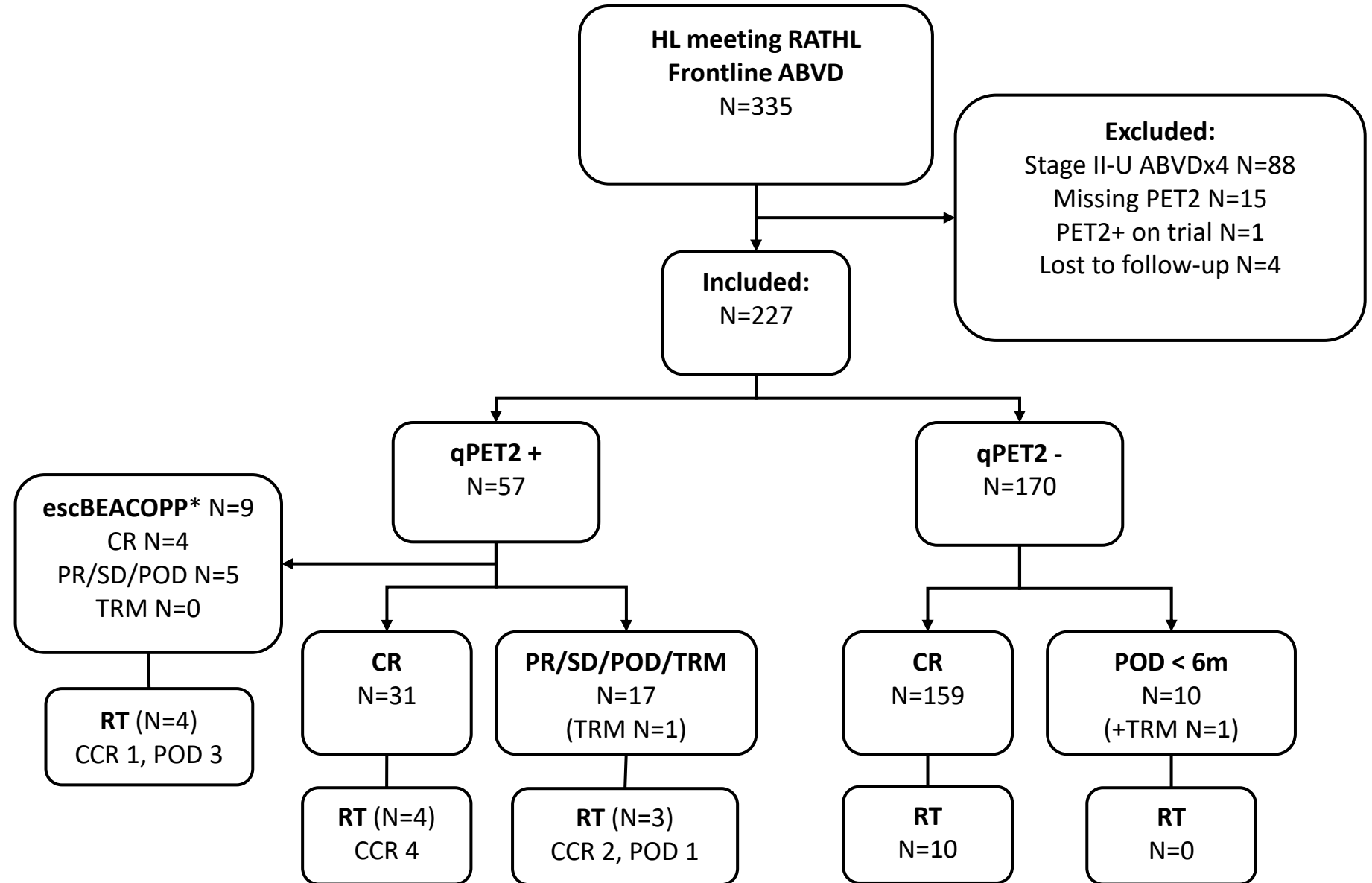


Supplementary Material

Supp. table S1 - Selected studies reporting PET2 on ABVD

Study	N	PET2+ (% , n)	Treatment failure PET2+ (POD + PR + SD)	3y PFS PET2-	3y PFS PET2+		Comment*
Gallamini (2007)	260	19%*	88%	2yPFS 95%*	13%		2y PFS (not 3y) PET2+ by IHP (~Deauville >2)
Biggi (2013)	260	17% (45)	NA	95%*	Central review 28%	Local review 54%	FFS <u>NOT</u> PFS; 25% of PET2+ were FP Excluded pts. w/ escalation based on PET2 8 patients (3+) w/o consensus were PET2+
Oki (2014)	174	17%	NA	83%	~48%		
Zinzani (2012)	304	22%	62%	NA	~40%*		Patient with EOT PET PR to further therapy w/o biopsy = event
Johnson (2016) RATHL	1119	16% (160)	25% * (on escBEACOPP)	85%	68% (on escBEACOPP)		24 pts. Received further chemo for consolidation after escBEACOPP and 43 RT
Zinzani (2016) HD0801	512	20% (103)* 14% DV >3 (72)	50%	2yPFS 81%	2yPFS 76% (IGEV+ASCT)		30% of PET2+ cases that went to intensive treatment were actually Deauville 3 15 PET2+ continued ABVD with 73% CR
Press (2016) SWOG S0816	331	18% (61)	45%* (on escBEACOPP)	2yPFS 82%	2yPFS 64% (on escBEACOPP)		38% PET-PR data not provided on RT additional therapy; 7% SD/UNK
Straus (2021) ECHELON-1	636	9%	NA	3yPFS 76%	3yPFS 55%*		Censoring mPFS at the modified event may have caused an underestimation of PFS. 5y-PFS recently published down to 46%
MSK	227	14%/25%	13%	86%	64%* (by dvPET2+)		If we were to include transition to escBEACOPP as event (PFS 23% if all cases = events PFS 61% if only POD on escBEACOPP = event)

Supp. figure S1 - CONSORT – Response and use of radiotherapy



Images and clinical data of interim PET+ cases (qPET2+ and/or dvPET2+)

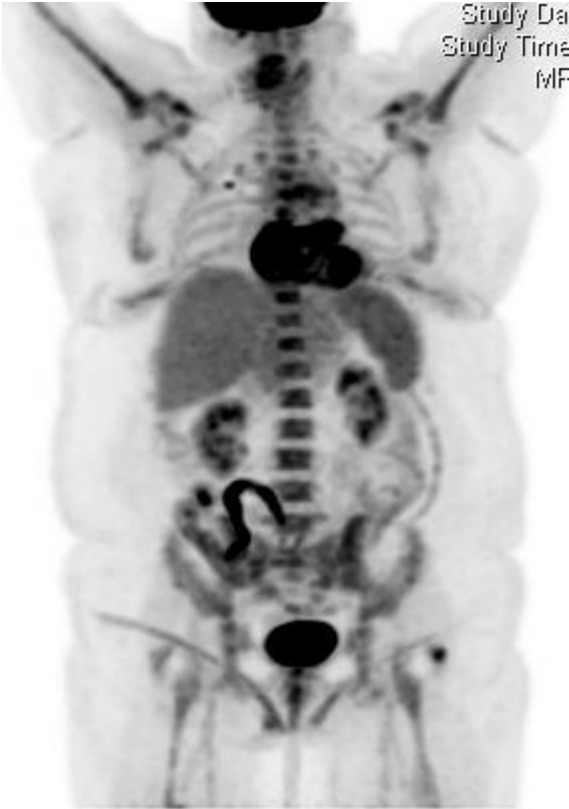
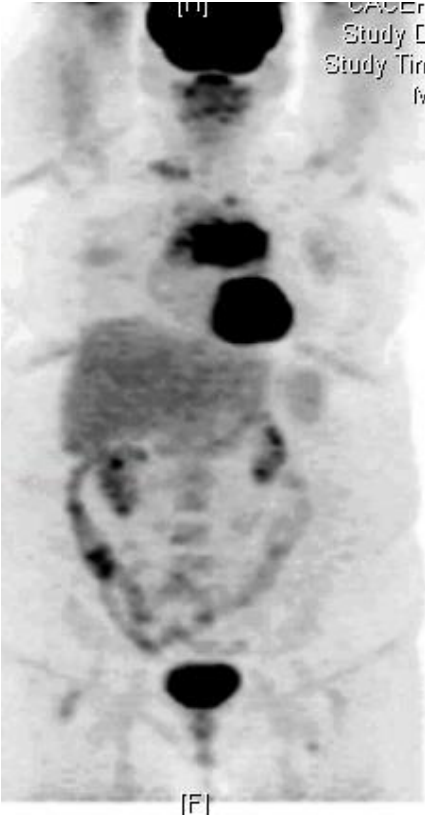
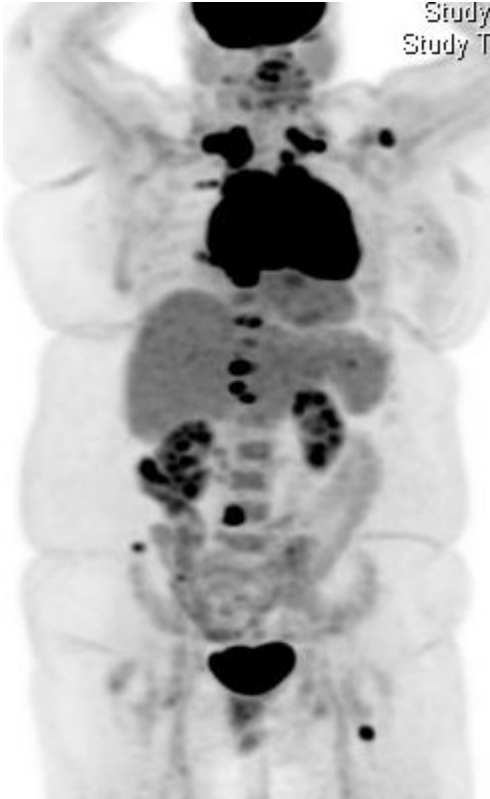
Images are ordered from left to right
Diagnosis – Interim – End of Therapy – [subsequent scans]

Abbreviations

- qPET2+ - lesion SUVmax to liver SUVmean ≥ 1.3
- dvPET2+ - Deauville score 4-5
- OSH – imaging loaded only to main radiology system (not available for download)
- POD – Progression of disease
- Tx 2 – second line treatment
- IPS – International prognostic score
- SUV ratio – lesion SUVmax to liver SUVmean
- SUV – SUV max @ lesion
- Liver – SUVmean @ liver
- Response – by imaging prior to any radiotherapy (imaging PR with negative biopsy or no progression on short follow-up -> CR)
- RT – radiotherapy after frontline therapy
- RT resp – response after radiotherapy
- Rel prev site – relapse at site of initial diagnosis
- Bx confirmed – was biopsy done to confirm relapse

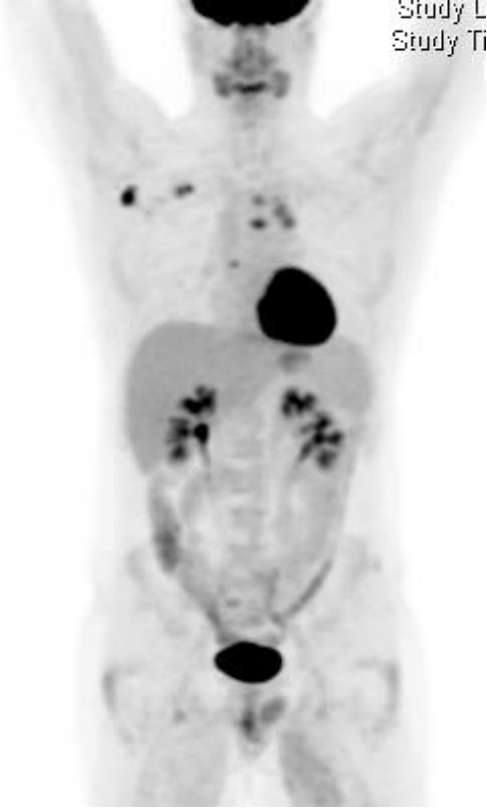
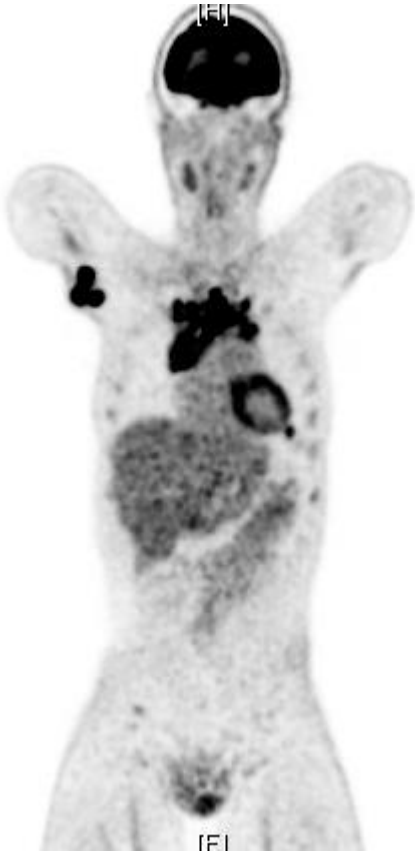
escBEACOPP
Patients with POD

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	53	4-7	BEACOPP	6	≥ 1.3	≥ 4	10	1.8	PR	1	POD	1	12	0	40	1	1



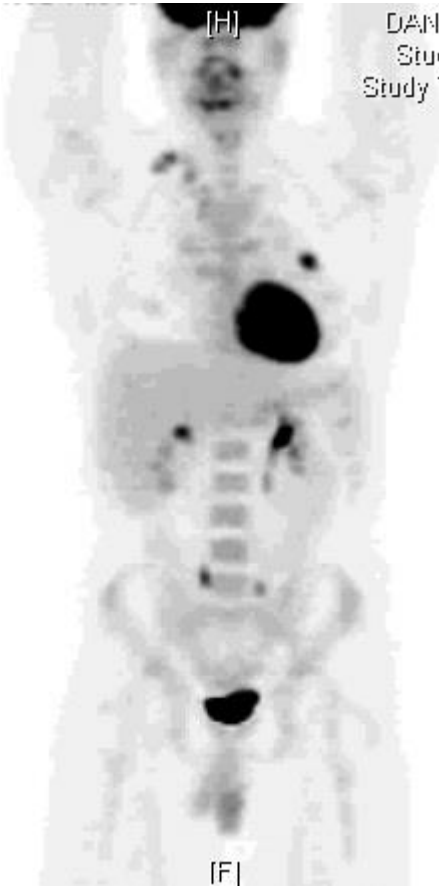
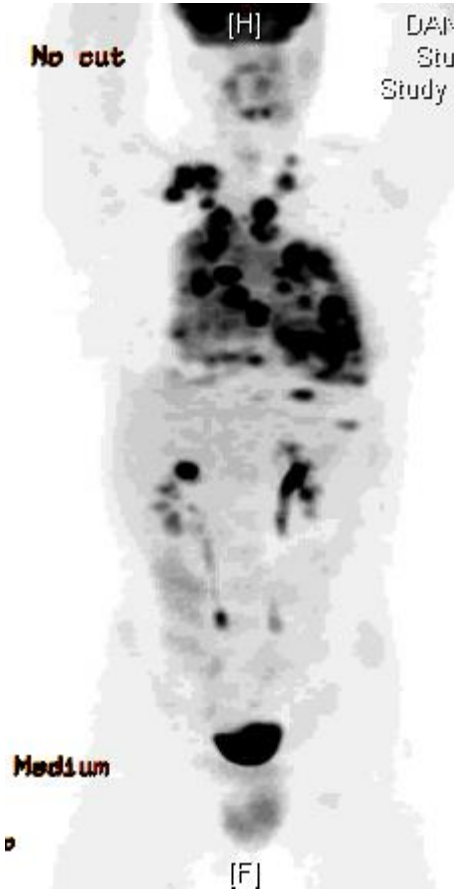
Tx-2
BV-ICE - ASCT

