

Δ SUV evaluation in DLBCL

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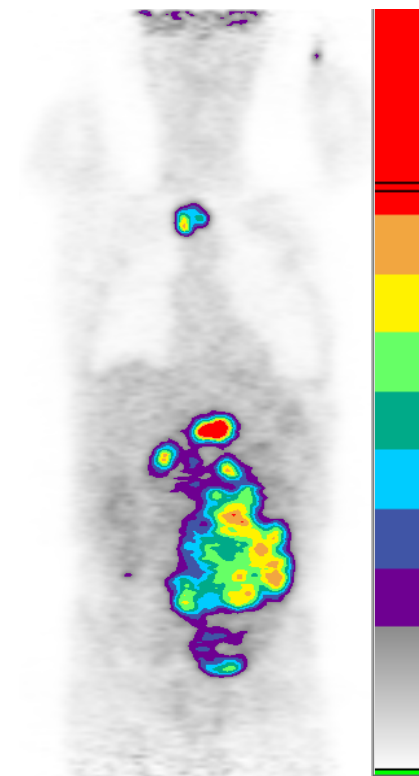
Menton April 8th, 2010



Criteria for interim PET assessment

Quantitative analysis in AOM00152

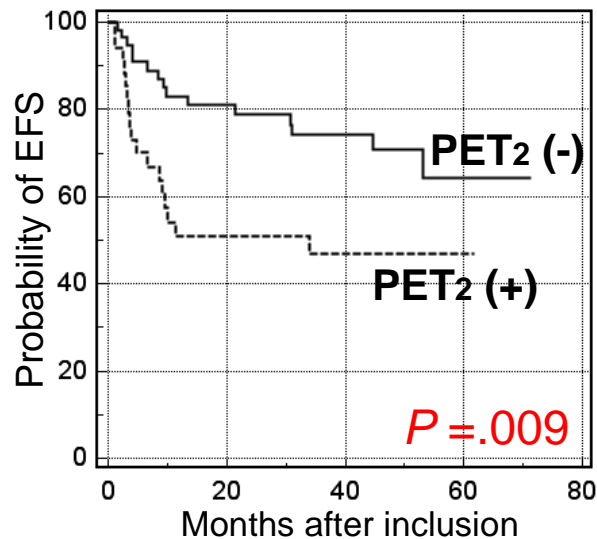
- Retrospective analysis
- 92 pts with DLBCL, median f-u 4 y
- Baseline PET :
 - SUV_{max} in the most active lesion
 - whichever CT size or location
- Interim PET :
 - if (+) → in the most active lesion
 - if (–) → in the area of PET₀ tumor
- Calculation of % of SUV_{max} reduction
- Optimal cut-offs determined by ROC



