Brief Case Report

Syndrome of Erythremia di Guglielmo after Lightning Injury with Autoimmune Antibodies and Terminating in Acute Monocytic Leukemia

By Lie-Injo Luan Eng and C. Sinnadurai

ALTHOUGH LEUKEMIA has been frequently recorded following exogenous influences, there is no conclusive evidence for the direct etiologic relationship of these to leukemia. Damage to the hemopoietic system ranging from total aplasia to different kinds of leukemia has been observed in survivors of atomic bomb explosions, \(^1\) including a case of monocytic leukemia. Leukemia and other hematologic disorders have been described following the medical use of x-radiation. \(^2\) Johnson \(^3\) reported a case of acute myelomonocytic leukemia after radioiodine therapy for hyperthyroidism. Leukemia following chronic exposure to benzene \(^4\) and following trauma \(^5\), \(^6\) has been described. Infectious agents have also been suggested as causes of leukemia. \(^7\)

In this paper we present a case in which a patient, after being struck by lightning, developed the syndrome of erythremia di Guglielmo, \(^8\) with autoimmune hemolytic anemia, which terminated in acute monocytic leukemia.

CASE REPORT

An Indian man, 60 years old, was admitted to the General Hospital, Kuala Lumpur on December 17, 1963, with the following history. He had been in good health and was able to work normally until approximately 6 to 7 months prior to his hospital admission, when he was suddenly struck by lightning in the open and fell unconscious. Unfortunately no further details were obtainable about the accident or his condition immediately following it by the time he came for treatment. Following the accident, he was unable to work due to generalized weakness and severe anorexia. After 2 months, he resumed work but did not feel the same as before. His wife noticed that he became progressively pale and developed swelling of the legs. On admission he was found to be very anemic, with generalized edema and ascites. Lymph glands were not enlarged. His temperature was subfebrile. His lungs were normal, heart slightly enlarged and systolic murmur was heard over the entire area. Spleen was enlarged 3 fingers and liver 4 fingers below costal margin. Hemoglobin level was 3.0 Gm. per cent, PCV, 11 per cent; MCHC, 27.3 per cent; reticulocytes, 4.6 per cent; nucleated cells, 9000 per cu. mm. There were nucleated red blood cells in the peripheral blood, some of them resembling megaloblasts. Indirect bilirubin was 2.1 mg. per cent, and liver function tests were normal. He was given a course of daily injections of Imferon, folic acid and vitamin B12; however, there was no response and because of his critical condition, he was given a transfusion with packed red blood cells. Rise of hemoglobin level was soon followed by a drop. Findings on

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January 6, 1964 were: Hb, 2.2 Gm. per cent; RBC, 0.70 million per cu. mm.; PCV, 7.5 per cent; MCV, 107.1 cu. μ.; MCH, 31.4 μg.; MCHC, 29.3 per cent; reticulocytes, 9.5 per cent; platelets 50,000 per cu. mm.; nucleated blood cells, 20,300 per cu. mm.; differential counts, PS 67, Eos. 1, Mono 5, Ly 27. Normoblasts were seen, many of them resembling megaloblasts. No malarial parasites were seen. Alkaline phosphatase activity of the leukocytes showed a scoring of 162 examined by the method of Kaplow. Serum bilirubin of the indirect type was increased. Direct Coombs test was weakly positive. Cold agglutinin at 5 C. using normal blood group O blood cells in saline showed a titer of 32. With trypsinized red cells, the titer was 512. At 20 C., the titer was respectively negative and 128. At 37 C. negative, with normal as well as with trypsinized red cells, hemolysins were not detected. Motulsky test for G-6-PD deficiency was negative. Hemoglobin analysis showed normal Hb A and normal alkali denaturation. Kahn test of the serum was negative. The bone marrow showed severe hyperactivity of erythropoiesis with basophilic stages predominating and many resembling megaloblasts. Myelopoiesis was relatively depressed and only a few megakaryocytes were seen. Intensive search for malignant tumors was unsuccessful. Because of the presence of autoimmune antibodies, 60 mg. prednisolone daily were tried and blood transfusion given. After a transient rise of the hemoglobin level to 3.6 Gm. per cent, it dropped again, so that another blood transfusion was necessary and prednisolone dosage was increased to 80 mg. daily. Although the hemoglobin level did not drop appreciably, it also did not rise. The blood was studied on numerous occasions with the following hematologic findings on February 3, 1964: Hb 4.2 Gm. per cent; RBC, 1.58 million per cu. mm.; PCV, 13.5 per cent; MCV, 84.8 cu.; MCH, 26.6 μg. per cent; MCHC, 31.1 per cent; nucleated blood cells 5700 per cu. mm.; differential counts, PS 47, Eos 3, Mono 7, Ly 43; 22 normoblasts per 100 WBC. Erythroid hyperplasia of the peripheral blood with bizarre-looking normoblasts and thrombocytopenia were the outstanding features of the disease. Although occasional myelocytes were seen in the peripheral blood, the picture was never leukemic at this stage. On February 24, 2 days before the patient expired, the blood picture was remarkable. There were numerous bizarre-looking normoblasts, some extremely large and many with multiple nuclei; some were difficult to classify (fig. 1). The number of monocytes was much increased so that monocytic leukemia was suspected. A few myelocytes were also present. On February 26, the day the patient died, the peripheral blood picture was that of a full-blown monocytic leukemia (fig. 2), confirmed by the finding of a bone marrow teeming with monocytes and monoblasts. Autopsy was unfortunately refused by the family. Spleen and liver puncture aspirates showed numerous monocytes and monoblasts.

