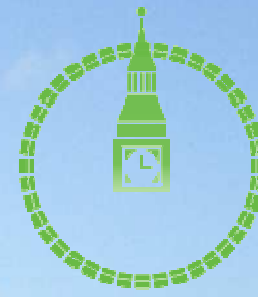


# Deauville Criteria for end of therapy



King's College London  
& Guy's and St Thomas'  
**PET Centre**

Sally Barrington



# 5 Point Scale /Deauville criteria

*Leukemia & Lymphoma*, August 2009; 50(8): 1257–1260

**informa**  
healthcare

## REVIEW

### **Report on the First International Workshop on interim-PET scan in lymphoma**

MICHEL MEIGNAN<sup>1</sup>, ANDREA GALLAMINI<sup>2</sup>, & CORINNE HAIOUN<sup>3</sup>

<sup>1</sup>*Nuclear Medicine Department, H. Mondor Hospital, AP-HP/Paris 12 University, Creteil, France,* <sup>2</sup>*Hematology Department and BMT Unit, Az. Ospedaliera S. Croce e Carle, Cuneo, Italy,* and <sup>3</sup>*Hematology Department, H. Mondor Hospital, AP-HP/Paris 12 University, Creteil, France*

*(Received 8 May 2009; accepted 12 May 2009)*

1. n
2. u
3. u
4. n
5. n
- ne

or

# Deauville Criteria

Concern that MBP  $\geq 2$ cm background  $< 2$ cm was too low a threshold

Take account of varying FDG uptake during tx

Graded method of assessment

Flexible suited for trial



## Role of Imaging in the Staging and Response Assessment of Lymphoma: Consensus of the International Conference on Malignant Lymphomas Imaging Working Group

*Sally F. Barrington, N. George Mikhael, Lale Kostakoglu, Michel Meignan, Martin Hutchings, Stefan P. Müller, Lawrence H. Schwartz, Emanuele Zucca, Richard I. Fisher, Judith Trotman, Otto S. Hoekstra, Rodney J. Hicks, Michael J. O'Doherty, Roland Hustinx, Alberto Biggi, and Bruce D. Cheson*

## Recommendations for Initial Evaluation, Staging, and Response Assessment of Hodgkin and Non-Hodgkin Lymphoma: The Lugano Classification

*Bruce D. Cheson, Richard I. Fisher, Sally F. Barrington, Franco Cavalli, Lawrence H. Schwartz, Emanuele Zucca, and T. Andrew Lister*

Published online 11/08/14

DOI 10.1200/JCO.2013.53.5229

DOI 10.1200/JCO.2013.54.8800

# Deauville criteria/5-PS for end tx

Good interobserver agreement

Score 3 in patients receiving standard treatment likely represents CMR

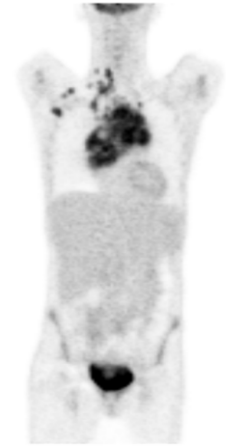
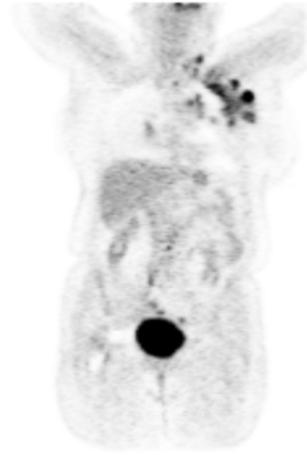
Especially with modern cameras

One method preferred for response assessment  
(score 2 similar to IHP criteria)

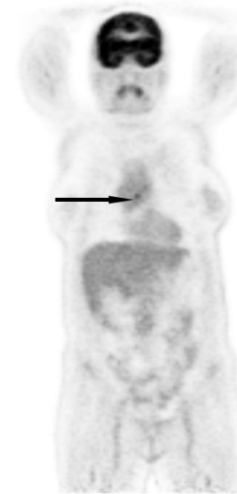
Barrington 2010 EJNMMI; 37(10):1824-33, Le Roux Eur J Nucl Med Mol Imaging 2011; 38:1064-71, Biggi J Nucl Med 2013; 54:683-90, Dupuis J Clin Oncol 2012;30:4317-22, Itti et al EJNMMI 2013; 40:1312-20, Mamot 2013 Hem Oncol 31:100 (s1;abst 15), Nols 2014 Leuk Lymphoma Apr;55(4):773-80, Pregno 2012 Blood 119:2066-73, Tychyj-Pinel EJNMMI2014 Mar;41(3):408-15

# Five Point Scale

Staging



Response



Score

1

2

3

4

5

No uptake

FDG < MBP

FDG > MBP ≤ liver

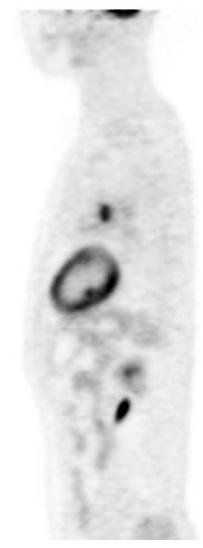
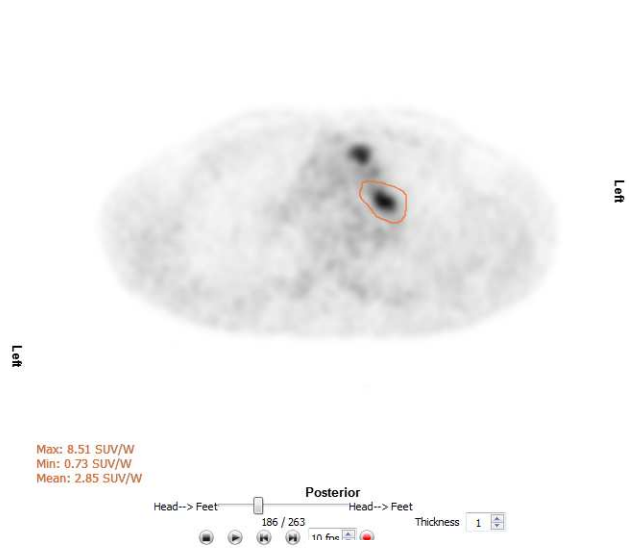
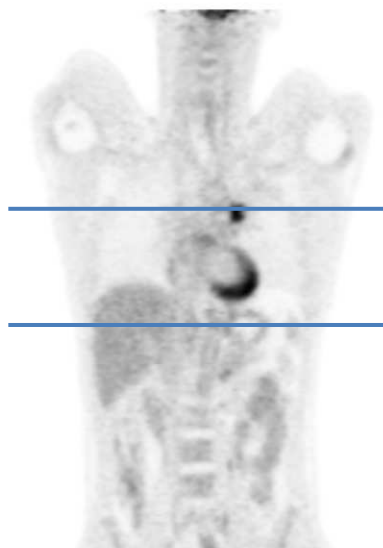
FDG > liver

FDG >> liver

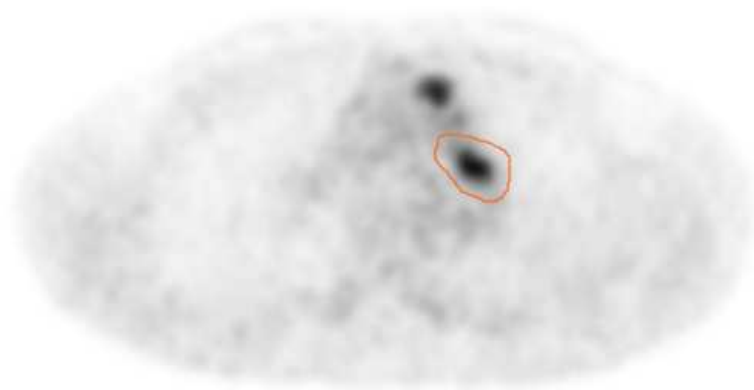
# 5 Point Scale (Deauville criteria)