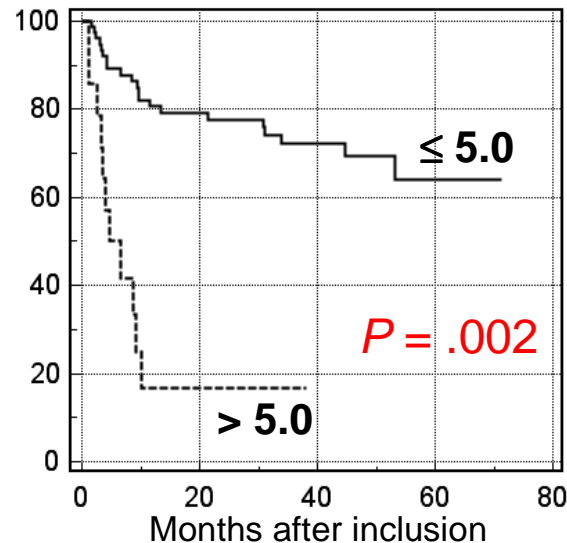
Visual vs. quantitative analysis

2 cycles, n=92

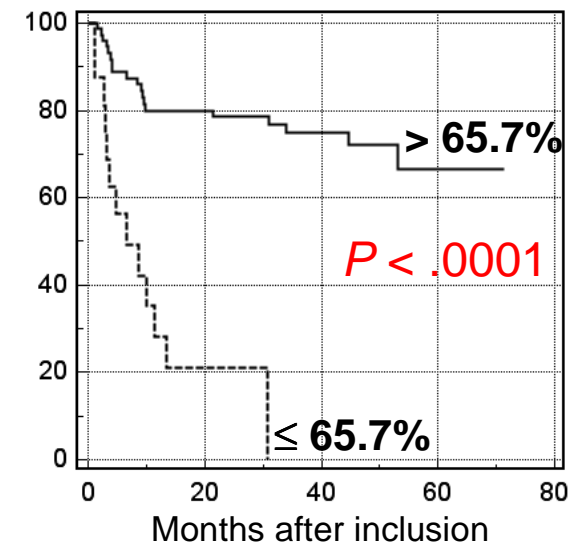
Visual analysis
(Créteil, MRU)



Quantitative analysis
(SUV_{max} at 2 cycles)



Quantitative analysis
(% reduction SUV_{max})

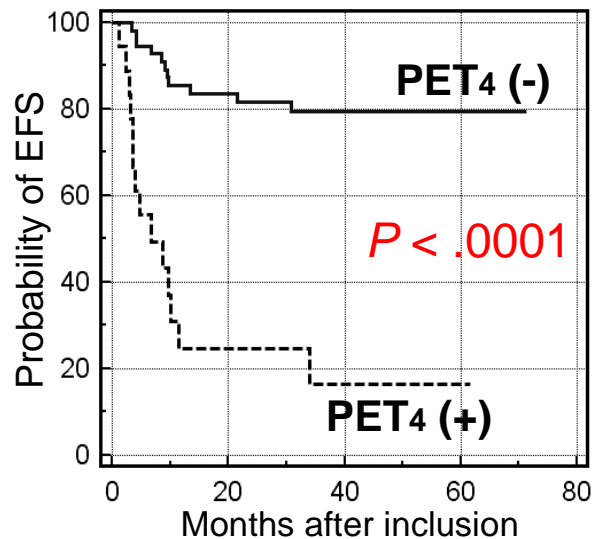


- Reduction of 14/17 false positives
- Cut-off may vary with histology, treatment, PET center

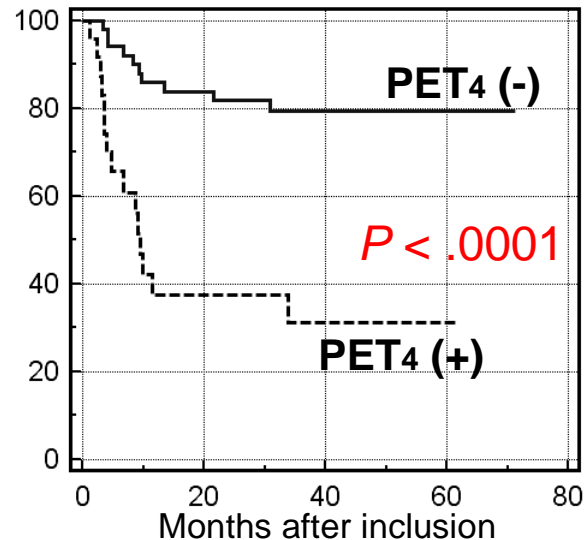
Visual vs. quantitative analysis

4 cycles, n=80

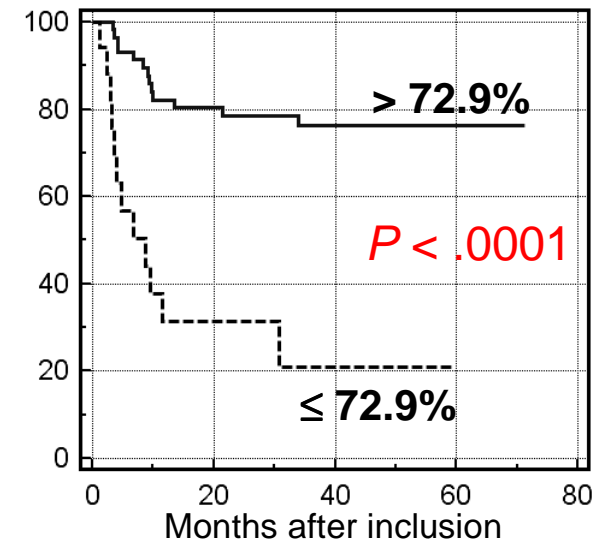
Visual analysis
(Créteil, MRU)