Fig. 1.—Peripheral blood 2 days before patient died, showing many bizarre-looking nucleated cells.
SYNDROME OF ERYTHEMIA DI GUGLIELMO

Fig. 2.—On the day the patient died the peripheral blood picture was that of a full-blown monocytic leukemia.

DISCUSSION

The symptoms in this patient resembled those found in the di Guglielmo syndrome, especially toward the end of the disease before it changed into acute monocytic leukemia—pronounced erythroblastosis, thrombocytopenia and hepatosplenomegaly. Megaloblastic changes in bone marrow and peripheral blood resistant to folic acid and vitamin $\text{B}_{12}$ seen in this patient have also been noticed by others in di Guglielmo disease.

Although it has not been proved that lightning caused or precipitated the blood changes, it is thought not to be improbable, since a striking change in the patient's condition was noticed by himself and his family only after he was struck by lightning.

The presence of autoimmune antibodies with hemolytic anemia in our patient is unusual. It might be explained, that injury to the patient's hemopoietic system resulted in the formation of blood cells which became foreign to his own body and led to the formation of autoimmune antibodies which acted against his own red blood cells.

The sudden change from a severe erythroblastosis with bizarre-looking nucleated red blood cells to a full-blown acute monocytic leukemia is noteworthy. The idea that the di Guglielmo syndrome is merely a phase in a myelo-proliferative disease usually ending in leukemia is gaining more and more ground. If no blood examination had been carried out the last few days before the patient expired, the case would have been classified simply as erythremia di Guglielmo and the true nature of the disease would not have been discovered. A transition from erythremia di Guglielmo to myelo- or monoblastic leukemia has also been observed by Block et al.

SUMMARY

A case of lightning injury is described associated with changes of the hemopoietic system resembling di Guglielmo's syndrome. Megaloblasts were found...
in the peripheral blood and bone marrow, resistant to folic acid and vitamin 
B<sub>12</sub> treatment. The presence of auto-immune antibodies in this patient was 
accompanied by a hemolytic condition not responding to treatment with high 
doses of prednisolone. Towards the end of the disease the most bizarre-looking 
erthroblasts were seen in the peripheral blood. Shortly before death the 
condition suddenly changed into an acute monocytic leukemia. Although it 
was not proved that the lightning injury caused or precipitated the blood 
changes, it is thought not to be improbable.

**SUMMARIO IN INTERLINGUA**

Es describite un caso de traumatismo per fulmine. Illo eseva characterisate 
per alterationes del systema hematopoietic resimilante le syndrome de di 
Guglielmo. Megaloblastos eseva trovate in sanguine periphenic e in medulla 
ossee. Illos se monstrava resistente contra tractamento con acido folièe e vita-
mina B<sub>12</sub>. Le presentia de anticorpore auto-immun in iste patiente eseva 
accompaniate de un condition hemolytic que non respondeva al tractamento 
con alte doses de prednisolona. Verso le fin del maladia, erythroblastos de 
bizzarissime apparentia eseva trovate in le sanguine peripheric. Brevemente 
ante le morte del patiente, su condition transiva subitemente ad in un acute 
leucemia monocytic. Ben que il non eseva provate que le traumatismo fulmi-
nal causava o precipitava le alteraciones in le sanguine, un tal nexo causal 
non pare improbabile.

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