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	36	0-1	BEACOPP	6	>=1.3	>=4	5.2	2.3	POD	0		1	7	0	35	1	1



Tx-2
BV + RT -> ASCT

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	21	0-1	BEACOPP	6	>=1.3	>=4	9	1.55	POD	1	POD	1	10	0	90	1	1

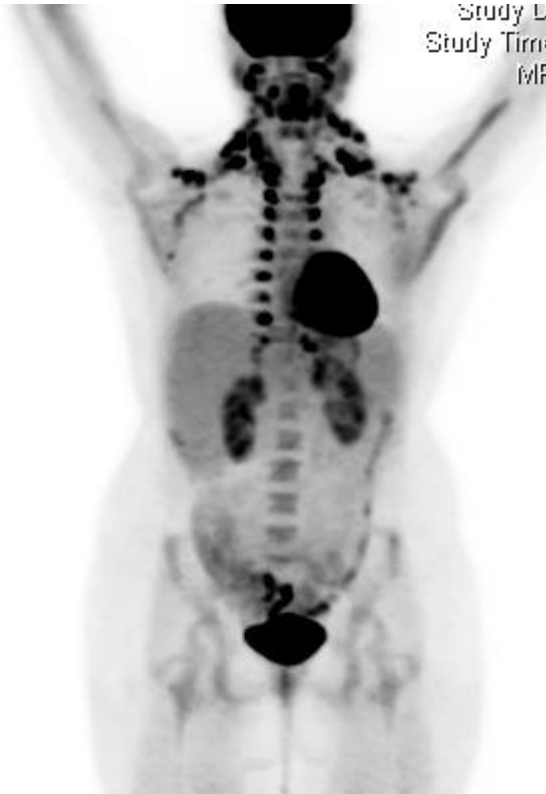


EOT



Relapsed w/in 6 mo

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	25	0-1	BEACOPP	6	>=1.3	>=4	7.8	1.63	POD	1	POD	1	9	0	27	1	1



EOT CR

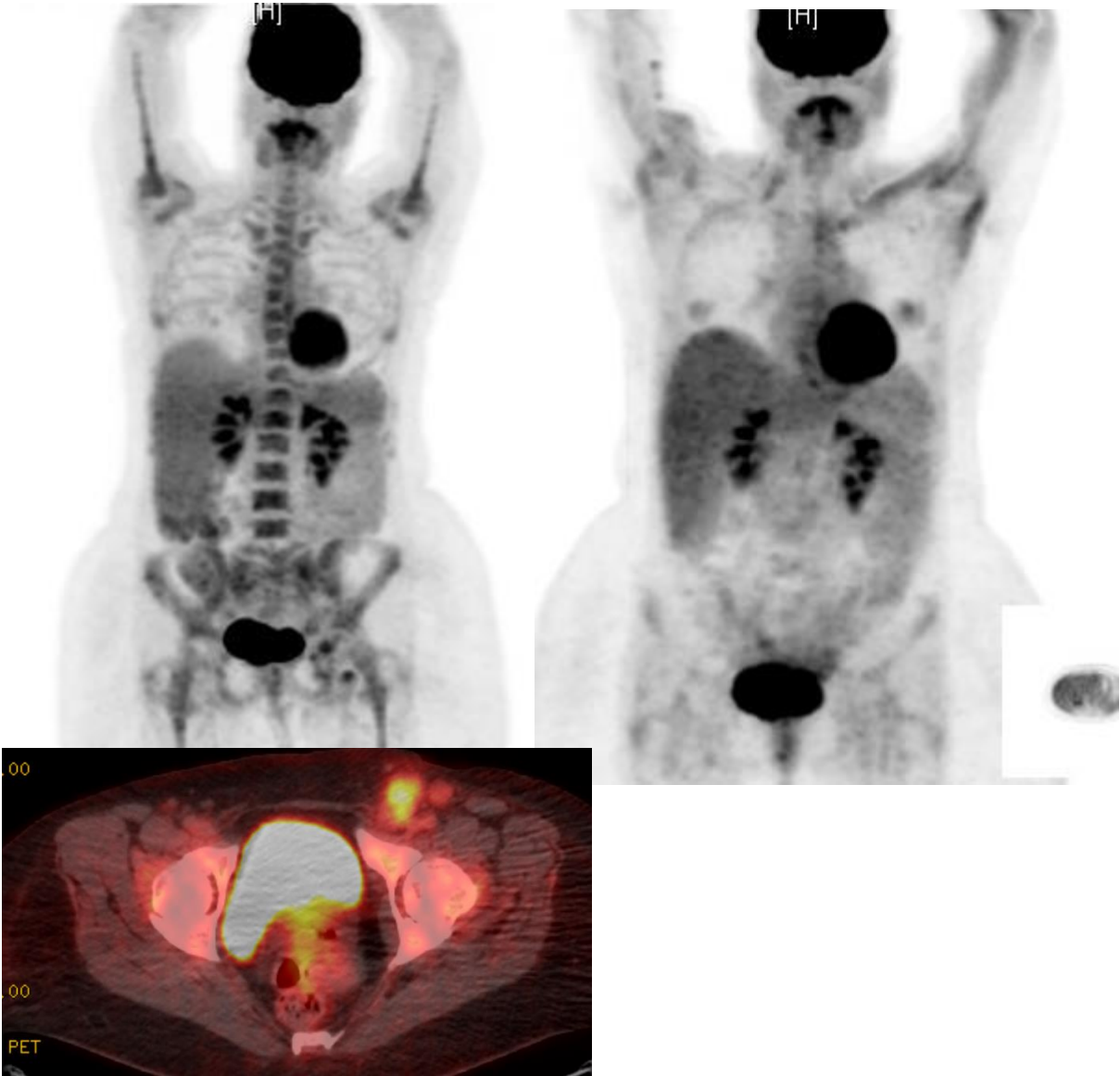


Relapsed within 5 mo

Tx-2
BV-ICE - ASCT

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
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OSH



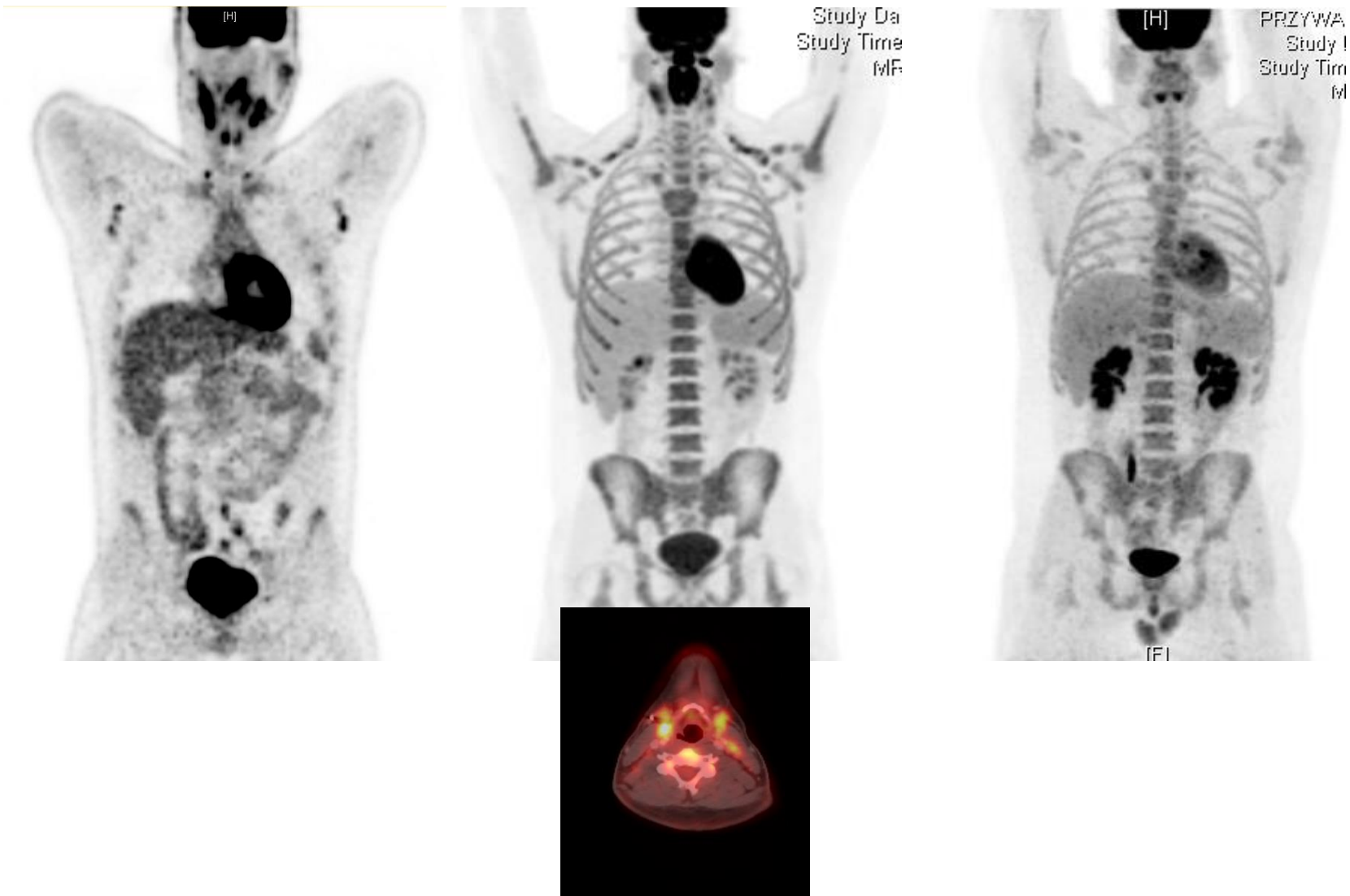
Note: relapse after 39m

Tx-2
BV + RT -> alloSCT

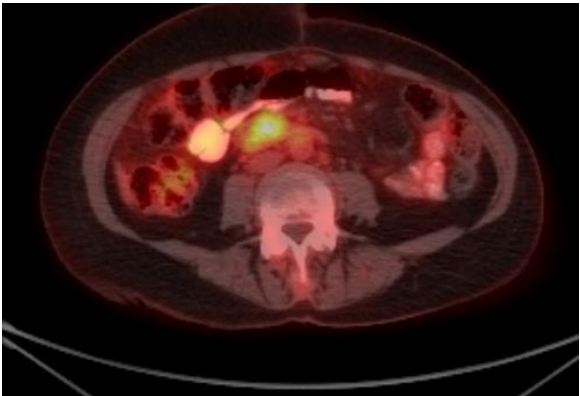
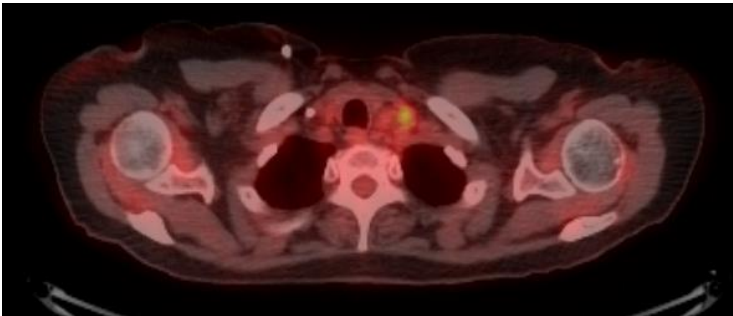
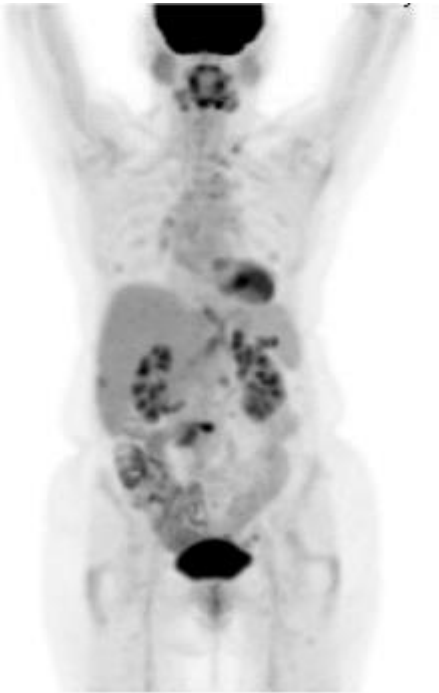
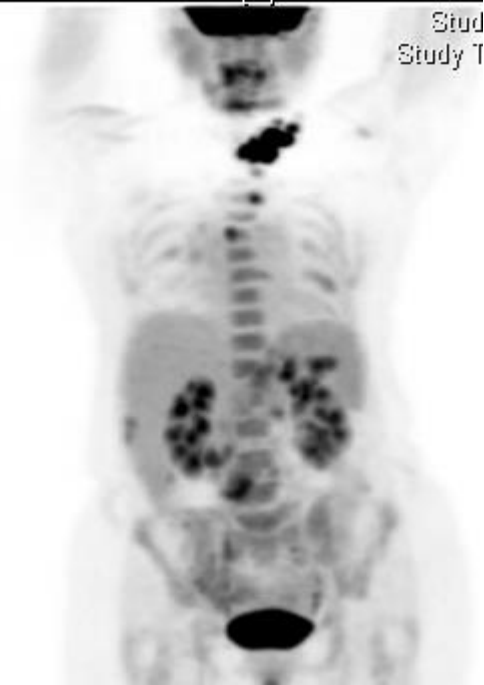
escBEACOPP

Patients with long-term remission

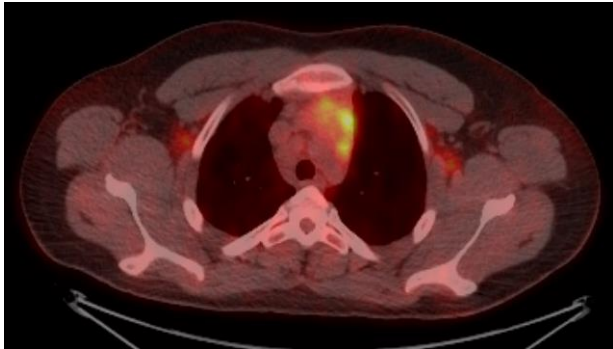
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	19	0-1	BEACOPP	6	>=1.3	>=4	8	2.4	CR	0		0	63	0	63		