Visual analysis
(Juweid, IHP)



Quantitative analysis
(% reduction SUV_{max})



- Reduction of false positives if we wait for 4 cycles
- Juweid criteria acceptable, Créteil slightly better
- Visual analysis reliable, ΔSUV more objective

Qualitative assessment at 4 cycles

Independent prognostic factor

Overall Model Fit

| | |
|------------------------------|------------|
| Null model -2 Log Likelihood | 154,74219 |
| Full model -2 Log Likelihood | 134,96769 |
| Chi-square | 19,7745 |
| DF | 5 |
| Significance level | P = 0,0014 |

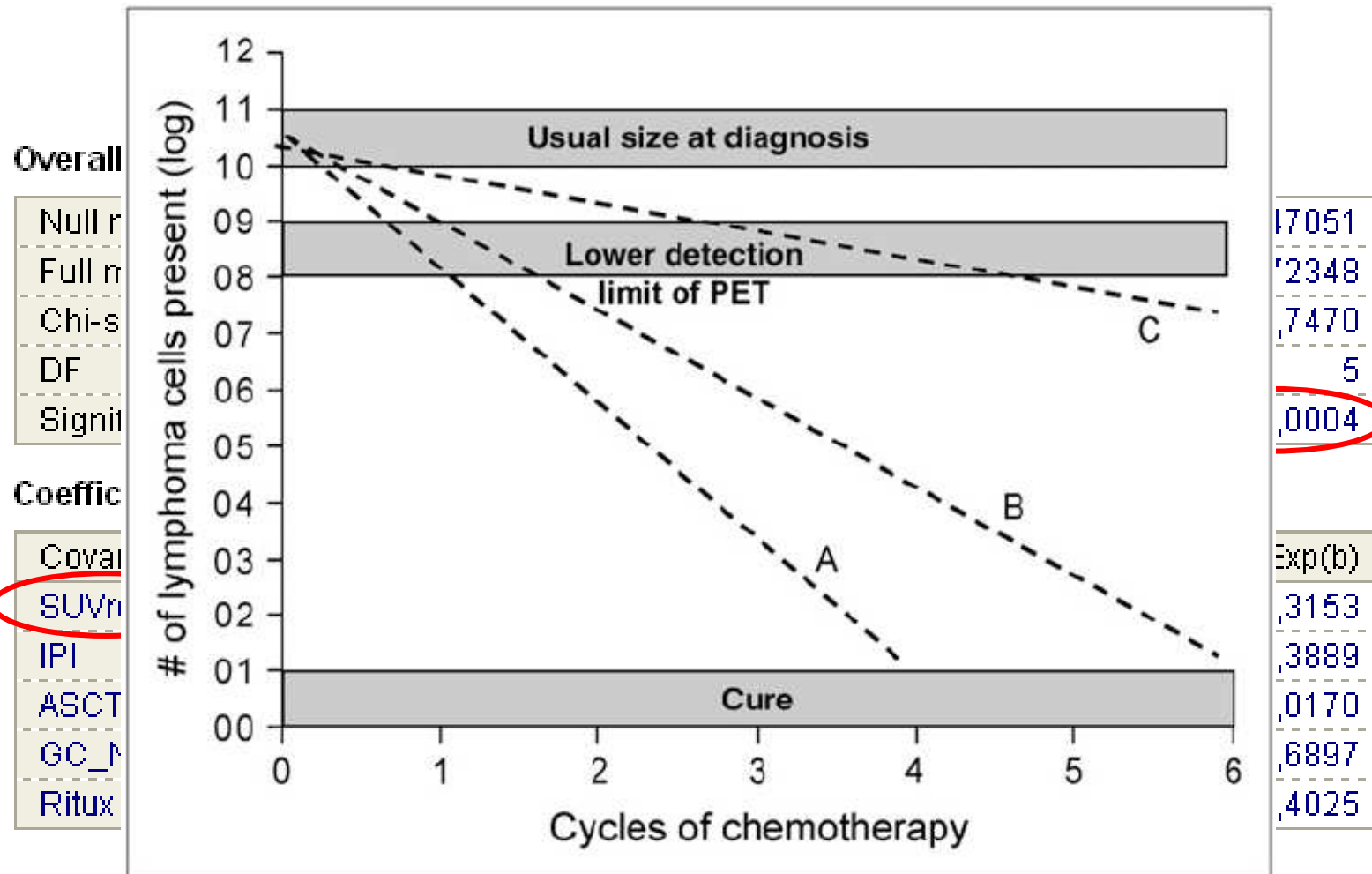
Coefficients and Standard Errors

| Covariate | b | SE | P | Exp(b) | 95% CI of Exp(b) |
|-----------|---------|--------|--------|--------|-------------------|
| PET4vis | 1,9252 | 0,4775 | 0,0001 | 6,8563 | 2,7023 to 17,3960 |
| IPI | 0,2145 | 0,1907 | 0,2606 | 1,2392 | 0,8545 to 1,7973 |
| ASCT | -1,1057 | 0,5192 | 0,0332 | 0,3310 | 0,1203 to 0,9110 |
| GC_NGC | 0,0465 | 0,4602 | 0,9196 | 1,0476 | 0,4270 to 2,5701 |
