

# The HPG Axis

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**Training Course in Reproductive Health Research – Geneva 2008**



# Hormones are messengers

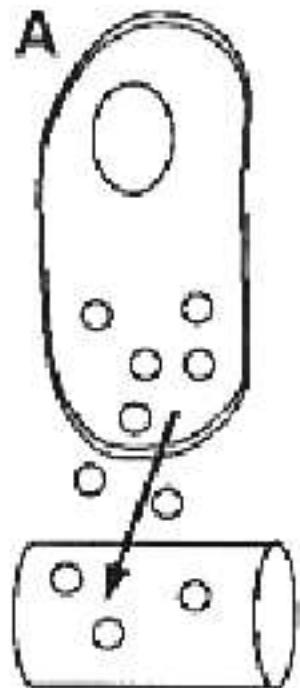
## Hormone:

Chemical signal secreted into the circulation and modulating the function of distant organs.

Particular example of a more generalized biological process

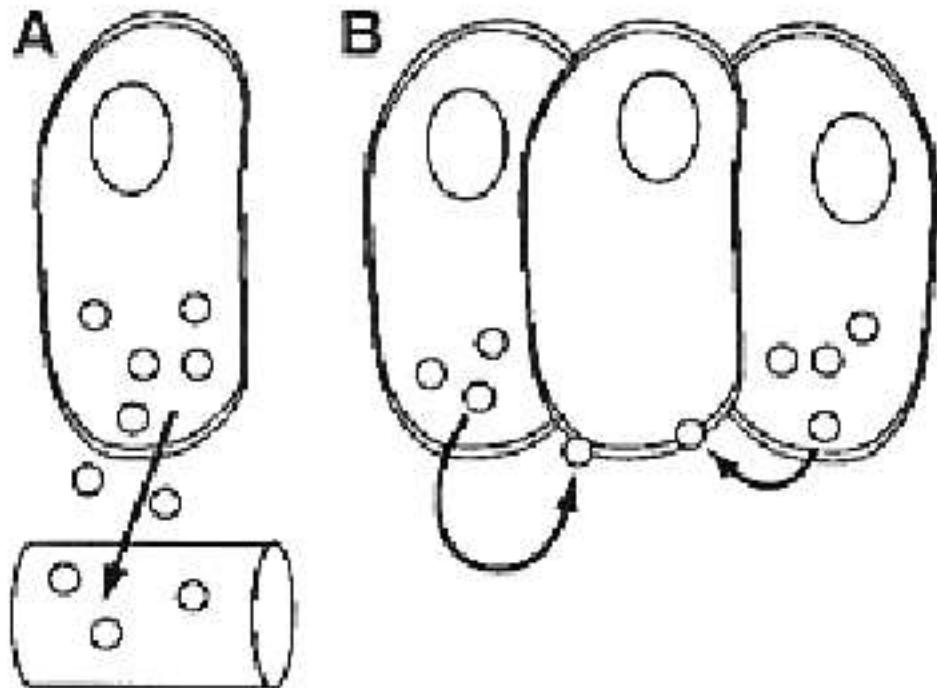
## Inter-cellular signalling

# Different types of hormonal interactions



**Endocrine**

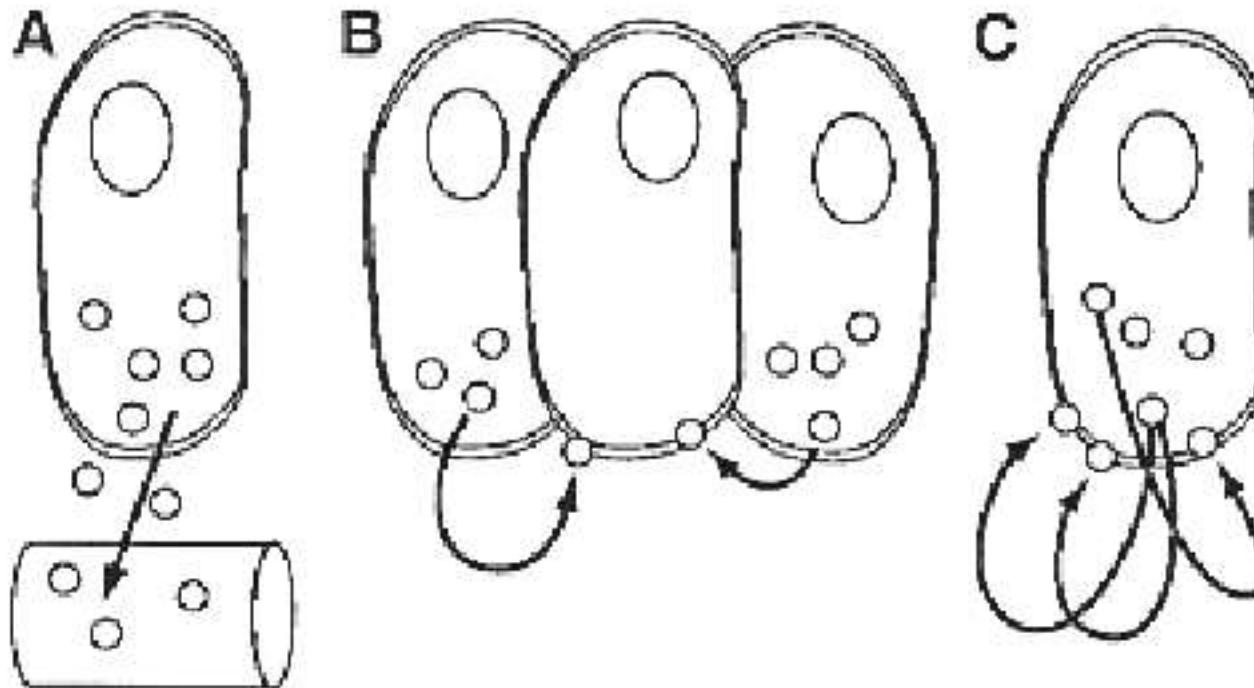
# Different types of hormonal interactions



**Endocrine**

**Paracrine**

# Different types of hormonal interactions

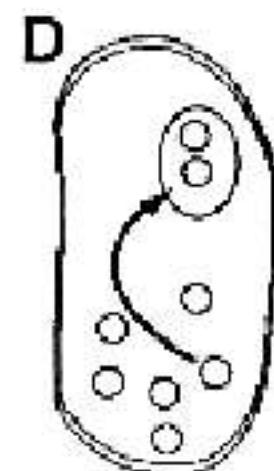
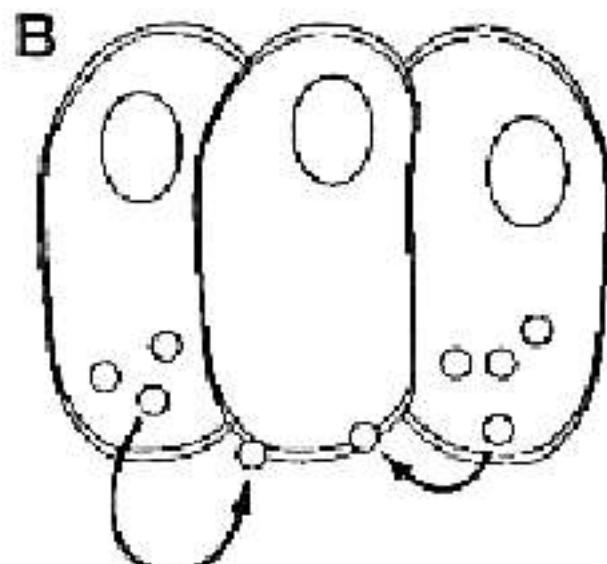
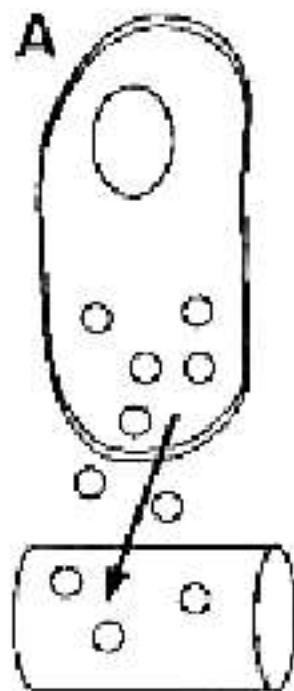


**Endocrine**

**Paracrine**

**Autocrine**

# Different types of hormonal interactions



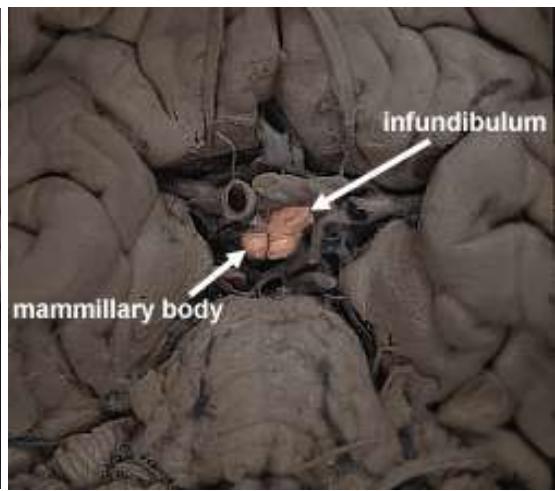
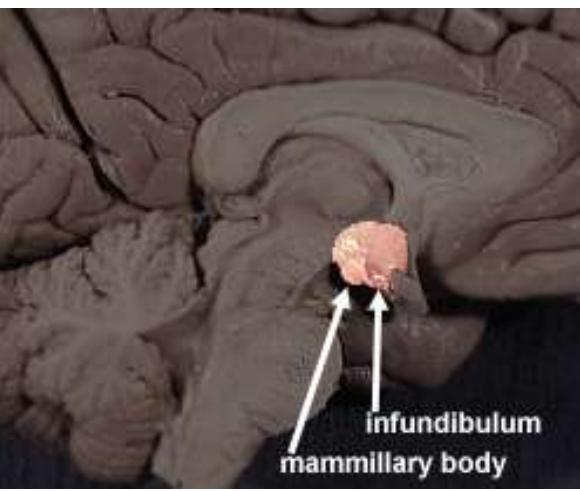
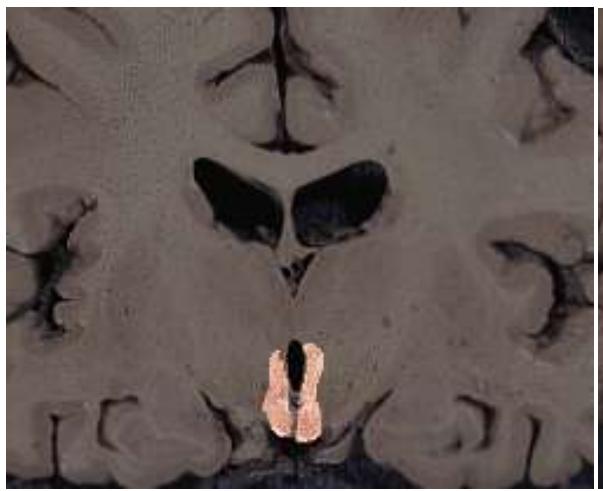
**Endocrine**