1. no uptake
2. uptake  $\leq$  mediastinum
3. uptake  $>$  mediastinum but  $\leq$  liver
4. **moderately** increased uptake compared to liver
5. **markedly** increased uptake compared to liver and/or new lesions

\*\* **markedly** increased uptake is taken to be uptake  $>$  2-3 times the SUV max in normal liver



Max: 8.51 SUV/W  
Min: 0.73 SUV/W  
Mean: 2.85 SUV/W



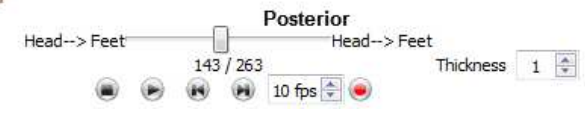
SUVmax lesion 8.51

SUVmax liver 2.50

Max: 8.51 SUV/W  
Min: 0.73 SUV/W  
Mean: 2.85 SUV/W



Max: 2.50 SUV/W  
Min: 1.46 SUV/W  
Mean: 1.94 SUV/W



Score 5



# High physiological FDG uptake

can occur in some sites...

e.g. Waldeyers ring , gut, bone marrow after chemotherapy or GCSF treatment with 'physiologic' uptake > normal liver

In this case, CMR may be inferred if uptake at sites of initial involvement is no greater than surrounding normal tissue

CATEGORY	PET – CT based metabolic response
CMR	Score 1,2,3* in nodal or extranodal sites with or without a residual mass using 5-PS
PMR	<p>Score 4 or 5, with reduced uptake compared with baseline and residual mass(es) of any size.</p> <p><i>At interim</i> , these findings suggest responding disease</p> <p><i>At end of treatment</i> these findings indicate residual disease</p> <p>Bone marrow: Residual marrow uptake &gt; normal marrow but reduced compared with baseline (diffuse changes from chemotherapy allowed). If there are persistent focal changes in marrow with a nodal response, consideration should be given to MRI, biopsy or interval scan.</p>
NMR	Score 4 or 5 with no significant change in uptake from baseline <i>At interim or end of treatment</i>
PMD	Score 4 or 5 with an increase in uptake from baseline and /or New FDG-avid foci consistent with lymphoma <i>At interim or end of treatment</i>

**\* Score 3 in many patients indicates a good prognosis with standard treatment. However in trials involving PET where de-escalation is investigated, it may be preferable to consider score 3 as inadequate response to avoid under-treatment** Cheson et al JCO 2014 on line

# Response according to 5-PS

Score 1, 2 is Complete Metabolic Response (CMR)

Score 3 is probably also CMR with standard treatment

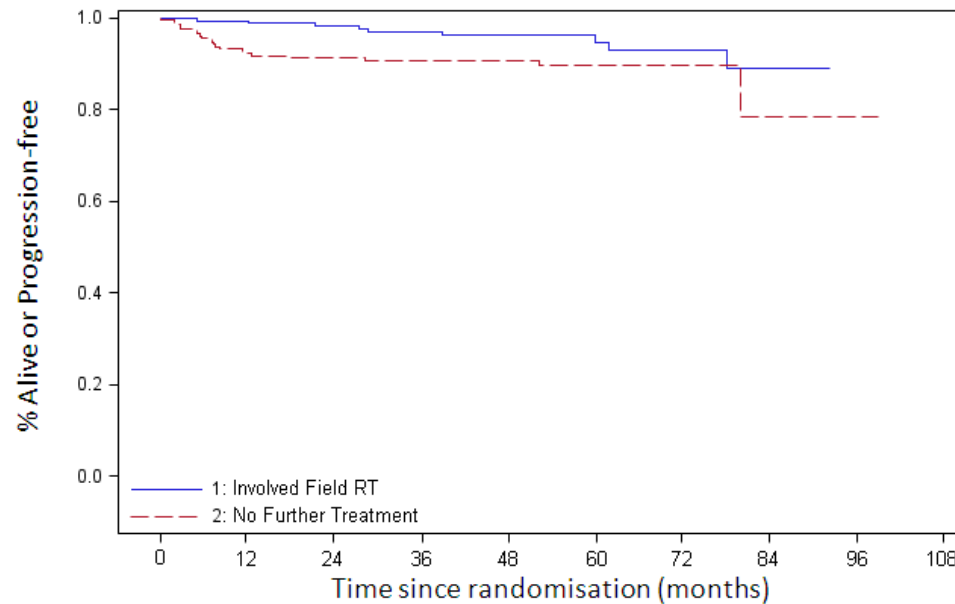
But in response-adapted trials exploring de-escalation, score 3 may be deemed inadequate response to avoid under-treatment

Interpretation of score 3 depends on timing of assessment, clinical context & treatment.

# Early stage HL



## RAPID : PFS in PET -ve population (per protocol analysis)



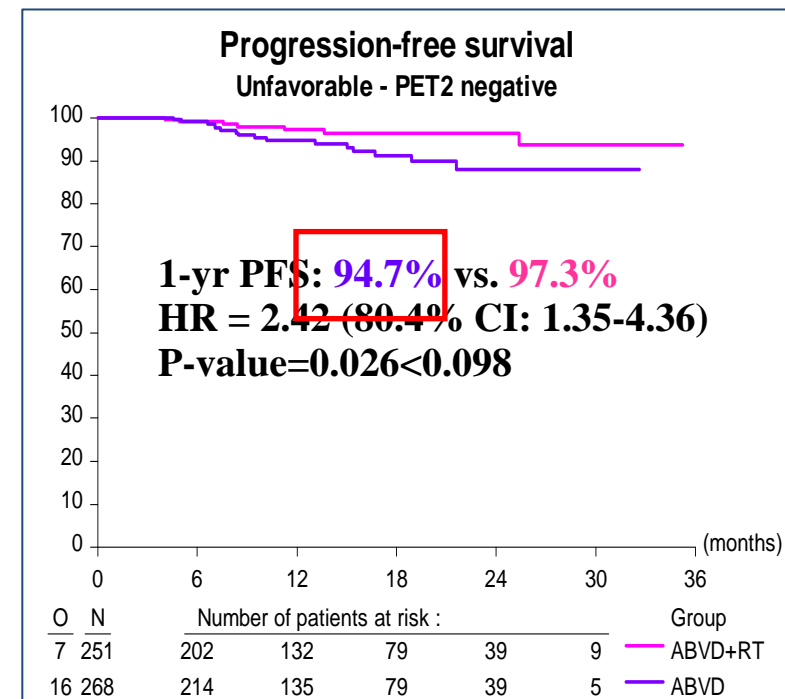
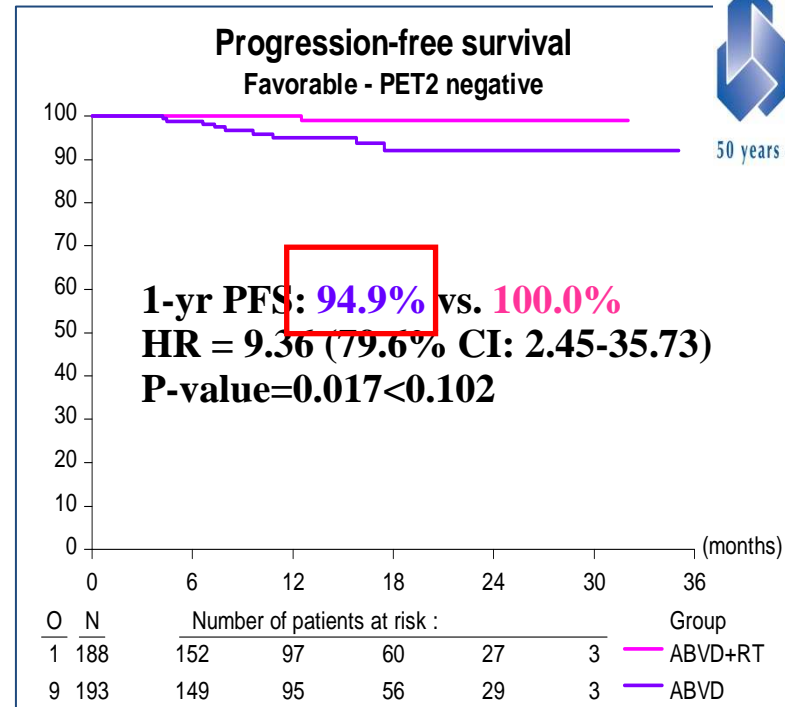
Number at risk:

