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The Ship Simulator
2008/2009 is an easy-
to-learn, attractive,
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objectives of this
proposal are to
develop a tumor-
targeted cytotoxic
agent utilizing a folate
receptor (FR) to tumor-
specific antibody
conjugate. We
propose that this

approach will be
useful for the
treatment of ovarian
and breast cancer. It is
thought that the FR-
expressing cancer cells
can be detected by the
anti-FR monoclonal
antibody (mAb) in the
periphery of the tumor
and therefore, using
the tumor-specific

antibody, cytotoxic agents can be targeted to the cancer cells. In the proposed Phase I SBIR study, we have demonstrated that anti-FR mAb-toxin conjugates can be selectively targeted to FR-positive tumor cells in vitro. In addition, anti-FR mAb

can selectively eliminate FR-positive tumor cells in mice bearing FR-positive xenografts. We propose to further optimize this approach by the preparation of anti-FR mAb-folate-DMBA conjugate. In addition, the sensitivity of FR-

positive tumor cells to anti-FR-folate-DMBA conjugate will be compared to that of FR-negative tumor cells. The *in vivo* antitumor activity of the conjugate will be evaluated in the A129-DMBA model, which closely mimics human ovarian and

breast cancer. Finally,
the immunogenicity of
the proposed
conjugate will be
evaluated in the
A129-DMBA model.
In the Phase II SBIR
study, the optimal anti-
FR mAb-folate-
DMBA conjugate that
is shown in the Phase I
study will be selected

for a clinical trial in
ovarian and breast
cancer patients.

Additionally, the in
vivo therapeutic
efficacy of this
conjugate will be
compared to that of
the free DMBA in
ovarian and breast
cancer patients.

[unreadable] [unreada

ble]Ribonucleic Acid (RNA) is an unstable molecule that tends to degrade over time. It can be stored at -80° C. or longer but remains stable for only short periods. This property has created

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