Ewing's sarcoma :long term favourable outcome in children with combinated modality therapy and conservative surgery

Standard treatment: chemotherapy and surgery avoiding radiotherapy

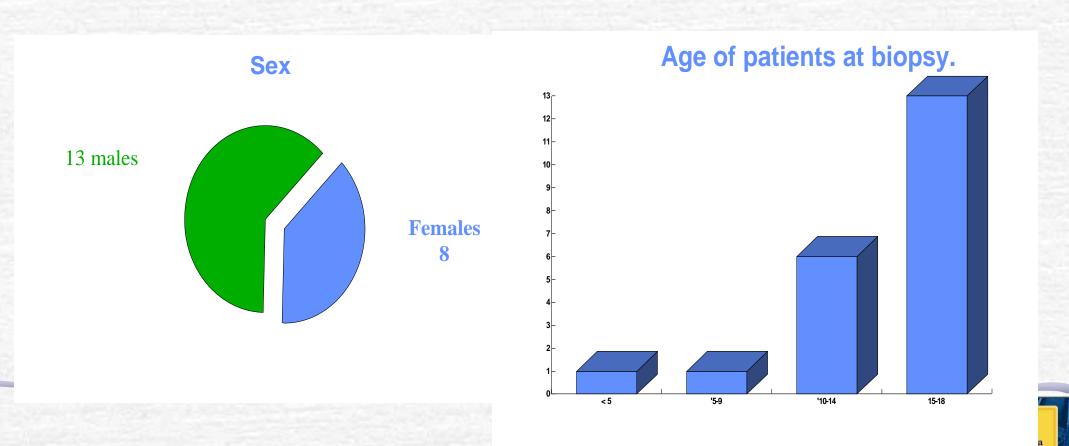
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Patients

From november 1985 to january 2000, 21 children aged de 4.5 to 18 were treated by EWDD protocol for localized Ewing's sarcoma



