

LETTERS TO THE EDITOR

## Triptolide and management of systemic malignancies besides pancreatic carcinomas

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### Abstract

The recent article by Zhou *et al* was highly interesting and thought provoking. The authors have clearly shown that triptolide administration is associated with up-regulation of the *Bax* gene, resulting in an attenuating effect on cell growth in gastrointestinal malignancies such as pancreatic carcinomas. The article by Zhou *et al* is all the more important because it highlights the rapidly increasing role of triptolide in the management of systemic malignancies. For instance, triptolide acts on the PI3K/Akt/NF- $\kappa$ B pathway, thereby enhancing apoptosis secondary to the administration of bortezomib in multiple myeloma cells. Similar synergisms are seen when triptolide is administered along with 5-fluorouracil for the management of colonic carcinomas. Similarly, triptolide causes down-regulation of the *Bcl-2* gene, resulting in control of cell growth in tumors, such as glioblastoma multiformes.

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**Key words:** Triptolide; *Bax* gene; *Bcl-2* gene; SDF-1/CXCR4 pathway; Acute T lymphocytic leukemias

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### TO THE EDITOR

The recent article by Zhou *et al*<sup>[1]</sup> was highly interesting and thought provoking. The authors have clearly shown

that triptolide administration is associated with up-regulation of the *Bax* gene, resulting in an attenuating effect on cell growth in gastrointestinal malignancies, such as pancreatic carcinomas<sup>[1]</sup>. The article by Zhou *et al*<sup>[1]</sup> is all the more important because it highlights the rapidly increasing role of triptolide in the management of systemic malignancies.

For instance, triptolide acts on the PI3K/Akt/NF- $\kappa$ B pathway thereby enhancing apoptosis secondary to the administration of bortezomib in multiple myeloma cells<sup>[2]</sup>. Similar synergisms are seen when triptolide is administered along with 5-fluorouracil for the management of colonic carcinomas<sup>[3]</sup>. Similarly, triptolide causes down-regulation of the *Bcl-2* gene resulting in control of cell growth in tumors, such as glioblastoma multiformes<sup>[4]</sup>. In fact, triptolide, when combined with ionizing radiation in the therapy of pancreatic carcinomas, decreases cell survival in these tumors by almost 21%<sup>[5]</sup>.

Triptolide also inhibits the SDF-1/CXCR4 pathway and thereby has an attenuating effect on lymphoid metastatic, as well as proliferative activity in non-Hodgkin lymphoma cell lines<sup>[6]</sup>. Similarly, triptolide demonstrates anti-proliferative effects in other hematological malignancies, such as acute myeloid leukemia. In fact, the anti-carcinogenic effects of triptolide in malignancies, such as acute myeloid leukemia are markedly enhanced by other agents such as AraC<sup>[7]</sup>. Recent studies also confirm that triptolide has a negative effect on proliferation in acute T lymphocytic leukemia<sup>[8]</sup>. These anti-carcinogenic functions of triptolide are in part secondary to its anti-angiogenic properties<sup>[9]</sup>.

More recently, Xu *et al*<sup>[10]</sup> have developed polymeric micelles of triptolide which appear to demonstrate anti-carcinogenic properties without affecting host immunity. These recent developments further highlights the immense therapeutic potential of triptolide and the need for further research to fully assess its anti-carcinogenic potential.

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