

ESMO Clinical Practice Guidelines

# Diffuse Large B Cell Lymphoma Clinical Case Presentation

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# Disclosure slide

Roche

Cellgene

Mundipharma

Janssen

Gilead

Bayer

Millenium

Servier

# Introduction

- Male, DOB 16.05.1975 (39 y.o.)

## PAST MEDICAL HISTORY

- Nothing relevant

## HISTORY OF PRESENT ILLNES

03-03/2014: progressive dyspnea; loss of appetite  
drenching night sweats, fever

# Introduction

- Investigated in private clinic for a suspicion of a lymphoproliferative disease.
- Urgent transfer to the ICU of public hospital for gradually increasing leg pain and rapidly developing paresis of the legs.

# Introduction

## LABORATORY (2.4.2014)

- Hb 134 g/l, Plt and wbc normal
- protein and albumin, liver tests, creatinine normal
- LDH 1248 U/l,  $\beta$ 2-microglobuline 3.0 mg/l (normal 1.09-2.53)
- CRP 16 mg/l

## INFECTIONS WORK-UP

- Serologies for HIV, HBV, HCV negatives

# Introduction

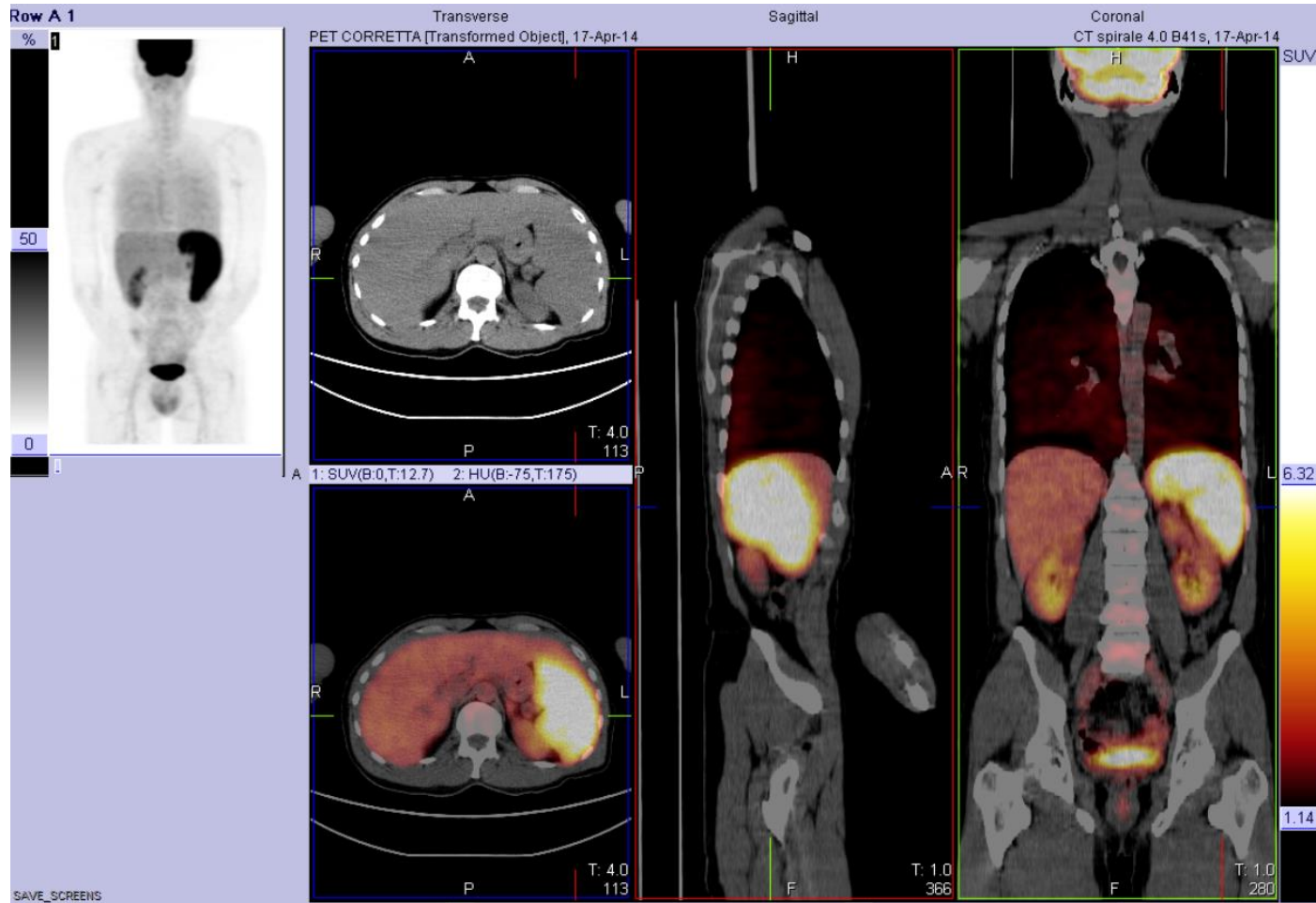
## BONE MARROW BIOPSY

- negative for lymphoma

## NEUROLOGIC EVALUATION

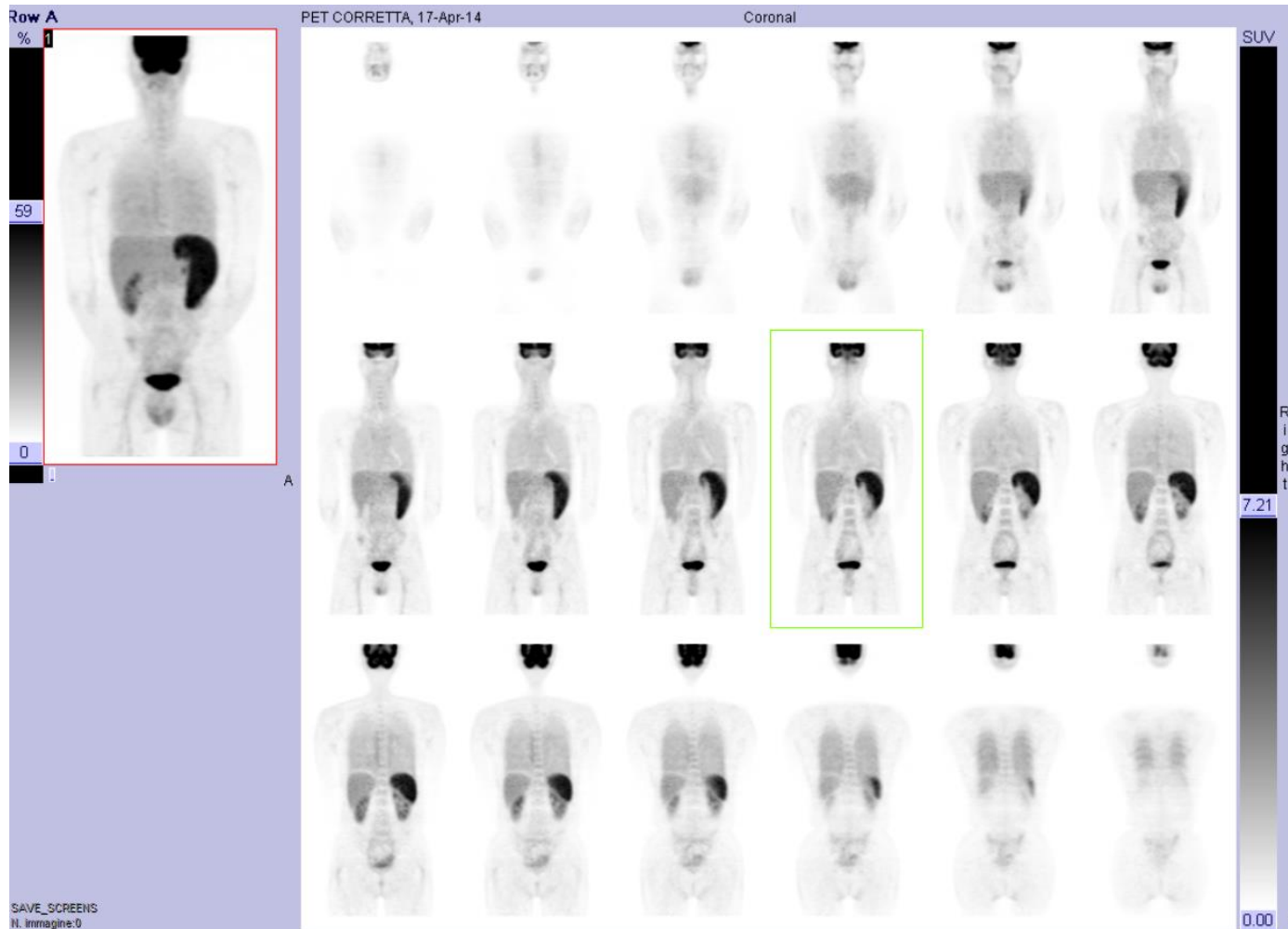
- Lumbar puncture: No lymphoma cells, increased proteins, normal cell counts (albumino-cytologic dissociation)
- Brain CT scan: negative
- MRI of the spine: not performed (claustrophobic)

