

April 2020

## Slide Summaries

## Slide 1

Myelocytes present and mild eosinophilia

## Slide 2

Generally normal film

## Slide 3

Borderline monocytosis  
Basophil identified

## Slide 4

Lymphocytosis with smear cells. Indicative of CLL

## Slide 5

Occasional atypical lymphocyte

## Slide 6

Acute Pro-Myelocytic Leukemia  
Neutropaenia  
Thrombocytopaenia  
Blasts +++  
(see case study opposite)

## This issue

Last Month's Slides **P.1**  
 Monthly Case study **P.1**  
 Scientific feature - APL **P.2-3**  
 Monthly Quiz **P.3**  
 Top Tips **P.3**

## Monthly Digital Case study

## April 2020 Slide 6



## Presentation

Male (73 years old)  
 Multiple Instrument alarms (with WBCs in particular)

WBC	51.9* ( $10^3/\text{mm}^3$ )
RBC	2.48 ( $10^6/\text{mm}^3$ )
HGB	8.0 (g/dL)
HCT	20.9 (%)
MCV	84 (fL)
MCH	32.0 (pg)
MCHC	38.1 (g/dL)
PLT	18* ( $10^3/\text{mm}^3$ )

## Differential

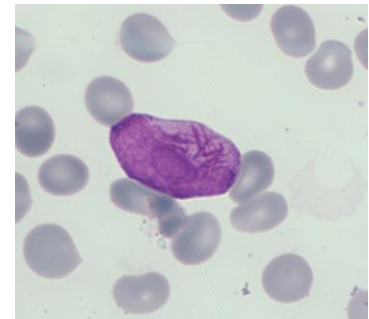
Neutrophils	0.7%
Lymphocytes	7.1%
Metamyelocytes	1.4%
Myelocytes	0.7%
Blasts	90.1%

## Diagnosis

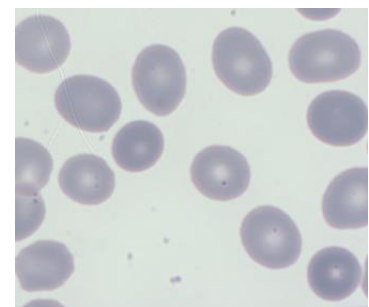
Acute promyelocytic Leukemia (variant)  
 The patient presented as a haematology emergency – sepsis and haemorrhagic disease

## Slide review

Hypergranular blasts.  
 Multiple Auer rods.  
 Convoluted and folded nuclei with fine chromatin strands:

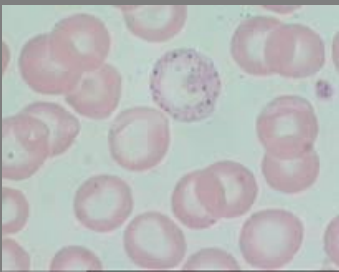
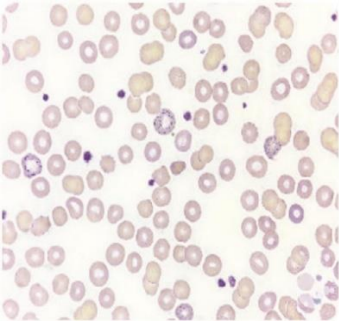


Examination of the blood film confirms the low platelet count and also shows the presence of spherocytes as indicated by the raised MCHC



## Monthly Morphology Quiz

Name these cells.



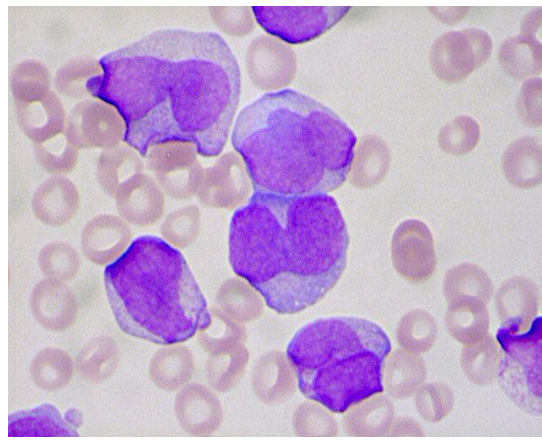
Look at the red cells in  
the images above.

What abnormalities can  
you see?

Name a condition  
where you might see  
this picture.

What other laboratory  
tests could be  
indicated?

Answers in the next  
edition.



## Acute Promyelocytic Leukaemia – In Brief

An overview of the  
clinical features,  
laboratory findings and  
treatment

### Introduction

**Acute Promyelocytic Leukaemia (APL)** is a differentiated variant of **Acute Myeloid Leukaemia (AML)** that usually (>95%) carries a **t(15:17)** chromosomal translocation.