**Paracrine**

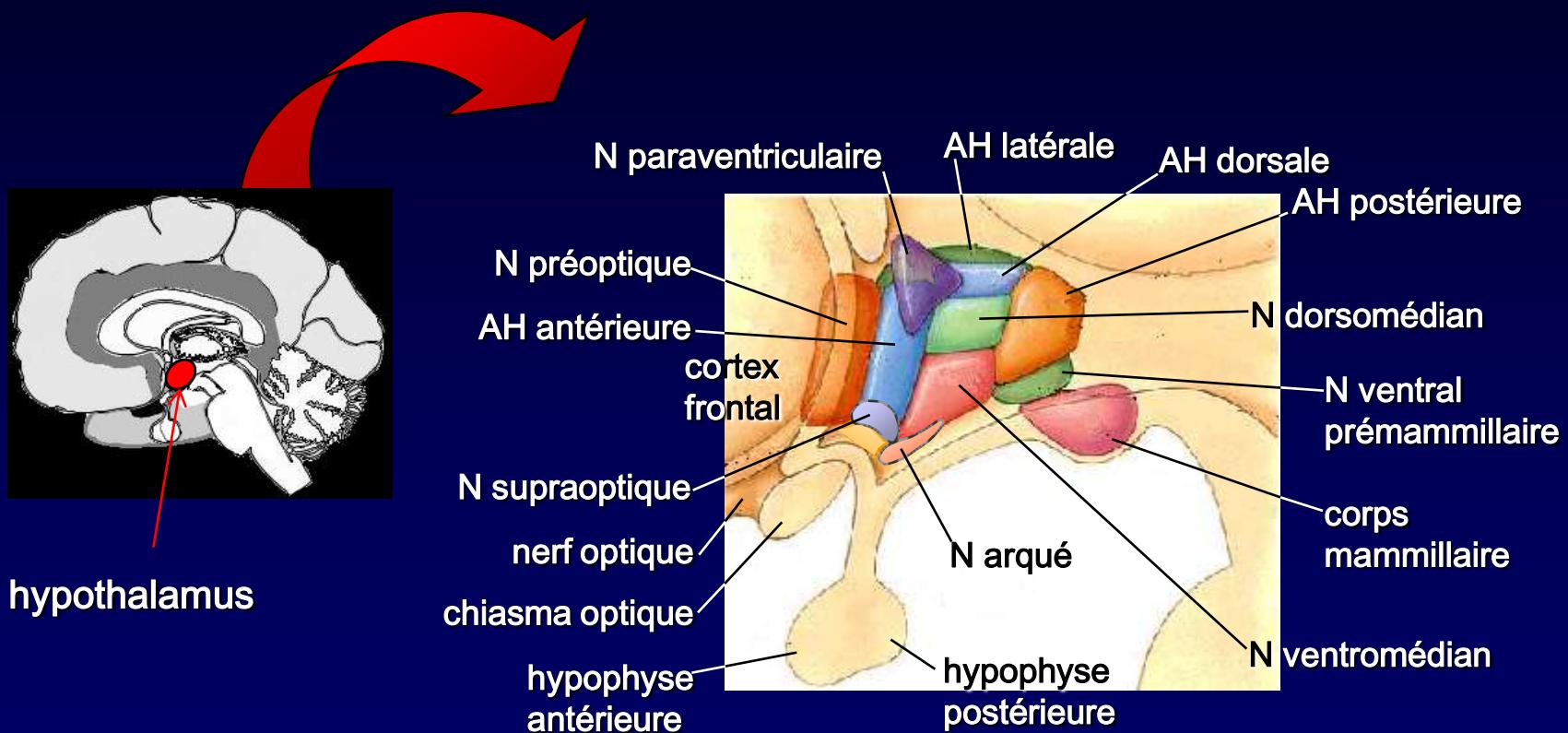
**Autocrine**

**Intracrine**

# **Functional anatomy of hypothalamo-pituitary unit**



# The Hypothalamus



# The Hypothalamus

## Structure

Different groups of neurones (nuclei) located in the walls of the inferior part of the third ventricle

## Fonction

Crossroad where numerous humoral signals are meeting and integrated, resulting in **metabolic and neuroendocrine** regulations.

# The Hypothalamus

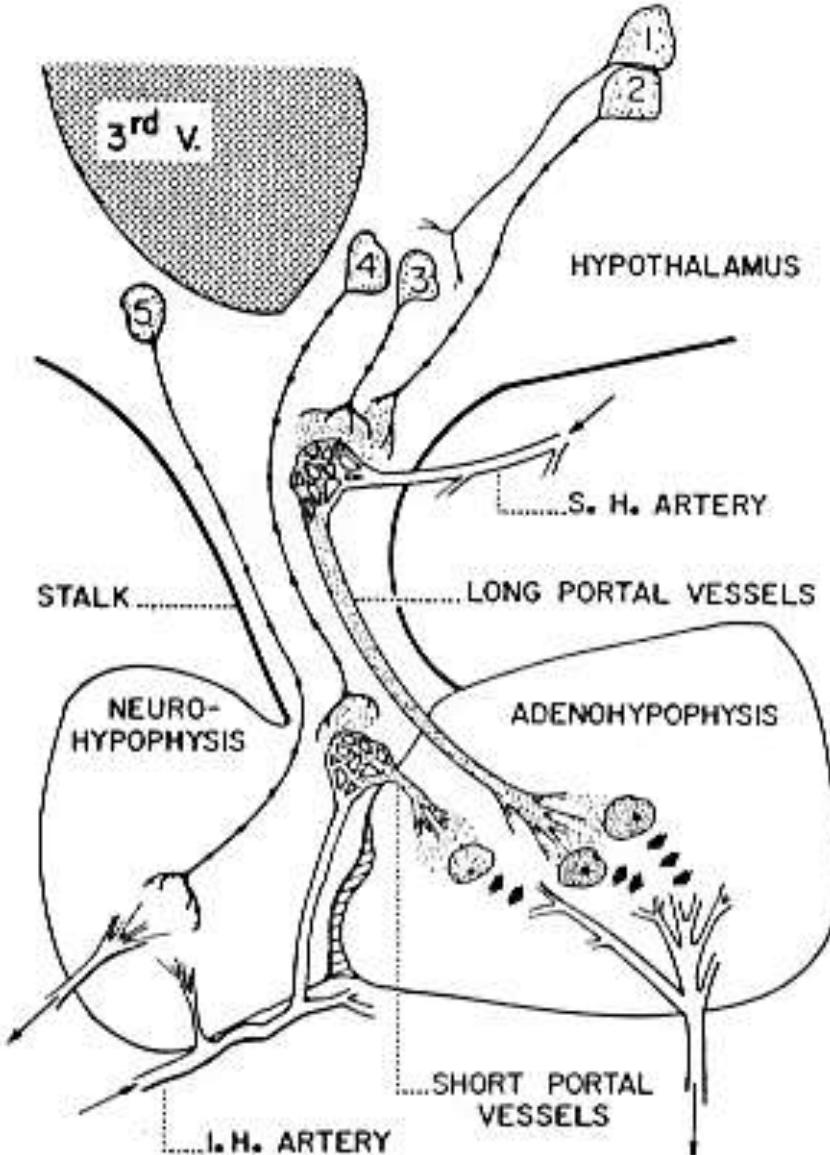
## Specific functions

- 1) Control of anterior pituitary, via *neurohormones (releasing factors)*
- 2) Synthesis of hormones to be stocked in posterior pituitary (then secreted into the peripheral circulation)
- 3) Modulation of food intake, metabolism and energy expenditure