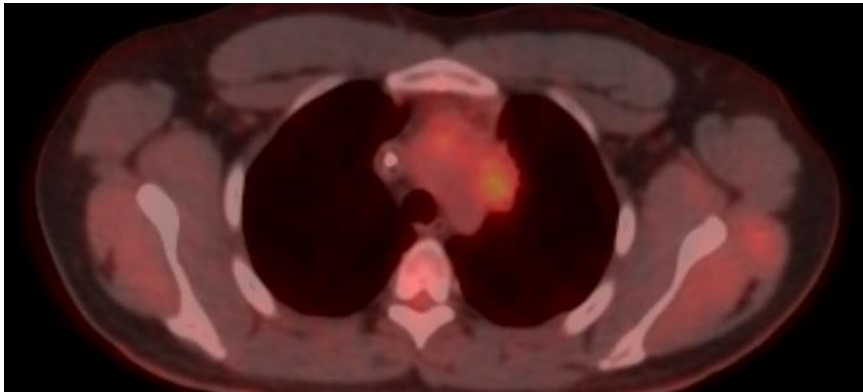
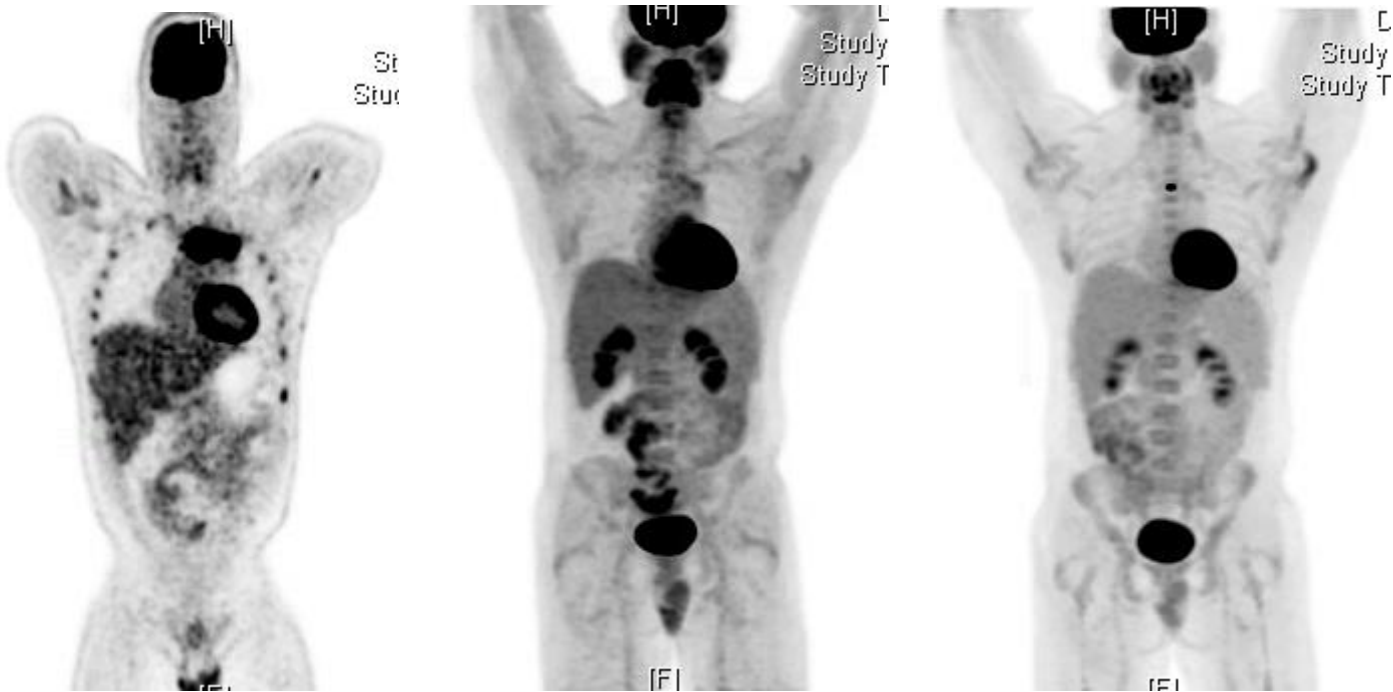
MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	59	4-7	BEACOPP	8	>=1.3	>=4	5.2	2.3	CR	0		0	24	0	24		



MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	19	2-3	BEACOPP	5	>=1.3	>=4	6.4	2.1	PR	1	CR	0	35	0	35		



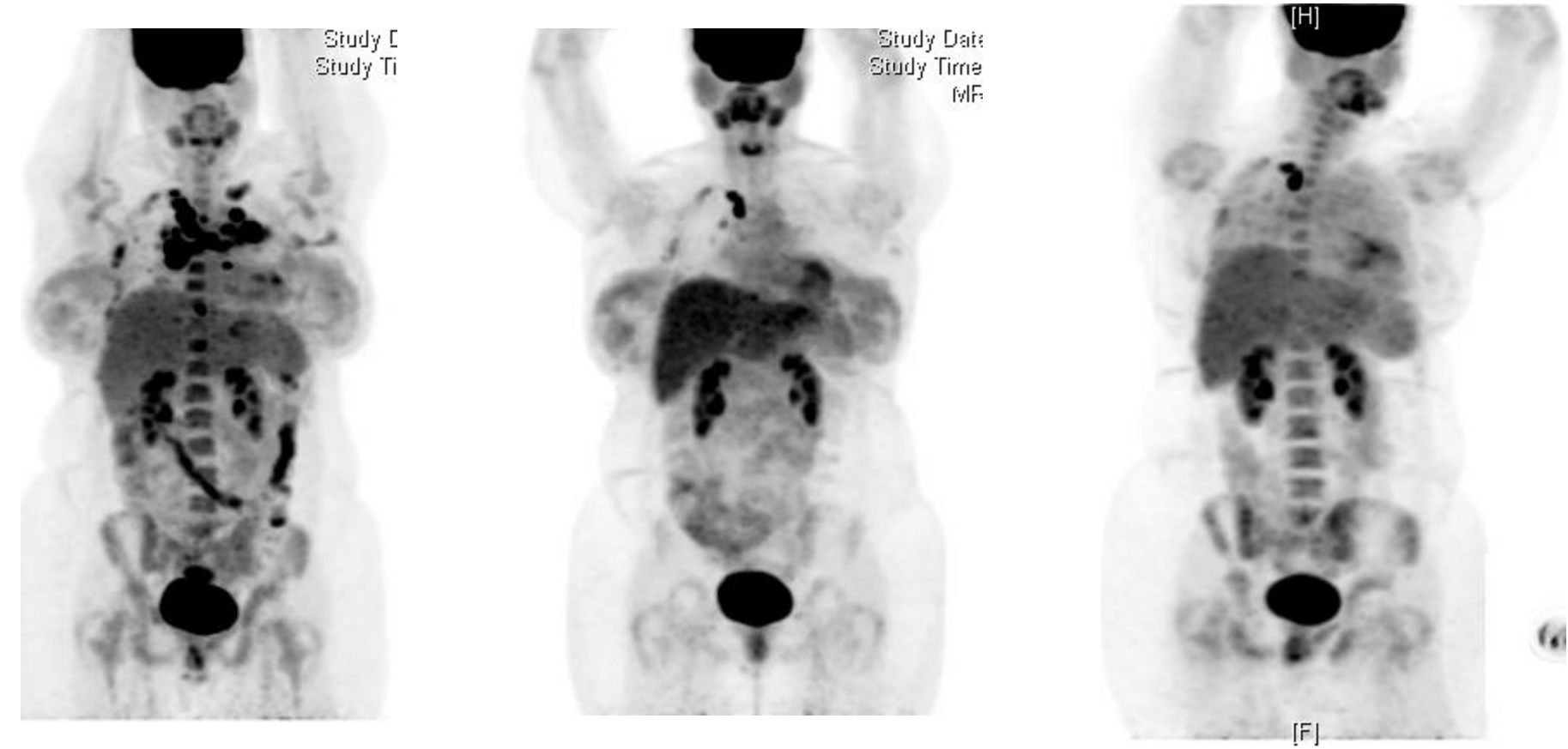
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dvPET2+ (Deauville)

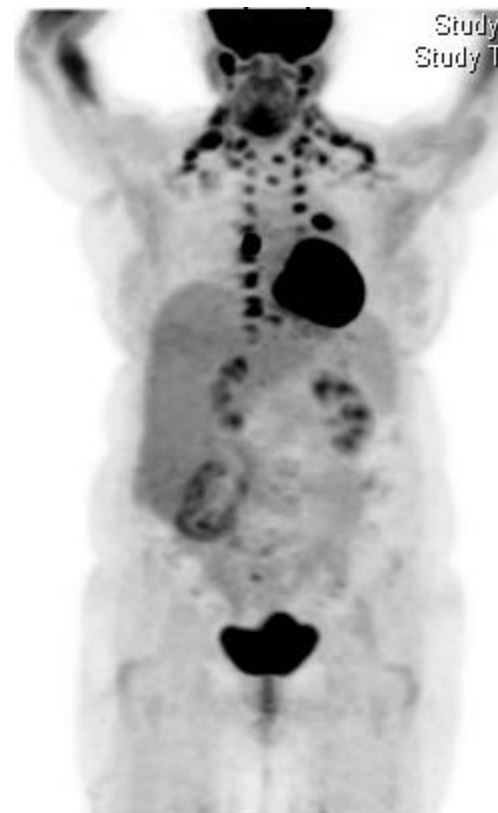
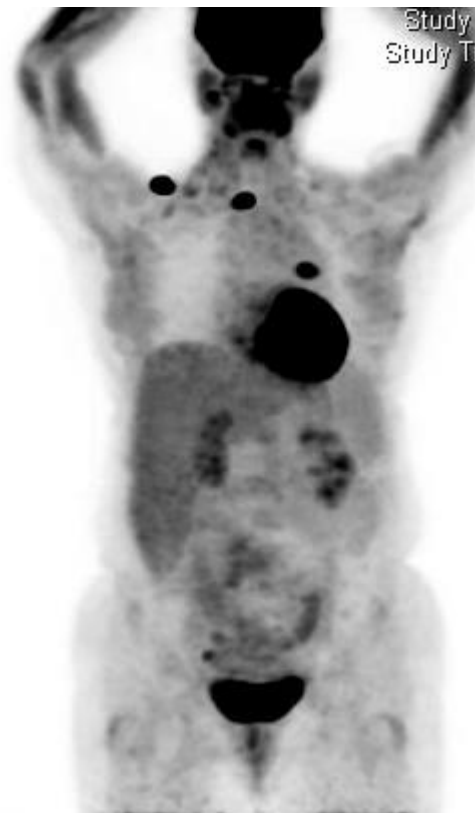
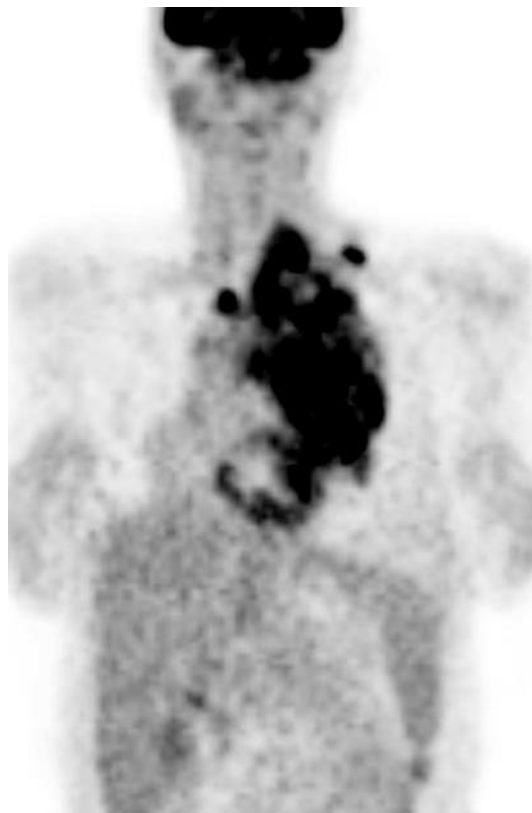
Patients with long-term Remission

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	39	0-1	ABVD	6	>=1.3	>=4	10.8	3.1	CR	0		0	42	0	42		NED



Biopsy w/o HL

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	47	2-3	ABVD	6	≥ 1.3	≥ 4	7.4	1.4	PR	1		0	84	0	84		NED



Persistent FDG avidity at EOT and after RT with biopsy x2
without evidence of HL (histiocytic infiltrate)

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	19	0-1	ABVD	6	≥ 1.3	≥ 4	5.5	1.7	CR	1		0	60	0	60		





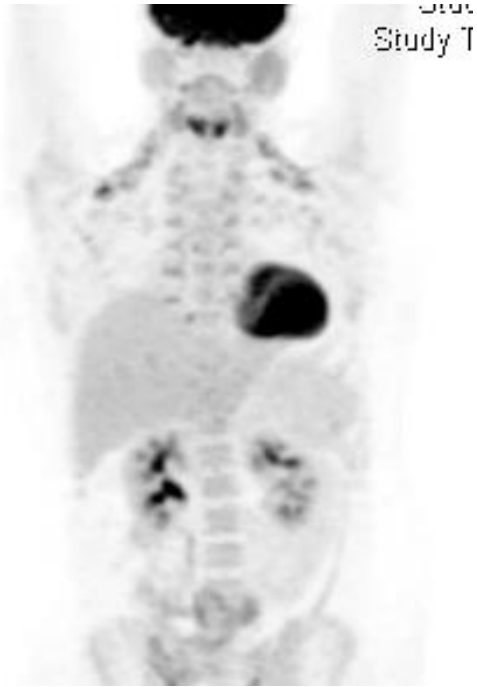
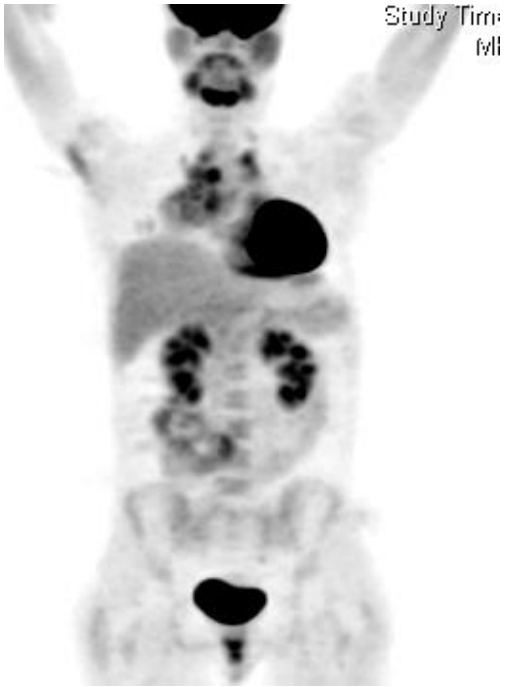