| Ritux | -0,3421 | 0,4812 | 0,4771 | 0,7103 | 0,2779 to 1,8152 |

→ Independent from IPI, treatment regimen, gene profiles

Quantitative assessment at 2 cycles

Independent prognostic factor



→ Δ SUV reflects tumoral destruction kinetics

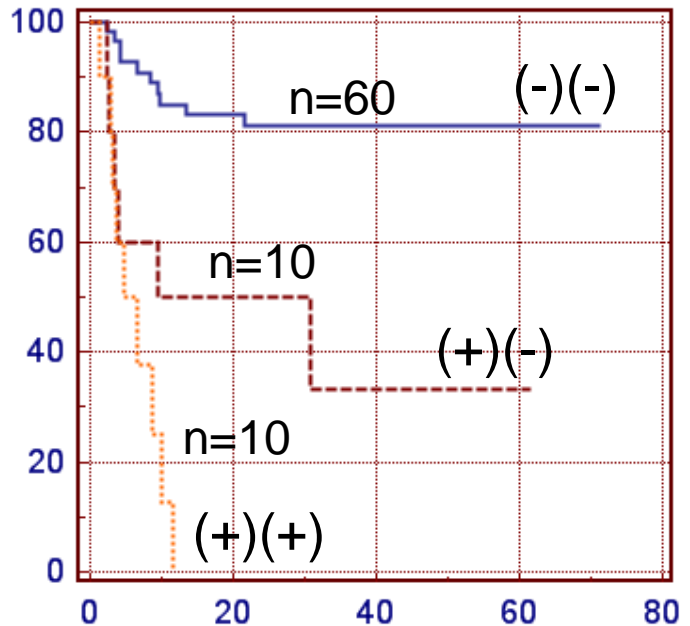
Association of both Δ SUV-PET2 and visual-PET4

Overall Model Fit

| | |
|------------------------------|------------|
| Null model -2 Log Likelihood | 154,74219 |
| Full model -2 Log Likelihood | 130,92093 |
| Chi-square | 23,8213 |
| DF | 5 |
| Significance level | P = 0,0002 |

Coefficients and Standard Errors

| Covariate | b | SE | P | Exp(b) | 95% CI of Exp(b) |
|-------------|---------|--------|--------|--------|------------------|
| PET2red4vis | 1,2301 | 0,274 | 0,0000 | 3,4215 | 2,0050 to 5,8389 |
| IPI | 0,1147 | 0,2053 | 0,5764 | 1,1215 | 0,75 to 1,69 |
| ASCT | -0,8722 | 0,5177 | 0,0920 | 0,4180 | 0,15 to 1,15 |
| GC_NGC | 0,2953 | 0,4776 | 0,5364 | 1,3435 | 0,52 to 3,42 |
| Ritux | -0,2408 | 0,4791 | 0,6153 | 0,7860 | 0,30 to 2,00 |

