Thu., 26 Apr., 4pm

Jukhyun Bio Auditorium(RM.121)

School of Life Sciences Seminar Series

2 0 1 8 Spring Semester

T Cell-mediated Immunopathogenesis during Viral Infection

Korean

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Speaker | Eui-Cheol Shin, Ph.D.



Affiliation | KAIST



Host | Prof. Sung-Gyoo Park



School of Life Sciences Seminar Series

Thursday, 26 April 2018, 4:00 PM

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No. 2018-12

Abstract

During hepatitis virus infection, liver injury is not directly caused by the hepatitis virus but is instead caused by immune-mediated mechanisms. Hepatitis A virus (HAV) is a hepatotropic RNA virus that preferentially infects hepatocytes and causes liver inflammation. HAV infection often causes severe liver injury in adults, whereas it results in asymptomatic subclinical infection in children. In the present study, we investigated the activation status and functions of CD8+T cells, and the mechanisms by which CD8+T cells mediate host injury in acute hepatitis A (AHA), which is caused by HAV infection. While it was known that HAV-specific CD8+T cells are responsible for liver injury in AHA, a recent study showed that HAV-specific CD8+T cells did not display effector function in HAV-infected chimpanzees. In this context, we examined HAV-specific and non-HAV-specific CD8+T cells in AHA patients, and studied a mechanism of immune-mediated liver injury in AHA. We demonstrate that (i) non-HAV-specific CD8+T cells are activated and proliferate during AHA, (ii) HAV-infected cells produce IL-15 which induced antigen-independent activation of CD8+T cells, (iii) CD8+T cells isolated from AHA patients exert innate-like cytolytic activity, (iv) the innate-like cytolytic activity is triggered by ligation of NKG2D and/or NKp30 without TCR engagement, (v) hepatocytes of HAV-infected liver overexpress NKG2D ligands, and therefore, could be cytolytic targets of bystander activated non-HAV-specific CD8+T cells, and (vi) liver injury during AHA is mainly associated with innate-like cytotoxic function of bystander activated CD8+T cells. Taken together, we report a human viral disease in which host injury is associated with innate-like cytotoxic function of antigen-independently activated CD8+T cells lacking specificity for the infecting virus.

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Speaker Eui-Cheol Shin, Ph.D.

Education/Experience

1990-1996 M.D., Medical School, Yonsei University 1996-2001 Ph.D., Medical School, Yonsei University

2002-2007 Postdoctoral fellow, NIDDK, NIH, USA

2007-2013 Assistant Professor, Medical Science and Engineering, KAIST

Associate Professor, Medical Science and Engineering, KAIST 2013-present

2017-present Chair Professor, KAIST

Research Interests

Our laboratory studies T cell immune responses in viral diseases and immunologic diseases. In order to understand T cell immune responses comprehensively, we study not only anti-viral immune responses but also immunopathologic mechanisms in host damage. In addition, we try to develop T cell vaccines and apply them in viral diseases.

