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# Graves' Disease in 100 Cases in Conakry: Epidemiological, Clinical, Therapeutic, and Evolutionary Aspects

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#### **Abstract**

Context and Objective: Graves' disease is an autoimmune disorder of the thyroid gland that occurs in genetically predisposed individuals. It represents the most frequent cause of hyperthyroidism with a clear female predominance. The objective of our work was to report the sociodemographic, clinical, therapeutic, and evolutionary characteristics of Graves' disease at the University Hospital of Conakry. Methods: This was a cross-sectional, descriptive study, over the period from December 2016 to June 2021, at the endocrinology consultation of Donka University Hospital. Epidemiological, clinical, therapeutic, and evolutionary variables of patients followed up for Graves' disease were collected and analyzed. The diagnosis of Graves' disease was based on the presence of clinical signs of thyrotoxicosis, diffuse goiter, exophthalmos, and or T-RAK positivity. Results: Graves' disease was related to 33% of thyroid consultations and 64% of hyperthyroidism. The sex ratio M/F was 0.07. The median age of the patients was  $39.4 \pm 13$  years. The main reason for consultation was thyrotoxicosis syndrome, dominated by cardiovascular signs (92%). TRAK was performed in 38 patients with a positive result in 89%, i.e., a mean level of 17.93 mUI/l. All patients were treated with synthetic antithyroid drugs, with a favorable clinical evolution. Surgery was considered in 4 patients after the stabilization of the thyroid function. The follow-up was considered regular in 49 patients (49%). Conclusion: Graves' disease is the most frequent hyperthyroidism in Conakry with a clear predominance of women, especially young women. Efforts should be focused on improving diagnosis and the access to treatment for better patient compliance.

## Keywords

Autoimmunity, Thyroid, Graves' Disease, Conakry

### 1. Introduction

Hyperthyroidism is one of the most common endocrine disorders. Graves' disease, is an autoimmune disorder of the thyroid gland that occurs in genetically predisposed individuals. It is the most frequent cause of hyperthyroidism, with a clear female predominance. It is characterized by the presence of anti-TSH receptor antibodies. It is classically manifested by signs of hyperthyroidism, homogeneous goiter, and sometimes ophthalmopathy [1] [2].

Graves' disease represents a variable percentage of diagnosed hyperthyroidism from one country to another, but it always remains high. In Cameroon, it represents 82% of hyperthyroidism [3] and 72% in Senegal [4]. Its diagnosis is often simple and its early management allows avoiding the evolution towards numerous complications, namely cardiovascular. Nevertheless, treatment remains difficult and relies on a therapeutic arsenal combining synthetic anti-thyroid drugs, radioactive iodine therapy, and surgery [5]. In Guinea, no recent study has been published on the prevalence of Graves' disease. Thus, the objective of this study was to report the sociodemographic, clinical, therapeutic, and evolutionary characteristics of Graves' disease at the University Hospital of Conakry.

#### 2. Method

This was a cross-sectional, descriptive study of patients followed for Graves' disease at Donka University Hospital from December 1, 2016, to June 30, 2021. All patients followed up in the department for Graves' disease were included. No patient was excluded.

Peripheral thyrotoxicosis was defined as the presence of clinical manifestations of thyrotoxicosis associated with increased thyroid hormone levels (free T3 and free T4) and decreased TSHus levels.

The diagnosis of Graves' disease was based on the presence of clinical signs of thyrotoxicosis, diffuse goiter, exophthalmos, and/or TSH receptor antibody positivity.

The patients were questioned (age, gender, profession, residence, family, and personal history) and systematically examined for goiter, signs of thyrotoxicosis, compression (dysphagia, dysphonia, cough, and dyspnea), and cardiothyreosis (heart failure and rhythm disorder).

According to the clinical context, a complementary exploration was proposed. These included hormonal exploration (TSHus, free T3 and free T4), inflammatory assessment (sedimentation rate, C-reactive protein, blood count), immunological assessment (anti-TSH receptor or TRAK), and imaging, particularly cervical ultrasound with determination of the TIRADS (Thyroid Imaging Reporting and Data System) classification [6] and thyroid scintigraphy.

