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Pharmacokinetics and biodistribution of a small radioiodine labeled nerve growth factor fragment

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Nerve growth factor (NGF) exerts various actions on neuronal and non-neuronal tissues and has potential therapeutic utility, but difficulties in using the whole protein have stimulated interest in small NGF fragments. We radioiodinated a small cyclic peptide derived from NGF using the Bolton-Hunter method [¹²⁵I-C(92-96)], and confirmed binding to high affinity NGF receptors by cross-linkage analysis. Pharmacokinetic characteristics in intravenously injected mice were T¹/₂ α 5.2 min, T¹/₂ β 121.3 min, clearance 11.8 ± 0.5 ml/min, and volume of distribution 69.7 ± 4.6 ml. Dose-proportionate increases in areas-under-curve and peak-concentrations indicated linear pharmacokinetics. Biodistribution data revealed that clinically relevant doses allowed C(92-96) accumulation sufficient to elicit biological responses in receptor expressing organs including the lungs, liver, spleen, and pancreas.

Key words: nerve growth factor, peptide, radioiodine, pharmacokinetics, biodistribution