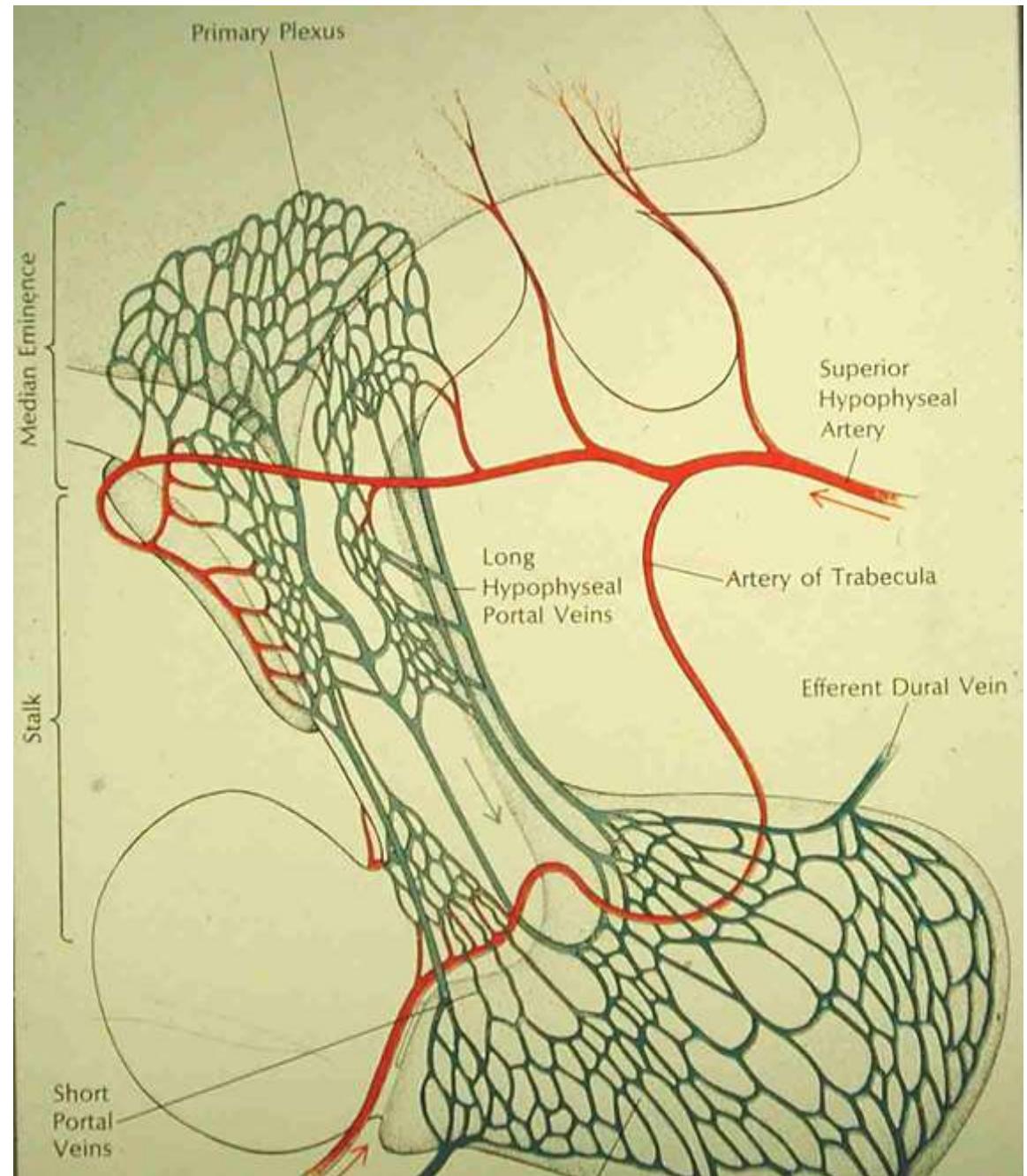
# **HORMONAL TRANSMISSION HT- HP**

***Anterior pituitary:  
vascular***

***Posterior pituitary :  
neuronal***



# The hypothalamo- hypophysial portal blood vessels system

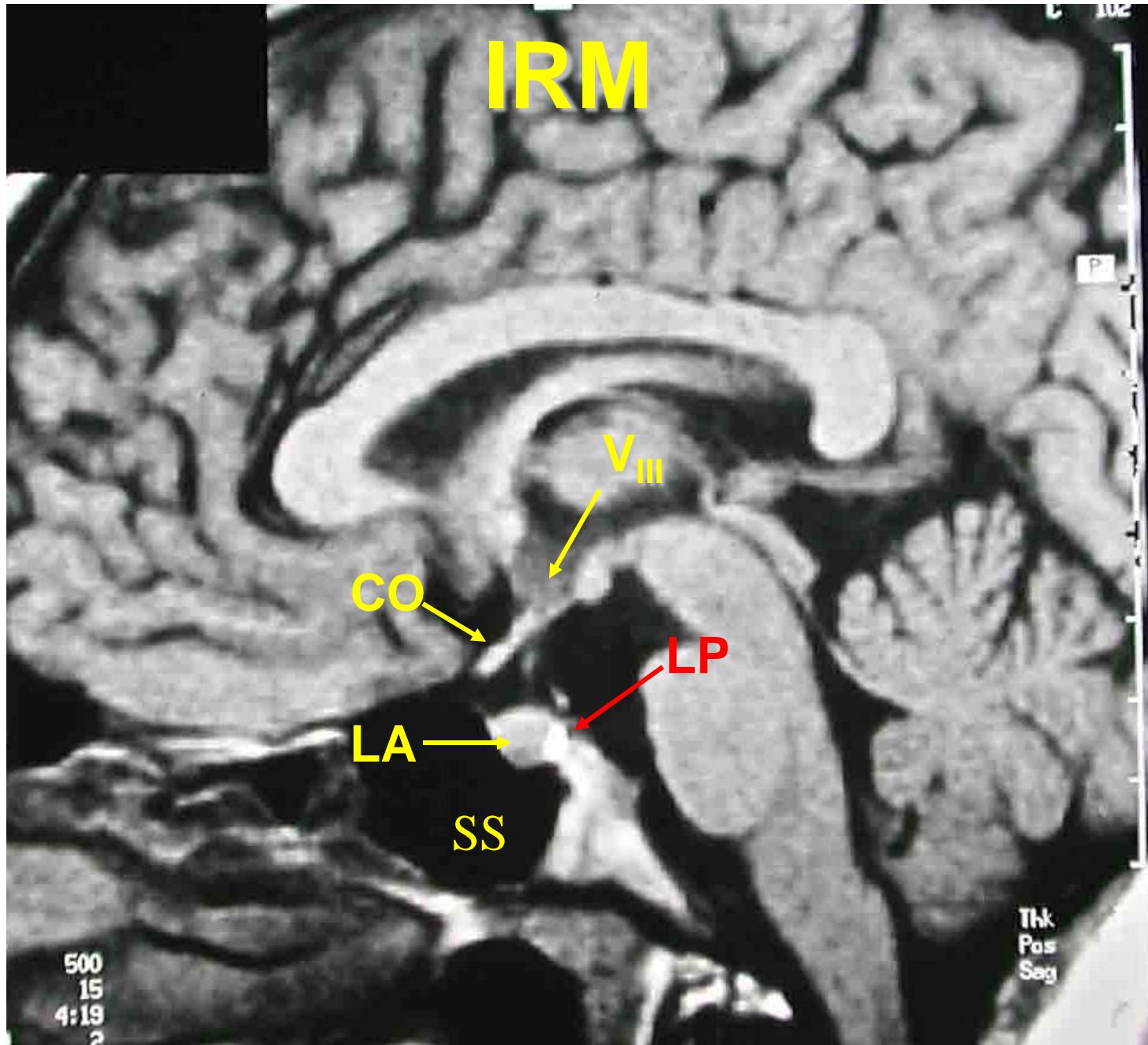


## **Anterior pituitary (adeno-hypophysis):**

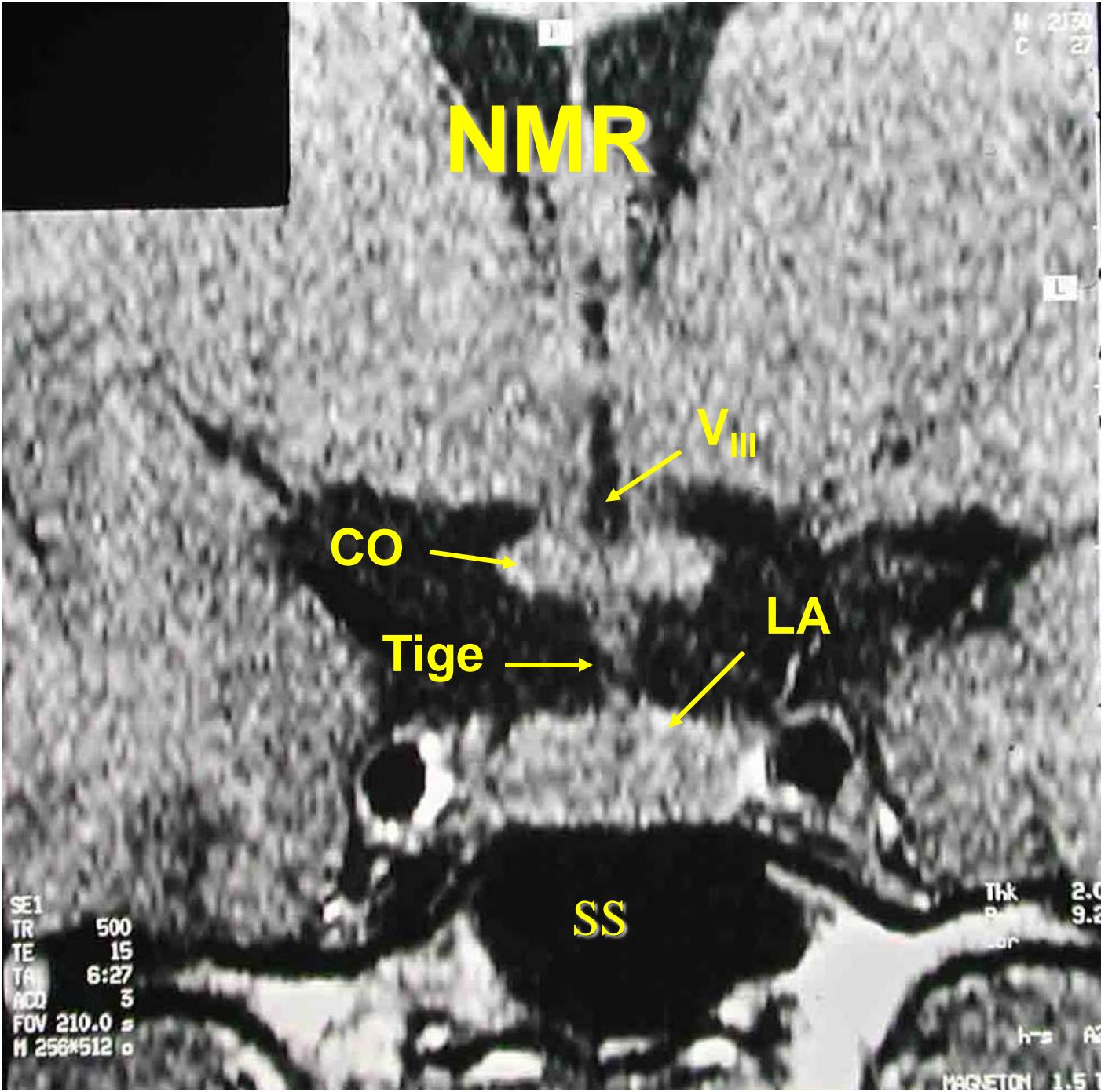
No nervous connection

## **Posterior hypophysis (neuro-hypophysis):**

Connected to the hypothalamus (nervous connections)