IFRT	183	179	162	129	98	65	38	17	0	0
NFT	209	188	163	132	100	60	18	4	2	0

3 year PFS 97% vs 90.7%

HR 2.39 in favour of IFRT, p=0.03

Radford et al, Blood 2012; 120: a547



Raemaekers J et al JCO 2014;32: 1188-94

# HL advanced stage HD15

n = 739 PR ≥ 2.5cm

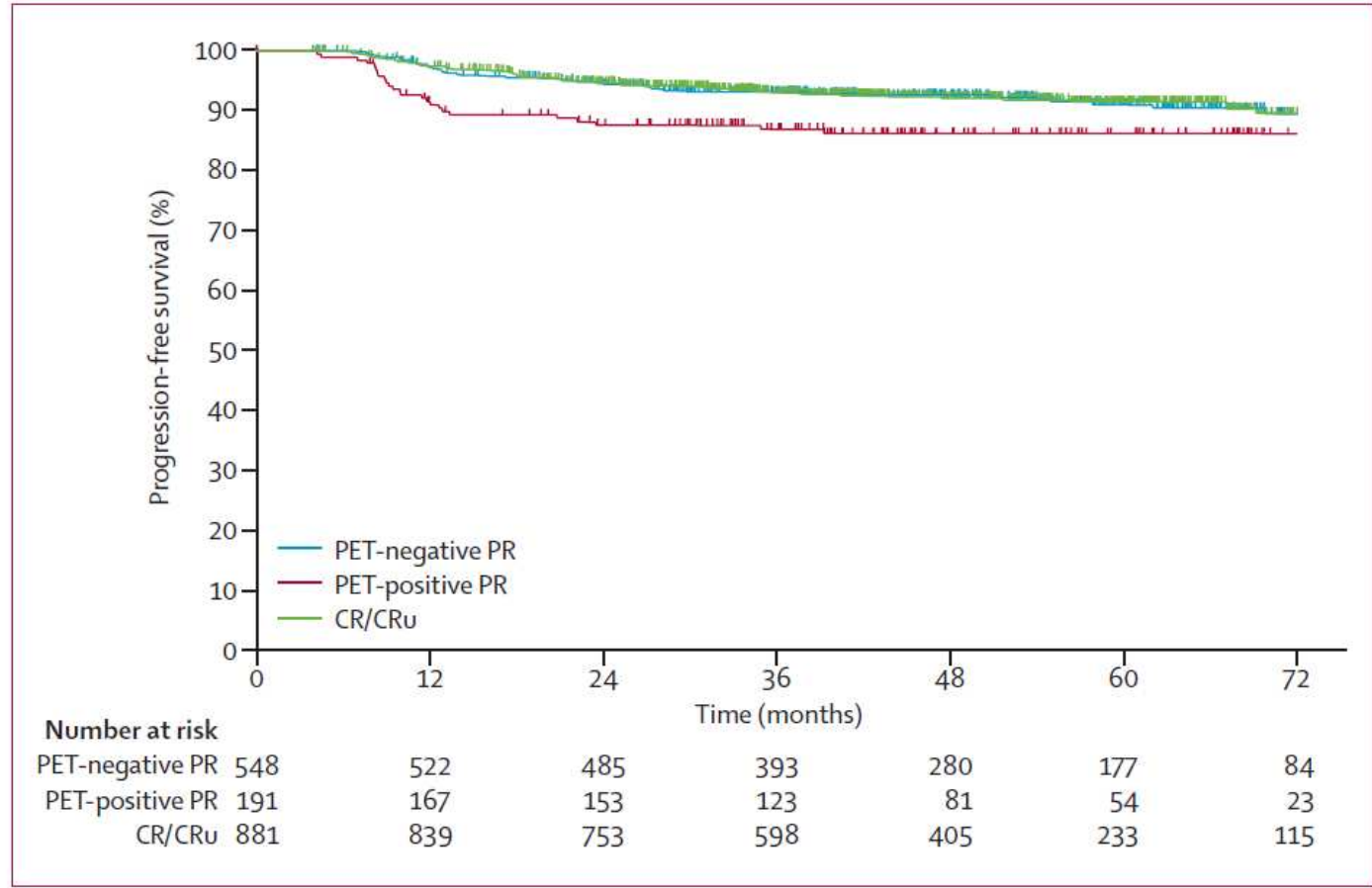
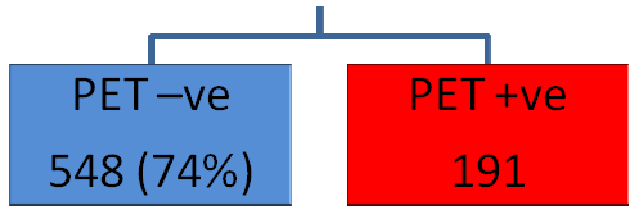


Figure 3: Progression free survival for PET study objective  
 PR=partial remission at the end of chemotherapy. CR/CRu=complete remission without or with residual abnormalities at the end of chemotherapy.

11% had RT

Originally published by the Lancet  
 [Engert A et al 2012 379(9828),May12 pp1791-9]



## Guidelines for the first line management of classical Hodgkin lymphoma

George A. Follows,<sup>1</sup> Kirit M. Ardeshtna,<sup>2</sup> Sally F. Barrington,<sup>3</sup> Dominic J. Culligan,<sup>4</sup> Peter J. Hoskin,<sup>5</sup> David Linch,<sup>2,6</sup> Shalal Sadullah,<sup>7</sup> Michael V. Williams<sup>8</sup> and Jennifer Z. Wimperis<sup>9</sup> for the British Committee for Standards in Haematology

### Recommendations:

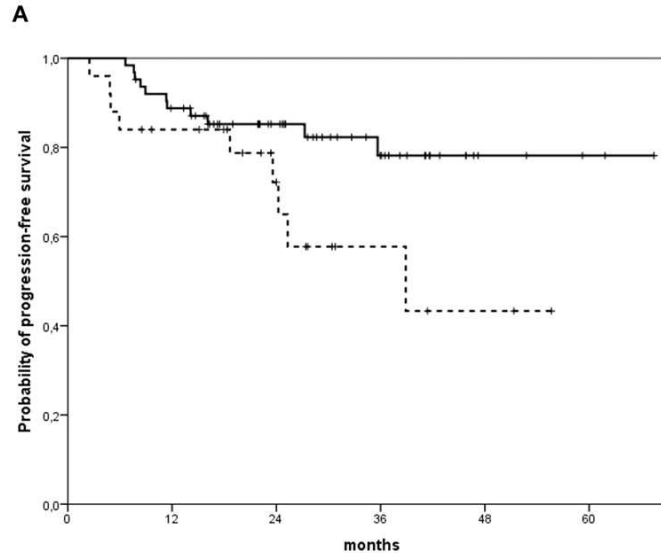
**The decision to omit RT from the management of IA/IIA non-bulky patients should involve discussion with a radiation oncologist (1B) and patients choosing to omit RT need to be aware of the balance of risks between RT and additional cycles of chemotherapy. (1B)**