TUMOR LOCATIONS

Femur: 4

✓ Iliac: 5

Rib: 4

rtibia:3

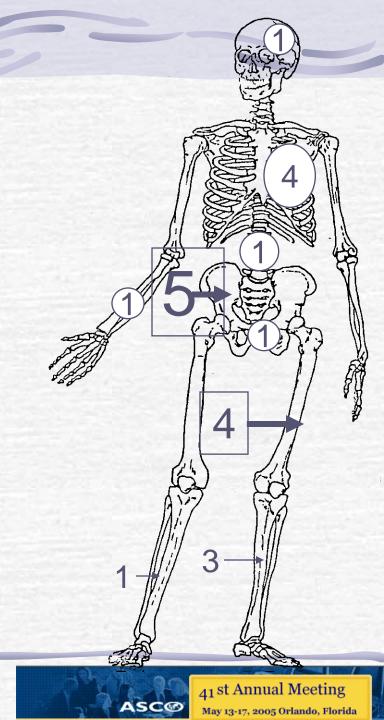
Fibula: 1

spine: 1

sacral: 1

radial: 1

mandibular: 1



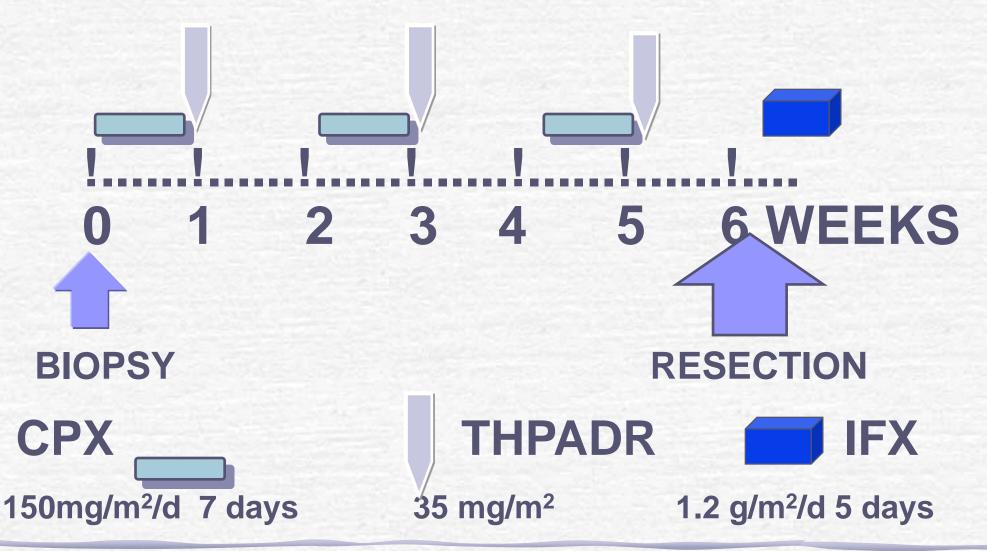
PROGNOSTIC FACTORS

18/21 PATIENTS HAD AT LEAST ONE CLASSICAL BAD PROGNOSTIC FACTOR (tumoral volume >100ml or central location).

13 INITIAL TUMOR VOLUMES > 100 ml 12 central **locations** 200-499 < 50 cc 50-99 cc 100-199 500-799 800-999 > 1000

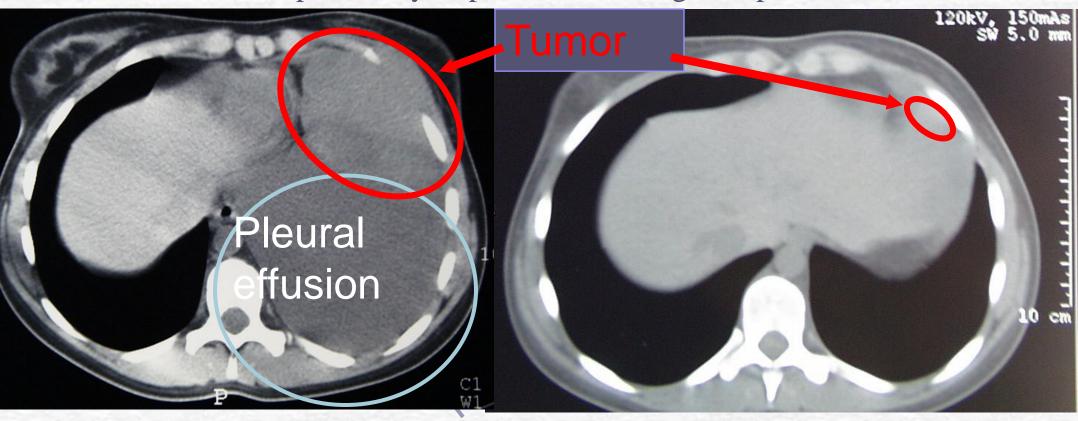
1 st Annual Meeting
May 13-17, 2005 Orlando, Florida

EW DD2 INDUCTION CHEMOTHERAPY





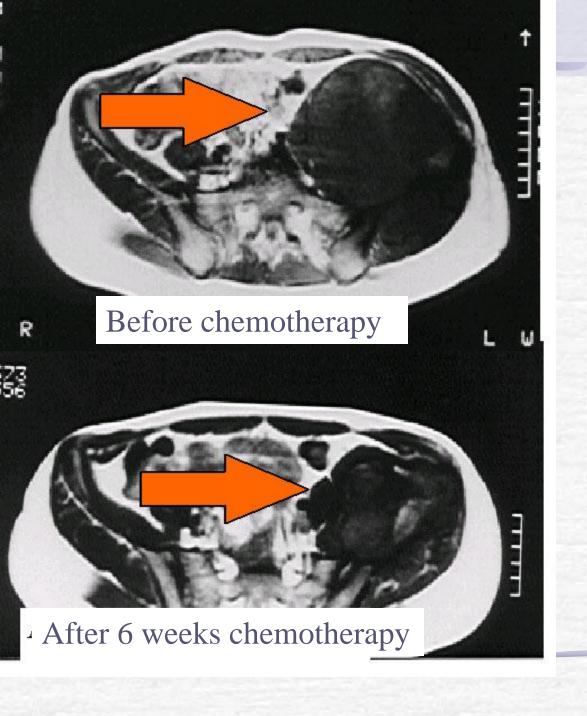
18 y old girl with a huge pleural infusion leading to discovery of a rib tumor with little pain. Very impressive radiologic response in 6 weeks.