NMR



# NMR, contrast

Limit of BBB



SE1  
TR 501  
TE 15  
TA 8:36  
ACQ 4  
FOV 210.0 s  
N 256x512 o

Thk 2.0  
Pos 9.2  
Cor

h-s A2

MAGNETOM 1.5 T

# The hypothalamus

## Specific functions

- 1) Control of anterior pituitary, via **neuro-hormones** (releasing factors)
- 2) Synthesis of hormones to be stocked in posterior pituitary (then secreted into the peripheral circulation)
- 3) Modulation of food intake, metabolism and energy expenditure

# **The hypothalamo-hypophyseal hormonal axes**

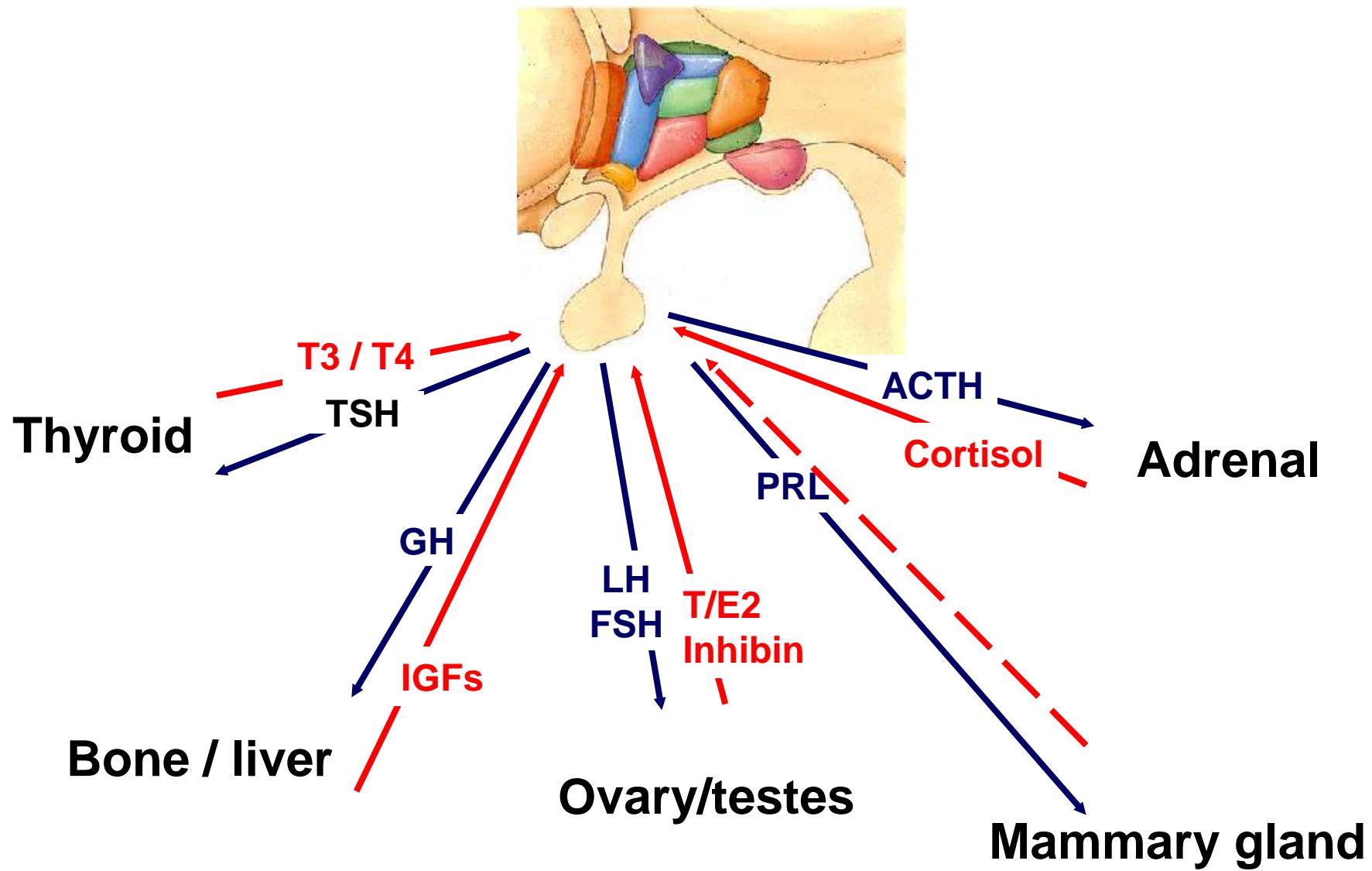
Functional units:

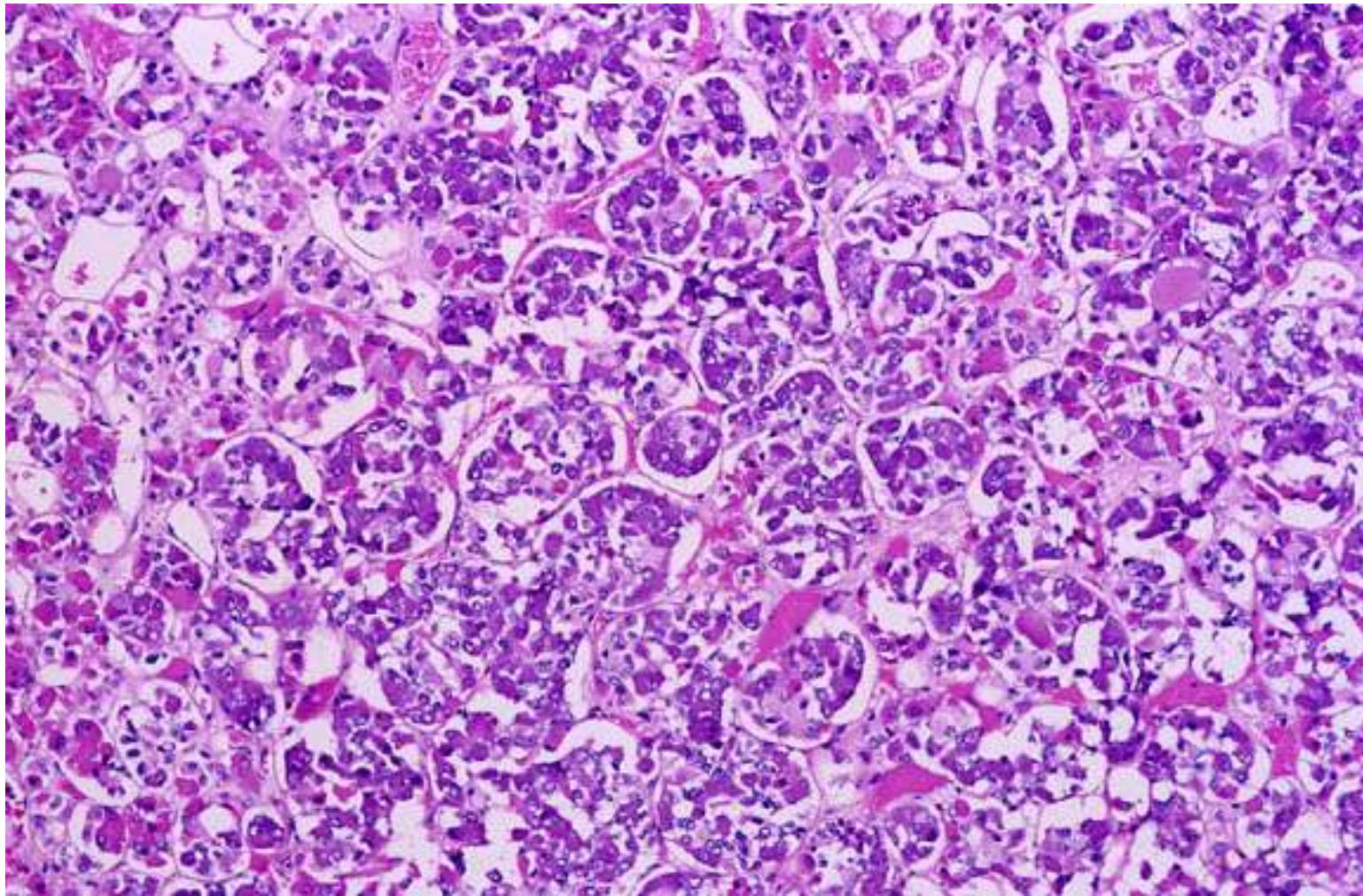
**HT - HP – peripheral glands**

Interdependent secretions

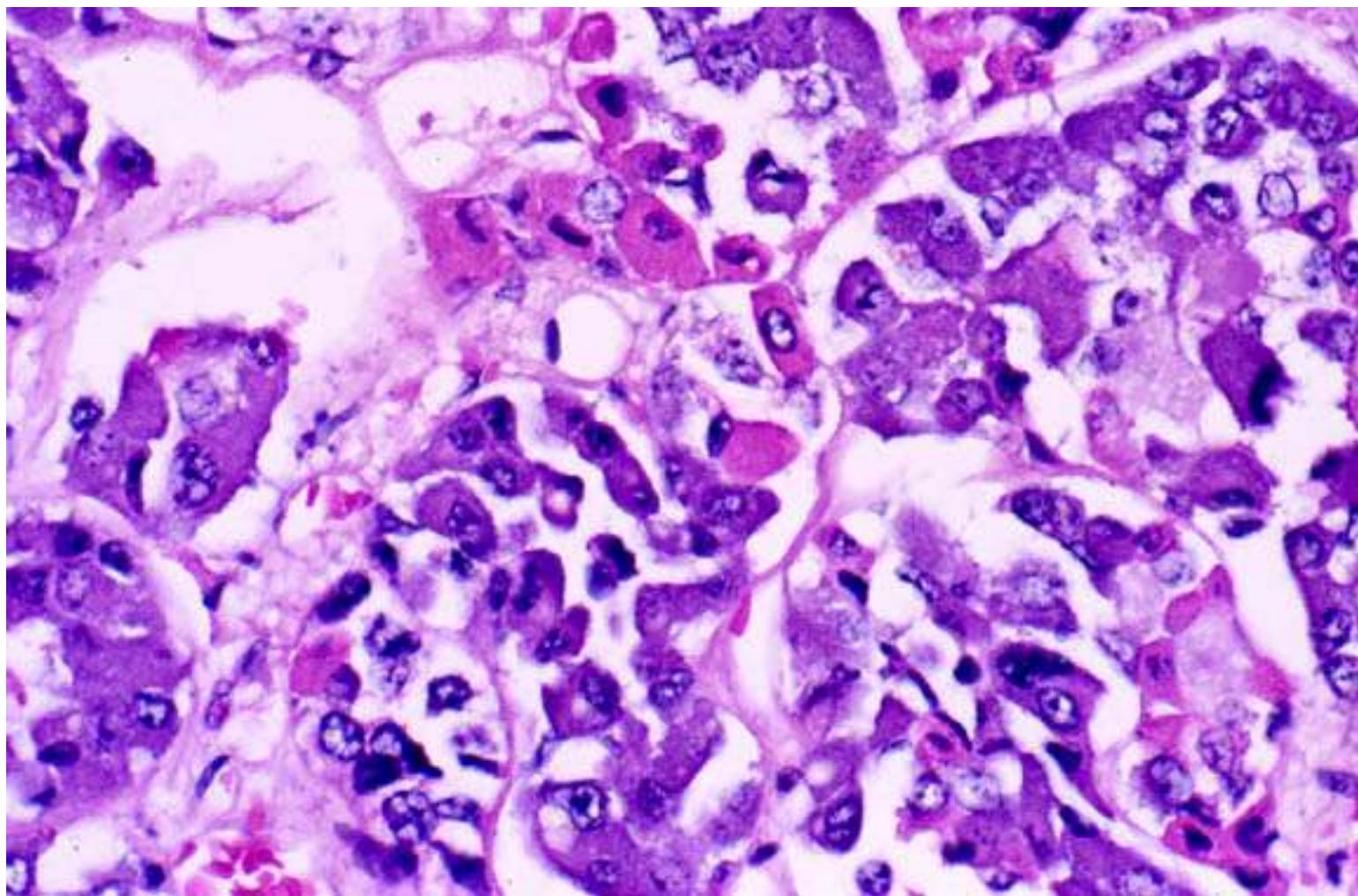
Feed-forward and feed-back effects

# The 5 classical neuroendocrine axes





Anterior pituitary, H&E, x66



Anterior pituitary, basophils, acidophils, and chromophobes, H&E,  
x132

## **Acidophil cells:**

GH, PRL

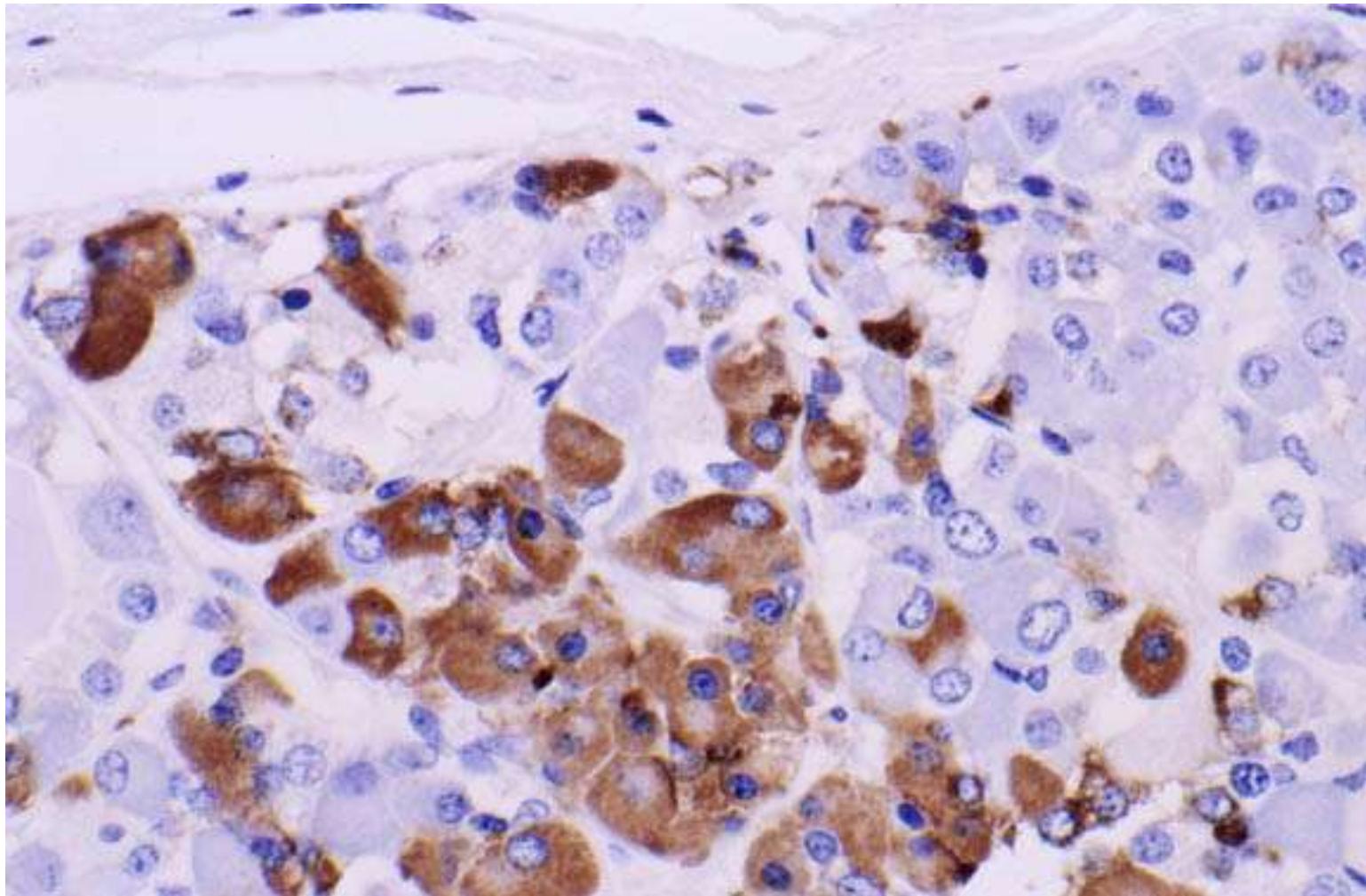
## **Basophil cells:**

TSH, LH et FSH, ACTH

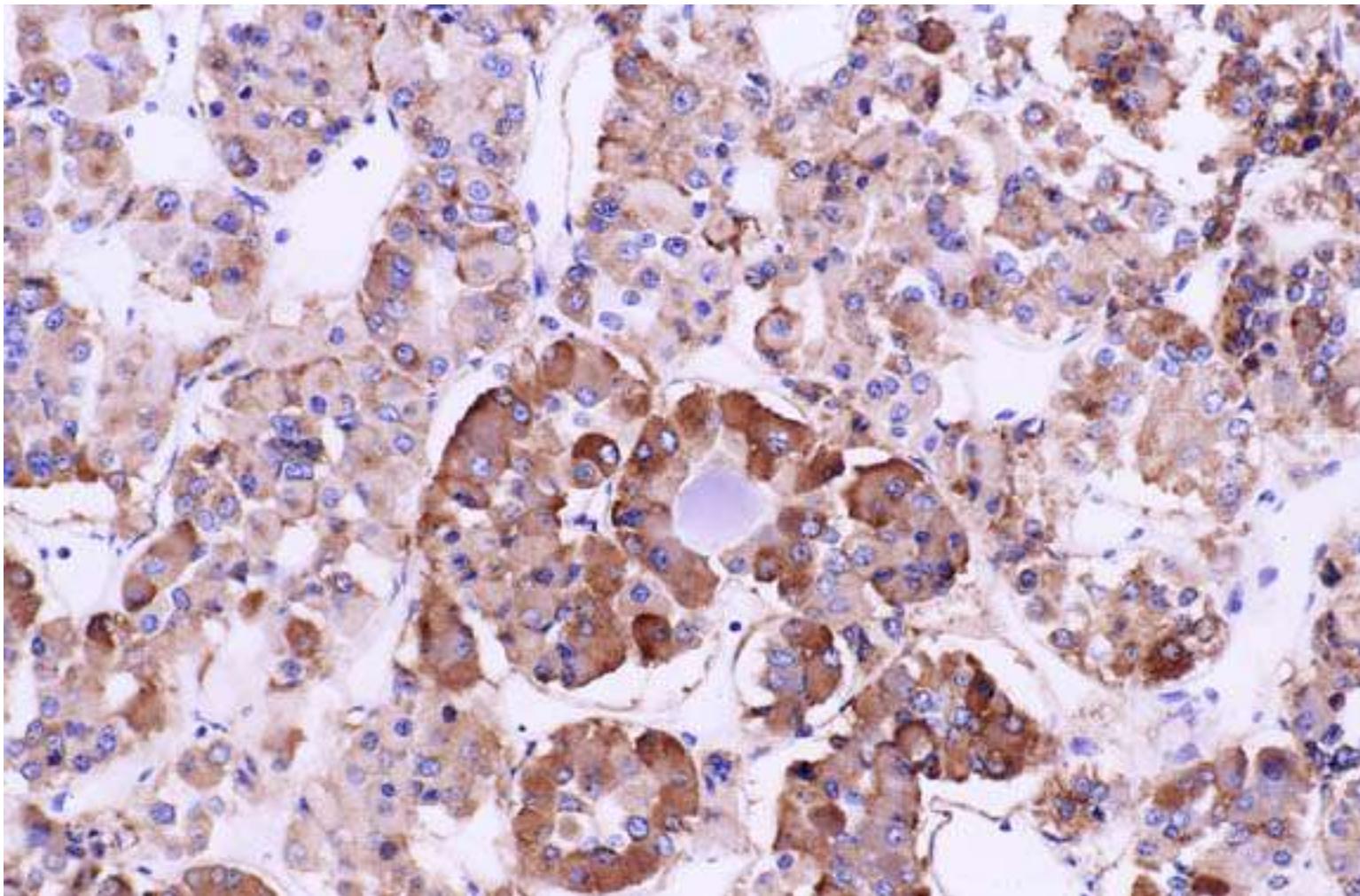
## **Chromophobe cells:**

No hormonal content. Stem cells? Structure cells?