**Patients treated with escalated BEACOPP who achieve an end-of-treatment PET-negative remission do not require consolidation RT to residual tissue (1A)**

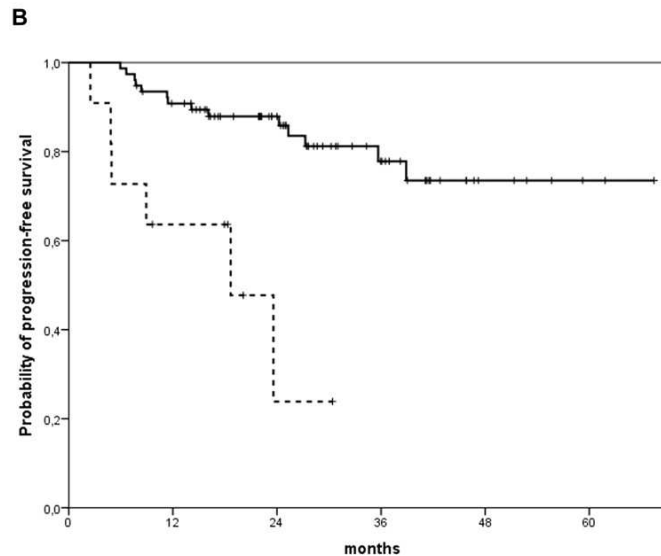
*‘It is recommended therefore that **score 1 or 2** is used to define a complete metabolic response (CMR) if omission of ‘standard’ radiotherapy treatment is being considered in discussion with patients.’*

# DLBCL

PFS according to response at I-PET and F-PET.



Interim



End

PPV 82 %  
NPV 100 %  
Using **Score 1,2 3**  
to define CMR  
At **END**

Pregno P et al. Blood 2012;119:2066-2073

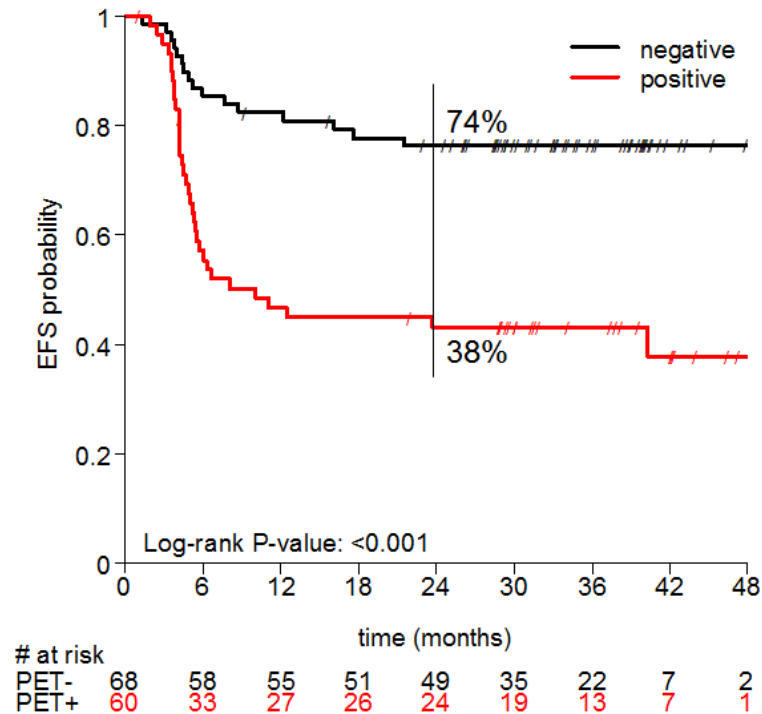


# Prospective evaluation of the predictive value of PET in 141 patients with DLBCL under R-CHOP-14 (SAKK 38/07)

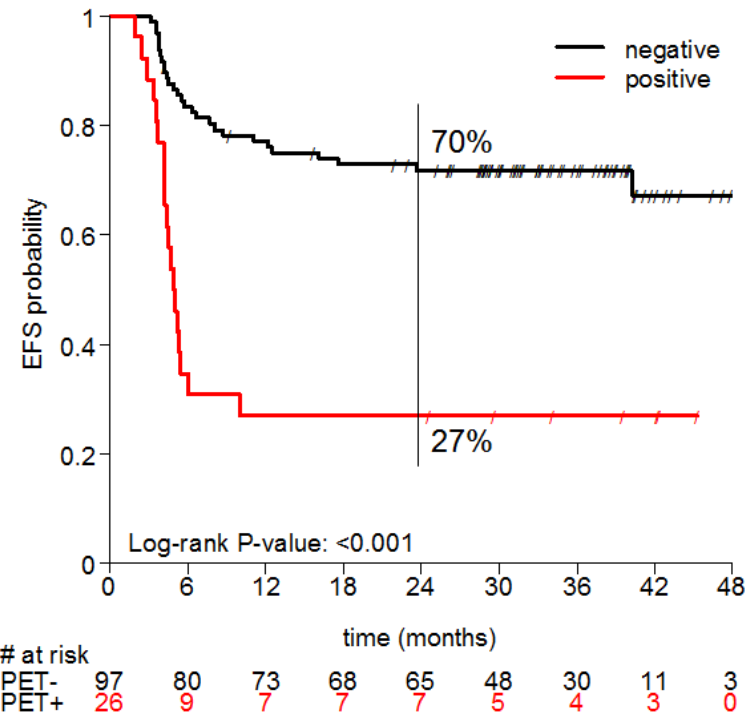
**Score 1,2 3** used to define CMR



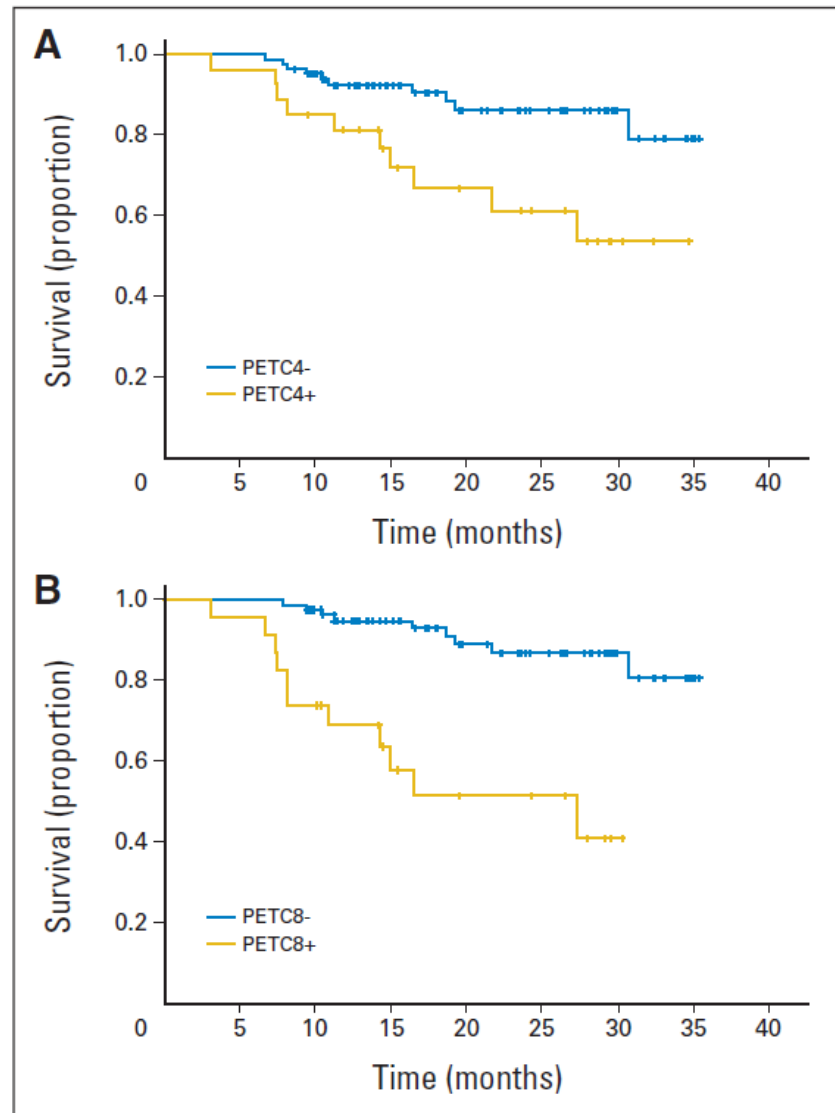
Interim PET, central review, 5-point scale



