Fibrin glue system for adjuvant brachytherapy of brain tumors with

188Re and 186Re-labeled microspheres.

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Abstract:

Brain tumors such as glioblastoma reappear in their original location in almost 50% of cases. To

prevent this recurrence, we developed a radiopharmaceutical system that consists of a gel applied

immediately after surgical resection of a brain tumor to deliver local radiation booster doses. The

gel, which strongly adheres to tissue in the treatment area, consists of fibrin glue containing the

beta-emitters rhenium-188 and rhenium-186 in microsphere-bound form. Such microspheres can be

prepared by short (2 h or less) neutron activation even in low neutron flux reactors, yielding a

mixture of the two beta-emitters rhenium-188 (E(max)=2.1 MeV, half life=17 h) and rhenium-186

(E(max)=1.1 MeV, half life=90.6h). The dosimetry of this rhenium-188/rhenium-186 fibrin glue

system was determined using gafchromic film measurements. The treatment efficacy of the

radioactive fibrin glue was measured in a 9L-glioblastoma rat model. All animals receiving the

non-radioactive fibrin glue died within 17+/-3 days, whereas 60% of the treated animals survived 36

days, the final length of the experiment. Control animals that were treated with the same amount of

radioactive fibrin glue, but had not received a previous tumor cell injection, showed no toxic effects

over one year. The beta-radiation emitting rhenium-188/rhenium-186-based gel thus provides an

effective method of delivering high doses of local radiation to tumor tissue, particularly to wet areas

where high adhesive strength and long-term radiation (with or without drug) delivery are needed.