Before RT

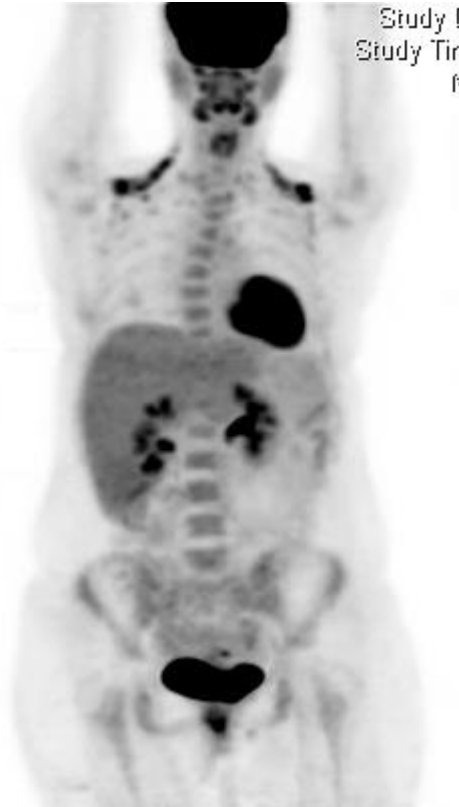
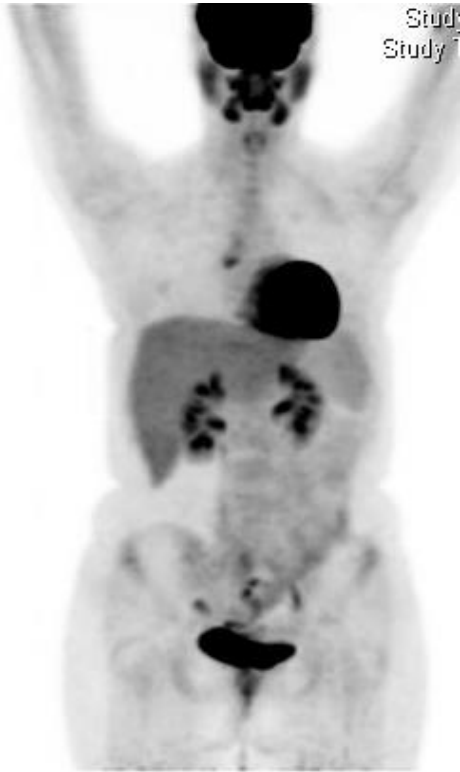
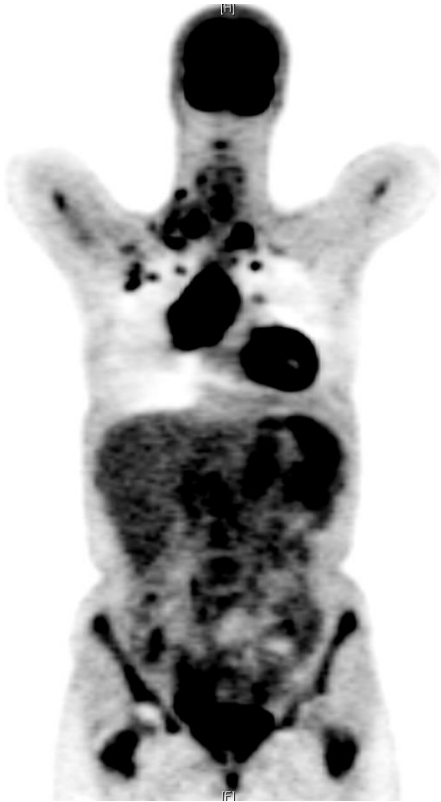


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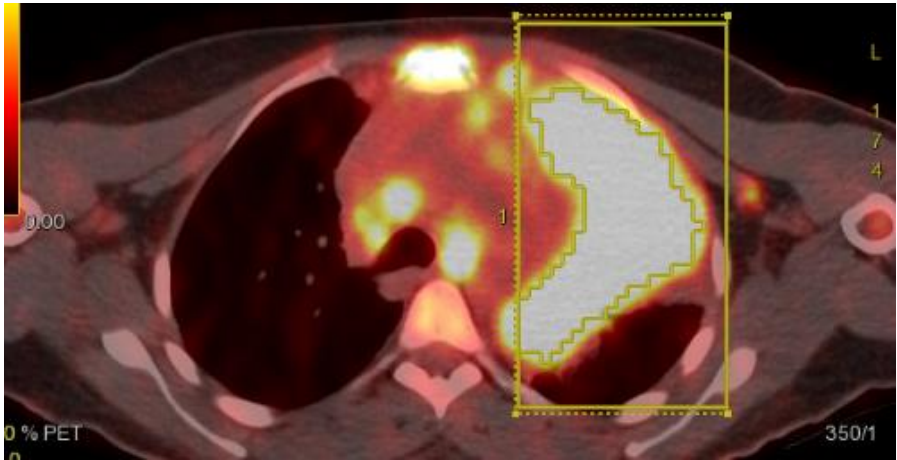
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	23	0-1	ABVD	6	>=1.3	>=4	3.6	1.6	CR	0		0	77	0	77		



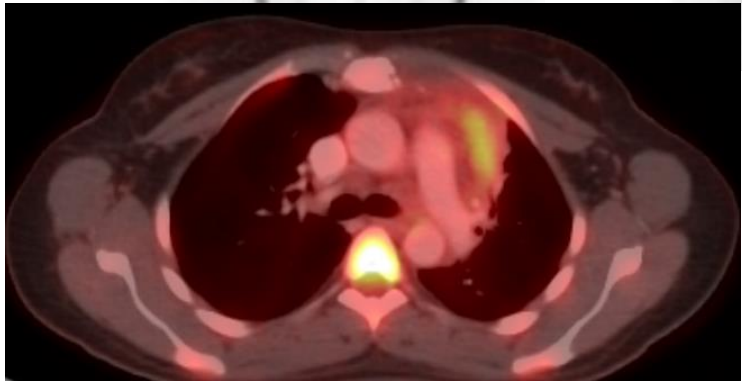
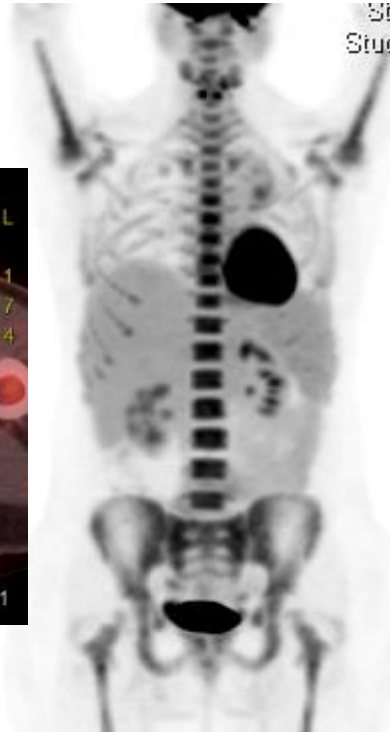
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	26	0-1	ABVD	6	≥ 1.3	≥ 4	2.8	1.68	CR	1		0	49	0	49		



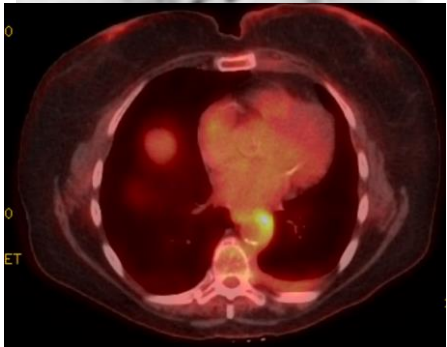
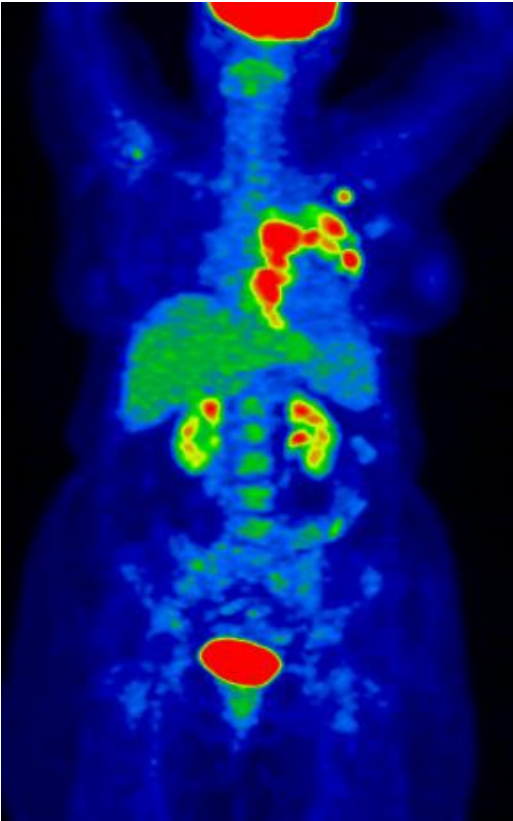
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	37	4-7	ABVD	6	≥ 1.3	≥ 4	4.6	2	CR	0		0	25	0	25		



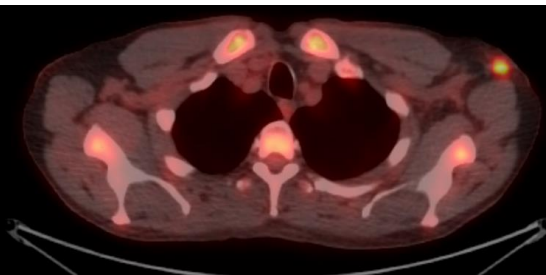
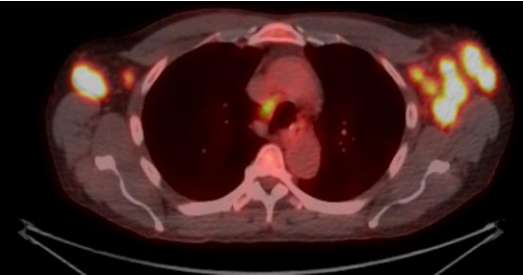
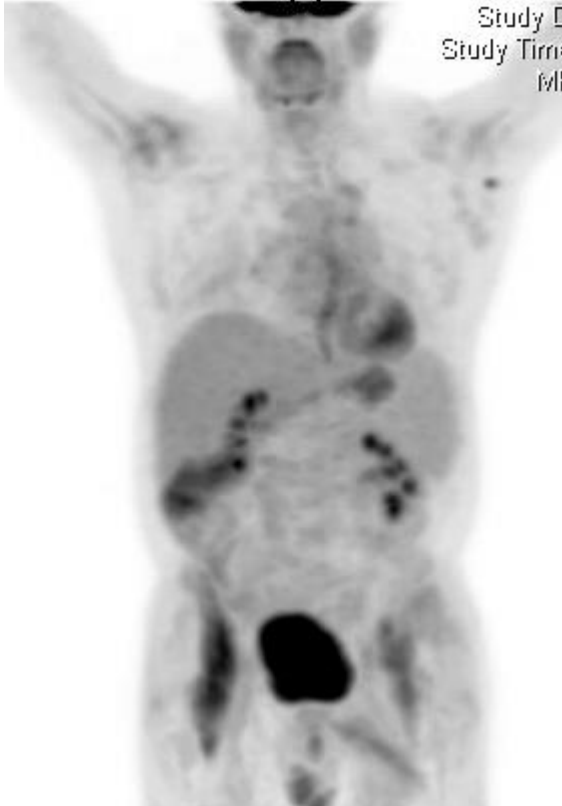
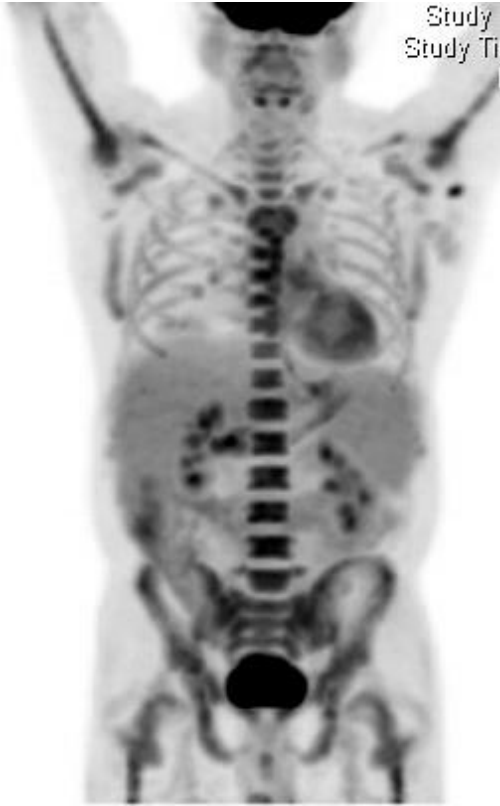
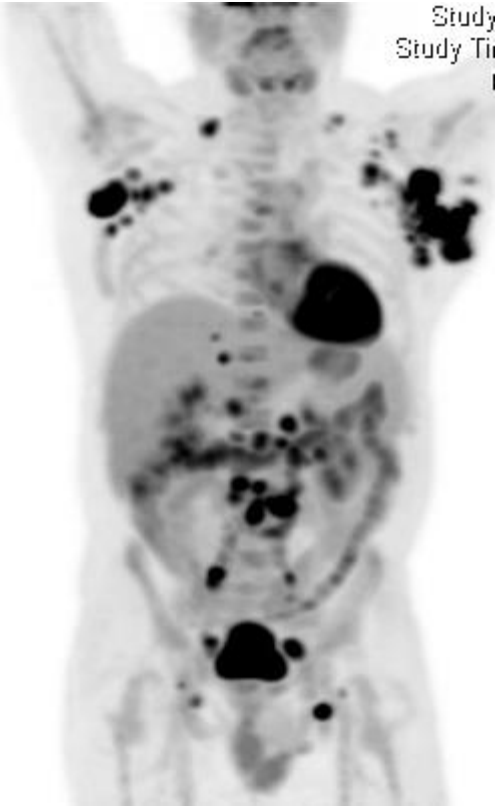
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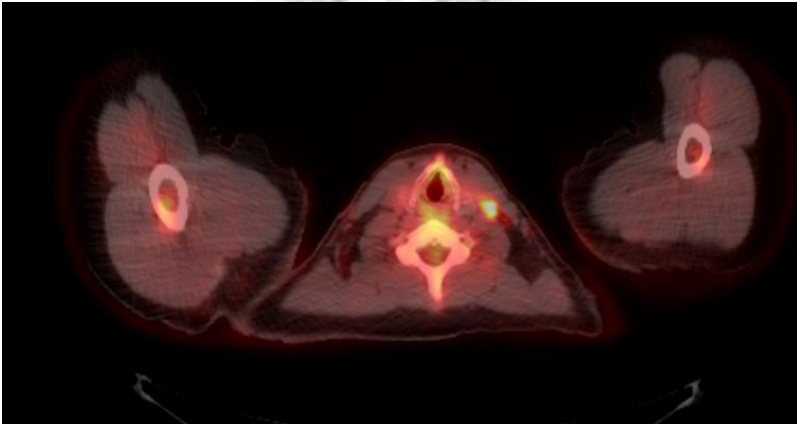
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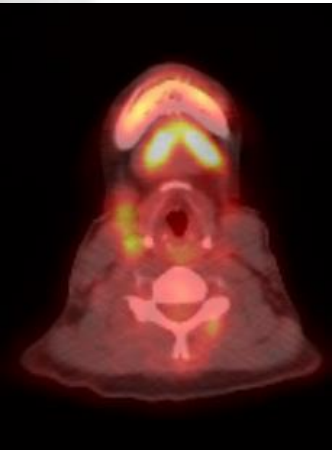
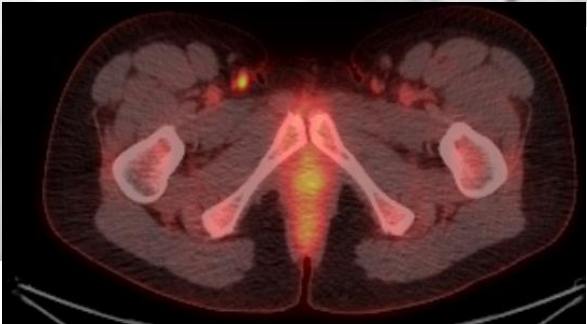
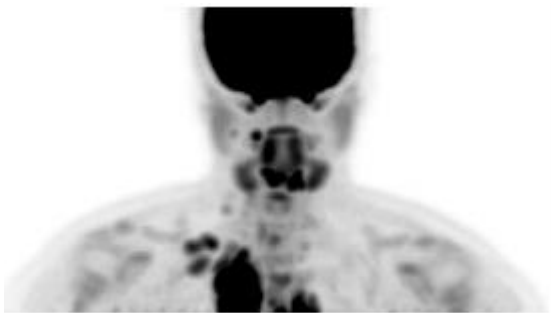
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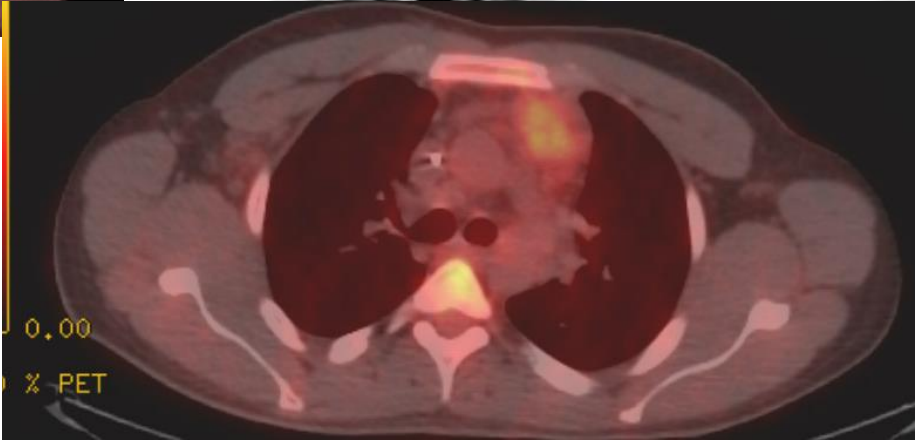
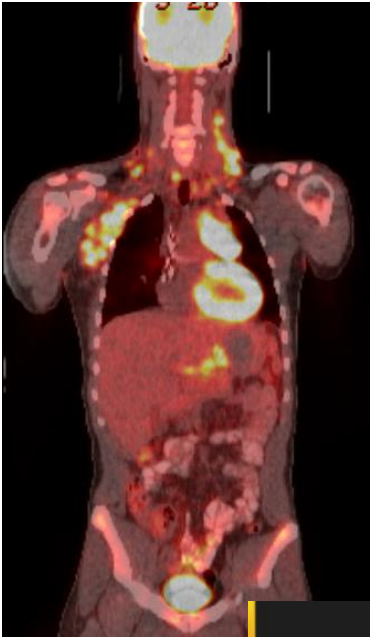
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	34	0-1	ABVD	6	>=1.3	>=4	8.1	2.6	CR	0		0	29	0	29		



