

Anti-apoptotic mechanism of silkworm hemolymph in staurosporine-induced apoptosis

이원중, 박태현*

Seoul National University, School of Chemical Engineering,

Cell & Microbial Engineering Lab.

Tel: (02) 880-8020, FAX (02) 875-9348

There exist some common pathways in apoptosis despite a wide range of inducing signals, and mitochondria play a crucial role especially by releasing cytochrome c into cytosol, which forms complex with Apaf-1 to turn on the caspase cascade reaction¹⁾. Silkworm hemolymph (SH) has shown anti-apoptotic activities in mammalian²⁾ and insect cell apoptosis³⁾, and five 30kDa proteins in SH are the major inhibitors of apoptosis. Here we developed anti-apoptotic mechanism of SH in staurosporine-induced HeLa cell apoptosis. SH did not directly inhibit caspase-3 and caspase-9 activities in cell-free reaction, but rather increased caspase activities by improving caspase reaction condition. This supports the claim that anti-apoptotic effect of SH lies in further upstream events than caspase activation. Cytochrome c release and the translocation of Bax to mitochondria after staurosporine treatment were blocked by SH. This indicates that SH affects a step above Bax translocation such as Bax conformational change by Bid in staurosporine-induced HeLa cell apoptosis. SH effects on cytosolic calcium concentration, generation of reactive oxygen species, and mitochondrial membrane potential were also determined.

References

1. Kirsten Lauber, Helga A. E. Appel, Stephan F. Schlosser, Michael Gregor, Klaus Schulze-Osthoff, and Sebastian Wesselborg (2001), The Adapter Protein Apoptotic Protease-activating Factor-1 Is Proteolytically Processed during Apoptosis, *JBC*, **276**, 29772-29781.
2. Shin Sik Choi, Won Jong Rhee, and Tai Hyun Park (2002), Inhibition of Human Cell Apoptosis by Silkworm Hemolymph, *Biotechnol. Prog.* **18**, 874-878.
3. Won Jong Rhee, Eun Jeong Kim, and Tai Hyun Park (2002), Silkworm hemolymph as a potent inhibitor of apoptosis in Sf9 cells, *BBRC*, **295**, 779-783.