Folliculo-stellate cells: involved in keeping the pituitary structure



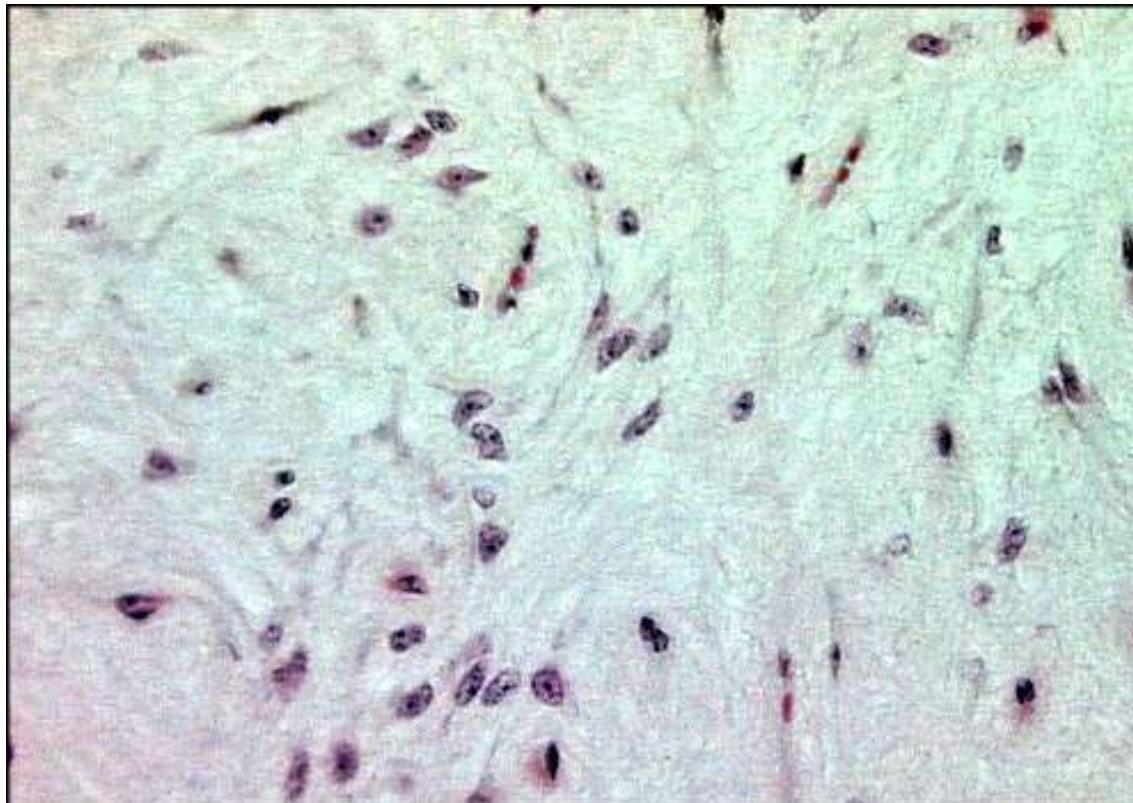
Anterior pituitary: immunohistochemistry **prolactin**  
(immunoperoxidase, x132)



Anterior pituitary: immunohistochemistry [synaptophysine](#)  
(immunoperoxidase, x66)

The posterior pituitary content essentially contains neurosecretory neurons non myelinated, originating from the hypothalamic supra-optic and paraventricular nuclei

They secrete oxytocin and vasopressin.



