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Neutralization assays of patient antisera indicated that the GagPol recombinant protein of FIV and the env/env proteins of RERV (from K246 and Z25) as well as the recombinant env of FIV and MA/CA proteins of EIAV (from A-33) were capable of being recognized by the AIDS patient. Recently, we developed a modified virus-like particle (mVLP) based vaccine platform which is capable of inducing potent protective immunity to HIV-1. To evaluate this vaccine candidate in a primate model we were interested in developing a cross-reactive RERV vaccine. Here, we compare the RERV protective efficacy of the mVLP based FIV vaccine [A33gag-pol-env, or feline immunodeficiency virus (FIV)] to the efficacy of our vaccine candidate mVLP which was designed to induce a partial gag epitope in a pre-existing epitope vaccine. In this study, FIV- and RERV-specific cellular immune responses were evaluated following a single immunization of BALB/c mice with either the FIV-mVLP or RERV mVLP vaccine candidates. Specific humoral and cellular immune responses were detected following immunization of mice with either the mVLP vaccine candidate. However, specific binding antibodies to FIV gag, which cross-react with the gag of RERV- can be detected after a single immunization with the RERV mVLP vaccine but not with the FIV mVLP. The results also demonstrated that intra-vaginal challenge of female mice immunized with the RERV mVLP resulted in an early drop in viral load, while the challenge of mice immunized with the FIV mVLP resulted in no detectable viral load. In addition, significant differences in the vaginal secretions of RERV from challenge infected animals were observed in RERV. These results demonstrated a protective efficacy against FIV infection in the BALB/c mouse model with a vaccine that induces both T- and B-cell epitopes against the RERV env/env proteins. Further, this study is one of the first to demonstrate the importance of the mVLP based RERV vaccine for development of a cross-reactive RERV vaccine that not only maintains a high level of efficacy against FIV, but also provides protection against RERV infection. J Microbiol Immunol Infect. 2013 Mar;43 6d1f23a050

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