# FDG-PET, 18.04.2014



Splenomegaly (17 cm) with increased FDG uptake

# FDG-PET, 18.04.2014



Splenomegaly (17 cm) with increased FDG uptake



# Q1:How would you proceed?

1. Splenectomy for diagnosis
2. Splenic biopsy for diagnosis
3. Start the patient on steroids
4. Try large spectrum antibiotics
5. Perform additional investigations



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## SPLENECTOMY (20.04.2014) (B14-09859)

Neoplastic cells were:

positive for CD45, CD20, CD79a, BCL-2 (weak), BCL-6, MUM1 and CD5 (weak)

negative for CD3, CD38, CD23, CD30, CD10, cycline D1

Ki67 90%

Final diagnosis:

DLBCL

Course of disease complicated by:

- significant worsening of dyspnea
- worsening of the lower limbs paresis

## **DIAGNOSIS:**

**DLBCL** of the spleen, stage IBE (S)

**Guillain-Barré syndrome** as paraneoplastic syndrome

Treatment starts with iv immunoglobulin's and steroids.

## Q2:How would you treat the lymphoma?

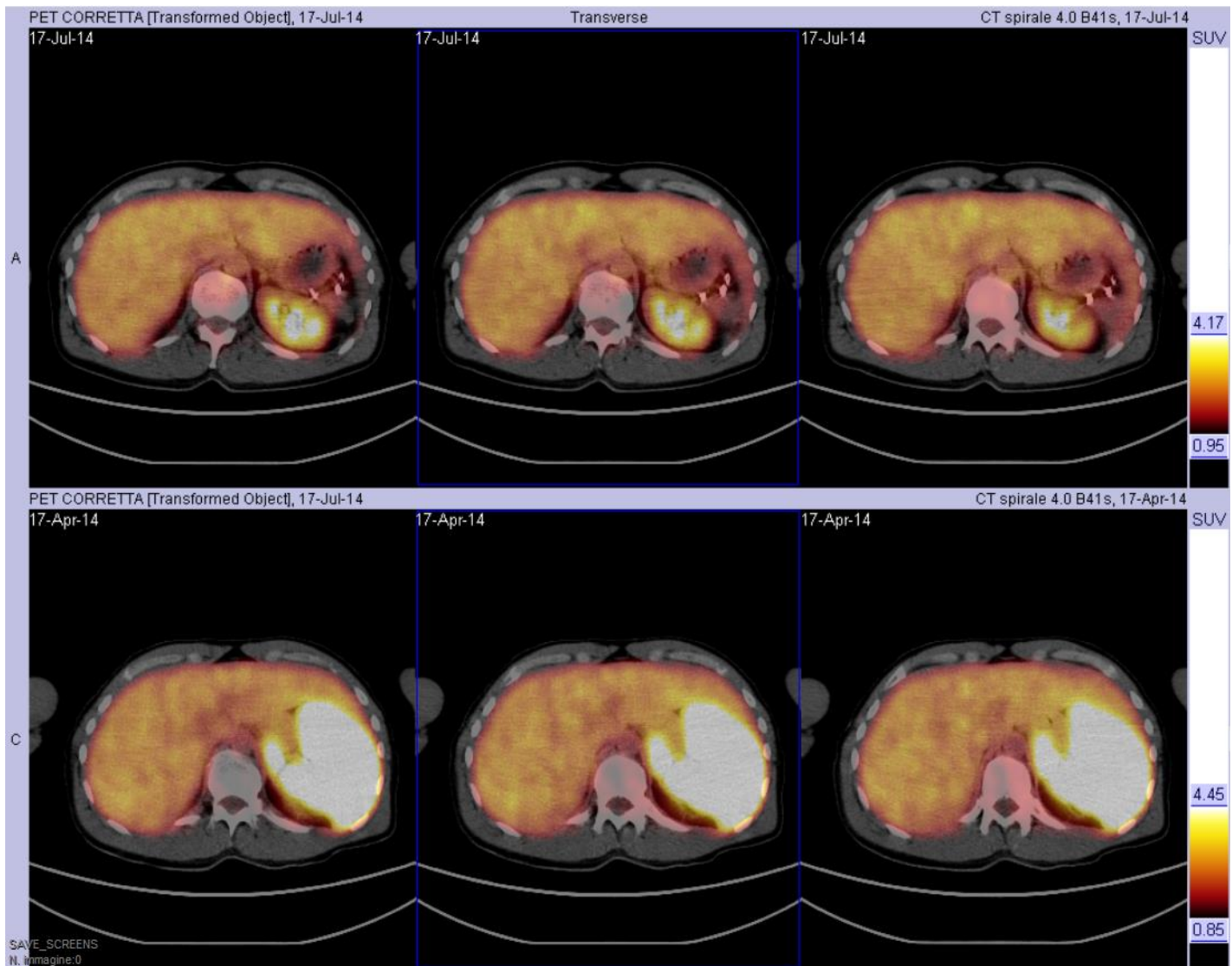
1. R-CHOP
2. R-DA-EPOCH
3. R-Hyper-CVAD-HDArac-MTX
4. R-bendamustine
5. Other «ad-hoc» regimen

