
The role of tumour markers in surgery

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This essay briefly outlines immunologic concepts as they relate to the aetiology, diagnosis and followup of tumours. The specific clinical indications for use of individual, tumour markers, particularly those currently in clinical use are discussed. The physical examination and standard diagnostic radiologic procedures have serious limitations in the early detection and localization of small tumour masses. In a neoplasm of 1 cc, a realistic limit of clinical screening, has already completed approximately 30 doublings or two thirds of its growth. It contains one billion cancer cells and viable cells are likely to have been shed into the blood stream or lymphatic system. In addition, in a human body composed of 10 to 14 trillion cells, harassed by environmental carcinogens, somatic mutations occur at conservatively estimated rate of 10⁻⁵ per gene per cell cycle. Assuming that only a minute fraction of aberrant cells will enter mitosis, the chance for the outgrowth of a potentially dangerous variant is nonetheless quite formidable. This ever-present threat is minimized by effective self-screening mechanisms.⁻¹¹³ There is a substantial body of evidence showing that some defect in immune surveillance permits the development of cancer or at least hampers their eradication. There have been sporadic attempts to influence the outcome of tumours by manipulating immunologic mechanisms. The rapid expansion of knowledge about immunology in recent years has been accompanied by increased interest in the role of immunology in the development and progress of cancer. Despite an early diagnosis and surgical removal, the patient may have many undetected microscopic metastases. In patients with intra-abdominal malignancy, it is often difficult to assess disease prognosis and response to treatment. Research has therefore been directed towards the identification of tumour-specific products in the body fluids, i.e., (Tumour markers, which are any chemical or biological factor that identifies the presence of a tumour or its recurrence). Markers are usually evaluated in terms of specificity and sensitivity. That is considered in light of whether they are elevated in the serum, body fluids, or tissues of patients with only a specific tumour or with any of a related malignancies.⁻¹¹⁴ If a marker is elevated above a normal level under conditions where the particular neoplasm is not present the result is called a false-positive value and the specificity of the assay is weakened, similarly, when the specific tumour being sought is present and a marker does not rise above a norm, a false negative result is obtained and the test is said to lack sensitivity. An ideal marker not only should signal the presence of microscopic tumour but also should define the site and morphologic type of the malignancy. The requirements for a marker vary depending on the specific application intended for the marker_ there are six major

areas of possible use of tumour markers. These are screening in the general population or high-risk groups, diagnosis or detection in symptomatic patients, staging or stratification, determination of adequacy of therapy, monitoring for recurrence and monitoring for response to radiation, chemotherapy or other treatment. Also tumour markers have been used as definitive tests through which a clinician can decide to start, continue or withhold treatment and it is influenced by serial determination of the levels of a tumour marker. There is a long list of possible useful markers and new ones are constantly appearing.¹¹⁵ There are many affirmative statements about their usefulness. In general, these tumour markers can be classified into 2 groups: hormonal products and protein products of the tumour. The hormonal markers tend to be more specific and are therefore usually more useful in the clinical setting. The protein products of tumours, many of them are identical to embryonal products and some are enzymes. Because of the unpredictable occurrence of an elevation of a single tumour marker in a particular individual, some researchers have suggested that multiple markers be evaluated in each tumour patient. So, the best biological diagnosis of the neoplastic process depends on the simultaneous measurement of a selected group of these markers.