Before chemotherapy

After 6 weeks bidrug chemotherapy

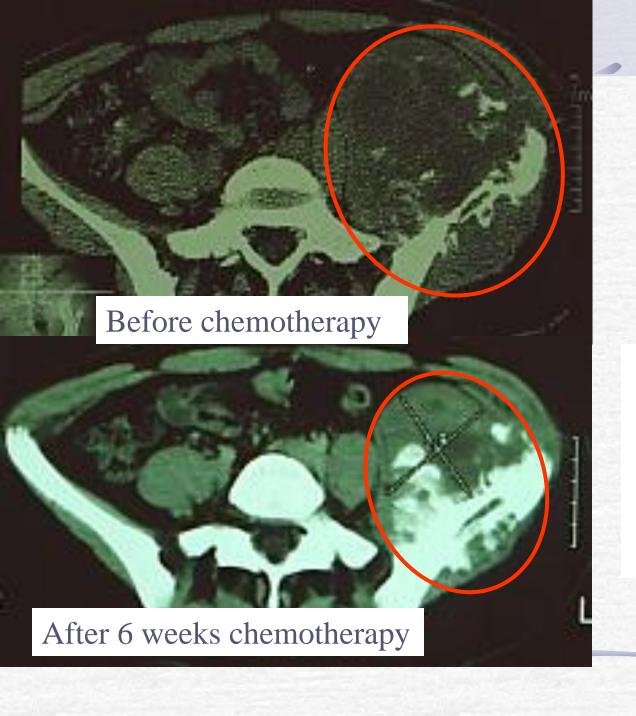




Decrease of tumoral diameter: 38 %

Decrease of tumoral volume: 76%



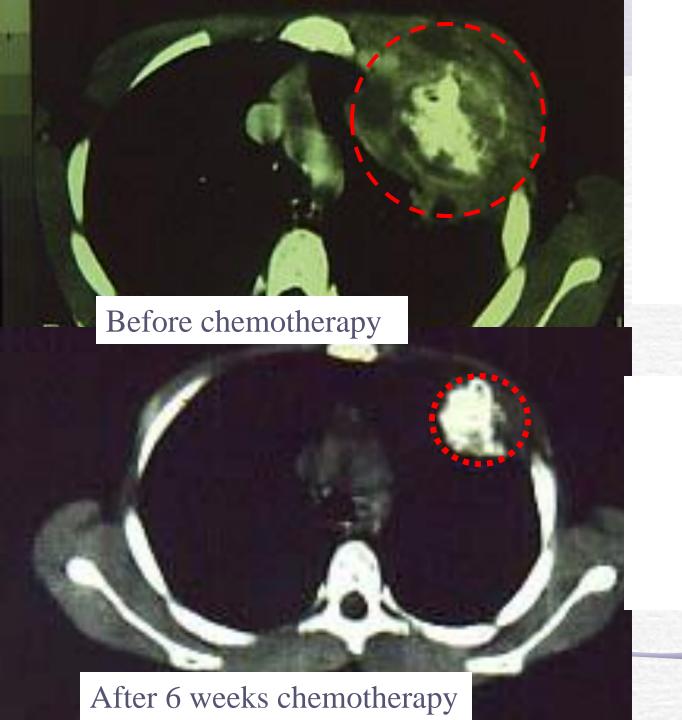


Decrease of tumoral diameter: 28 %

Decrease of tumoral

volume : 63%



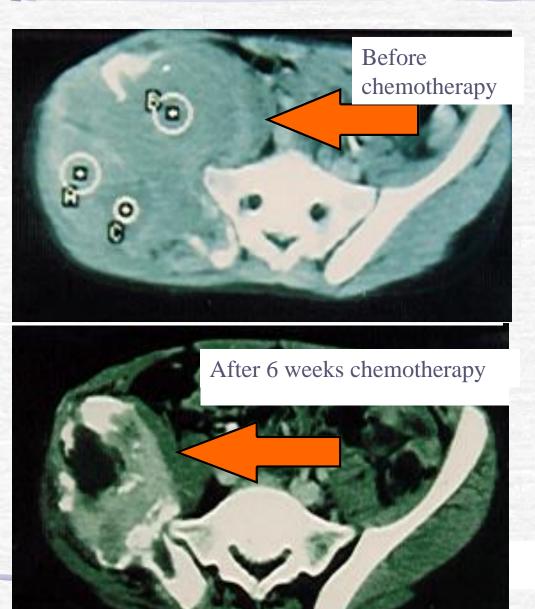


Decrease of tumoral diameter: 50 %

Decrease of tumoral

volume: 88%





Decrease of tumoral size: 66%

Decrease of tumoral volume: 95%



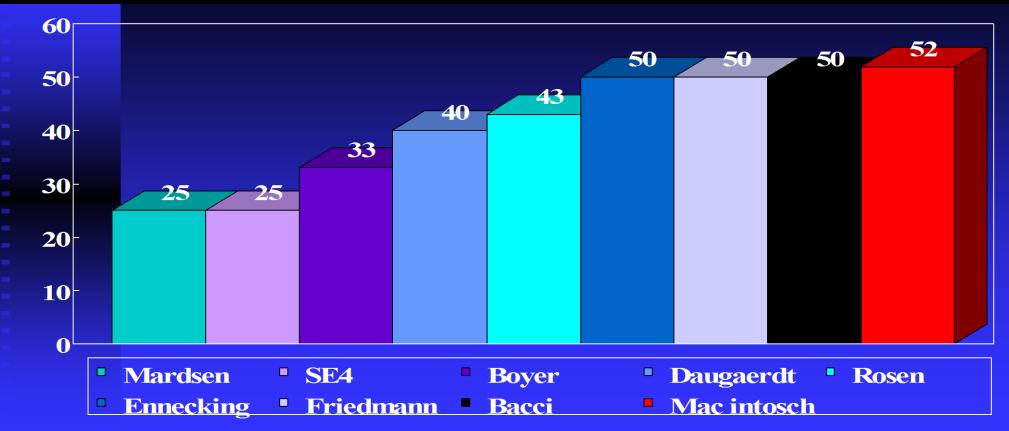
Standard treatment excluded radiotherapy

- In this protocol early extra tumoral en bloc resection without radiotherapy was the standard local treatment.
