

1, Abstract

2 Shrimp exhibit a diverse response to viral infection, that is manifested in drastic up- and
3 down-regulations of a variety of genes. In our previous work, we identified syntenin of the
4 shrimp *Penaeus monodon* (Pm) as a dynamic responder to White Spot Syndrome Virus
5 (WSSV) infection, its message being greatly upregulated in the acute phase of the infection. In
6 order to further explore the link between Pm-syntenin and viral infection, we performed a yeast
7 two-hybrid screening of a *P.monodon* cDNA library, using Pm-syntenin as bait. One of the
8 molecules that specifically interacted with Pm-syntenin was the receptor-binding domain of
9 alpha-2-macroglobulin (α_2M). A GST pull-down assay showed that GST- α_2M , but not GST
10 alone, was capable of co-precipitating syntenin. Another GST pull-down assay showed that
11 GST-syntenin, but not GST alone, was capable of co-precipitating α_2M . In addition, mutant
12 analyses showed that the N-terminal 131 amino acids of syntenin were both necessary and
13 sufficient to bind the C-terminus receptor-binding domain of α_2M . Furthermore, WSSV-
14 infected Pm showed a significant upregulation of the α_2M message, suggesting that both
15 syntenin and its protein partner α_2M are upregulated in the acute phase of a WSSV infection.
16 Taken together with a previous report showing the co-localization of α_2M and syntenin in the
17 exosome of a dendritic cell line, it is likely that syntenin, through its interaction with α_2M ,
18 plays an important role in the immune defense mechanisms of viral infections of shrimps.

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20 Key words: Syntenin; Alpha-2-macroglobulin; Yeast two-hybrid; Immune; White Spot
21 Syndrome Virus