

# **HYPOTHYROIDISM IN THE DOG**

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Michael Schaer, DVM

Diplomate ACVIM, ACVECC

University of Florida, College of Veterinary Medicine

Gainesville, Florida 32610

Hypothyroidism is one of the most commonly diagnosed hormone disorders in the dog. Some cases are clinically very obvious while others are more difficult to diagnose.

## **Types**

The causes of hypothyroidism are listed in Table 1 (pg. 4). Primary acquired hypothyroidism is the most common type of hypothyroidism in the dog. It is usually the result of lymphocytic (autoimmune) thyroiditis or thyroid atrophy. Congenital hypothyroidism is rare in occurrence. Central hypothyroidism results from damage or dysfunction to the anterior pituitary gland. The secondary form of hypothyroidism is rare.

## **Incidence**

Estimates of the incidence of canine hypothyroidism ranges from 1:156 to 1:500. Middle-aged dogs (4 to 10 years of age) of mid-to large-sized breeds are at increased risk. Commonly affected breeds include Golden Retriever, Doberman Pinscher, Irish Setter, Miniature Schnauzer, Dachshund, Cocker Spaniel, Airedale Terrier, Great Dane and Old English Sheepdog. German Shepherd Dogs and mixed-breeds are at low risk.

## **Physiologic Effects of Thyroid Hormones**

Most of the varied effects of thyroid hormones result from the stimulation of oxygen utilization (calorigenic action) by almost all metabolically active tissues. Tissues which do not depend on T<sub>3</sub> and T<sub>4</sub> for oxygen consumption include the adult brain, testes, uterus, lymph nodes, spleen, and anterior pituitary. As a consequence to the increased metabolic rate induced by T<sub>3</sub> or T<sub>4</sub>, nitrogen excretion is increased, internal protein and fat stores are broken down, and body weight is decreased. In young hypothyroid animals, small doses of thyroid hormones induce a positive nitrogen balance because they stimulate growth, but excessive doses will cause protein utilization.

Large doses of thyroid hormones cause excessive body heat production and a slight rise in body temperature. This, in turn, activates heat-dissipating mechanics. Excessive levels of thyroid hormone in conjunction with epinephrine secretion lead to an increased cardiac output. In

addition to these effects, thyroid hormones influence other physiologic processes as listed in Table 2 (pg. 4).

### **ADVERSE SYSTEMIC EFFECTS OF HYPOTHYROIDISM**

Hypothyroidism is defined as a clinical condition characterized by inadequate quantities of circulating thyroid hormone. Table 3 lists the various causes of hypothyroidism in the dog. With the addition of iodized salt to commercial pet foods, hypothyroidism is usually associated with the nongoitrous (without cystic enlargement) form.

The clinical signs of hypothyroidism that are related to decreased metabolic rate and calorogenesis include lethargy, cold intolerance, decreased cerebral function, and a mild increase in body weight. Dermatologic changes are characterized by hair loss, skin thickening, and increased pigmentation.

Cardiovascular changes associated with a decreased cardiac output include a weak apex beat and a weak peripheral pulse. The electrocardiogram can show low amplitude R waves with or without a slow heart rate. Impaired peripheral circulation is suspected when the extremities seem cool to the touch.

Other clinical signs associated with hypothyroidism are decreased libido, gonad underdevelopment, anestrus, diarrhea, constipation, anemia, muscle weakness, muscle and nerve dysfunction (including cranial nerves VII, VIII and X dysfunction), and mammary milk production.

Myxedema is the extreme form of hypothyroidism. The signs in the dog are characterized by severe mental depression terminating in coma and hypothermia. Signs of hypothyroidism are usually present, but hypoventilation, hypotension, slow heart rate, and profound hypothermia are usually present as well. Often acute decompensation is triggered by an anesthetic episode. For these reasons, great care should be taken when anesthetizing a hypothyroid dog.

### **Endocrinologic Diagnosis**

The serum T<sub>4</sub> determination is still the most commonly run test for initial thyroid evaluation. A normal value (1.5-4.3  $\mu$ /dl, 20-55 nmol/L) essentially rules out hypothyroidism. A very low T<sub>4</sub> level in conjunction with appropriate clinical signs and the absence of confounding factors is usually sufficient for making the diagnosis of hypothyroidism.

The serum T<sub>3</sub> determination (normal 0.7-2.3 nmol/L, 45-150 ng/dl) is commonly run but it is not as diagnostic as T<sub>4</sub> measurement. It is not uncommon to find normal T<sub>3</sub> concentrations in dogs with hypothyroidism.

The thyroid stimulating hormone (TSH) response test is used to identify hypothyroidism when the T<sub>4</sub> results are questionable. A reduced or no response to TSH is expected in hypothyroidism. It is important to consider that although the test will distinguish many borderline situations, the results are not always clear in dogs with nonthyroid illness or those treated with certain drugs.

