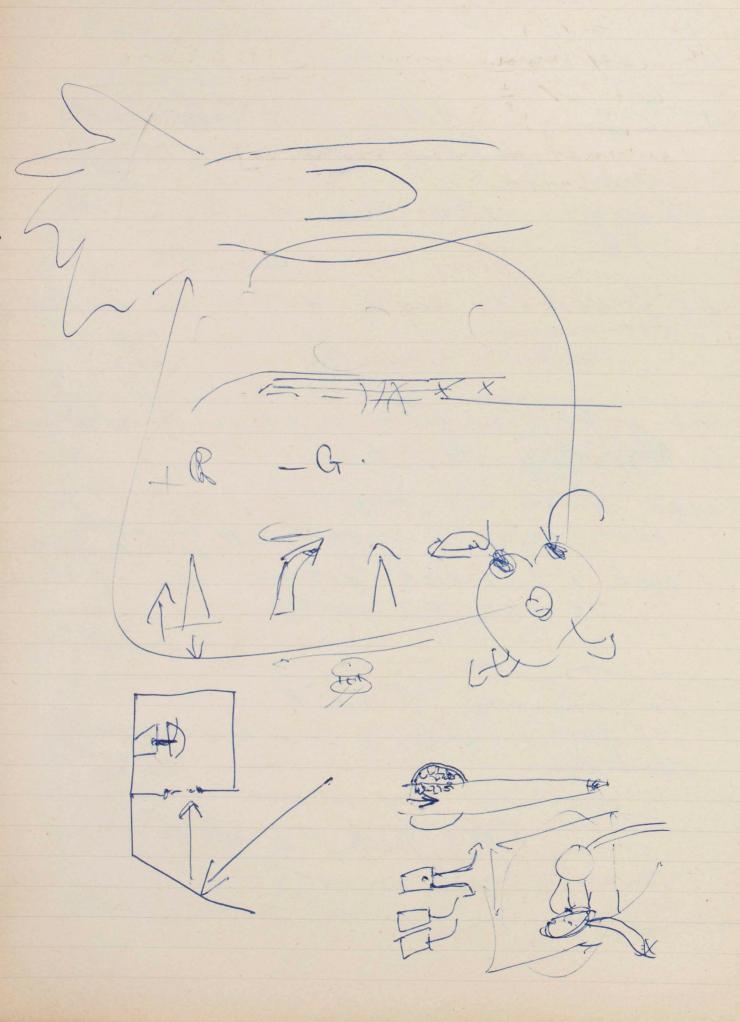
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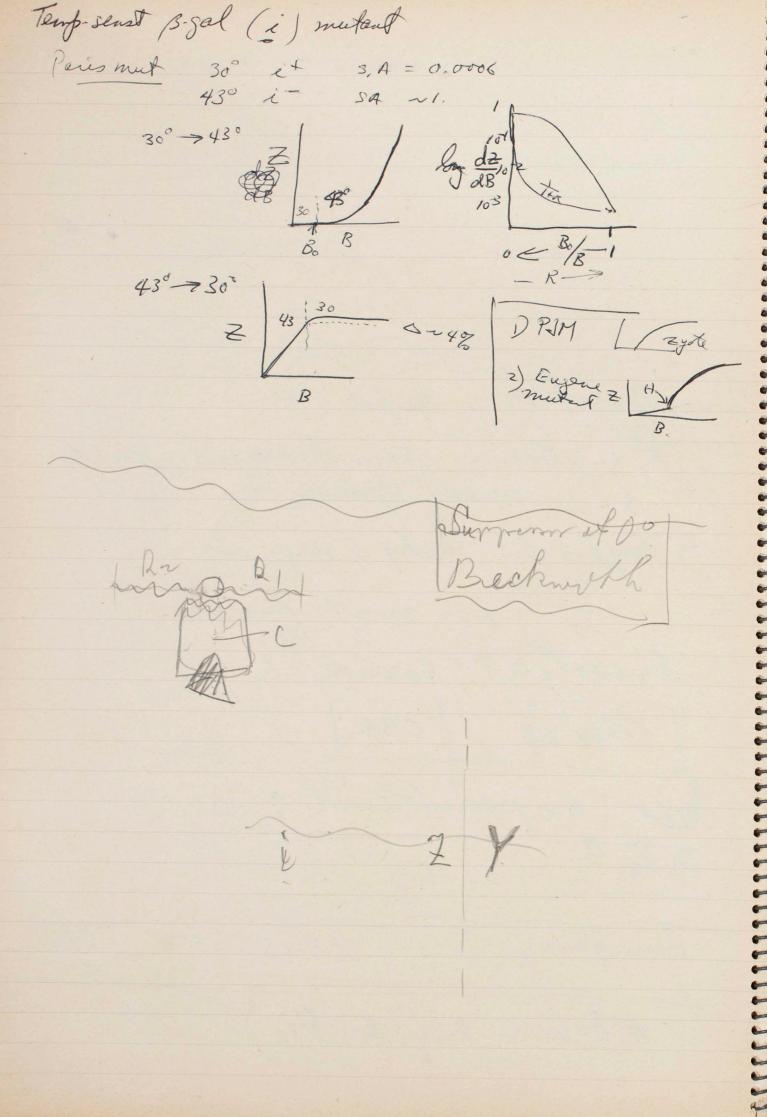
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