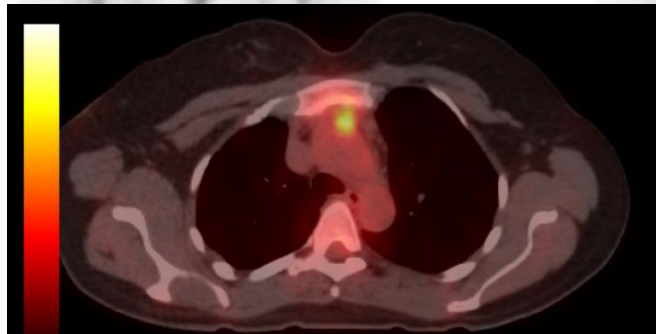
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	35	2-3	ABVD	6	>=1.3	>=4	9	2.3	PR	0		0	57	0	57		



MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	25	2-3	ABVD	6	≥ 1.3	≥ 4	2.9	1.8	CR	0		0	42	0	42		

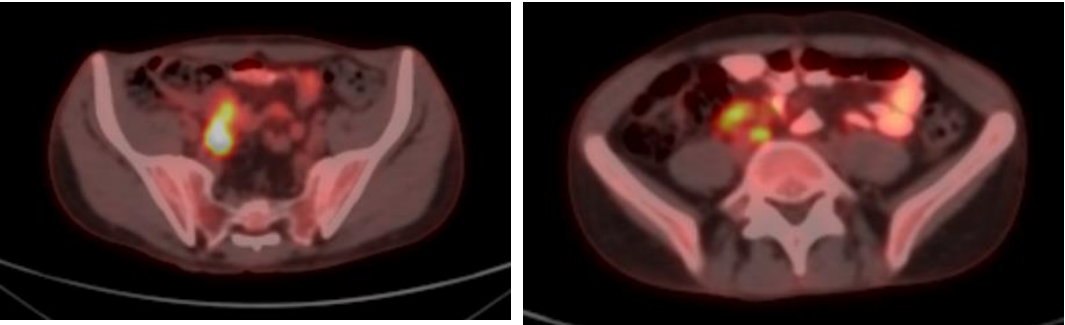
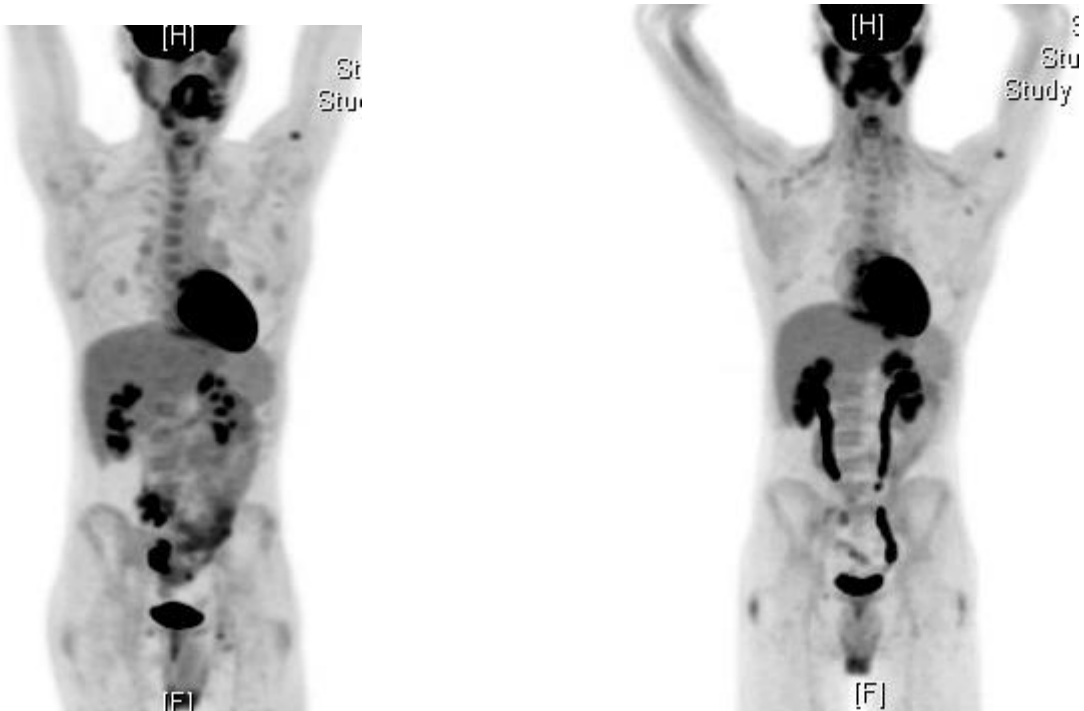


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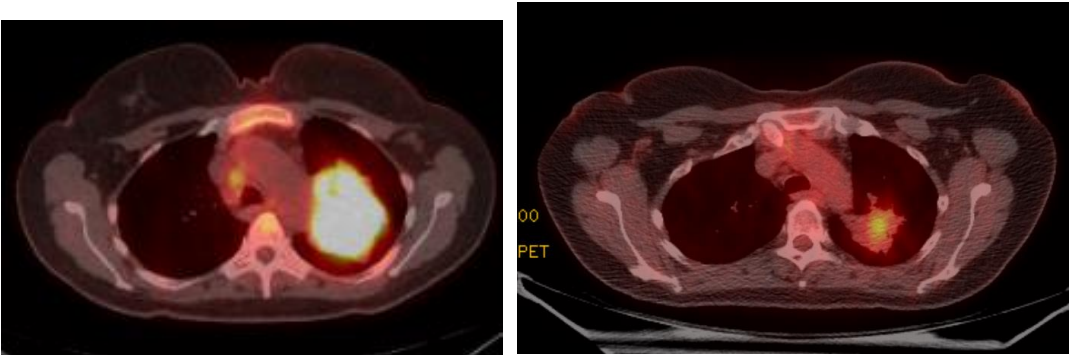
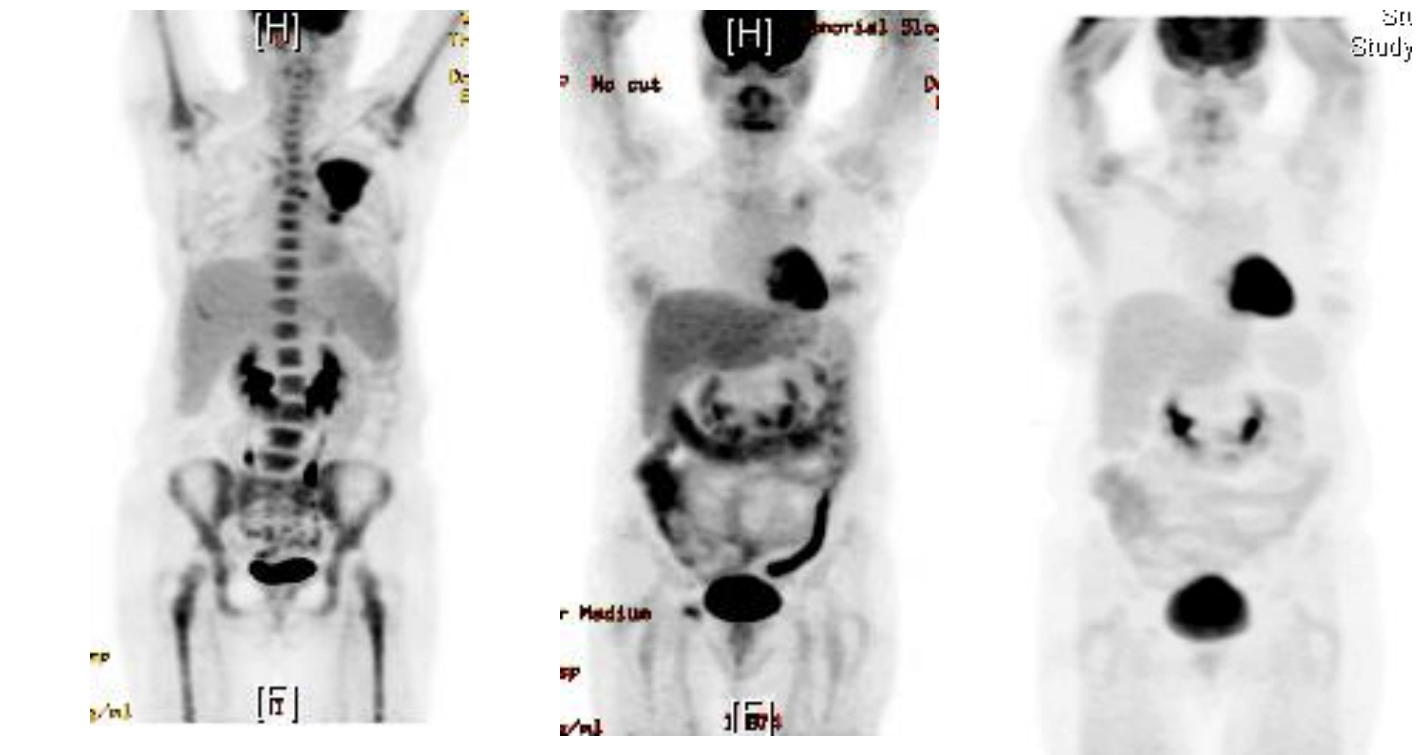


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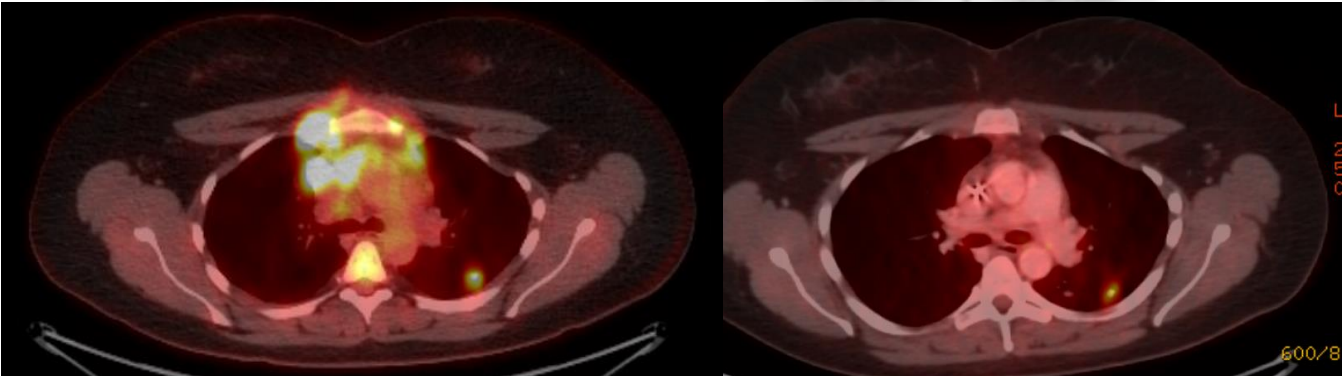
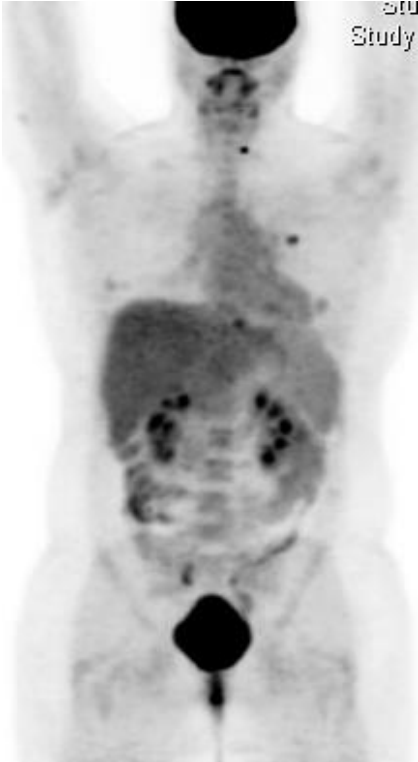
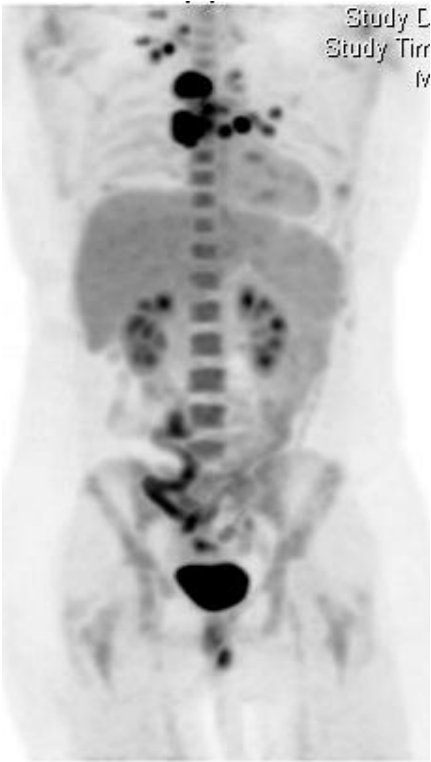
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MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
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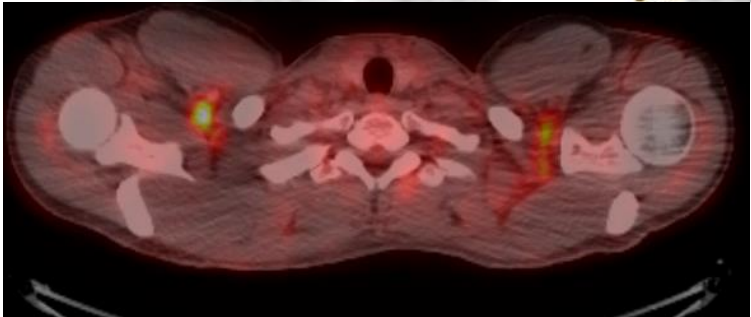
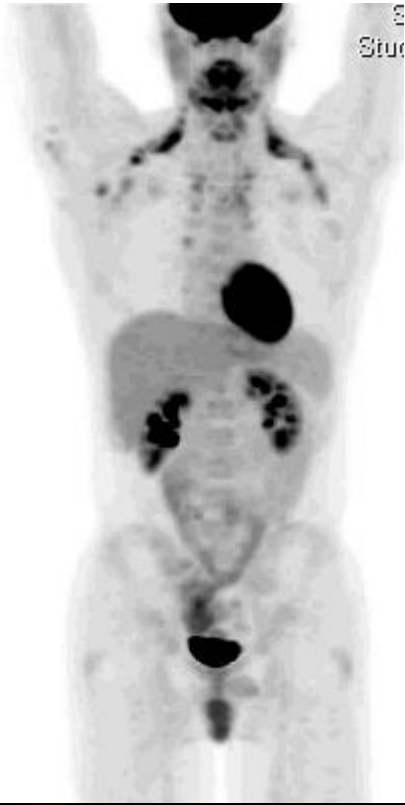
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	39	0-1	ABVD	6	>=1.3	>=4	4.2	2.9	CR	0		0	61	0	61		



MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	34	0-1	ABVD	6	≥ 1.3	≥ 4	4.2	3.32	CR	0		0	30	0	30		



MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
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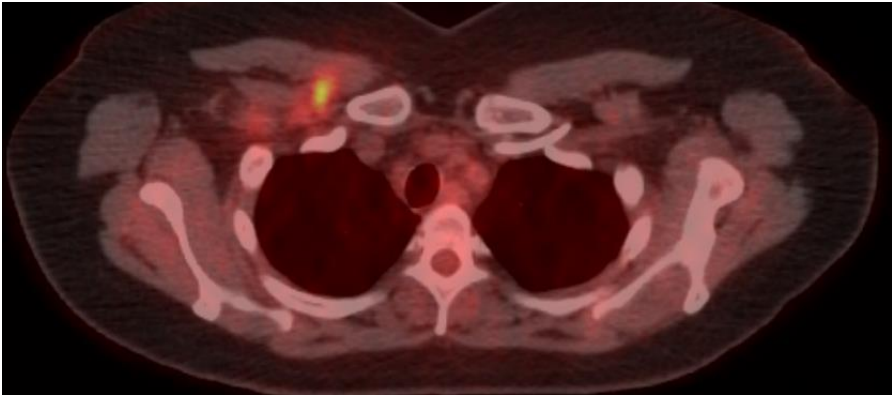


dvPET2+ (Deauville)
Patients with POD

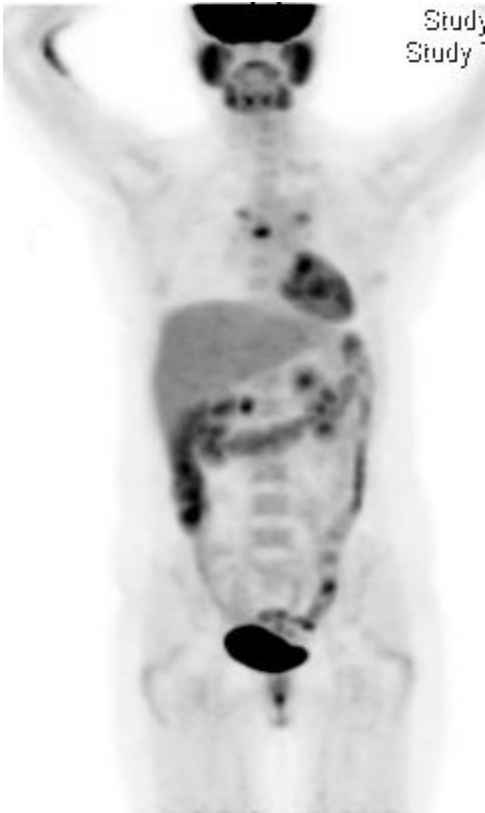
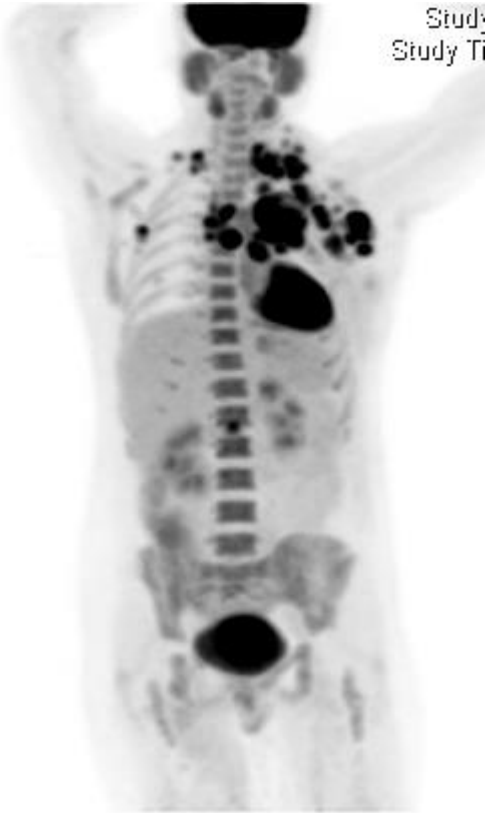
MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	37	2-3	ABVD	4	>=1.3	>=4	7.2	3.4	PR	0		1	4	0	39	1	1



Tx 2
BV+nivolumab - ASCT



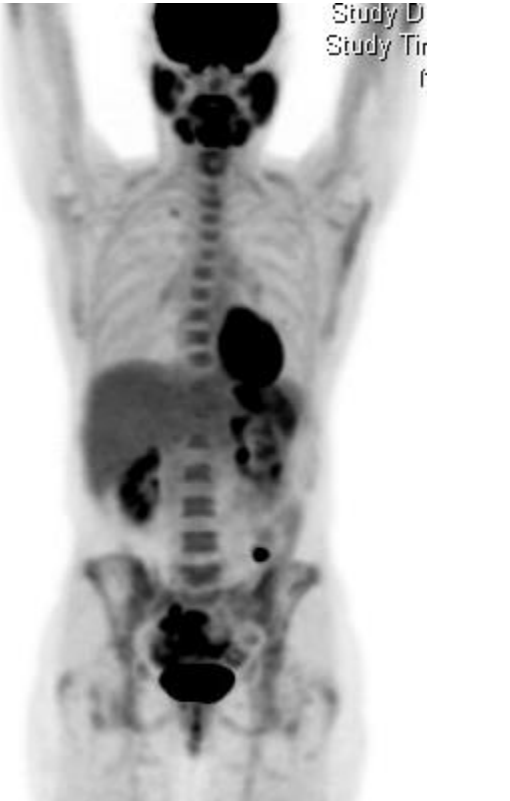
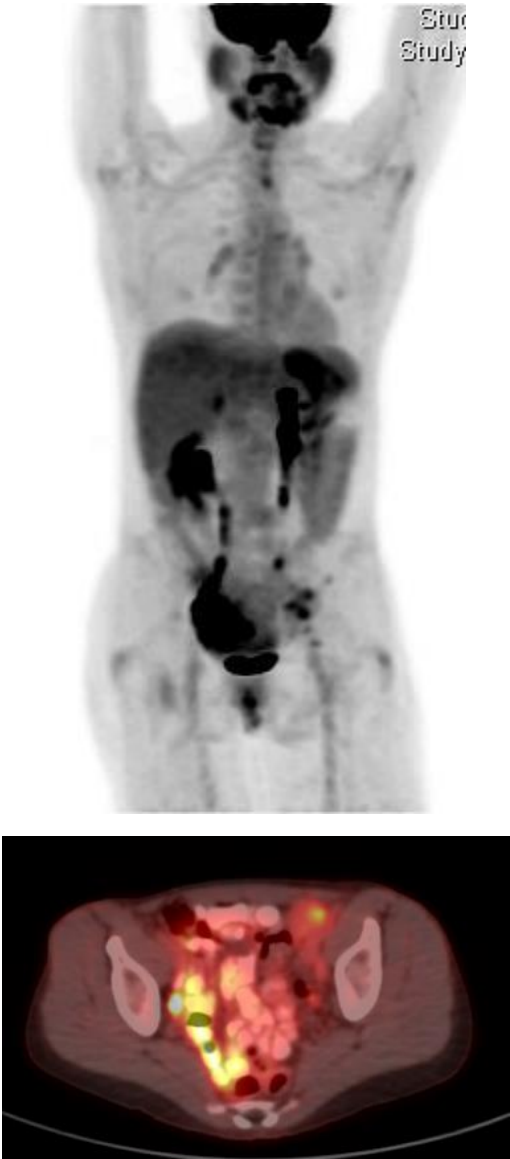
MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	36	0-1	ABVD	6	≥ 1.3	≥ 4	4.3	1.6	POD	0		1	6	1	79	1	1



Tx-2
BV-ICE-ASCT

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	57	0-1	ABVD	6	>=1.3	DV-4	3.7	2.29	POD	0		1	6	0	28	1	1

OSH



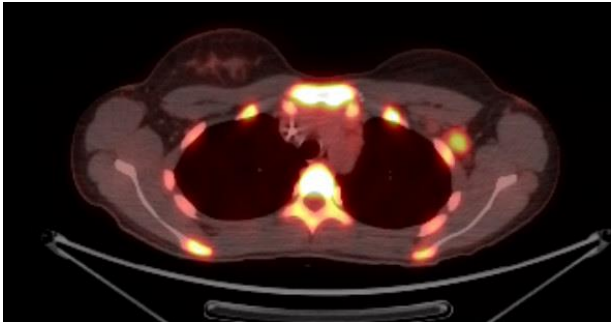
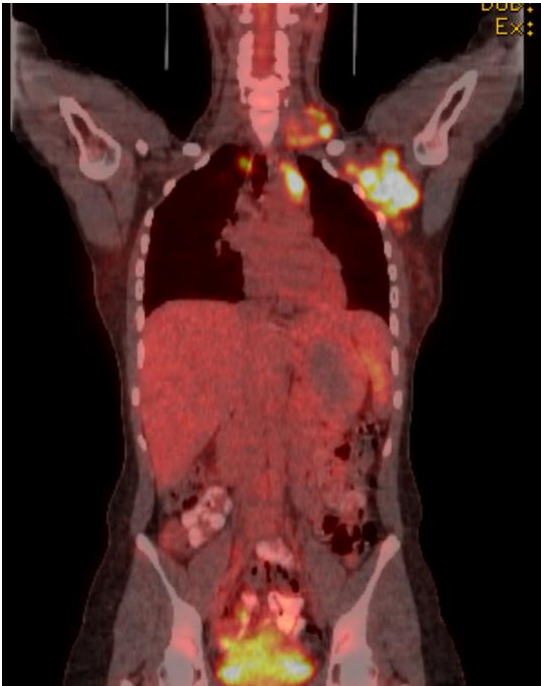
Tx-2
BV-ASCT

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	31	0-1	ABVD	6	>=1.3	>=4	6.9	3.04	POD	0		1	6	1	102	1	1



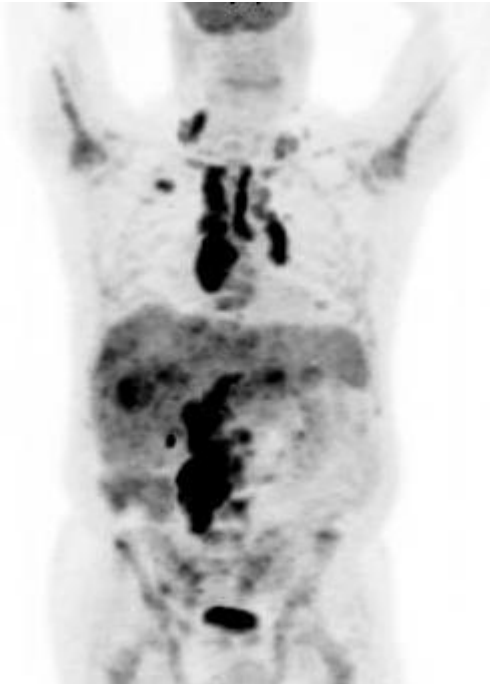
Tx-2
IVG + allo SCT

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	22	0-1	ABVD	4	>=1.3	>=4	4	1.7	PR	0		1	4	0	54	1	1



Tx-2
BV-ICE-ASCT

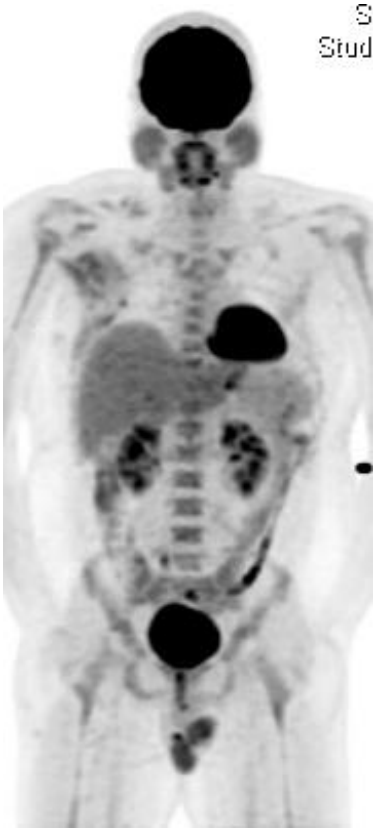
MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	65	4-7	ABVD	6	>=1.3	>=4	3.7	2.55	CR	0		1	25	1	86	1	1



Note: relapse after 25m

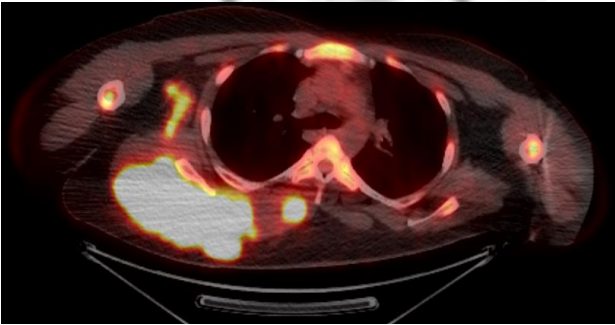
Tx-2
ICE-ASCT

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	32	4-7	ABVD	6	≥ 1.3	≥ 4	3.7	2.5	CR	0		1	22	0	54	1	1