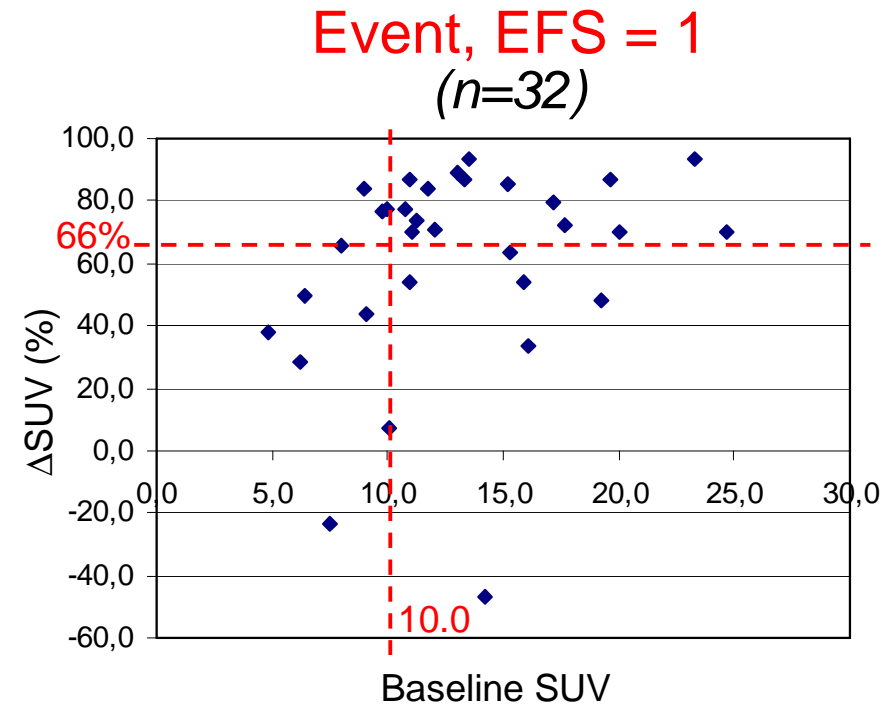
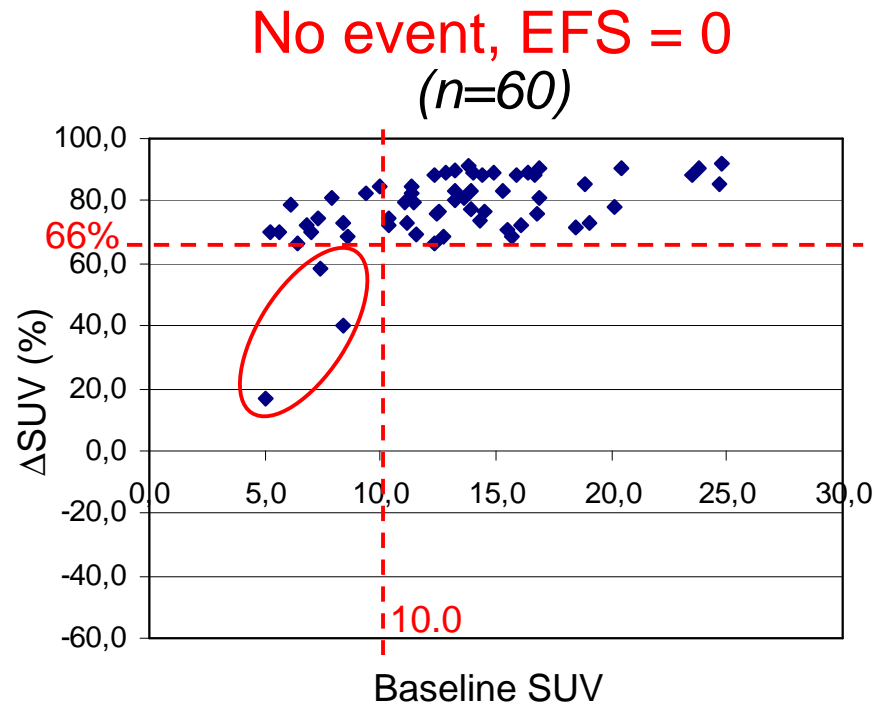