It accounts for around **10% of AML cases** and has a different treatment protocol from AML.

### Clinical Presentation

The disease can be seen in either gender and at any age but is most common in middle-age. It frequently presents with a severe haemorrhagic syndrome.

- The following features may be seen:
- Symptoms of anaemia- leading to fatigue, pallor, tachycardia, dyspnoea
- Fever and infections due to neutropaenia
- Symptoms relating to abnormal platelet function and or coagulopathy – severe bleeding and or ecchymosis
- Non-specific such as pain, slow healing and unexplained weight loss

### Laboratory Findings

Laboratory investigations, besides FBC/Differential and blood film morphology, include the following:

- Bone Marrow Aspirate and Trepine
- Immunophenotyping
- Cytogenetic and mutation analysis
- Biochemistry (LFTs, U+E, Calcium, LDH)
- Coagulation Screening/ D-Dimer

Haematological investigations normally show normochromic normocytic anaemia and thrombocytopenia on presentation.

WBC is usually increased with a variable number of blast cells. The condition is morphologically identified as M3 by the FAB classification.

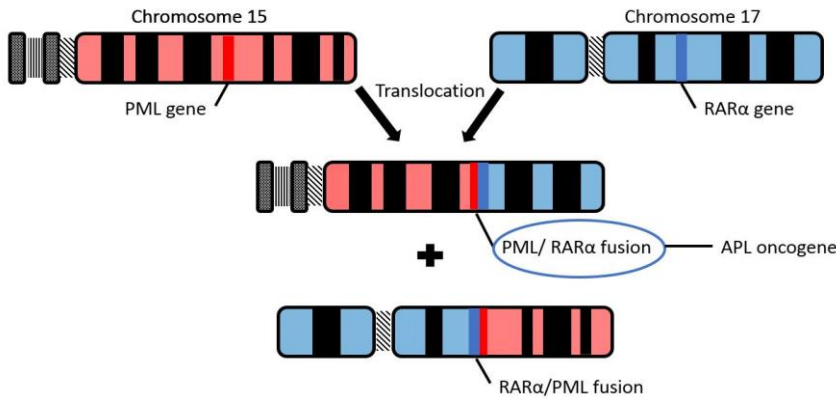
The bone marrow is hypercellular, typically containing many leukaemic blasts.

Morphology, immunophenotyping and genetic analysis confirm the diagnosis.

Coagulopathy is often present in APL.

Biochemistry tests are performed as a baseline prior to treatment. Raised Uric Acid and/or LDH may be present.

## APL t(15;17) chromosomal translocation – formation of PML/RAR $\alpha$ protein



## Acute Promyelocytic Leukaemia Continued

### Genetics

The somatic mutation that causes acute promyelocytic leukemia involves two genes, the PML gene on chromosome 15 and the RAR $\alpha$  gene on chromosome 17.

A rearrangement of genetic material (translocation) between chromosomes 15 and 17, written as t(15;17), fuses part of the PML gene with part of the RAR $\alpha$  (retinoic acid receptor  $\alpha$ ) gene. The protein produced from this fused gene is known as PML-RAR $\alpha$ .

The RAR $\alpha$  gene controls the maturation of WBCs which is blocked in APL

### Treatment

Treatment consists of supportive therapy (maintaining haemoglobin and platelet count, treatment of infection and treatment of DIC (FFP)) and specific chemotherapy.

Treatment differs from that of other acute myeloid leukaemia with the addition of all-trans retinoic acid (ATRA), often combined with arsenic trioxide or anthracycline

### Prognosis

APL can be aggressive on presentation but cells are relatively sensitive to chemotherapy. The addition of ATRA and other targeted therapies gives a >90% prospect of remission.

## This Month's Q&A Morphology Tips

### Quick tips for making manual blood films. Problems and answers

- P Blood film extends right to the end of the slide
- A Blood drop is too large or too small an angle between slide and spreader
- P Blood film is too short
- A Blood drop too small or too large an angle between slide and spreader
- P The blood film is irregular with ridges and a long tail
- A The edge of the spreader is dirty or chipped
- P Holes appearing in the spread
- A The slide is dirty/ greasy. Nb. Sometimes it can be lipaemia in the sample

## Other News

### Hematovision 5

Hematovision 5 is a free haematology atlas that is available via the HORIBA Medical website or offline, as part of the HORIBA Medical app downloadable for IOs or Android devices

## Bibliography

Hoffbrand's Essential Haematology 7<sup>th</sup> Edition Ch.13. Wiley Blackwell 2015

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Acute promyelocytic leukemia: from highly fatal to highly curable  
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Zhu Chen  
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