End of treatment, central review, 5-point scale, liver cut-point



# Follicular Lymphoma



Interim

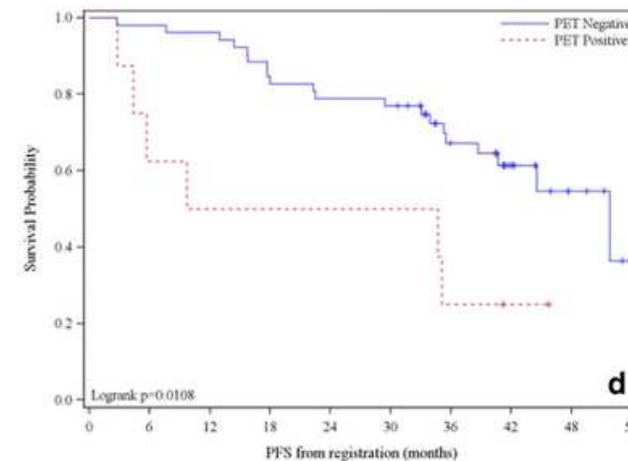
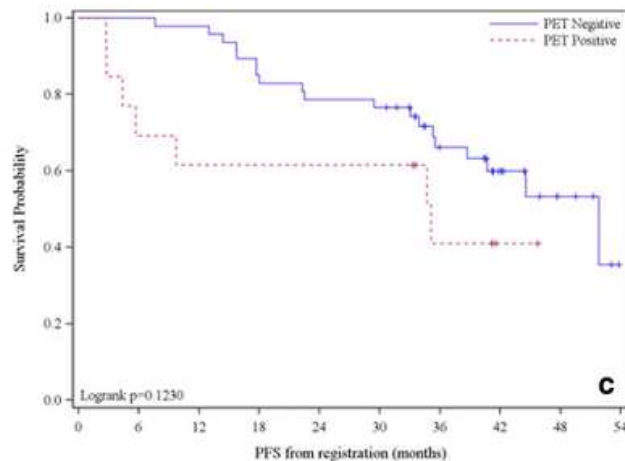
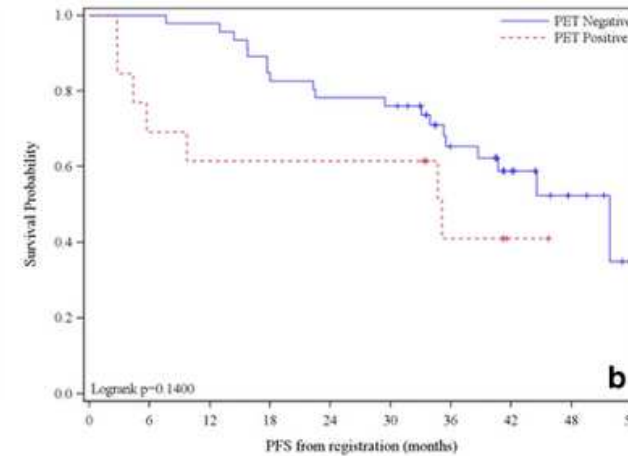
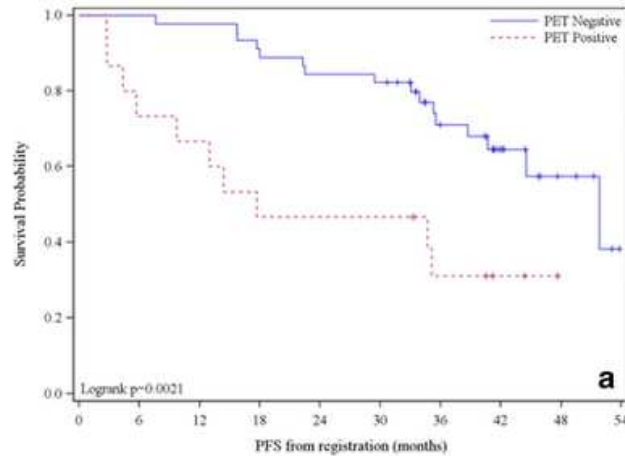
End

