

内在性分泌型RAGEタンパクの病原微生物に対する 防御作用

メタデータ	言語: jpn 出版者: 公開日: 2021-02-22 キーワード (Ja): キーワード (En): 作成者: Karasawa, Tadahiro メールアドレス: 所属:
URL	https://doi.org/10.24517/00060392

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Research Project

Project/Area Number	16659116	All
Research Category	Grant-in-Aid for Exploratory Research	
Allocation Type	Single-year Grants	
Research Field	Bacteriology (including Mycology)	
Research Institution	Kanazawa University	
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Project Period (FY)	2004 – 2006	
Project Status	Completed (Fiscal Year 2005)	
Budget Amount *help	¥3,500,000 (Direct Cost: ¥3,500,000) Fiscal Year 2005: ¥1,700,000 (Direct Cost: ¥1,700,000) Fiscal Year 2004: ¥1,800,000 (Direct Cost: ¥1,800,000)	
Keywords	内在性分泌型RAGE / Listeria monocytogenes / ノックアウトマウス	
Research Abstract	<p>本研究の目的は、「内在性分泌型RAGE(esRAGE)タンパクは、自然免疫・粘膜防御を担う重要な液性因子である」という仮説を検証することである。</p> <p>1.esRAGEと病原微生物、特にListeria monocytogenesとの相互作用の解析</p> <p>(1)表面プラスモン共鳴法を用いた実験において、esRAGEとL.monocytogenesが直接結合することが明らかになり、そのリガンドは鞭毛タンパクflagellinであった。その結合解離定数は約10⁻⁸Mであった。</p> <p>(2)esRAGEタンパクが鞭毛タンパクflagellinと結合することで、L.monocytogenesの運動を抑制した。増殖阻害活性、殺菌作用、オブソニン効果は認めなかつた。</p> <p>(3)flagellinはesRAGBのみならず膜型RAGEにも結合することがわかり、その結合によってRAGE細胞内シグナルであるNFkBが活性化が生じた。</p> <p>2.RAGE/esRAGEを共に欠損するRAGEノックアウトマウスを用いた感染実験</p> <p>(1)RAGE+/+マウス由来腹腔マクロファージの方が、RAGE-/マウス由来腹腔マクロファージよりL.monocytogenesの処理能力が高かった。</p> <p>(2)RAGE/esRAGE系はマウスのリストアリア感染に対して防御的に働いていた。</p> <p>a.2種類の遺伝背景のマウスで感染実験を行った。B6, CD-1ともにRAGE+/+マウスの方がRAGE-/マウスに比べて有意に生存率が高かった。</p> <p>b.定量的に臓器単位面積当たりのL.monocytogenesの病巣部数を数えたところ、RAGE-/マウスの方が有意に多かった。</p> <p>c.組織学的炎症の程度もRAGE-/マウスの方が有意に重症であった。</p> <p>以上の結果から、esRAGEのみならず膜型RAGEを含めたRAGE/esRAGE系は、病原微生物に対する防御作用を担う生体因子であると考えられた。</p>	

Report (2 results)

2005 Annual Research Report

2004 Annual Research Report

Research Products (42 results)

All 2006 2005 2004

All Journal Article Book Patent(Industrial Property Rights)

- [Journal Article] Detection of cfxA and cfxA2, the β-lactamase genes of Prevotella spp., in clinical samples from dentoalveolar infection by real-time PCR. 2006 ▾
- [Journal Article] A severe diabetic nephropathy model with early development of nodule-like lesions induced by megsin overexpression in the RAGE/ iNOS transgenic mice. 2006 ▾
- [Journal Article] Proteinase activity of Prevotella species associated with oral purulent infection. 2006 ▾
- [Journal Article] Occurrence of the fimbria gene hifA in clinical isolates of nonencapsulated Haemophilus influenzae. 2006 ▾
- [Journal Article] Development of an ELISA for esRAGE and Its Application to Type 1 Diabetic Patients. 2006 ▾
- [Journal Article] Identification of mouse ortholog of endogenous secretory receptor for advanced glycation endproducts : structure, function and expression. 2006 ▾
- [Journal Article] Isoflavones inhibit nicotine C-oxidation catalyzed by human CYP2A6. 2006 ▾

[Journal Article] 糖尿病性腎症とRAGE	2006 ▼
[Journal Article] グリケーションと糖尿病腎症	2006 ▼
[Journal Article] 合併症におけるRAGE	2006 ▼
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[Journal Article] Roles of the receptor for advanced glycation endproducts in diabetes-induced vascular injury.	2005 ▼
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URL: <https://kaken.nii.ac.jp/grant/KAKENHI-PROJECT-16659116/>

Published: 2004-03-31 Modified: 2016-04-21