

Table S1. Initial cleavage sites in human IAPP determined from LP/MS experiments using thermolysin (TH) and human neutrophil elastase (HNE) at a substrate to enzyme ratio of 100:1 in 10 mM sodium phosphate buffer (pH 5.5) at 25 °C.

Cleavage Site	Enzyme	Fragment	Observed Mass, Da	Calculated Mass, Da	Δ (Observed Mass - Calculated Mass), Da
Arg11-Leu12	TH	1-11	1194.7	1194.4	0.3
		12-37	2726.0	2727.0	-1.0
Asn14-Phe15	TH	1-14	1429.8	1492.7	0.1
		15-37	2428.4	2428.7	-0.3
Phe15-Leu16	TH	1-15	1639.9	1639.9	0
		16-37	2280.5	2281.5	-1.0
Asn22-Phe23	TH	1-22	2392.4	2391.7	0.7
		23-37	1529.9	1529.7	0.2
Val17-His18	HNE	1-17	1852.1	1852.2	-0.1
		18-37	2068.9	2068.9	0