

phoblastic leukaemia/lymphoblastic lymphoma (precursor B-cell acute lymphoblastic leukaemia). In: Jaffe ES, Harris NL, Stein H, Vardiman JV (Eds): Pathology and genetics of tumors of hematopoietic and lymphoid tissue. IARC Press, Lyon, 2001, p 111.

3. Kishbaum JD, Preuss FA. Leukemia: a clinical and pathological study of 123 fatal cases in a series of 14,400 necropsies. *Arch Intern Med* 1943; 71: 777-792.
4. Porcaro AB, D'Amico A, Novella G et al. Primary lymphoma of the kidney. Report of a case and update of the literature. *Arch Ital Urol Androl* 2002; 74: 44-47.
5. Moral P, Dupriez B, Herbrecht R et al. Aggressive lymphomas with renal involvement: a study of 48 patients treated with the

LNH-84 and LNH-87 regimens. Group d'Etude des Lymphomes de l'Adulte. *Br J Cancer* 1994; 70: 154-159.

S. Serefhanoglu¹, B. Bitik², A. Aybal³, D. Ertoy Baydar⁴, A. Gurlek⁵

¹Department of Internal Medicine, Division of Hematology, ²Department of Internal Medicine, ³Department of Internal Medicine, Division of Nephrology, ⁴Department of Pathology, ⁵Department of Internal Medicine, Division of Endocrinology and Metabolism, Hacettepe University, Faculty of Medicine, Ankara, Turkey

Correspondence to: Songul Serefhanoglu, MD. E-mail: dr.songul1978@yahoo.com

Vitamin D intake may be effective in the management of triple-negative breast cancer

Dear Editor,

Triple-negative breast cancers are defined by lack of expression of estrogen, progesterone and HER-2 receptors. This subgroup accounts for 15% of all types of breast cancer. Since there are no specific-targeted treatment guidelines for this subgroup, triple-negative breast cancer is managed with standard treatment, leading to a high rate of local relapse and systemic dissemination [1]. Checkpoint mechanisms are essential for the maintenance of genomic integrity. In vertebrates when cells experience replication arrest or undergo DNA damage by UV irradiation, the ATR kinase [ataxia telangiectasia mutated (ATM)- and Rad3-related kinase] phosphorylates and activates the checkpoint kinase 1 (Chk1). The activated Chk1 inhibits Cdc25 phosphatases, which control inhibitory phosphorylation sites on cyclin-dependent kinases, the latter being critical regulators of cell cycle transitions [2]. Full activation of Chk1 by ATR requires Claspin, which may act as a scaffolding protein that brings together ATR and Chk1 [3]. A recent study showed that the E2F-regulated gene Chk1 is highly expressed in triple-negative breast carcinomas [4]. Furthermore, Chk1 and Claspin were quickly downregulated in mouse MC3T3-E1 and in mouse mammary carcinoma cells by treatment with 1,25-dihydroxyvitamin D₃, which is a known inhibitor of cell proliferation [5]. In the light of the above infor-

mation, we suggest that vitamin D intake may be effective in the management of triple-negative breast cancer. This proposal should be verified in large clinical trials including patients with triple-negative breast cancer.

References

1. Cleator S, Heller W, Coombes RC. Triple-negative breast cancer: therapeutic options. *Lancet Oncol* 2007; 8: 235-244.
2. Kastan MB, Bartek J. Cell-cycle checkpoints and cancer. *Nature* 2004; 432: 316-323.
3. Kumagai A, Kim SM, Dunphy WG. Claspin and the activated form of ATR-ATRIP collaborate in the activation of Chk1. *J Biol Chem* 2004; 279: 49599-49608.
4. Verlinden L, Vanden Bempt I, Eelen G et al. The E2F-regulated gene Chk1 is highly expressed in triple-negative estrogen receptor /progesterone receptor /HER-2 breast carcinomas. *Cancer Res* 2007; 67: 6574-6581.
5. Verlinden L, Eelen G, Van Hellefont R et al. 1 α ,25-Dihydroxyvitamin D(3)-induced down-regulation of the checkpoint proteins, Chk1 and Claspin, is mediated by the pocket proteins p107 and p130. *J Steroid Biochem Mol Biol* 2007; 103: 411-415.

H. Harputluoglu, O. Dizdar, F. Karaahmet, K. Altundag

Department of Medical Oncology, Hacettepe University Institute of Oncology, Ankara, Turkey

Correspondence to: Kadri Altundag, MD. E-mail: altundag66@yahoo.com