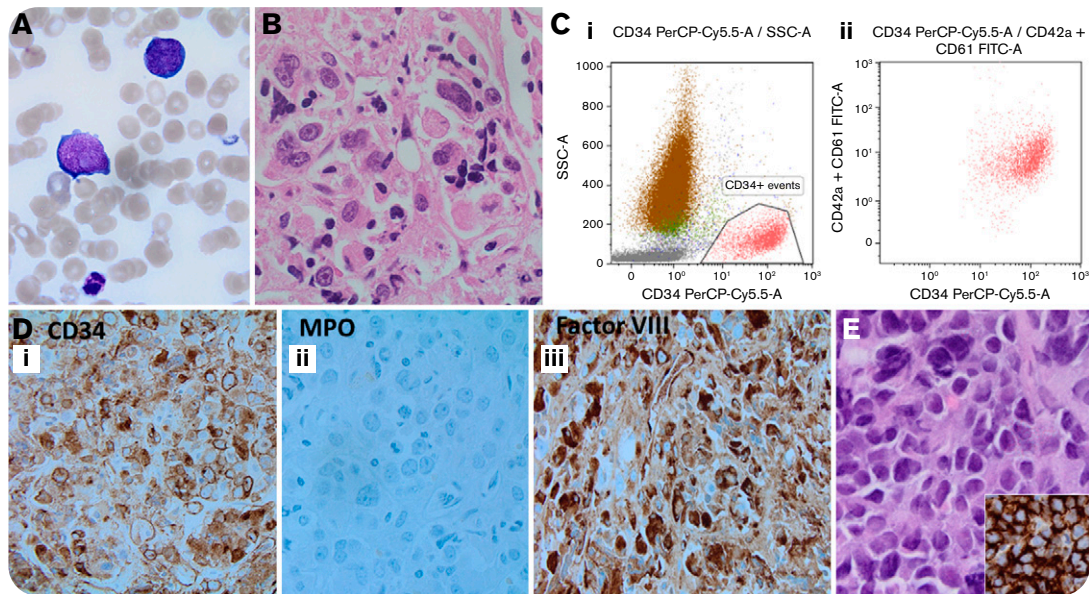


## CML in megakaryocytic transformation and extramedullary infiltration

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A 78-year-old man with a diagnosis 1 month ago of asymptomatic treatment-naive chronic myeloid leukemia (CML) presented with back pain and dyspnea. Peripheral blood revealed a white blood cell count of  $23.8 \times 10^9/L$  (neutrophils,  $22.8 \times 10^9/L$ ; basophils,  $0.0 \times 10^9/L$ ; lymphocytes,  $0.8 \times 10^9/L$ ), hemoglobin count of 68 g/L, and platelet count of  $6 \times 10^9/L$ , with scattered large mononuclear cells with open chromatin, prominent nucleoli, and deep basophilic cytoplasm with cytoplasmic blebs (panel A; original magnification  $\times 100$ , May-Grünwald-Giemsa stain). The marrow aspirate was hemodilute; trephine biopsy showed hypercellularity with extensive fibrosis and sheets of blasts (panel B; original magnification  $\times 60$ , hematoxylin and eosin stain). Flow cytometry (panel C i-ii) and immunohistochemistry (panels D i-D iii; original magnification  $\times 40$ ) showed that these blasts were positive for CD45 (dim), CD34, CD33, CD42a, CD61,

and factor VIII and negative for myeloperoxidase. In addition, a biopsy of a lytic lesion of the ilium bone revealed sheets of large blasts (panel E; original magnification  $\times 40$ ) that were positive for CD34 (panel E inset, original magnification  $\times 40$ ) and CD45 (dim), consistent with a myeloid sarcoma. Cytogenetics revealed  $t(9;22)$  and a complex karyotype. Next-generation sequencing detected *TP53* mutation only. A diagnosis of CML in megakaryoblastic crisis was rendered, and the patient succumbed to his illness 9 days later.

The most common blast crisis in CML is myeloid and then lymphoid, but very rarely, megakaryocytic, especially with extramedullary manifestation, which has an even poorer prognosis. This case also emphasizes the importance of performing megakaryocytic markers in the workup for acute leukemia, using flow cytometry and/or immunohistochemistry.