

Materials and Methods: The present study investigates BIBW 2992 in models derived from HNSCC.

Results: In vitro, BIBW 2992 inhibited the proliferation of transfected Ba/F3 cells expressing the wild-type receptor with EC50 = 0.8 nM. Importantly, the compound showed similar potency on Ba/F3 cells expressing the EGFRvIII mutant receptor (EC50 = 0.5 nM). BIBW 2992 inhibited the proliferation of the HNSCC cell line FaDu with an EC50 of 7 nM. Cell cycle analysis by propidium iodide staining of treated cells showed a reduction of S-phase cells and a concomitant increase of G0/G1 cells at concentrations that match the EC50 values observed in the proliferation assays. In vitro combination experiments BIBW 2992 shows at least additive activity when added to standard chemotherapeutics used in HNSCC patients (e.g. 5FU or taxanes). In vivo, potent, dose-dependent and long-lasting growth suppression and even tumor regressions were observed when mice carrying subcutaneous FaDu xenografts were treated daily p.o. with 20 mg/kg BIBW 2992. Short term treatment (3 days) of mice with BIBW 2992 before a single 20 Gy dose did not result in significant sensitization to radiotherapy. However, long term treatment with BIBW 2992 after a 20 Gy dose of radiation resulted in a tumor volume doubling time of 104 days thus slowing tumor growth by more than 3-fold. Combination of BIBW 2992 at suboptimal doses with the triple angiokinase inhibitor, BIBF 1120, resulted in improved efficacy in the FaDu model.

Conclusion: BIBW 2992 shows efficacy in human HNSCC models in vitro and in vivo. Clinical studies in this tumour type seem warranted.

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POSTER

Multicomponent coatings improve the biocompatibility of load-bearing implants

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Aim and Innovation: At this work was to estimate the influence of new multicomponent nanostructured coatings on the implant's osseointegration process. Titanium implants commonly used in orthopedics and dentistry integrate into host bone by a complex and coordinated process. The results of their application are completely satisfactory in many instances; however, are encountered the cases of the complications, which can be treated as the consequences of the insufficient biocompatibility of pure titanium. The signs of inflammation, thinning the skin, threat of the formation of sore and even fistulae can be seen. The danger of the similar complications development makes it necessary the search for titanium implants coatings, which would improve their biocompatibility and osseointegration. Osseointegration is a direct connection between living bone and the titanium implant at the level of the light microscope. Comparing with the previous studies new implants are studied in vivo under the conditions, when the replaced defect is located on the bone, which accomplishes motions with the large amplitude and with the large load.

Methods and Materials: of this investigation 48 rats femur model (250–300 g) were used. 3 types of implants were placed: one type had pure titanium core with $\text{TiC0.5} + 10\%(\text{Ca}_{10}(\text{PO}_4)_6(\text{OH})_2)$ composition coated on the surface. The average grain size is 10–40 nm. Another type had pure titanium core with $\text{TiC0.5} + \text{CaO}$ composition coated on the surface. The average grain size is 10–40 nm as well. The control was a pure titanium implant. An osteotomy was performed, and a 3 mm length of femur was removed. The implant was placed into the animal's tissue. Four screws fasten the implant to the femur's fragments fixing them. The animals were allowed full weight bearing without any mobility restrictions immediately postoperatively. Standard plain radiographs of the dissected bones were taken in lateral projections to ensure implant's stability. The rats were sacrificed and tissues investigated 5, 10, 15 and 30 days postoperatively. The degree of osseointegration correlates with the presence of osteocalcine, a differentiation marker of mature bone cells. The more rapidly increases osteocalcine concentration on the boundary between the bone and the implant, the more biocompatible implant appears.

Results: the effectiveness of implants considered in comparison with a control group of implants without nanostructured modification was proved by experimental models not only for stable, but also for moving with the large amplitude load-bearing implants.

