SYNTHESIS AND ASSEMBLY OF HUMAN PAPILLOMAVIRUS TYPE 16 L1 VIRUS-LIKE PARTICLES

Bazan, S.B.^{1,4}, Aires, K.A.^{2,4}, Cianciarullo, A.M.³, Ho, P.L.^{1,2,4}

- Departamento de Bioquímica, Instituto de Química, Universidade de São Paulo, São Paulo, Brazil;
 Instituto de Ciências Biomédicas, Universidade de São Paulo, São Paulo, Brazil;
 Departamento de Genética, Instituto Butantan, São Paulo, Brazil;
 Centro de Biotecnologia, Instituto Butantan, São Paulo, Brazil.
- Human Papillomaviruses (HPVs) are important human pathogens that infect epithelial surfaces. The high-risk types, especially HPV-16, are the primary etiologic agents of cervical cancer, which is the second cause of cancer death in women worldwide. Thus HPV-associated malignancies might be prevented by vaccine inducing virus-specific immune responses. The papillomavirus major capsid protein L1 self assembles into virus-like particles (VLPs) that elicit high titers of neutralizing antibodies. The aim of this work is to produce and to purify the recombinant HPV-16 L1 major capsid protein in yeasts, focusing on the development of a prophylactic vaccine free of charge for public health system in Brazil. Codon-optimized HPV-16 L1 gene was cloned into expression vectors under the regulation of a methanol-inducible promoter and the constructions were used to transform competent Pichia pastoris cells. After PCR screening of transformants, veasts were grown, induced for 48h and lysed. Purification of L1 protein from yeast extracts was performed using a column of Heparin-Sepharose[®]. The eluted samples were subjected to a procedure of disassembly and reassembly of VLPs. Electron microscopy confirmed the presence of intact recombinant particles. Supported by: Fapesp, CNPq, Fundação Butantan.