- It was performed in all cases without risk of neurological sequellaes.
- 3 patients with contaminated margins received postoperative radiotherapy

Why all our patients were operated?

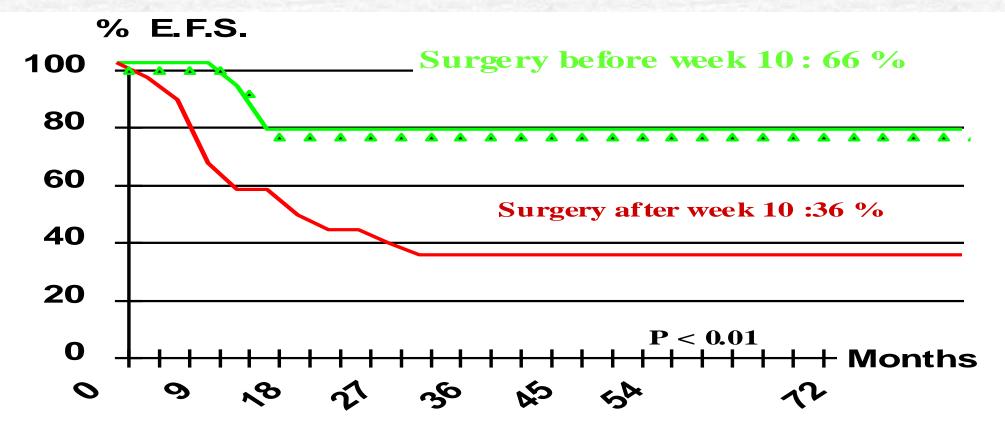
In the litterature, risk of local reccurence following radiotherapy reaches 20% to 50%





All patients were operated EARLY

In our surgical experience (1) early resection is a significant prognostic factor



(1) Analysis of a monocentric experience of surgery in Ewing's sarcoma.N. Delepine and all

Med. and Ped. Oncology. Vol. 27 n°4, 316, octobre 1996.

