# **Amphibian Metamorphosis**

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#### Urodeles

resorption of the tail fin the destruction of the external gills change in skin structure

#### Anurans

More complicated

Every organ is subject to modification

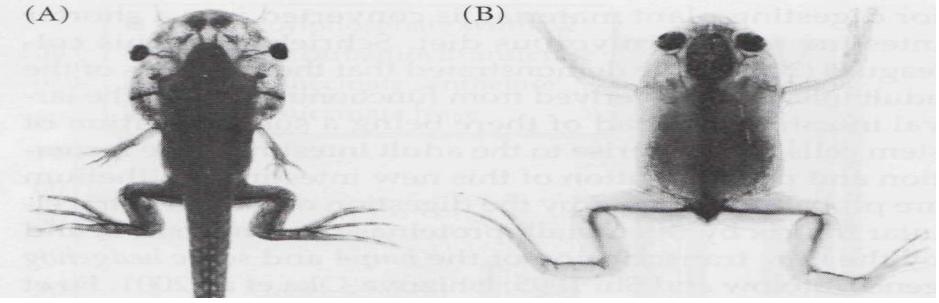
Larval structures respond to T4 and T3 in four ways

growth, death, remodeling, and respecification

#### **Growth of New Structures**

The limbs, nictitating membranes and eyelids emerge

- T3 induces the proliferation and differentiation of new neurons to serve these organs
- Blocking T3 activity prevents these neurons from forming and causes paralysis of the limbs
- Eyes move to the front of the head from their originally lateral position
- Ipsilateral pathways emerge, enabling input from both eyes to reach the same area of the brain



### **Cell Death during Metamorphosis**

T3 causes the degeneration of the paddle-like tail and the oxygen- procuring gills

- First part of tail resorption is caused by suicide, but that the last remnants of the tadpole tail must be killed off
- T3 tells the muscle cells to kill themselves by apoptosis Later in metamorphosis, the tail muscles are destroyed by phagocytosis
- Larval red blood cells are specifically digested by macrophages in the liver and spleen

#### **Remodeling during Metamorphosis**

Larval intestine is converted into a shorter intestine for a carnivorous diet

- Much of the nervous system is remodeled as neurons grow and innervate new targets
- The lateral line system of the tadpole degenerates, and the ears undergo further differentiation
- The middle ear develops, as does the tympanic membrane characteristic of frog and toad outer ears
- Tadpoles experience a brief period of deafness as the neurons change targets

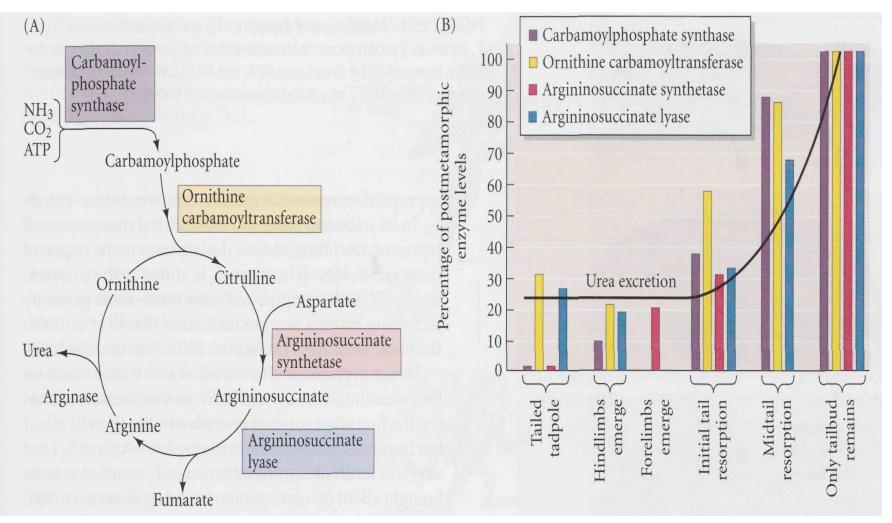
The shape of the anuran skull also changes significantly

#### **Biochemical Respecification**





#### ureotelic

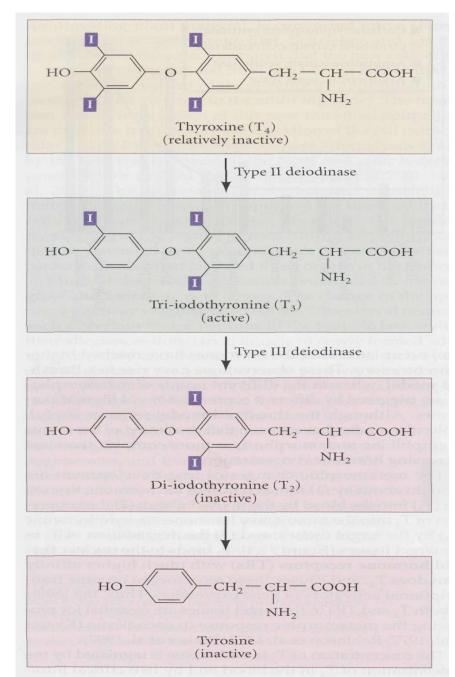


Hormonal control of amphibian metamorphosis Metamorphic changes are due to

- 1. the secretion of the hormone thyroxine (T4)
- 2. the conversion of T4 into the more active hormone, tri-iodothyronine (T3)
- 3. the degradation of T3 in the target tissues
- T3 binds to the nuclear **thyroid hormone receptors (TRs)**

Thus T3 and TRs are essential in each tissue

- Conc. of T3 depends on T4 and two imp. Enzymes
- Type II deiodinase removes an iodine atom from the outer ring of the precursor hormone (T4) to convert it into the more active hormone T3
- 2. Type III deiodinase removes an iodine atom from the inner ring of T3 to convert it into an inactive compound that will eventually be metabolized to tyrosine



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#### **Receptor types**

Two types

- 1. TR α 2. TR β
- TR  $\alpha$  is widely distributed and is present before the organism has a thyroid gland
- TR  $\boldsymbol{\beta}$  is gene product which is activated by hormone
- TR binds and forms dimer with RXR. These dimers bind T3 and effect transcription
- TR-RXR is associated with gene promoters and enhancers and repress transcription
- When T3 added to this complex, gene activation takes place

#### Premetamorphosis

During this stage the thyroid gland has begun to mature and is secreting low levels of T4

- The initiation of T4 secretion may be brought about by corticotropin releasing hormone (CRH)
- CRH may act directly on the frog pituitary, instructing it to release thyroid stimulating hormone (TSH), or it may act generally to make the body cells responsive to low amounts of T3

The tissues that respond earliest to the thyroid hormones are those that express high levels of deiodinase II, and can thereby convert T4 directly into T3

- As the thyroid matures to the stage of prometamorphosis, it secretes more thyroid hormones
- Many major changes (such as tail resorption, gill resorption, and intestinal remodeling) must wait until the **metamorphic climax** stage
- The concentration of T4 rises dramatically, and TRβ levels peak inside the cells

#### Prometamorphosis

During prometamorphosis, the rising levels of thyroid hormones induce higher levels of TRβ

- At metamorphic climax, deiodinase II is expressed, and the tail begins to be resorbed
- Thus the tail undergoes absorption only after the legs are functional

The frog brain also undergoes changes during metamorphosis, and one of the brain's functions is to downregulate metamorphosis once metamorphic climax has been reached

## Thanks