Limitations of Δ SUV

- Necessity of a baseline PET
- Tumors with baseline uptake <10.0
- SUV variability/normalization to internal bkg
- No external validation

Tumors with baseline uptake <10.0

influence of baseline SUV on Δ SUV

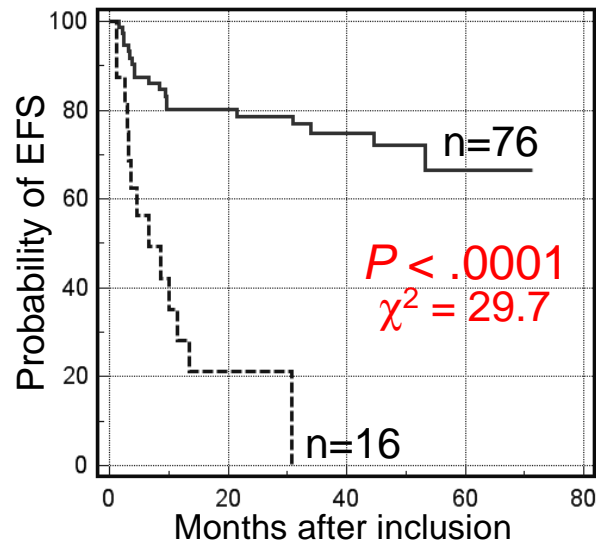


→ 3 FP pts w/ baseline SUV < 10.0, Δ SUV < 66%, no event

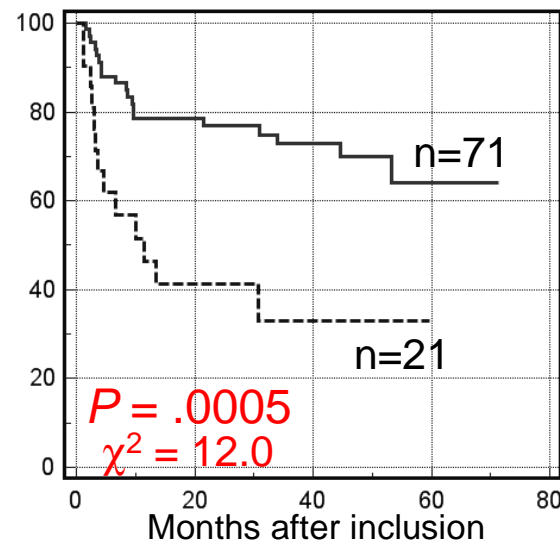
SUV variability

normalization to liver activity

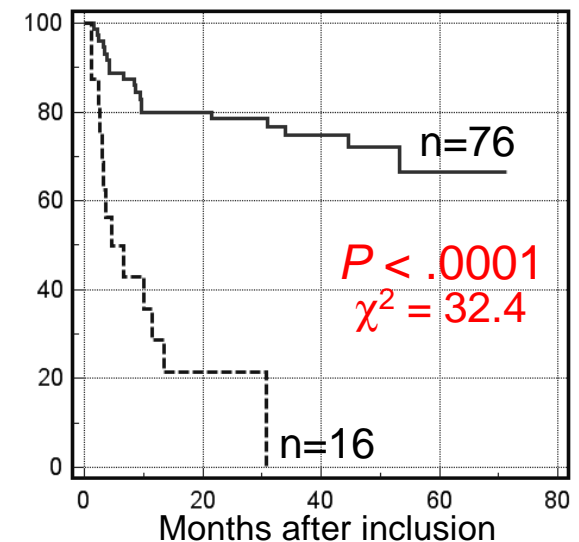
Raw Δ SUV
(cut-off 66%)



Δ SUV / liver
(cut-off 66%)



Δ SUV / liver
(cut-off 60%)

