Abstract

Title of thesis:

"Preparation of immunogen from prevalent strains of diarrheagenic Escherichia coli and

evaluation of its protective efficacy in animal model to reduce diarrheagenic Escherichia

coli mediated clinical health burden"

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In developing countries, diarrhea-induced morbidity and death in children can be attributed

primarily to diarrhoeagenic Escherichia coli (DEC) pathotypes. There are currently no licensed

vaccines available that offers broad-spectrum protection against infections caused by DEC.

Several immunogenic proteins are found in outer membrane vesicles (OMVs), which are

microvesicles secreted by gram-negative bacteria during the growth phase. We have developed

a pentavalent outer-membrane vesicles (POMVs) based immunogen that targets the five

primary pathotypes of DEC that cause diarrheal diseases, taking into account the prevalence of

infections. Adult BALB/c mice were administered with 10µg of the POMVs intraperitoneally

after OMVs were isolated from five DEC pathotypes and combined in equal amounts to

develop POMVs immunogen. In adult mice, three doses of POMVs significantly increased the

cellular immune response in addition to inducing a strong humoral immune response against

whole cell lysates (WCLs), outer membrane proteins (OMPs), and lipopolysaccharides (LPS)

isolated from DEC pathotypes. Neonatal mice were effectively protected against DEC

infections by the passive transfer of POMVs immunized sera of adult mice. Overall, this study

reveals that POMVs are immunogenic, providing neonatal mice with broad-spectrum passive

protection against five major DEC pathotypes. Taken together, our results indicate that POMVs

could serve as a potent vaccine candidate to reduce the health burden caused by DEC.

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