Early diagnosis and prediction of therapeutic responses are crucial in cancer patients to tailor and optimize the treatment, increase the likelihood of cure, reduce side-effects, and avoid overtreatment. This special issue compiles relevant articles focused on the development of innovative cancer biomarkers and their validation.

New molecular biology tools, including genome-wide analysis, deep-sequencing, and RNA-seq, are currently available for the identification of novel unique biomarkers with great potential for developing more sensitive and specific diagnostic tools as well as discovering new candidate therapeutic targets for personalized medicine.

In this special issue the relevance of systemic biomarkers for early diagnosis and/or cancer characterization/monitoring has been emphasized by several articles focused on blood tumor markers with an overview by S. Holdenrieder et al. The great variety of blood tumor markers is shown by the wide range of biomarkers spanning from the basic parameter changes of blood count in renal cell carcinoma (reported by G. Prokopowicz et al.) to the detection of circulating tumor cells in gastric cancer (observed by K. Zou et al.), the level of antibodies to the hidden IgG antibodies to the tumor-associated Thomsen-Friedenreich antigen (described in gastric cancer patients by O. Kurtenkov and K. Klaamas), and the levels of plasma circulating tumor DNA, observed in melanoma patients by B. Busser et al. Moreover, the diagnostic relevance of systemic miRNAs (in particular miRNA 21) in biliary tract cancer seems even greater than current markers (i.e., CA19-9) and has been reported by C. Mayr et al.

Most articles, however, have been focused on tissue biomarkers in the early discovery and/or characterization stage, with a great potentiality of becoming relevant cancer biomarkers for molecular characterization/diagnosis, prognostic evaluation, and therapeutic responsivity. The articles on the predictive positive clinical significance of novel long noncoding RNA in HCC (by L. Zhang et al.) whose expression levels are directly related to overall survival, the low proliferation level of the human lung cancer cell lines A549 in presence of high expression of circular RNA-ITCH (by L. Wan et al.), and limited disease in SCLC of patients with miRNA-related polymorphisms in P13K/Akt/mTOR pathway genes (by W. Jiang et al.) are establishing a new paradigm.

Moreover, high EF24 levels have been associated with suppression of invasion and migration of HCC cells (by R. Zhao et al.), while high expression levels of Cadherin 17 have been associated with high frequency of bone marrow metastasis in a murine breast cancer model (by T. Okada et al.) and high expression of meiotic recombination II homolog A (MRE11) oncprotein with clinical breast cancer progression (by C.-H. Yang et al.). The overall characterization of the primary tumor has been described for renal cell carcinoma
by S. H. Kim et al., glioblastoma by W. Szopa et al., and
nasopharyngeal carcinoma by A. Lu et al. The latter study
includes the primary tumor immune-evaluation showing that
in patients with elevated neutrophil to lymphocyte ratio
(NLR) the balance is tipped in favor of tumor-promoting
inflammation resulting in tumor cell proliferation and cancer
metastasis.

Finally specific gene signatures have been proposed to
optimize personalized cancer treatment; in particular the
overall higher responsiveness to PARPi of patients carrying
BRCA2 mutations has been reviewed by S. Murata et al. and
specifically for pancreatic cancer by J. Martinez-Useros and J.
Garcia-Foncillas.

We hope that our current paper collection will enrich
our readers and researchers, giving them an overview of the
current broad range of cancer biomarkers being pursued by
research teams around the world.

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