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POSTER

Heparanase expression in the differentiation of follicular thyroid lesions: from laboratory to clinical practice

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The papillary, follicular, medullary and anaplastic variants of thyroid carcinoma can be promptly diagnosed by cytological criteria in material obtained by fine-needle aspiration (FNA) ultrasonography-guided. However, the distinction between follicular carcinoma and benign follicular adenoma needs histological demonstration of vascular or capsule invasion; therefore, they are cytologically grouped as undetermined tumors or suspect follicular neoplasm ("follicular pattern"). The aim of this study was to evaluate the immunohistochemical expression of heparanase, an endo-beta-glucuronidase, implicated in the process of tumor invasion in histological fields of thyroid follicular adenomas and carcinomas in an attempt to make a differential diagnosis of these neoplasms. Forty-nine thyroid follicular adenomas and 11 thyroid follicular carcinomas were evaluated, using the monoclonal antibody anti-heparanase by immunohistochemical reactions through the LSAB-peroxidase technique. The analysis was made by a quantitative digital computer-assisted method (Imagelab[®]). The immunostaining analysis obtained showed a distinct pattern between follicular adenomas and carcinomas: while carcinomas showed positive immunostaining on neoplastic cells and negative immunostaining on colloid, adenomas showed an inverse pattern. This test presents sensibility of 91%, specificity of 86% and negative predict value of 98%. In conclusion, the association of positive heparanase on neoplastic cells and negative heparanase on colloid is a good immunohistochemical test in the exclusion diagnosis of thyroid follicular carcinoma when compared to adenomas, with high sensibility, specificity and negative predict value.

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POSTER

The use of a radiophotoluminescent glass rod detector for the determination of cyberknife stereotactic radiosurgery system output factors

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Background: The Cyberknife (CK) radiosurgery system can deliver single or several fractions of radiation doses to a well-defined small intracranial or extracranial target with a high precision. A radiophotoluminescent glass rod detector (GRD) system has recently become commercially available. The purpose of this study is to evaluate the possibility of the GRD as a new detector for dose measurement in small fields and high dose gradient regions. We introduce a novel method for measurement of 5 mm output factor for the CK using GRD. Although the concept of using GRD to determine output factors is not new, they have not gained measured output factor in water phantom. The GRD holder is specially designed for this study to put into the water phantom for the irradiations.

Materials and Methods: In this study, the model GD-301 glass rod dosimeter (Asahi Techno Glass Corporation, Japan) and FGD-1000 automatic reader are used. The size of the model GD-301 is 1.5 mm in diameter and 8.5 mm in length. The relative output factor of CK collimators (5, 7.5, 10, 12.5, 15, 20, 25, 30, 35, 40, 50 and 60 mm) measurements with the GRD were compared with those with a PTW 60008 diode detector, PTW 31006 pinpoint type ionization chamber and a Gafchromic film (Type MD-55). The output measured with GRD, pinpoint chamber and diode was performed at a depth of 1.5 cm in water phantom. The GRD was irradiated in a water phantom using a holder stand, which was specially designed for this study. The holder is composed of the PMMA tube with a hole for GRD at 1.5 cm from its top. The water level was set precisely to the top of the holder and the axis of the beam aligned with holder axis in a way that the radiation beams pointed down vertically.

Results: The measured relative output factors with four dosimeters shown very similar results except for three smallest collimators (5, 7.5 and 10 mm). The mean value of the output factor for GRD in the 5 mm collimator is 0.705. Each dose point of GRD is presented by an average of 5GRD readings and their one standard deviation of each dose point is within $\pm 1.0\%$. The pinpoint chamber output is approximately 11% lower than the corresponding GRD values at the 5 mm collimator. Because the pinpoint chamber had a larger effective volume, this most likely contributed to these differences. The GRD is 5.2% and 4.1% lower than diode in the 7.5 mm and 5 mm collimators, respectively. It is not obvious whether the difference