

Letter to the Editor

Dear Sir,

There is no standard palliative treatment of hepatocellular carcinoma. Various modalities have been employed and these include local infarction, regional chemotherapy, systemic chemotherapy and intra-lesional alcohol. We would like to report the use of high-dose chemotherapy consisting of ifosfamide, carboplatin and etoposide (ICE), followed by peripheral blood progenitor cell (PBPC) reinfusion. This treatment has resulted in disease stabilisation.

A 57-year-old woman presented with complaints of a gradually increasing lump in the epigastric region for 6 to 7 months. A CT scan of the abdomen revealed an irregular heterogenous mass in the left lobe suggestive of hepatocellular carcinoma. She underwent a left lobectomy. The histopathology was consistent with hepatocellular carcinoma and cirrhosis of the liver. Six months later, the mass recurred in the left lobe of the liver along with a small focal hypodense area in the right lobe. Serum alpha-feto protein had increased progressively from 762 ng/mL to 3477 ng/mL over the last two months (normal limits 0 to 10 ng/mL). She was only seen by us at this stage. On examination, she had normal vital signs with a scar from a previous surgery in the abdomen and a hard mass in the epigastric region, 4 cm below the right coastal margin; the spleen was not palpable. Laboratory investigations showed a haemoglobin of 14.3g/dL, Hct 45.9% WBC was $14.7 \times 10^9/L$; 66% neutrophils and 29% lymphocytes, platelets were $170 \times 10^9/L$. Serum chemistries revealed serum creatinine of 0.5 mg/dL, LDH 843 IU/L, uric acid 5.3 mg/dL; Serum total bilirubin 0.7 mg/dL; SGPT 38 IU/L; alkaline phosphatase 112 IU/L.

Various treatment options including transarterial chemo-embolisation (TACE) were discussed with the patient and family members. The patient refused TACE and opted for high-dose chemotherapy.

The patient underwent priming chemotherapy with cyclophosphamide 1.5 grams intravenously, followed two days later by granulocyte-monocyte colony stimulating factor (GM-CSF or leucomax supplied by Sandoz Corporation) and granulocyte colony stimulating factor (GCSF or Neupogem supplied by Roche Corporation) for three days each. On the seventh day, the patient underwent a 9 stem-cell apheresis. The CD-34 positive cells were calculated at around 2 million cells per kg of the patient's body weight. This was followed by conditioning with ICE chemotherapy over the next 3 days, consisting of ifosfamide 6 gm/m^2 over 24 hours for 2 days, carboplatin 600 mg/m^2 over 2 hours on days 1 and 2 and etoposide 250 mg/m^2 over 3 hours on days 1 and 2. This was followed by re-infusion of the PBPC product 24 hours after completion of chemotherapy. She developed a fever at this time, requiring ceftazidime, amikacin, and empiric itraconazole for fungal prophylaxis. Her platelet counts fell to $72 \times 10^9/L$. The TLC decreased to a low of $0.4 \times 10^9/L$. On the seventh day after conditioning, GM/CSF and GCSF were re-instituted for a total of 10 days, beginning on the next day of infusion of apheresis product. The total duration of neutropenia was 7 days. At the time of discharge, the patient was afebrile, feeling well, and had a WBC of $3.4 \times 10^9/L$, with 64% neutrophils and 33% lymphocytes. Following her discharge, the alpha-feto protein normalised and the patient was on regular follow-up in the outpatient

clinic. She is currently well and alive, in excellent functional status and it has been 20 months since she underwent chemotherapy.

We do not believe that high-dose chemotherapy with stem cell support has been previously reported as a treatment option in hepatocellular carcinoma. We feel that in highly selected cases with preserved liver function tests and non-localised disease, as in this case, high-dose chemotherapy with PBPC can be used as a treatment modality for disease stabilisation purpose.

Yours sincerely,
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