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## 幽门螺杆菌尿素酶B亚单位优势Th表位的系统筛选到:

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Title: Systemic identification of immunodominant Th epitopes from urease B subunit of *Helicobacter pylori*

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关键词: 幽门螺杆菌; 尿素酶B亚单位; Th表位

Keywords: *Helicobacter pylori*; urease B subunit; T helper lymphocyte epitopes

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摘要: 目的 筛选幽门螺杆菌 (*Helicobacter pylori*, Hp) 尿素酶B亚单位 (UreB) 中CD4<sup>+</sup> 细胞优势应答表位, 并确定其MHC限制性。 方法 以重组UreB蛋白 (rUreB) 与弗氏佐剂联合皮下多点注射免疫BALB/c小鼠, 从小鼠脾脏中体外扩增UreB特异性T淋巴细胞, 采用步移重叠合成肽法筛选CD4<sup>+</sup> T细胞优势应答表位, 分析其MHC分子限制性。 结果 18mer短肽筛选发现UreB<sub>403-420</sub>和UreB<sub>409-426</sub>可显著刺激UreB特异性CD4<sup>+</sup> T细胞产生IFN-γ。13mer短肽鉴定实验结果显示, UreB<sub>409-421</sub>可刺激产生与18mer短肽UreB<sub>403-420</sub>和UreB<sub>409-426</sub>等同的应答强度。同时抗体阻断实验表明, 抗H-2<sup>d</sup>(I-A)单克隆抗体能有效阻断该表位的应答。 结论 UreB<sub>403-420</sub>和UreB<sub>409-426</sub>为UreB抗原中的优势Th表位, 其刺激免疫反应的核心位置为UreB<sub>409-421</sub>, 且存在H-2<sup>d</sup>(I-A)限制性。

Abstract: Objective To identify systematically MHC-restricted immunodominant T helper lymphocyte (Th) epitopes from urease subunit B of *Helicobacter pylori* (*H. pylori*) and to confirm the MHC-restricting profile. Methods BALB/c mice were immunized with recombinant UreB protein (rUreB) and Freund's adjuvant

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by subcutaneous injection. Antigen-specific CD4<sup>+</sup> T cells were amplified *in vitro* from spleens of immunized mice. The MHC-restricted immunodominant Th epitopes were identified by synthesizing overlapping peptides and antibody-blocking assay. Results UreB<sub>403-420</sub> and UreB<sub>409-426</sub> induced the strongest secretion of IFN-γ in all the 93 18mer peptides, while 13mer peptide UreB<sub>403-426</sub> within UreB<sub>403-420</sub> and UreB<sub>409-426</sub> induced equivalent responses compared with the 18mer peptides. The results of antibody-blocking assay showed that the H-2<sup>d</sup> (I-A) monoclonal antibody efficiently blocked the T cell activation induced by peptide UreB<sub>403-426</sub>. Conclusion UreB<sub>403-420</sub> and UreB<sub>409-426</sub> are immunodominant Th epitopes of urease subunit B antigen. The core reactive sequence is UreB<sub>409-421</sub> and its restriction molecule is H-2<sup>d</sup> (I-A).

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