All limb tumors were resected



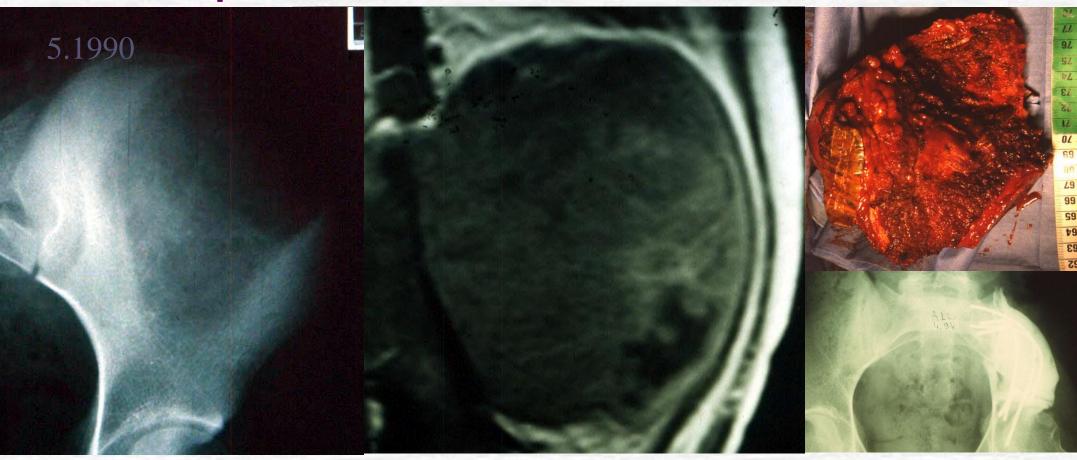




Some cases of reconstructive procedures used after resection of femoral Ewing

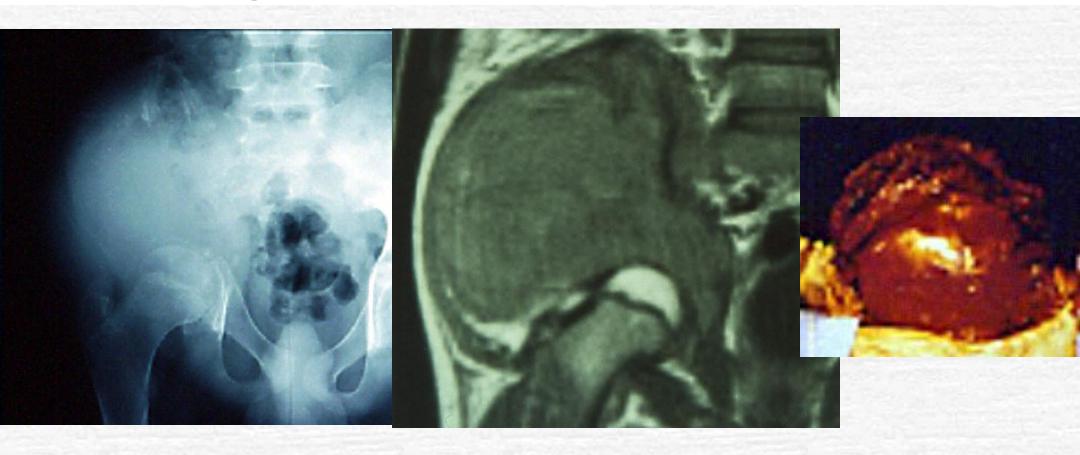


All pelvic tumors were resected



- 16 Y old girl with a huge IIB tumor of iliac wing.
- En bloc resection after 6 weeks chemotherapy.
- Skeletal reconstruction using cement. Excellent oncologic and fonctionnal results with 15 years follow up