Axonal prolongations, mixed with glial cells and capillaries



989762  
15-DEC-1954  
14:01  
30-MAR-2001  
IMAGE 48  
SER 1-5

MF 2.00

H

MAGNETOM Symphony  
H-SP VAI3E  
+ : F A L

RP

SD 47 d

09:13  
22-FEB-2002  
IMAGE 25  
SER 1-3

NF 2.00

RA



05:23  
22-FEB-2002  
IMAGE 48  
SER 1-5

MF 2.00

RA

SP 19.2

09:07  
22-FEB-2002  
IMAGE 12  
SER 1-2  
MF 1.15

AL

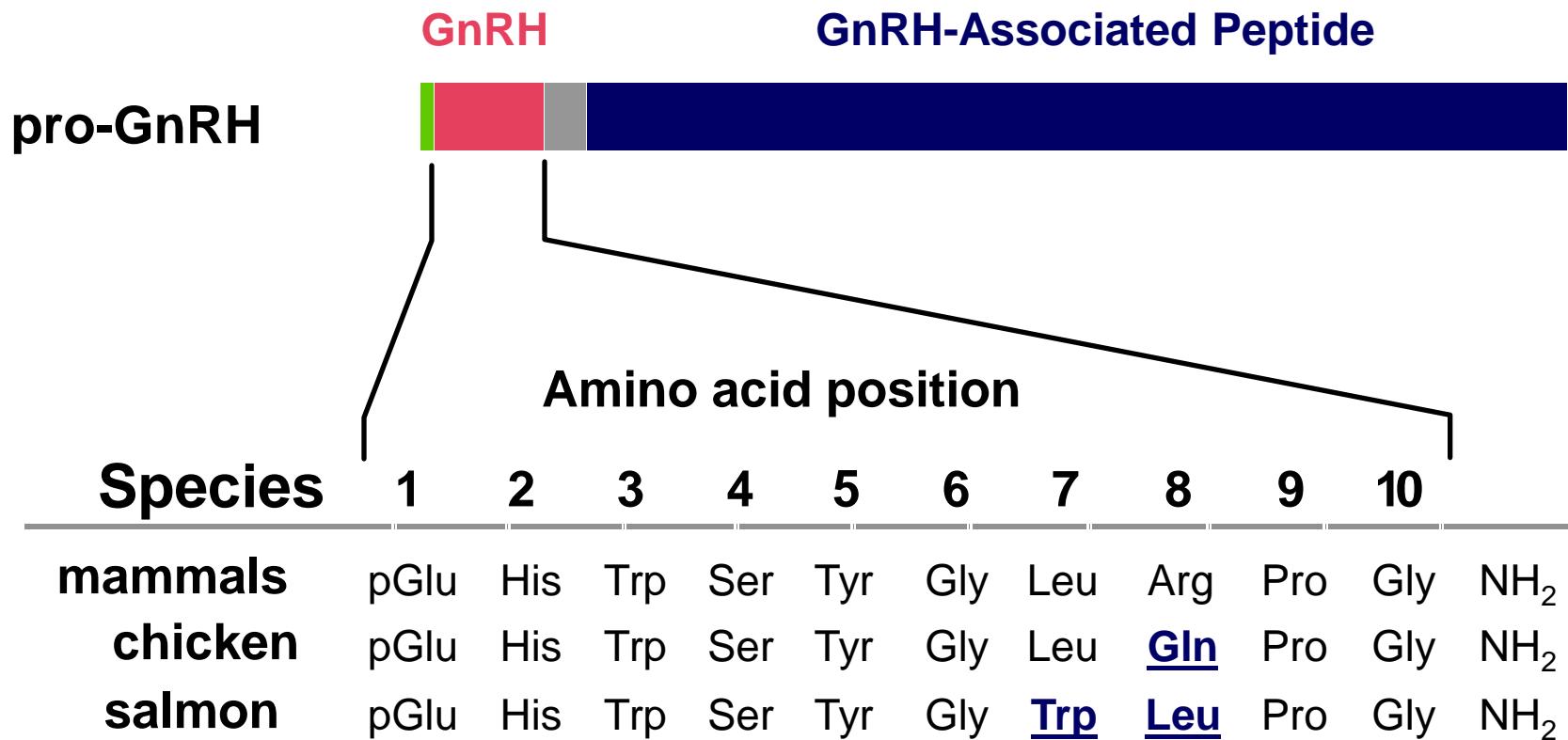
se1

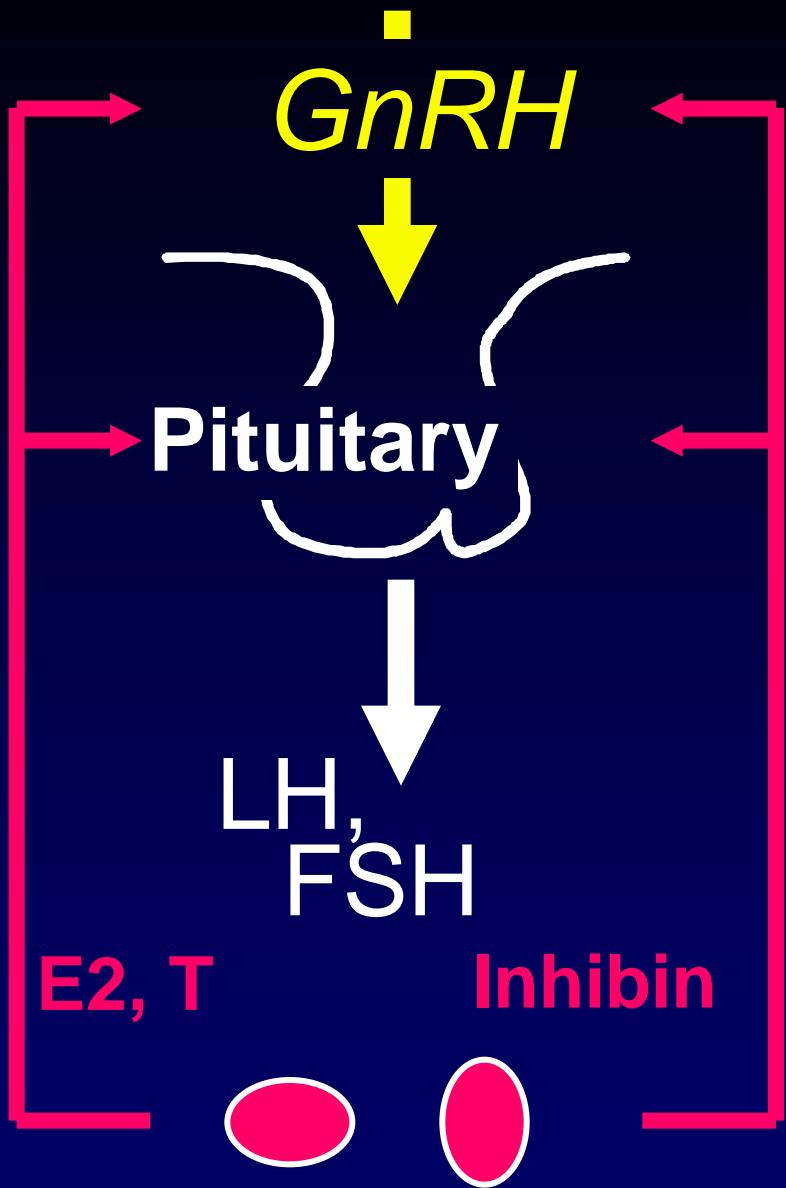
65

SP -0.9  
SL 3.0  
FoV 230\*230

# **The neuroendocrine reproductive axis and hypothalamic GnRH neurons**

# Conservation of