#### **IMAGES IN HEMATOLOGY**

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### Acute Promyelocytic Leukemia with Basophilic Differentiation A Rare Variant

#### Arias A.F.P. et al.: Acute Promyelocytic Leukemia with Basophilic Differentiation A Rare Variant

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A 46-year-old man presented after a colon surgery with hemoglobin, neutrophils and platelet counts of 106g/L, 0.63x10<sup>9</sup>/L, and 102x10<sup>9</sup>/L, respectively. Coagulation tests showed aPTT, PT, Clauss fibrinogen and D-dimer of 26.9s, 15.4s, 164mg/dL, and 36,727ng/mL. Peripheral blood smear revealed anisopoikilocytosis, leukoerythroblastic reaction, and blasts with azurophilic granules, without Auer rods. Bone marrow aspirates showed 60% cells being abnormal promyelocytes with azurophilic granules peroxidase-staining positive (Figure 1). Inmunophenotyping detected 76% of CD117+, CD13+(dim), CD33+(bright), CD56+, CD123+(dim), CD9+(dim), MPO+, CD34-, HLA-DR-, and CD203c- abnormal promyelocytes, 4% of these cells showed CD34+(10%), CD35+, CD22+, CD203c+, CD9+, CD123+(bright), MPO+, CD117-, and HLA-DR- (Figure 2). Tryptase levels were 7.55μg/L. RT-PCR demonstrated a bcr3 PML/RARA transcript without additional alterations in conventional karyotype or FISH. An acute promyelocytic leukemia (APL) with basophilic-differentiation was diagnosed. Treatment with ATRA and ATO was initiated, reaching morphologic CR, negative MRD, and 0.1% PML/RARA gene rearrangement. No bleeding complications or hyperhistaminemia-related symptoms occurred during treatment.

APL with basophil-maturation represents one-third of APL. Basophil-associated markers, CD203c and CD22, have predictive ability for increased bleeding risk and reduced overall survival. Severe bleedings are still the main cause of treatment failure and early mortality in APL patients, making the diagnosis of this variant important 1-5.

## Ethics

All procedures performed in studies involving human participants were in accordance with the ethical standards of the institutional and/or national research committee and with the 1964 Helsinki declaration and its later amendments or comparable ethical standards.

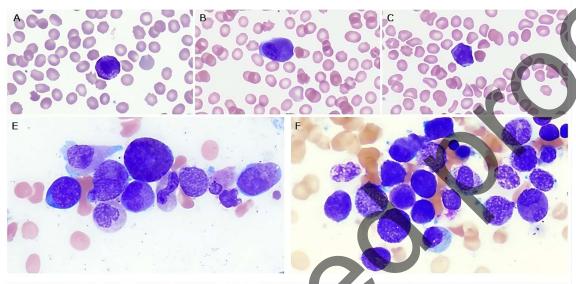
**Informed Consent:** Informed consent was obtained from the participant included in the study. **Authorship Contributions** 

AFMA and SES contributed equally as co-first authors. AFMA, SES and FAGF designed the research, analyzed the data and wrote the paper. MPZ and CBC requested the studies. CGS and MAG performed the flow cytometry analysis. EA accomplished the molecular and genetic studies. All authors read and approved the manuscript.

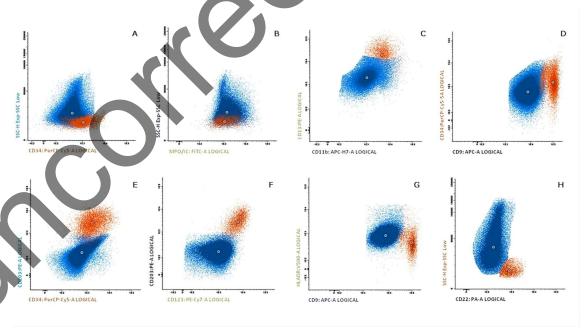
# References

- 1. Matarraz S, Leoz P, Fernández C, et al. Basophil-lineage commitment in acute promyelocytic leukemia predicts for severe bleeding after starting therapy. Modern Pathology. 2018;31(8):1318–31. doi:10.1038/s41379-018-0038-2
- 2. Shameli A, Jamani K. Acute promyelocytic leukemia presenting with atypical basophils. Clinical Case Reports. 2020;8(3):584–5. doi:10.1002/ccr3.2686

- 3. Ghimire A, Liesveld J, Wallace D, et al. Case of acute promyelocytic leukemia with basophilic differentiation and an ETV6 mutation. Journal of Hematopathology. 2021;14(4):333–6. doi:10.1007/s12308-021-00469-1
- 4. Iwakiri R, Inokuchi K, Dan K, et al. Marked basophilia in acute promyelocytic leukaemia treated with all-trans retinoic acid: Molecular analysis of the cell origin of the basophils. British Journal of Haematology. 1994;86(4):870–2. doi:10.1111/j.1365-2141.1994.tb04845.x
- 5. Piva E, Tosato F. Promyelocytic leukemia with basophil-like granules. Blood. 2020;136(3):372. doi:10.1182/blood.2020006060



**Figure 1.** Panels A-C: Wright-Giemsa stain; x100 objective, original magnification x1000. Peripheral blood smear showing blasts with intense azurophilic granules. Panels E-F: Wright-Giemsa stain; x100 objective, original magnification x1000. Bone marrow smear revealing blasts with irregular nuclei, some of them binucleated, dispersed chromatin, prominent nucleof, and frequent cytoplasmatic azurophilic granules.



**Figure 2.** Panels A-H. Bone marrow flow cytometry analysis. Two populations are identified, which are marked with blue and orange. The blue population shows abnormal promyelocytes positive for CD117, CD13 (dim), CD33, CD56, and MPO which represents the 76% of the total cellularity. The orange population reveals 4% of abnormal cells with basophilic differentiation positive for CD34, CD35, CD22, CD203c, CD9, CD123 (bright), and MPO.