**Score 1,2 3** best to define CMR

# Comparison of criteria for end tx in FL

Local review

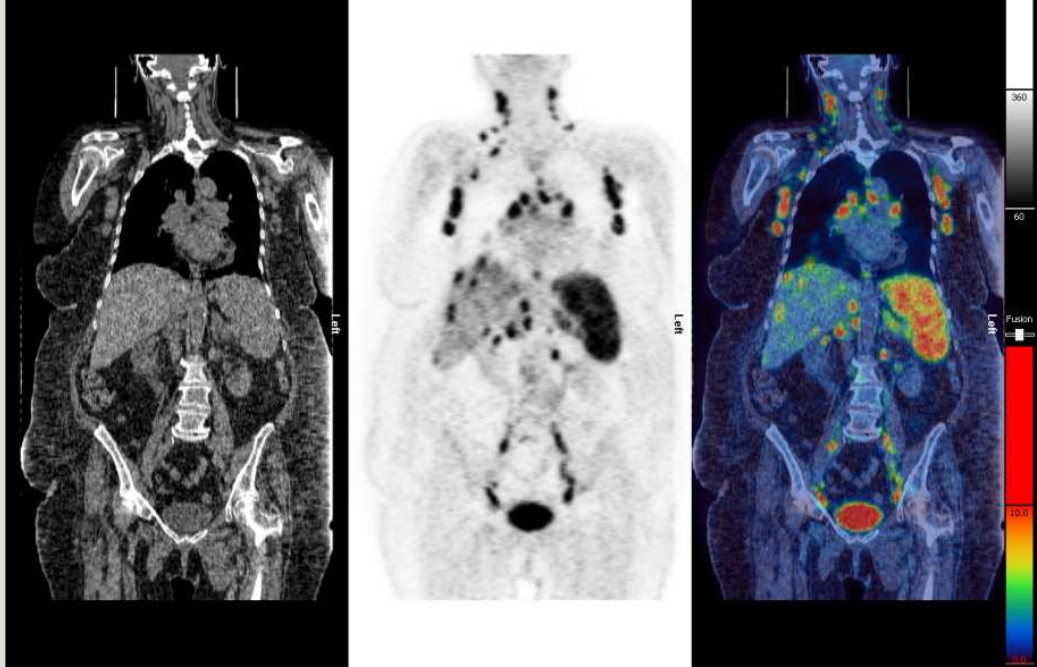
IHP



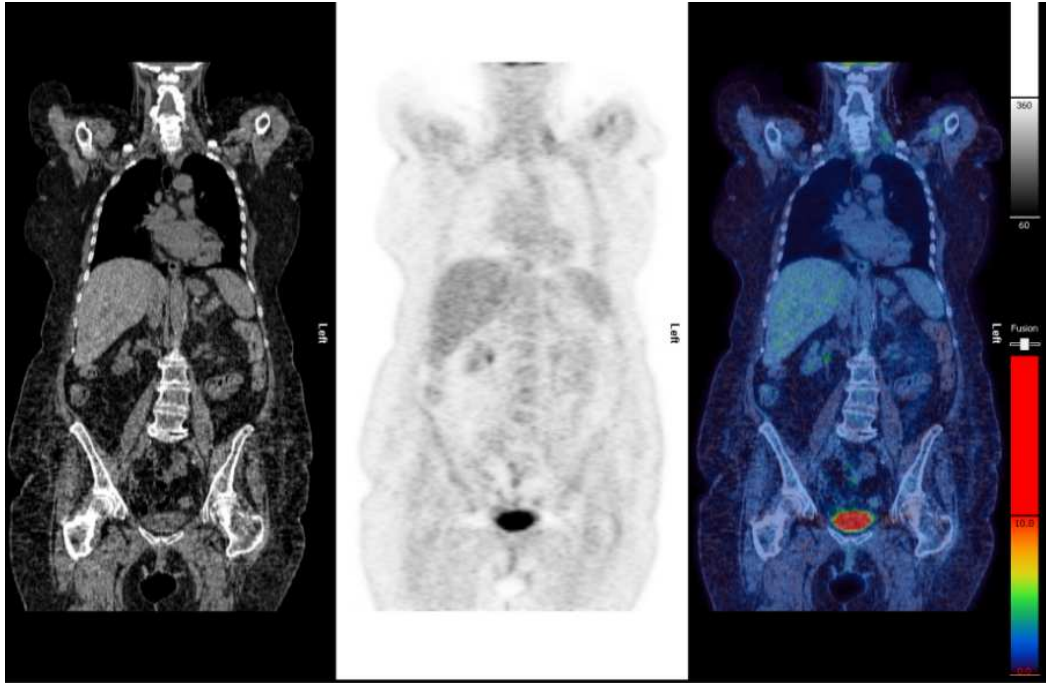
MBP (CMR score 1, 2) Liver (CMR score 1, 2, 3)



Baseline



Response



Score 1

CMR

# Response according to 5-PS

Score 4, 5 with reduced uptake from baseline is partial metabolic response (PMR)

-At interim this suggests responding disease

-At end of treatment this suggests residual metabolic disease

Baseline



Response  
End



Score 5

PMR

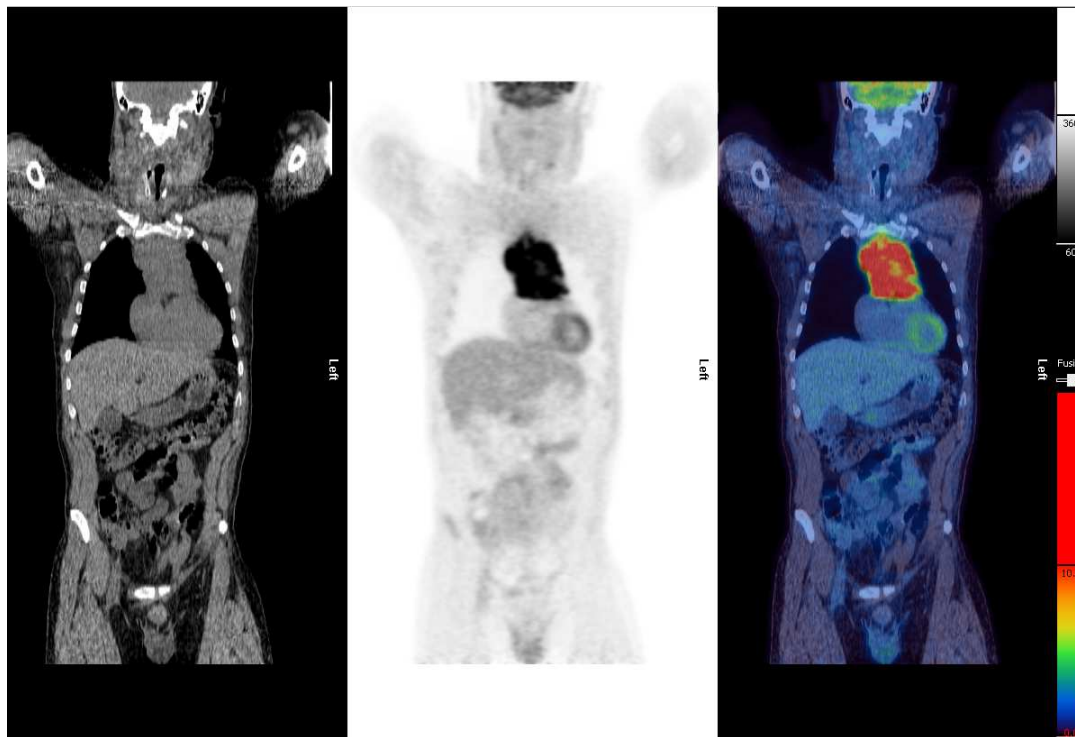
# Response according to 5-PS

Score 4, 5 with no change in uptake from baseline means no metabolic response (NMR)

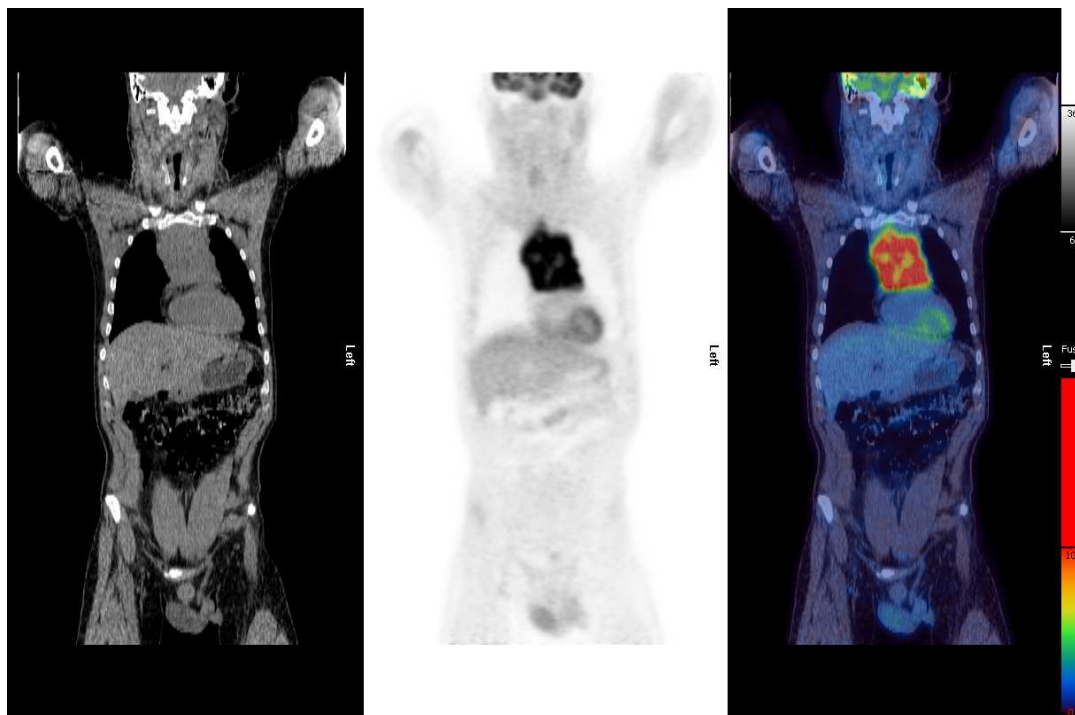
Score 4, 5 with an increase in uptake from baseline &/or new lesions is progressive metabolic disease (PMD)