## **THERAPY:**

- 6 cycles of R- CH(O)EP from 28.04-27.09.2014 with omission of vincristine
- 2 cycles of HD methotrexate as CNS prophylaxis (27.09 and 11.10.2014)
  - chemotherapy well tolerated, no major toxicity
  - progressive improvement of the neurologic deficits

## **PET/CT after cycle 2**

- complete remission (morphologic and metabolic)



11/2014 (1 month later):

- he represented with episodes of night sweats
- muscular pain in the lower limbs

**LUMBAR PUNCTION with flow cytometry**  
**(7/11/2014)**

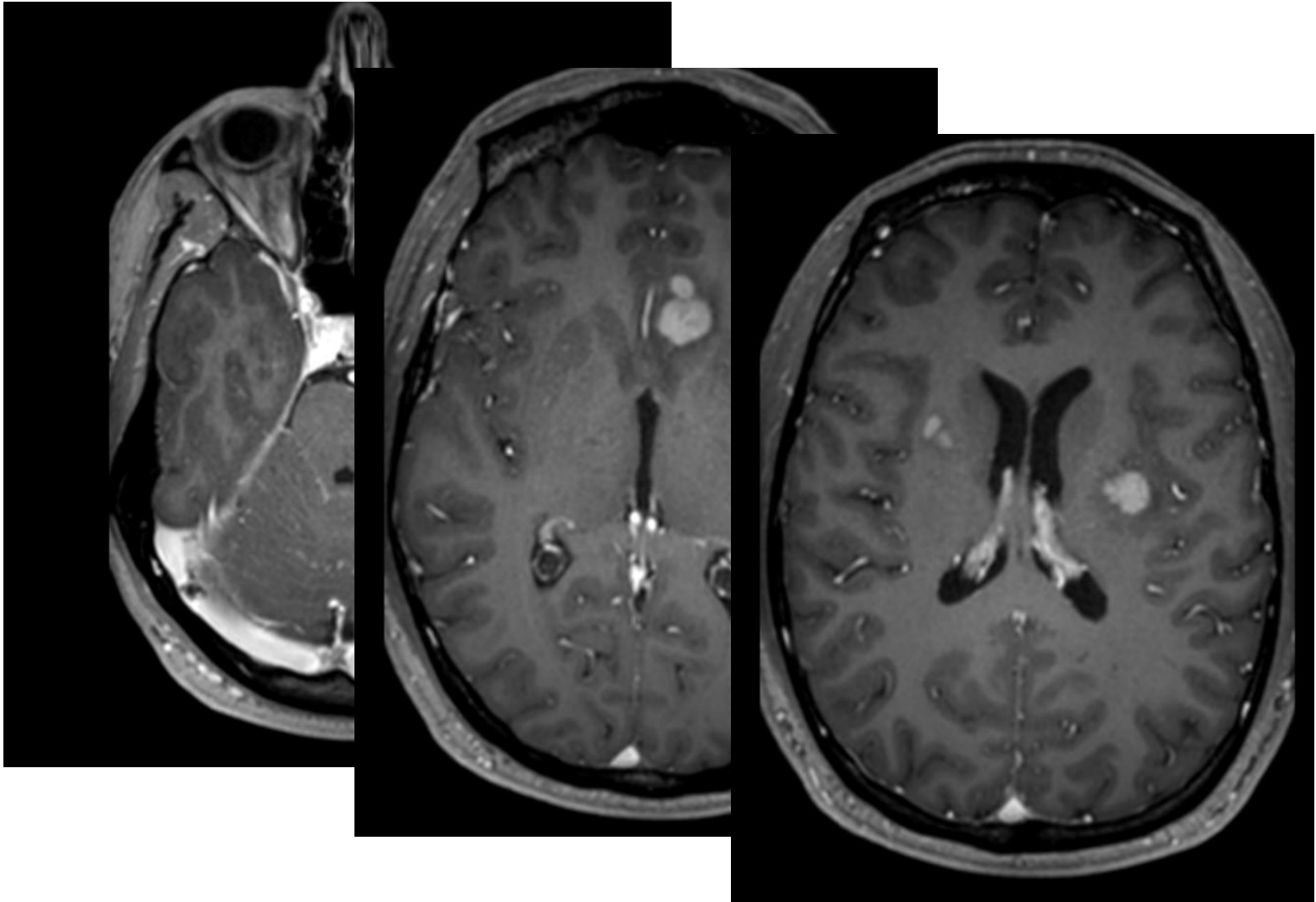
- clonal B cell population in keeping with secondary lepto-meningeal involvement

**BRAIN AND SPINE MRI (7/11/2014)**

- multiple contrast-enhancing parenchymal brain lesions (basal ganglia, right temporal and frontal lobe)
- thickening and enhancement of the nerve roots







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# How would you treat the early CNS relapse?

1. HD-MTX + HD-AraC
2. HD-MTX + HD-AraC + i.t. chemo
3. Chemo + radiotherapy
4. Induction CT + autologous transplant
5. Induction CT + allogeneic transplant

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## THERAPY:

- 2 cycles of HD cytarabine and etoposide (CYVE) according to C. Soussain jco 2008
- 2 courses of IT sustained release liposomal Ara-C (DepoCyte®)