However, post TSH T<sub>4</sub> levels greater than 45 nmol/L rule out hypothyroidism while levels less than 15 nmol/L indicate a need for replacement therapy. On the negative side, the commercial availability of TSH is unreliable and it is relatively expensive.

The free T<sub>4</sub> level represents the fraction of total hormone available for target cell entry. It comprises less than 1% of total T<sub>4</sub>. The equilibrium dialysis technique is the preferred method with normal values ranging from 12-33 pmol/L (Nachreiner, MSU).

Many laboratories are now offering the TSH assay. It is essential that the appropriate standards are run with this test. Normal TSH levels range from 7-40 mU/L (Nachreiner, MSU), while the hypothyroid dog typically has elevated serum levels.

Antithyroid globulin antibody and antithyroid hormone antibody titers can be increased with lymphocytic or autoimmune thyroiditis. These dogs can be either euthyroid or hypothyroid in the presence of an elevated antibody titer. Some dogs with elevated antithyroid hormone antibody titers can also have increased measured serum thyroid hormone levels while still being clinically euthyroid. Therefore, elevated antithyroid globulin and antithyroid hormone antibodies are not reliable tests for diagnosing canine hypothyroidism.

### **Treatment**

Primary hypothyroidism is initially treated with thyroxine (levothyroxine) at a dosage of 22 µg/kg (0.1 mg/10lb) every 12 hours. This same dose can be decreased to once daily treatment after the first month. Periodic blood level monitoring should be done 4-7 hours post-thyroxine administration, and the treatment should be adjusted accordingly.

Improved mentation and activity levels should become apparent over the first 2-7 day period. Skin and neurological improvement should occur after 1-3 months of treatment. Reproductive abnormalities might hopefully improve over a 3-10 month period.

Sodium liothyronine (synthetic T<sub>3</sub> Cytomel) is not the initial thyroid hormone supplement of choice. While liothyronine will raise the T<sub>3</sub> level, it will also lower the T<sub>4</sub> level through negative feedback inhibition. Synthetic T<sub>3</sub> therapy is indicated when levothyroxine treatment fails to achieve a desired clinical response in a confirmed hypothyroid dog. This can arise if there is impaired thyroxine absorption from the bowel.

### **Prognosis**

The prognosis is generally good for primary hypothyroidism so long as there are no complications associated with the hyperlipidemia (coronary artery disease, acute pancreatitis) or the neuropathy.

**Table 1. Causes of Hypothyroidism in the Dog**

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<b>I.</b>	<p><b><u>Primary hypothyroidism</u></b>          congenital          acquired – atrophy of unknown etiology          autoimmune – lymphocytic thyroiditis          neoplastic – bilateral thyroid carcinoma</p>
<b>II.</b>	<p><b><u>Pituitary hypothyroidism (Secondary hypothyroidism)</u></b>          pituitary neoplasms, e.g. chromophobe adenoma          congenital hypopituitarism          defective TSH molecule          Iatrogenic            drugs (glucocorticoids)            radiation            hypophysectomy</p>
<b>III.</b>	<p><b><u>Tertiary hypothyroidism</u></b>          congenital hypothalamic malformation          acquired destruction of the hypothalamus</p>
<b>IV.</b>	<p><b><u>Hypothyroidism associated with goiter</u></b>          iodine deficiency          dyshormonogenesis (congenital)          ingestin of various goitrogens, i.e., thiocyanates</p>

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**Table 2. Physiologic Effects of Thyroid Hormones**

<b>Effects of Excessive Levels of Thyroid Hormones</b>	<b>Effects of Inadequate Levels of Thyroid Hormones</b>
<ul style="list-style-type: none"> <li>■ rapid mentation, irritability, and restlessness</li> <li>■ increased CNS sensitivity to circulating catecholamines</li> <li>■ shortened stretch reflex time</li> <li>■ increased rate of carbohydrate absorption from the intestinal tract</li> <li>■ increased lipid metabolism</li> <li>■ increased requirements of water soluble vitamins and fat soluble vitamins</li> </ul>	<ul style="list-style-type: none"> <li>■ anemia resulting from decreased erythropoiesis</li> <li>■ depressed lactation</li> <li>■ infertility</li> <li>■ decreased mentation</li> <li>■ increased stretch reflex time</li> <li>■ depressed cholesterol and lipid metabolism</li> <li>■ impaired growth and skeletal maturation in young animals</li> </ul>

**Table 3. Common Clinicopathologic Findings in the Hypothyroid Dog**

<b>Test</b>	<b>Abnormalities</b>
Hemogram	Normocytic normochromic anemia
Serum chemistries	Increased cholesterol Increased triglycerides Increased creatine kinase