-At interim and end of treatment NMR and PMD indicates treatment failure

Baseline



Response



Score 5

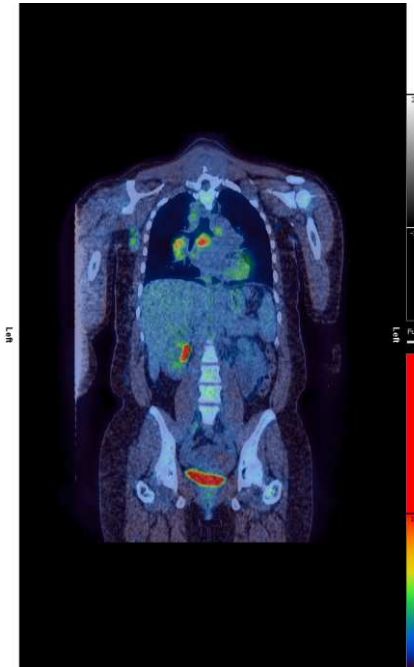
NMR



Baseline

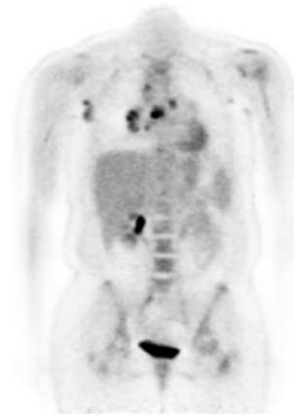


Post 2

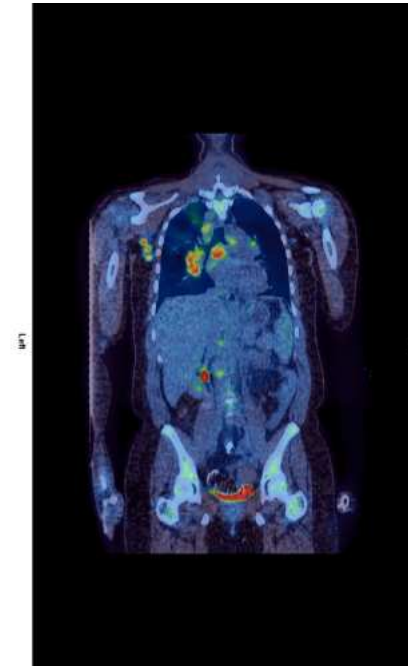


PMD ?

Interim  
Post 2



Post 3



PMD  
confirmed

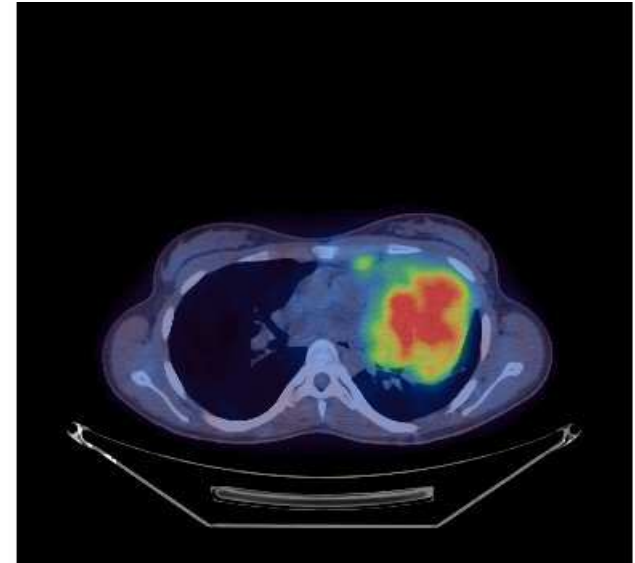
# Residual masses

Biopsy of residual metabolically active tissue is recommended if salvage treatment is considered

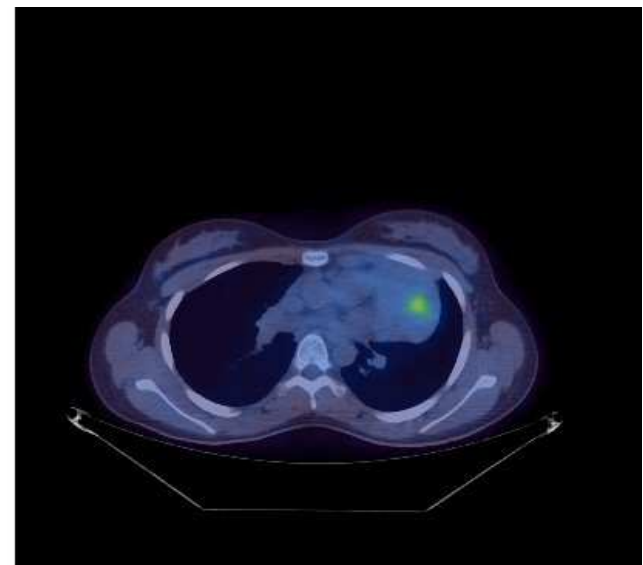
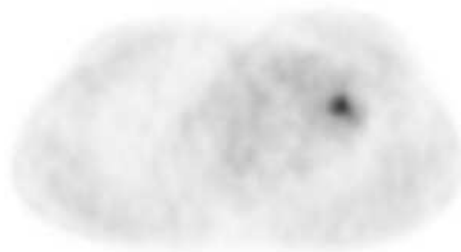
or an interval scan where clinical likelihood of disease is low to decide on treatment (or not)