primary structure of GnRH in vertebrates





**GnRH**

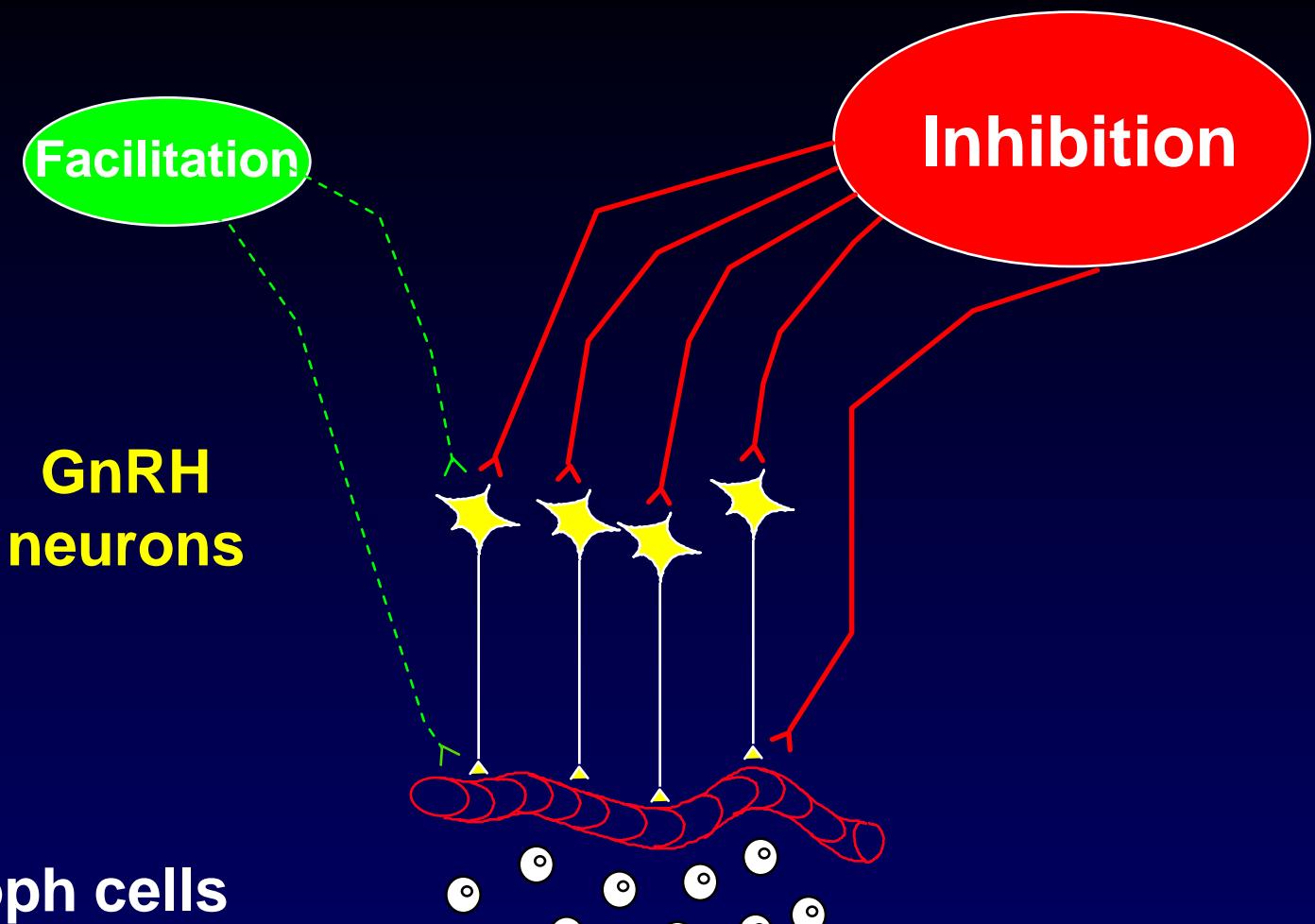
*Gonadotropin Releasing  
Hormone*

Luteinizing Hormone

Follicle Stimulating  
Hormone

E2, T

Inhibin



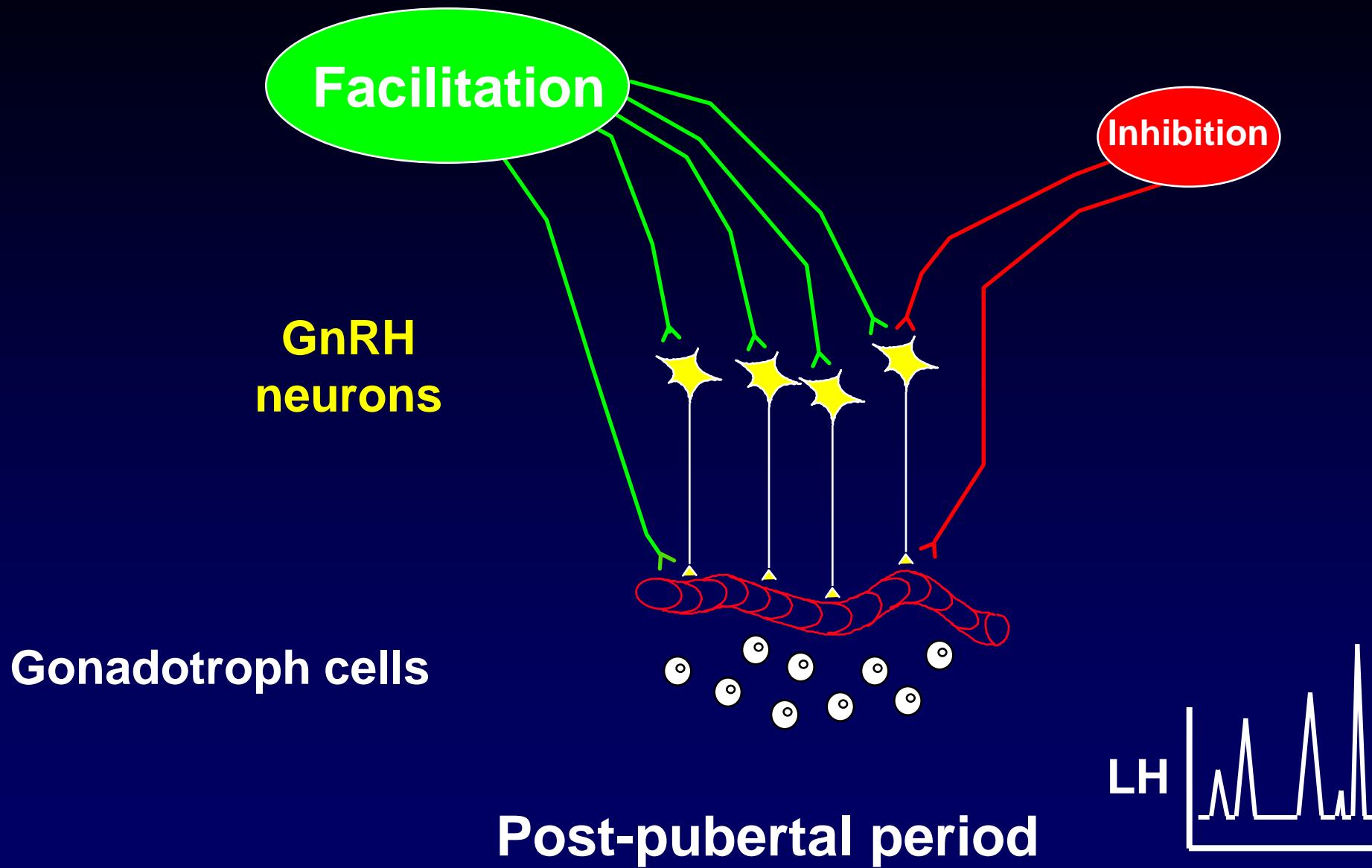
GnRH  
neurons

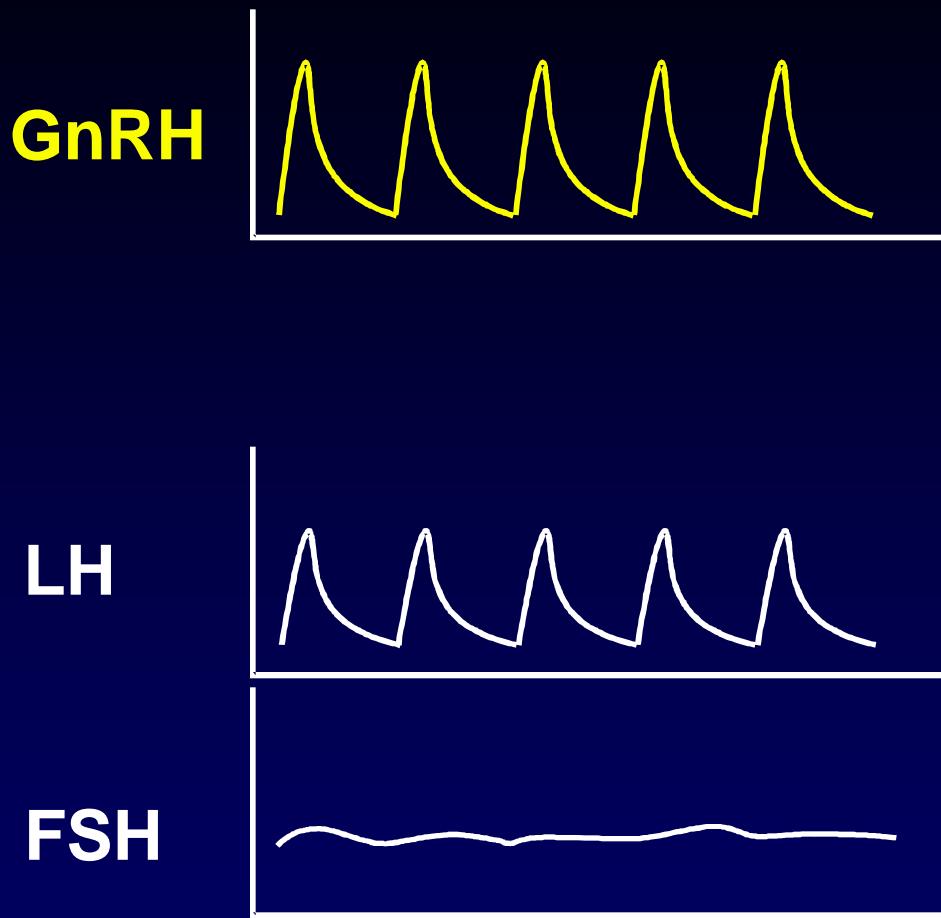
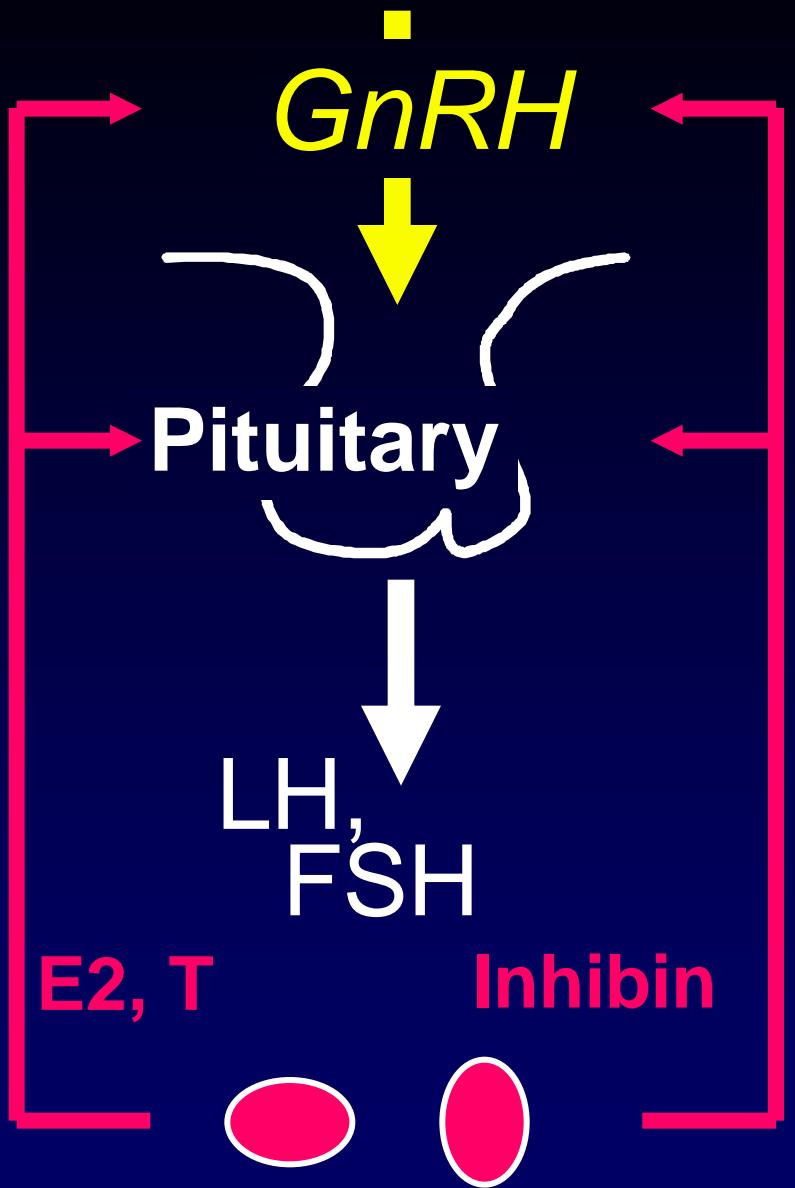
Gonadotroph cells

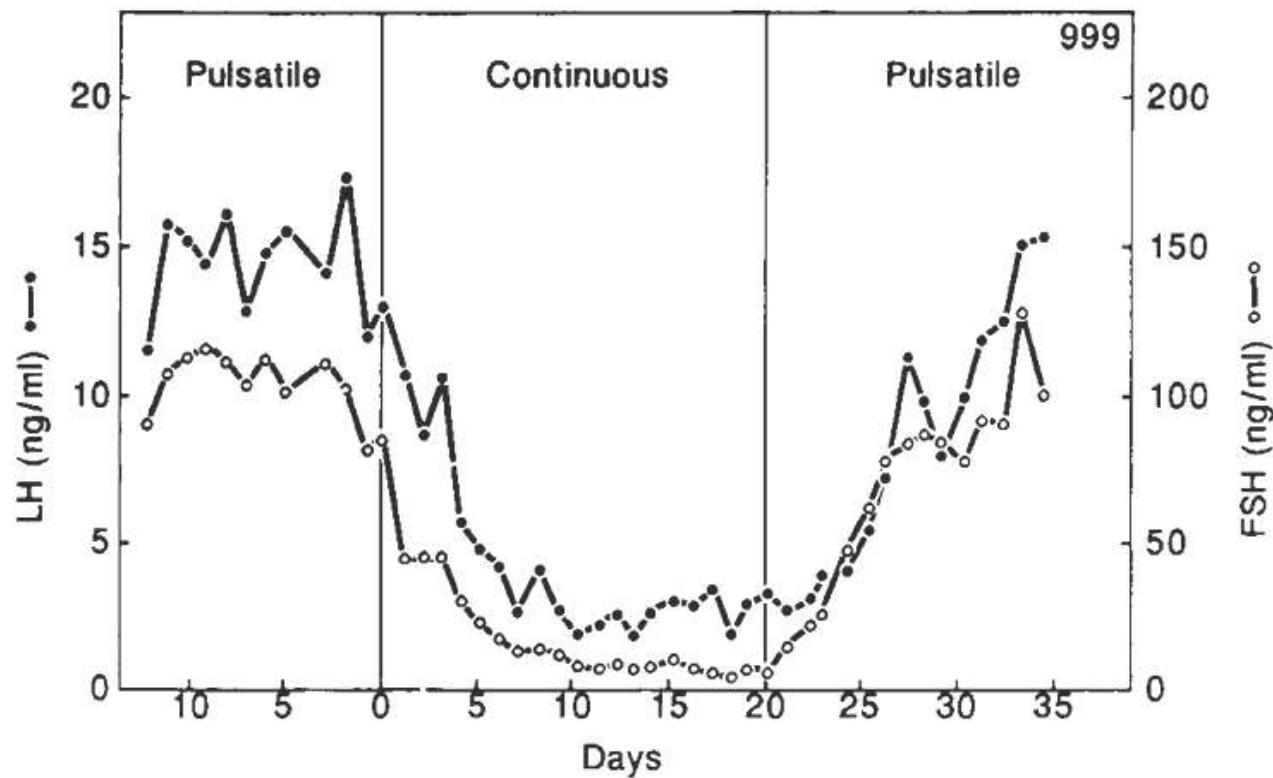
Inhibition

Pre-pubertal period

LH







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**Belchetz et al, Science 1978;**

# Re-awakening of pulsatile GnRH secretion at puberty

