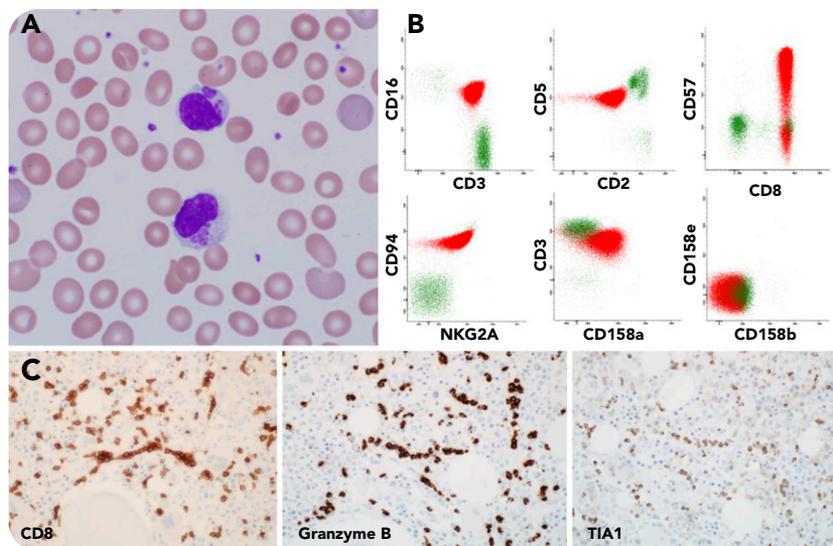




Giant intracytoplasmic inclusions in a T-cell large granular lymphocytic leukemia patient with acute severe anemia

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A 54-year-old man presented with shortness of breath and light-headedness. Complete blood count (CBC) showed anemia (hemoglobin, 5.0 g/dL) and lymphocytosis (lymphocyte count, $7.9 \times 10^9/L$). Peripheral blood smears (panel A; original magnification $\times 1000$, hematoxylin and eosin stain) showed increased large granular lymphocytes ($7.3 \times 10^9/L$) with giant reddish-purple cytoplasmic inclusions. Flow cytometry (panel B) revealed $CD3^+CD8^+$ T cells (77% of total events) with dim expression of CD2 and CD5 and co-expression of natural killer cell-associated markers CD16, CD57, CD94, and NKG2A. Clonality was established by restricted killer-cell immunoglobulin-like receptor CD158a and clonal T-cell receptor gene rearrangement. STAT3 mutation was detected. Bone marrow biopsy (panel C; original magnification $\times 400$, immunohistochemical stain) showed intrasinusoidal infiltrates of $CD8^+$, granzyme B⁺,

and TIA1⁺ T cells. T-cell large granular lymphocytic leukemia (T-LGLL) was diagnosed, and cyclophosphamide was administered. CBC normalized upon 5 months of treatment and no cytoplasmic inclusions were identified in the subsequent smears, indicating an excellent response to therapy.

T-LGLL is an indolent disorder of persistent cytotoxic T-cell expansion with coarse/fine cytoplasmic azurophilic granules containing cytolytic proteins. The giant cytoplasmic inclusions are rarely seen in T-LGLL, likely representing a substantial accumulation of cytolytic proteins in the setting of acute presentation. The inclusions subsided after therapy, underscoring their association with disease status. Recognizing this phenomenon in T-LGLL is important for accurate diagnosis and treatment.



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