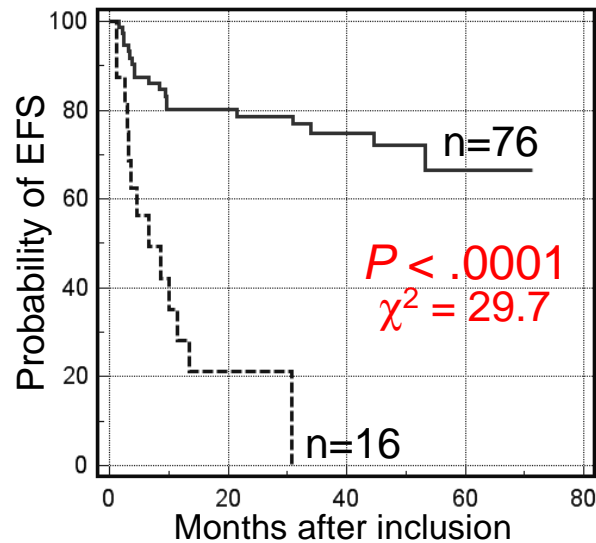


$$\Delta\text{SUV} = 100 \times \frac{\text{SUV}_{T1}/\text{SUV}_{L1} - \text{SUV}_{T2}/\text{SUV}_{L2}}{\text{SUV}_{T1}/\text{SUV}_{L1}}$$

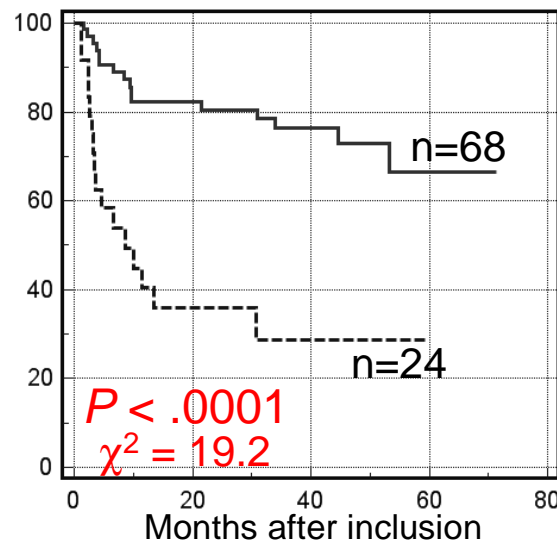
SUV variability

normalization to MBP activity

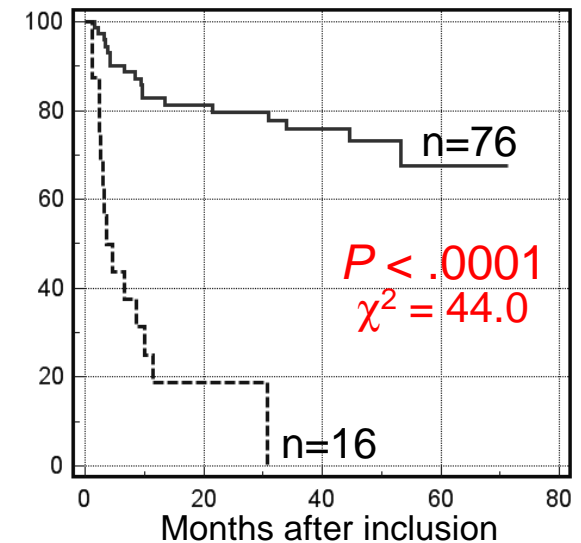
Raw Δ SUV
(cut-off 66%)



Δ SUV / MBP
(cut-off 66%)



Δ SUV / MBP
(cut-off 61%)



$$\Delta\text{SUV} = 100 \times \frac{\text{SUV}_{T1}/\text{SUV}_{M1} - \text{SUV}_{T2}/\text{SUV}_{M2}}{\text{SUV}_{T1}/\text{SUV}_{M1}}$$

Conclusions

- Must follow strict procedure for injection, delay between injection and scanning, glucose level
- Same procedure to identify SUVmax, with help of the MIP, graded color scale
- No need for an internal reference
- External validation : ongoing (PETAL, IVS)