Note: relapse after 22m

Tx-2
BV-ASCT



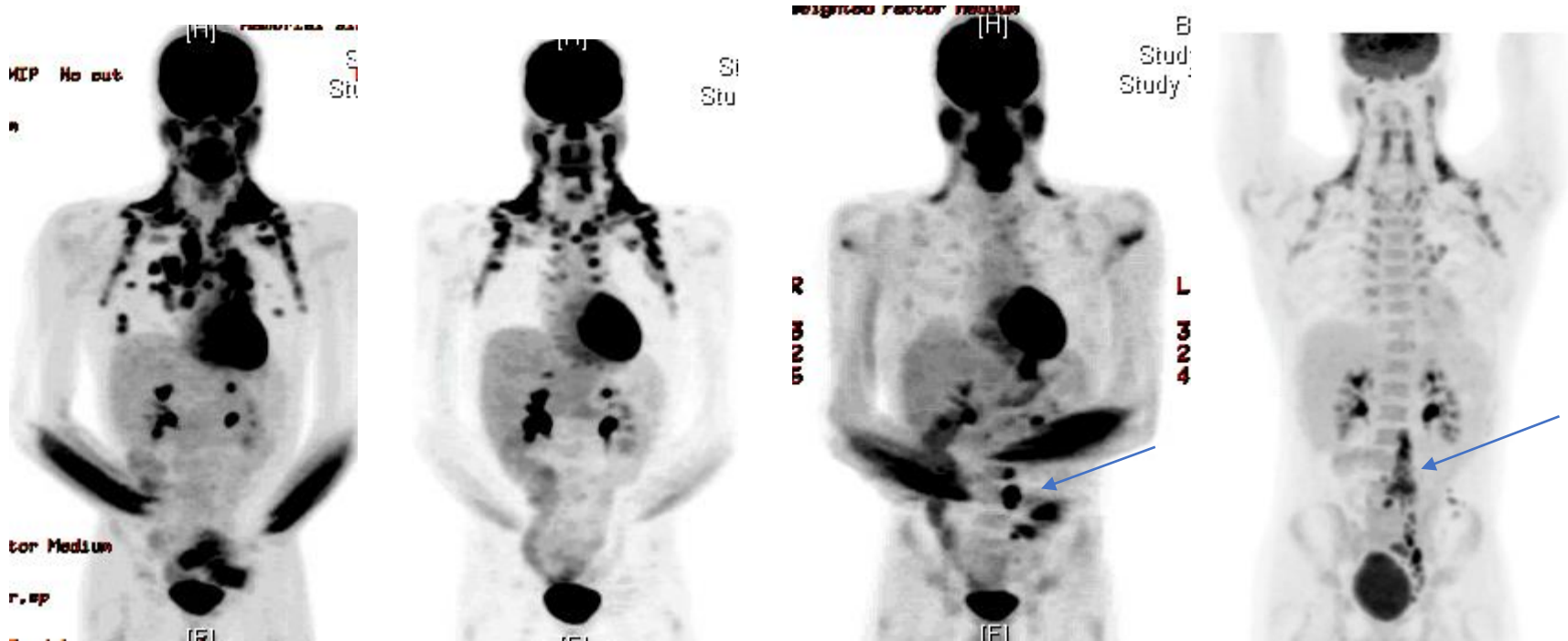
MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	22	0-1	ABVD	6	>=1.3	>=4	3.7	2	POD	0		1	6	0	76	1	1



Tx-2
BV-ICE-ASCT

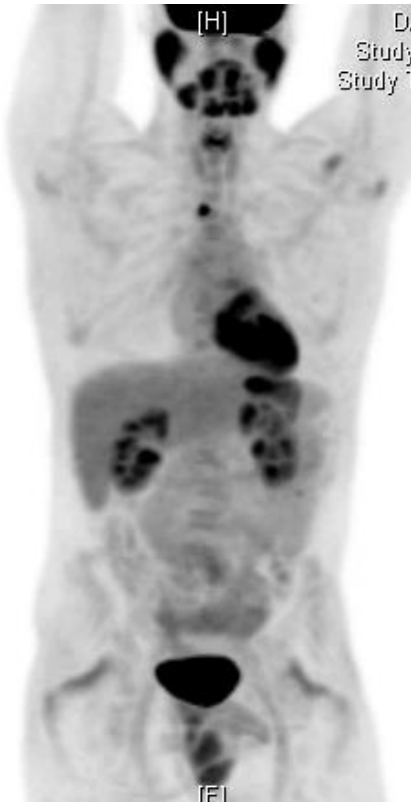
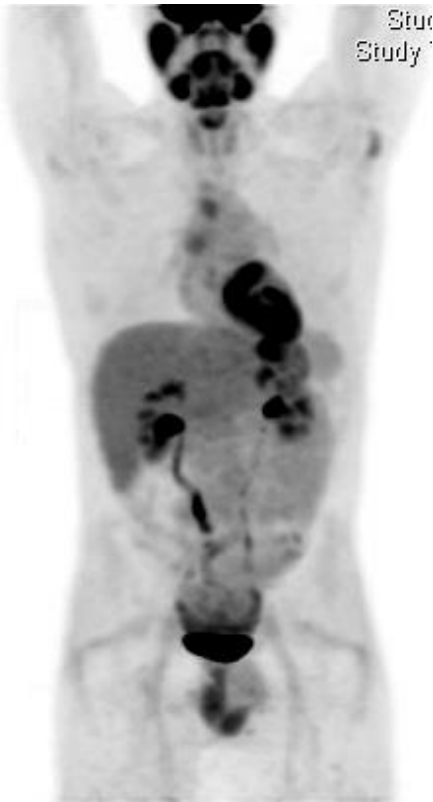
qPET2+ by LLS only (dvPET2 -)
Patients with POD

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	20	2-3	ABVD	6	≥ 1.3	DV-1-3	2.7	1.8	POD	0		1	6	0	66	1	1



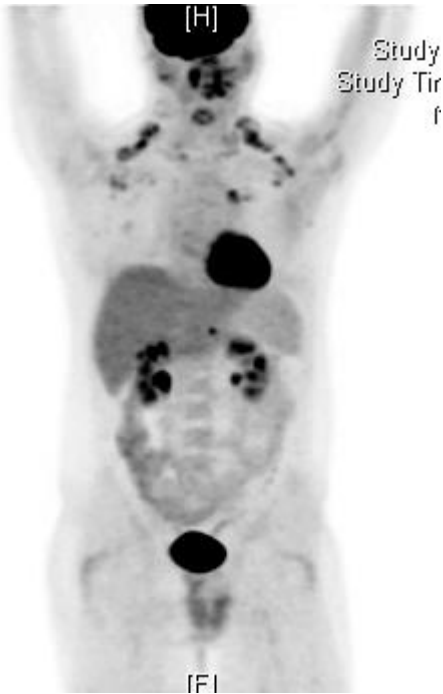
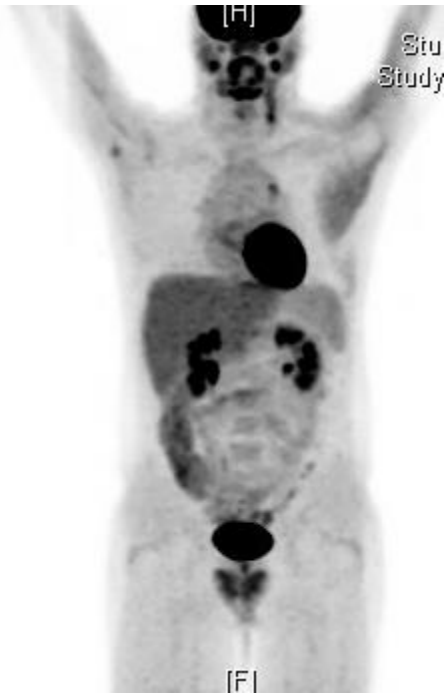
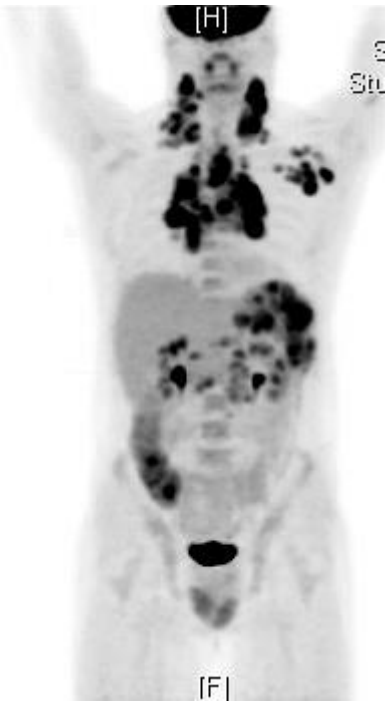
Tx-2
BV-ICE-ASCT

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	34	4-7	ABVD	6	>=1.3	DV-1-3	2.3	1.8	POD	0		1	10	0	27	1	1



Tx-2
ICE-ASCT

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	31	0-1	ABVD	6	>=1.3	DV-1-3	2.8	1.9	POD	0		1	9	0	47	1	1



Tx-2
ICE-ASCT

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	36	0-1	ABVD	6	>=1.3	DV-1-3	3.3	2.5	PR	1	CR	1	16	0	51	1	1

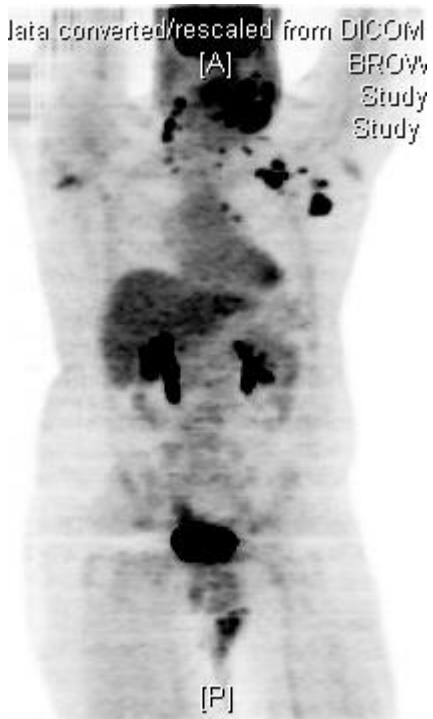


Tx 2
BV+nivolumab - ASCT



Residual mild tiny focal areas of hypermetabolism in the spleen with maximum SUV 3.3 compared to 26.

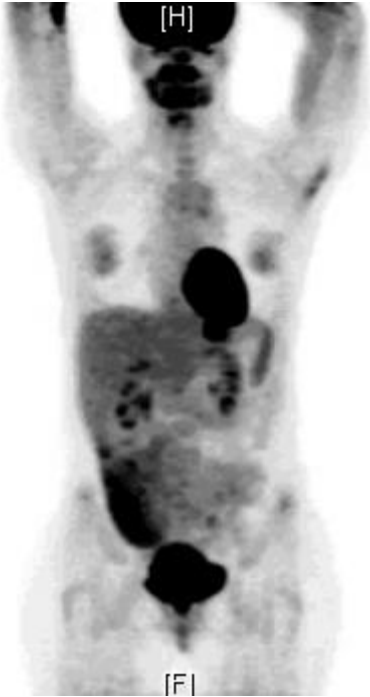
MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	32	0-1	ABVD	6	≥ 1.3	DV-1-3	3.8	2.4	PR	0		1	9	0	91	1	1



Tx 2
R-da-EPOCH

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	43	2-3	ABVD	6	>=1.3	DV-1-3	2.2	1.68	PR	0		1	49	0	88	1	1

OSH

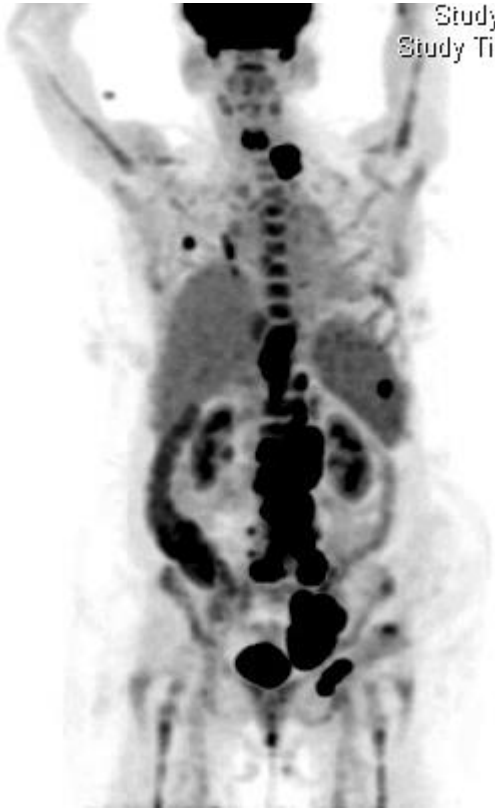


Interim only

EOT - CT

Tx-2
ICE-ASCT

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	64	4-7	ABVD	6	>=1.3	DV-1-3	3.53	2.9	TRM	0		0	6	1	6		



TRM

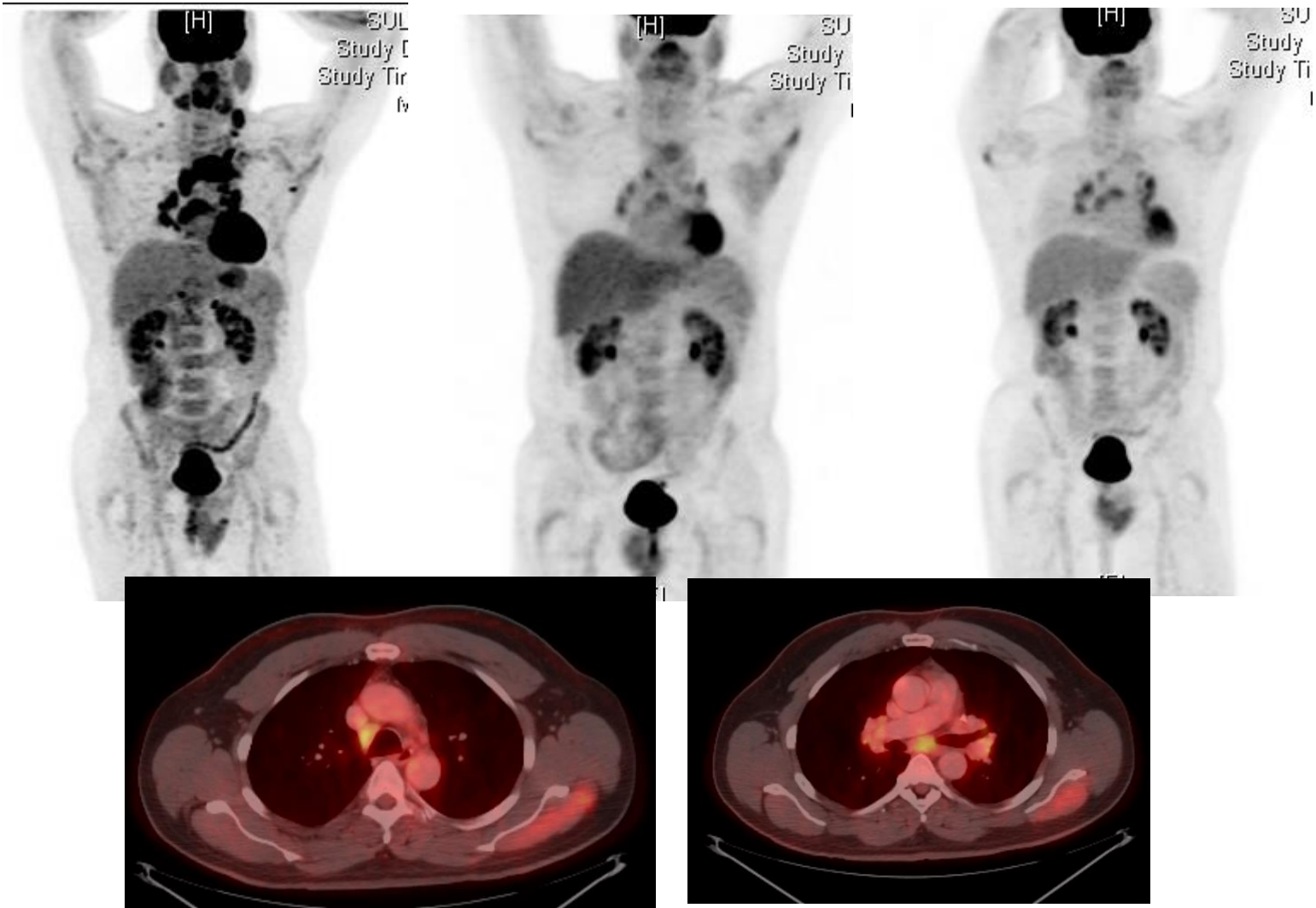
qPET2+ by LLS only (dvPET2-)
Patients with long term remission