Even huge pelvic tumors were resected

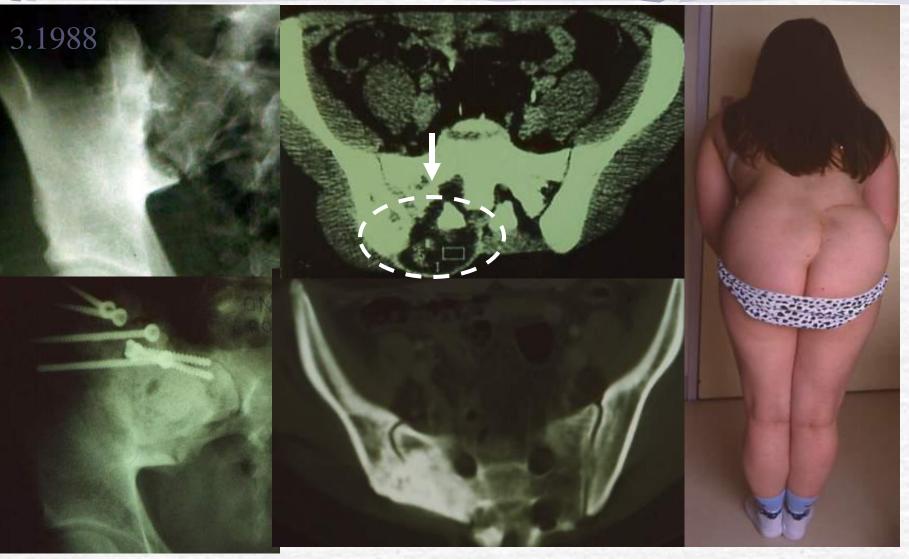


Resection of huge tumors are challenges but most important for prognosis.

Early resection of huge tumoral mass can prevent chemoresistance



Even sacroiliac tumors



17 years remission after en bloc resection and reconstruction with cement



POSTOPERATIVE CHEMOTHERAPY

Postoperative chemotherapy <u>alternated</u>:

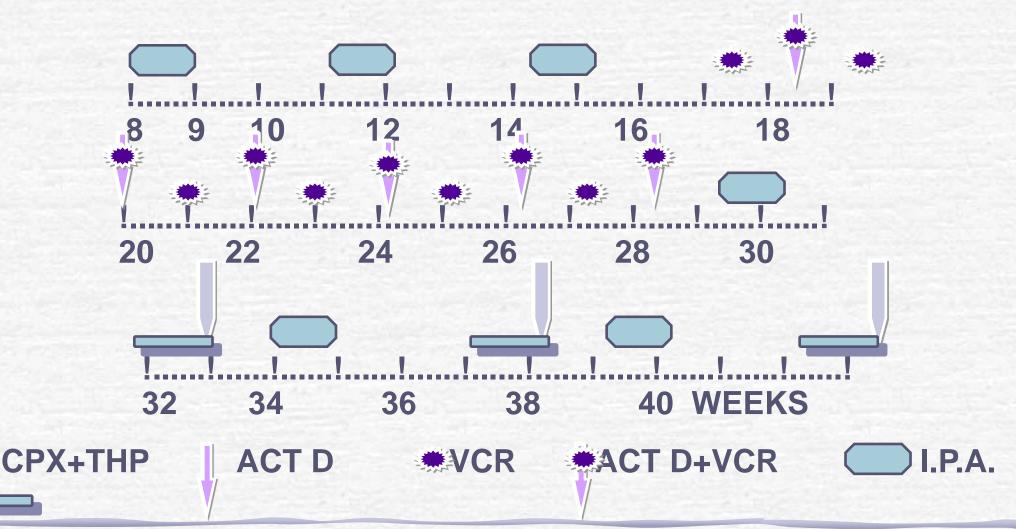
6 IPA (Ifosfamide 6 gr/sqm, Cisplatinum 125 mg/ m², Pirarubicine 35 mg/ m²)

- 3 Cyclophosphamide(150 mg/m² 7d)
- +Pirarubicine (35 mg/m² d 8)

12 weeks of Vincristine (2 mg/ m² x 12), Actinomycine D (2 mg/ m² x 6)



EW.DD2 PROTOCOL. POST OPERATIVE CHEMOTHERAPY





TREATMENT TOLERANCE

Tolerance of preoperative chemotherapy is excellent and permits best conditions for the surgery.

Toxicity of postoperative chemotherapy is heavier

- 1. 20% Vincristine induced multinevritis.
- 2. 1 Vesical grade 4 toxicity induced by Ifosfamide.
- 3. 1 Renal toxicity following a course of IPA.
- 4. Anthracyclines (pirarubicine) low toxicity: only two ECG transitory anomalies.



Late complications related to treatment

We observed two colic cancers in patients treated by this protocol.