# Guiding a biopsy

Baseline



End PET



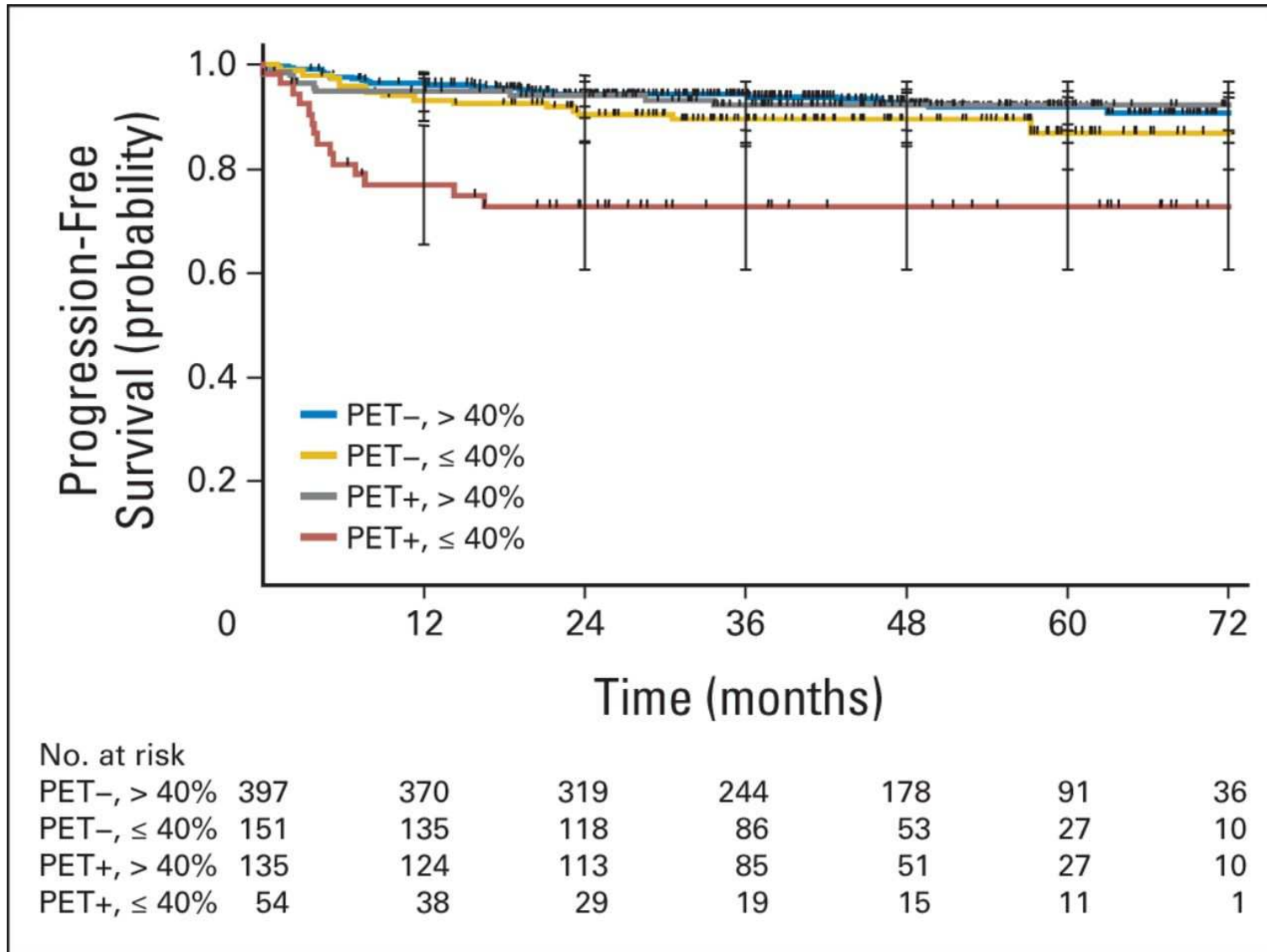
# Residual masses

Residual size mass and location should be recorded in PET-CT reports where possible

*as significance of the size of masses is unclear but may be complementary to metabolic information and data should be collected prospectively in clinical trials*

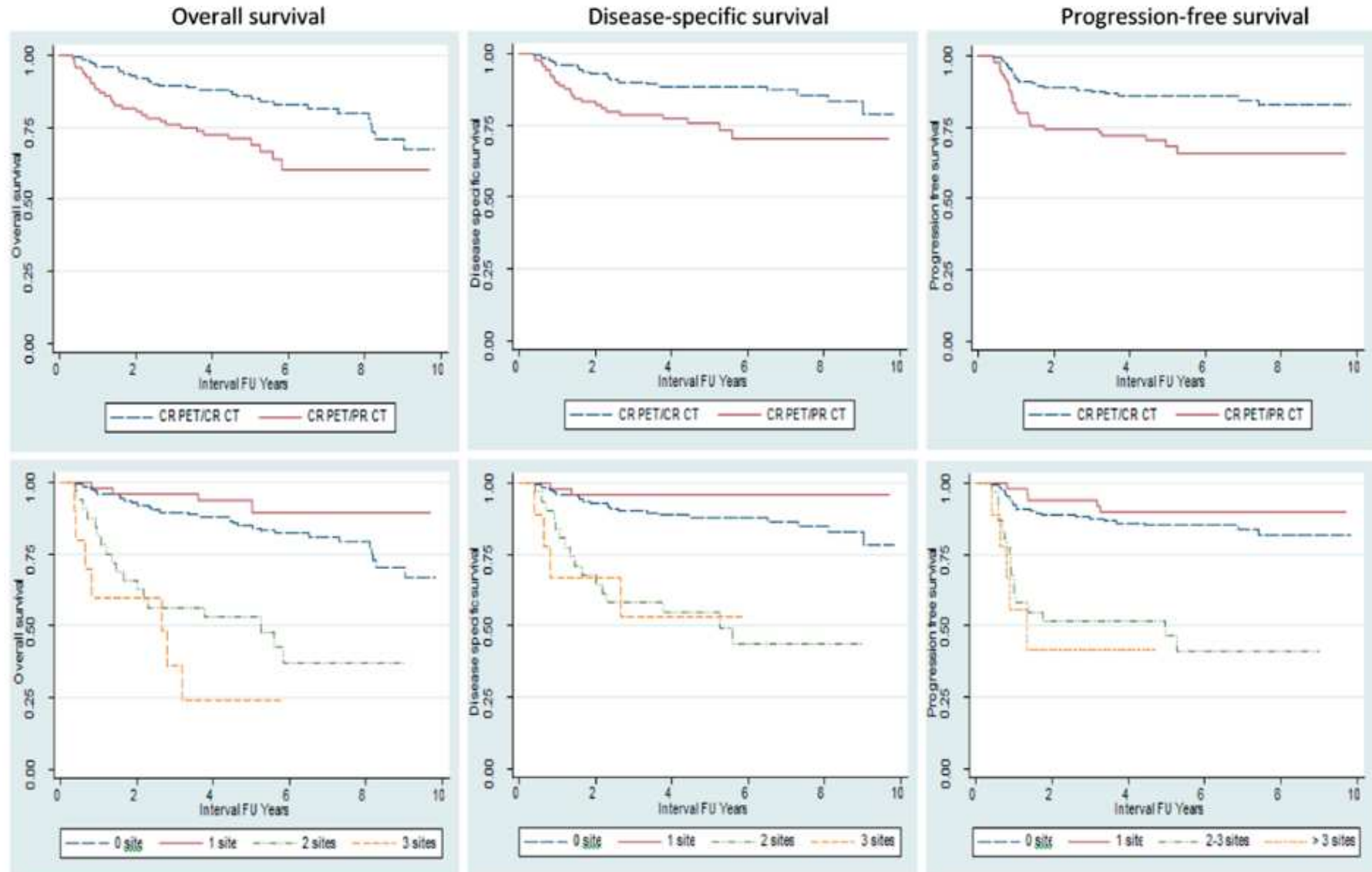


**Progression-free survival in positron emission tomography (PET) –positive and PET-negative patients and 40% reduction of the largest tumor diameter.**



Kobe C et al. JCO 2014;32:1776-1781

# Influence of residual mass ?



# Timing of PET-CT scans

Should be:

as long as possible after the last chemotherapy administration for interim scans

6-8 weeks post chemotherapy at end of treatment ideally (but a minimum of 3 weeks)

≥ 3 months after radiotherapy

# Summary

- DC are recommended for response assessment in 2014 ICML guidelines
- Can be used to assign **metabolic response categories**
- **Score 3** likely represents CMR in patients receiving standard therapy **BUT score 1,2** may be preferred to define CMR using de-escalation strategies to avoid undertreatment
- End of treatment residual or new metabolic disease requires biopsy confirmation before salvage therapy where feasible or an interval scan if clinical index of residual disease is low



# With special thanks to co-authors

N George Mikhaeel

Rodney Hicks

Lale Kostakoglu

Michael O'Doherty

Michel Meignan

Roland Hustinx

Martin Hutchings

Alberto Biggi

Stefan Müller

Franco Cavalli

Lawrence Schwartz

Andrew Lister

Emanuele Zucca

Bruce Cheson

Richard Fisher

Judith Trotman

Otto Hoekstra

And all attendees at Deauville & Menton workshops  
2009-current