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BIOSÝNTHESIS:

- * The oxytocin peptide is synthesized as an inactive precursor protein from the *OXT* gene.
- This precursor protein also includes the oxytocin carrier protein neurophysin I.
- * The inactive precursor protein is progressively hydrolyzed into smaller fragments (one of which is neurophysin I) via a series of enzymes. The last hydrolysis that releases the active oxytocin nonapeptide is catalyzed by peptidylglycine alpha-amidating monooxygenase (PAM).
- The activity of the PAM enzyme system is dependent upon vitamin C (ascorbate), which is a necessary vitamin cofactor.
- By chance, sodium ascorbate by itself was found to stimulate the production of oxytocin from ovarian tissue over a range of concentrations in a dose-dependent manner.
- Many of the same tissues (*e.g.* ovaries, testes, eyes, adrenals, placenta, thymus, pancreas) where PAM (and oxytocin by default) is found are also known to store higher concentrations of vitamin C.

METABOLISM:

- Oxytocin is known to be metabolized by the oxytocinase, leucyl/cystinyl aminopeptidase.
- × Other oxytocinases are also known to exist.
- * Amastatin, bestatin (ubenimex), leupeptin, and puromycinhave been found to inhibit the enzymatic degradation of oxytocin, though they also inhibit the degradation of various other peptides, such as vasopressin, met-enkephalin, and dynorphin A.

Vertebrate Oxytocin Family			
Cys-Tyr-Ile-Gln-Asn-Cys-Pro-Leu-Gly-NH	Oxytocin (OXT)	Most mammals, <u>ratfish</u>	
Cys-Tyr-Ile-Gln-Asn-Cys-Pro-Pro-Gly-NH	Prol- <u>Oxytocin</u>	Some <u>New</u> <u>World</u> <u>monkeys</u> , northern tree <u>shrews</u>	
Cys-Tyr-Ile-Gln-Asn-Cys-Pro-Ile-Gly-NH	Mesotocin	Most marsupials, all birds, reptiles, amphibians, lungfishes, coelacanths	
Cys-Tyr-Ile-GIn-Ser-Cys-Pro-Ile-Gly-NH2	Seritocin	Frogs	
Cys-Tyr-Ile-Ser-Asn-Cys-Pro-Ile-Gly-NH	Isotocin	Bony fishes	
Cys-Tyr-Ile-Ser-Asn-Cys-Pro-Gln-Gly-NH2	Glumitocin	skates	
Cys-Tyr-Ile-Asn/Gln-Asn-Cys-Pro-Leu/val-Gly-NH2	Various tocins	Sharks	

Vertebrate Vasopressin Family			
Cys-Tyr-Phe-Gln-Asn-Cys-Pro-Arg-Gly-NH	Argipressin (AVP, ADH)	Most <u>mammals</u>	
Cys-Tyr-Phe-Gln-Asn-Cys-Pro-lys-Gly-NH ₂	Lypressin (LVP)	Pigs, hippos, warthogs, some marsupials	
Cys-Phe-Phe-Gln-Asn-Cys-Pro-Arg-Gly-N৮	Phenypressin	Some marsupials	
Cys-Tyr-Ile-Gln-Asn-Cys-Pro-Arg-Gly-NH	Vasotocin†	Non-mammals	
Invertebrate VP/OT Superfamily			
Cys-Leu-Ile-Thr-Asn-Cys-Pro-Arg-Gly-N닍	Inotocin	Locust	
Cys-Leu-Ile-Thr-Asn-Cys-Pro-Arg-Gly-NH2 Cys-Phe-Val-Arg-Asn-Cys-Pro-Thr-Gly-NH2	Inotocin Annetocin	Locust Earthworm	
Cys-Phe-Val-Arg-Asn-Cys-Pro-Thr-Gly-NH2	Annetocin Lys-	Earthworm Geography & imperial <u>cone</u> snail, <u>pond</u> snail, <u>sea hare</u> ,	
Cys-Phe-Val-Arg-Asn-Cys-Pro-Thr-Gly-NH2 Cys-Phe-Ile-Arg-Asn-Cys-Pro-Lys-Gly-NH2	Annetocin <u>Lys-</u> Connopressin Arg-	Earthworm Geography & imperial cone snail, pond snail, sea hare, leech Striped cone	
Cys-Phe-Val-Arg-Asn-Cys-Pro-Thr-Gly-NH ₂ Cys-Phe-Ile-Arg-Asn-Cys-Pro-Lys-Gly-NH ₂ Cys-Ile-Ile-Arg-Asn-Cys-Pro-Arg-Gly-NH ₂	Annetocin <u>Lys-</u> Connopressin <u>Arg-</u> Connopressin	Earthworm Geography & imperial <u>cone</u> snail, <u>pond</u> snail, <u>sea hare</u> , <u>leech</u> Striped cone snail	

Vasotocin is the evolutionary progenitor of**a**the vertebrate neurohypophysial hormones^[40]