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	87	2-3	ABVD	6	>=1.3	DV-1-3	4.5	2.56	CR	0		0	11	0	11		

OSH

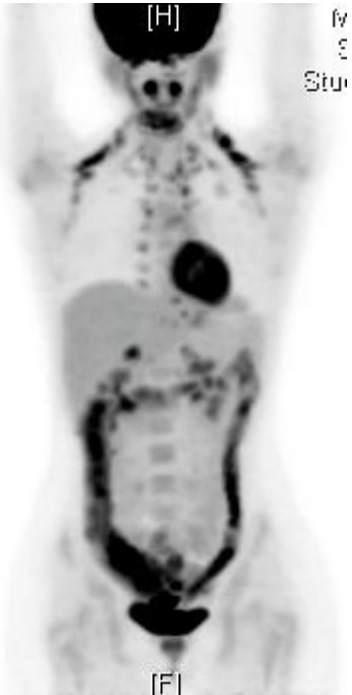


MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	56	2-3	ABVD	6	>=1.3	DV-1-3	2.9	2.2	CR	0		0	83	0	83		

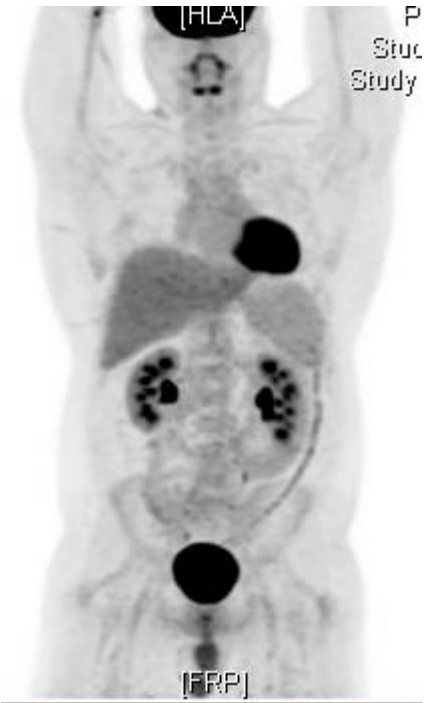


MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	19	0-1	ABVD	6	>=1.3	DV-1-3	2.8	1.6	CR	0		0	51	0	51		

OSH



MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	32	2-3	ABVD	6	>=1.3	DV-1-3	3.6	2.8	CR	0		0	32	0	32		



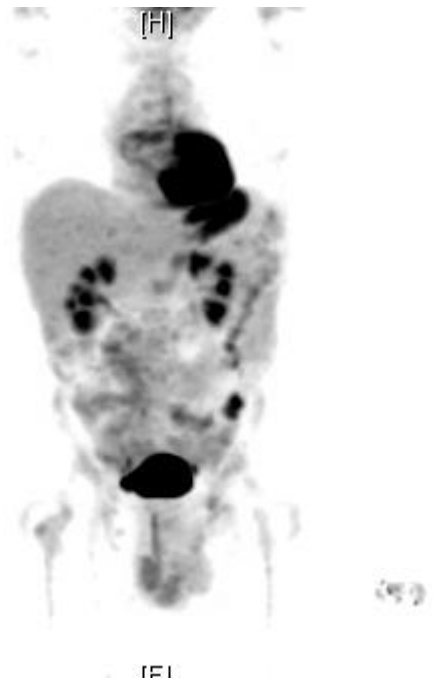
MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	35	2-3	ABVD	6	>=1.3	DV-1-3	2.5	1.9	CR	0		0	50	0	50		



EOT - CT

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	47	4-7	ABVD	6	>=1.3	DV-1-3	2.3	1.6	CR	0		0	27	0	27		

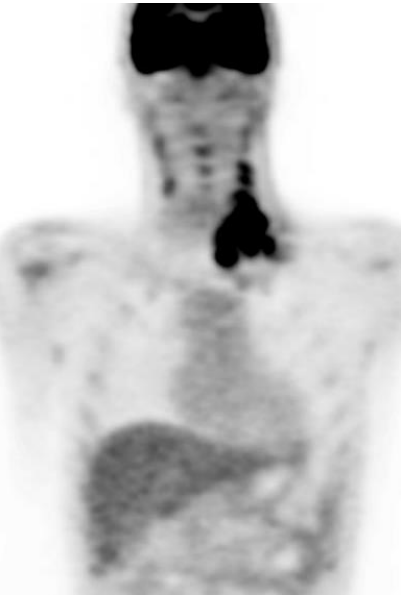
OSH



OSH

Only interim PET is available

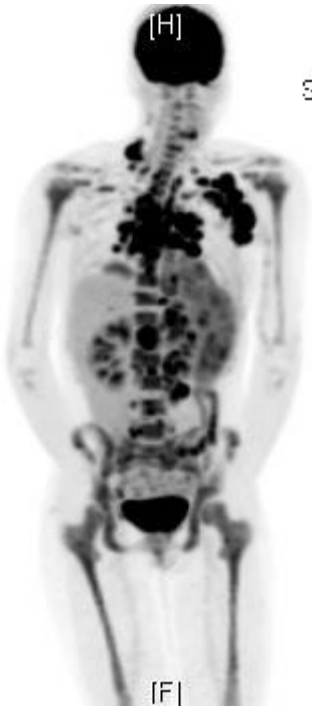
MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	19	0-1	ABVD	6	>=1.3	DV-1-3	4.4	1.8	CR	0		0	66	0	66		



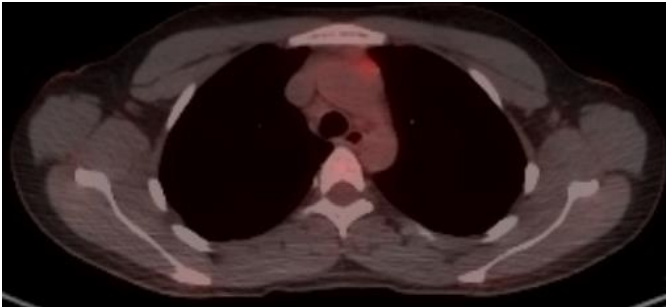
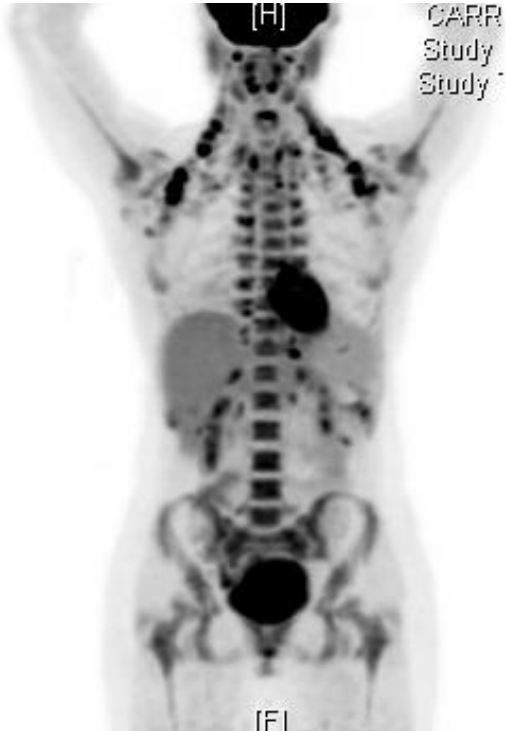
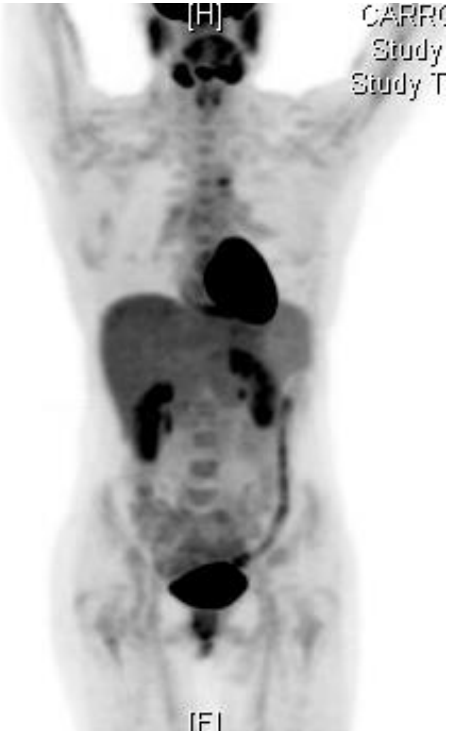
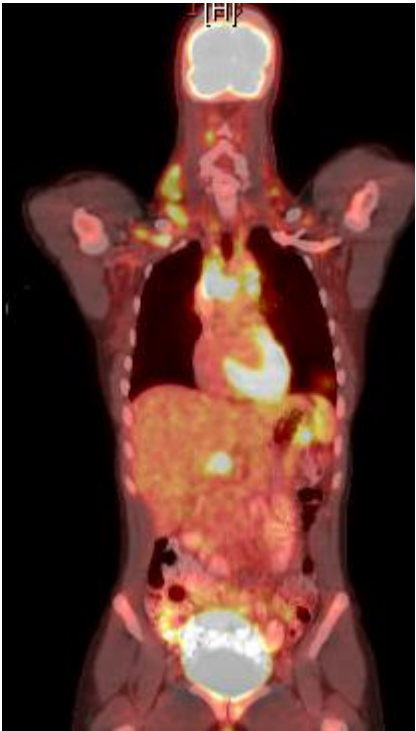
+ splenic
lesion not
visible on this
section



MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	36	4-7	ABVD	6	>=1.3	DV-1-3	1.4	2	CR	0		0	66	0	66		



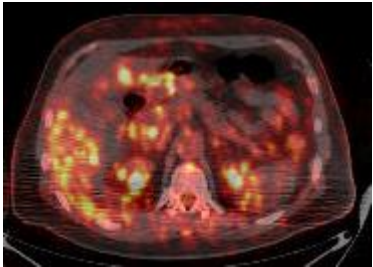
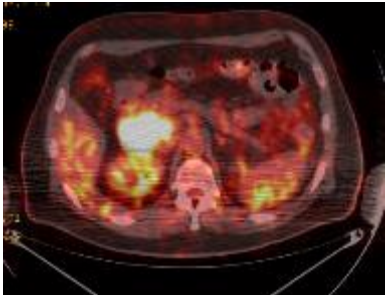
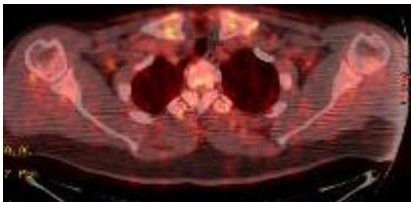
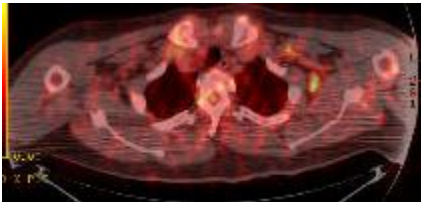
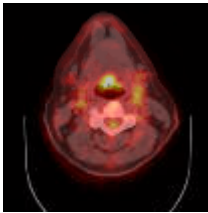
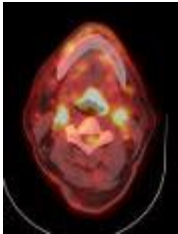
MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	27	2-3	ABVD	6	≥ 1.3	DV-1-3	2.8	2	CR	0		0	38	0	38		



MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	32	2-3	ABVD	6	≥ 1.3	DV-1-3	3.1	2.4	CR	0		0	32	0	32		



MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	59	2-3	ABVD	6	>=1.3	DV-1-3	9.6	3.1	CR	0		0	81	0	81		



portocaval mass anterior to the right kidney 5.7cm SUVmax 9.6
aortocaval lymph node, which has increased SUV of 7.1



Hypermetabolic left masseter and pterygoid muscles SUV 6.9. All sites of initial disease resolved

MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	36	4-7	ABVD	6	>=1.3	DV-1-3	3.6	2.2	CR	1		0	42	0	42		

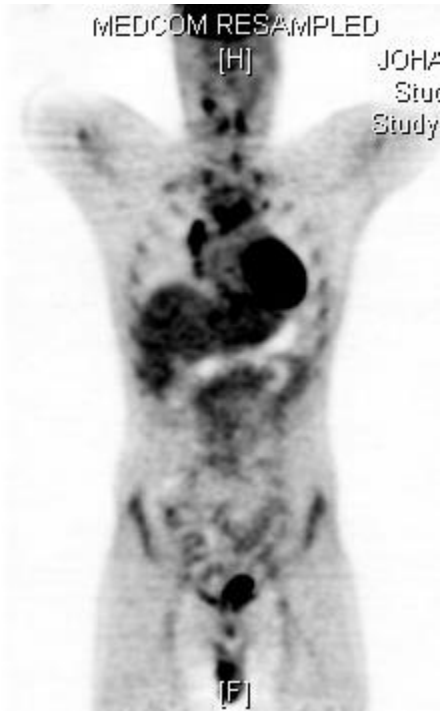
OSH



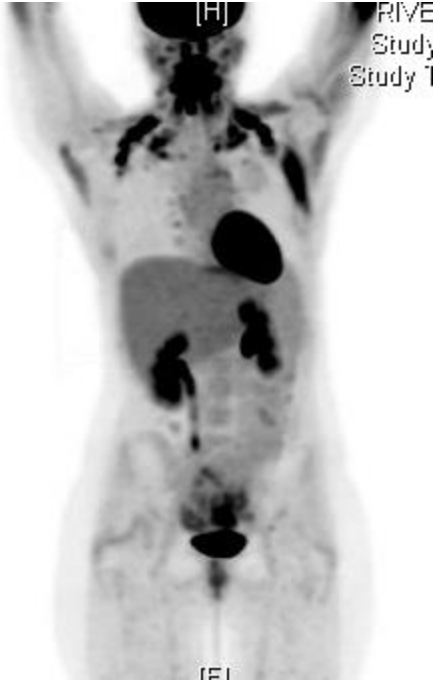
MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	44	2-3	ABVD	6	>=1.3	DV-1-3	3.9	1.9	PR	1	CR	0	23	0	23		



MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	24	0-1	ABVD	5	≥ 1.3	DV-1-3	3	1.9	CR	1	CR	0	68	0	68		



MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	19	2-3	ABVD	6	>=1.3	DV-1-3	2.8	1.9	CR	0		0	40	0	40		



MRN	Age	IPS	Escalated	Cycles	SUV Ratio	Deauville	SUV	Liver	Response	RT	RT resp	Relapse	PFS	Death	OS	Rel prev site	Bx confirmed
	59	4-7	ABVD	6	>=1.3	DV-1-3	3	2.1	CR	0		0	9	0	9		