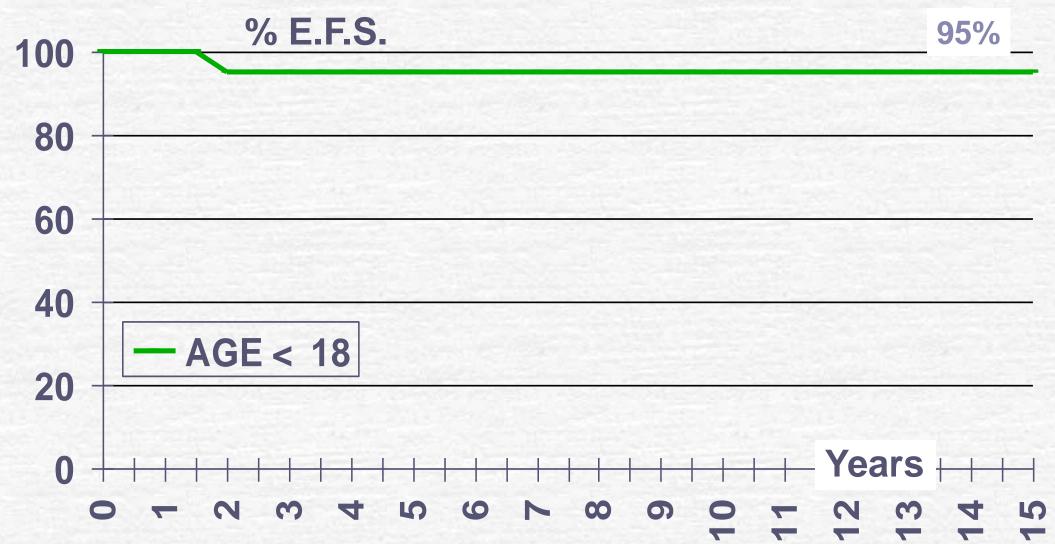
Patients were even free survivors when colic cancer occured, respectively 13 years and 15 years following treatment.

Both patients received post operative radiotherapy after contaminated resection of sacroiliac tumors.

Both patients are in CR for the second cancer



Oncologic results



EW DD. 21 children with localized Ewing . Median follow up : 10 years (min 5, max 19,5). 20/21 patients are DFS.(15 / 04 / 2005)



- This serie shows the excellent long term prognosis of children with localized Ewing's sarcoma treated by this protocol derived from Hayes, with shorter (6 weeks) bidrug induction.
- and systematic conservative surgery.
- followed by six drugs postoperative chemotherapy



This serie confirms the results of Hayes

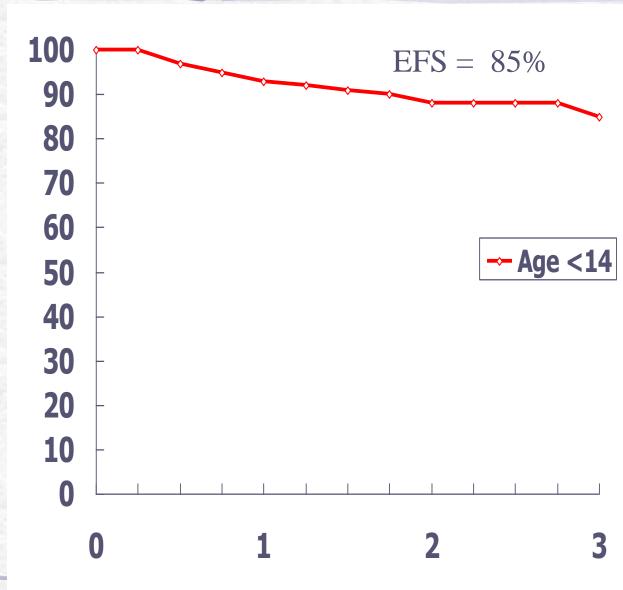
on patients treated by surgery: 10/11 operated patients were DFS

Therapy for localized ewing's sarcoma. Hayes J.Clin.oncol. 1989, 7, 2, 208-213.



This serie also confirms the good prognosis of young patients observed in the multicentric italian SE 91 study published by Rosito

Rosito P. et al Cancer,1999;86:421-428.



- It emphases the low toxicity of this chemotherapy.
- And underlines the risks of radiotherapy even at mild doses (2 second cancers out of 3 patients irradiated) and the need to avoid it as often as possible.