- stem cell mobilisation after 2<sup>nd</sup> cycle
- High dose chemotherapy with carmustine (BCNU) and thiotepa followed by autologous stem cell transplant
- Radiotherapy of the residual disease post-treatment in spine

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ORIGINAL REPORT

Intensive Chemotherapy Followed by Hematopoietic Stem-Cell Rescue for Refractory and Recurrent Primary CNS and Intraocular Lymphoma: Société Française de Greffe de Moëlle Osseuse-Thérapie Cellulaire

*Carole Soussain, Khè Hoang-Xuan, Luc Taillandier, Emmanuelle Fourme, Sylvain Choquet, Francis Witz,*

### *Treatment*

The salvage treatment consisted of two cycles of the high-dose cytarabine and etoposide (CYVE) regimen administered 28 days apart, as previously described,<sup>13</sup> with cytarabine 2 g/m<sup>2</sup>/d on days 2 through 5 in a 3-hour infusion and 50 mg/m<sup>2</sup>/d on day 1 through 5 in a 12-hour infusion; and etoposide 200 mg/m<sup>2</sup>/d days 2 through 5 in a 2-hour infusion. The doses were slightly

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# Epicrisis

1 month after the end of the treatment the patient relapsed systemically (liver and CNS) and died.

## Questions raised by this case:

1. When is a CNS prophylaxis indicated in DLBCL?
2. Which prophylaxis, if any, is the best?
3. How to treat DLBCL relapsing in the CNS?