Epidemiological, clinical, therapeutic, and evolutionary variables of the followedup patients were collected and analyzed. Informed consent was obtained for the collection and analysis of patient and anonymity was in accordance with the principles of scientific writing. The data were collected using a questionnaire, entered, and analyzed using SPSS version 23 software.

#### 3. Results

Between December 2016 and June 2021, 300 patients were consulted in the Endocrinology Department of Donka University Hospital for a thyroid problem. Among them 156 patients (52%) were hyperthyroid. Graves' disease was the main etiology and concerned 100 patients (64%). Women represented 93% of the cases, with a sex ratio M/F of 0.07. The medium age of the patients was  $39.4 \pm 13$ years and the age group 31 to 40 years was the most affected (30%). According to the socio-professional categories, civil servants were the most represented (33%), followed by housewives (31%), workers in the informal sector (20%), and pupils/students (12%). More than half of the patients (65%) resided in Conakry, the capital city, as opposed to 35% who came from the cities inside the country. The average delay of consultation of the patients in our service was 2 years. The thyrotoxicosis syndrome dominated symptoms (Table 1), with cardiovascular signs being the most frequently found, namely palpitation and dyspnea, respectively in 92% and 84% of cases. Goiter (Table 2) was noted in 94 patients, i.e., 94% of cases, and a vascular murmur were found in 62% of cases. Exophthalmos was found in 67 patients (67%) of which 10% were unilateral. Pretibial myxedema was noted in 28 patients (28% of cases). Graves' disease was associated with other autoimmune pathologies in 3 cases: 2 cases of vitiligo and one case of rheumatoid arthritis.

Graves' disease was associated with diabetes mellitus in 8% and hypertension in 22%. Cardiothyreosis was the main complication found (6%). The hormonal assessment was performed in 91 patients (91%) and TSH was lower in all patients. The mean TSH level was 0.09 ųUI/ml with extremes from 0.001 to 0.14 ųUI/ml.

**Table 1.** Distribution of functional signs in the 100 patients with Graves' disease in the endocrinology department from December 2016 to June 2021.

Symptoms	Percentage (%)
General:	
- Weight loss	89
- Physical asthenia	89
Cardiovascular:	
- Palpitations	98
- Dyspnea	98
Neuropsychiatric:	
- Nervousness	91
- Insomnia	68
- Thermophobia	81
Digestive:	
- Acceleration of the transit	58

**Table 2.** Distribution of physical and associated signs in the 100 patients with Graves' disease in the endocrinology department from December 2016 to June 2021.

Signs	Percentage (%)
Thyrotoxicosis	
- Hand Wetness	91
- Osteotendinous hyperreflexia	91
- Tremors	83
- Tachycardia	92
Associated Signs	
- Goiter	94 cases, vascular 62 cases
- Bilateral exophthalmos	67 cases
- Pre-tibial myxedema	28 cases

Anti TSH receptor antibodies were performed by 38 patients, the result was positive in 89%, *i.e.*, a mean level of 17.93 mIU/l for extremes from 1.7 to 44 IU/l. Thyroid ultrasound was performed in 77 patients (77%). TIRAD III was the most represented (51%).

All patients were treated with synthetic antithyroid drugs, with a favorable clinical course. Beta-blockers were prescribed in 77 patients (77%) and anxiolytics were prescribed in 65 patients (65%).

Thyroid surgery was considered in 4 patients after stabilization of thyroid function. One patient with iatrogenic hypothyroidism was substituted with levothyroxine until thyroid function stabilized. Follow-up was considered regular in 49 patients (49%).

Six patients (6%) relapsed when the treatment was stopped. This relapse was observed on average after 18 months. These patients had a large goiter. A total of 27 patients (27%) were lost to follow-up during treatment.

## 4. Discussion

This study reflects the hospital data on Graves' disease in our country. However, it had limitations related to the sample size, the high cost, and the local unavailability of some complementary explorations, especially immunology. As a result, diagnostic certainty could not be established in all cases. Furthermore, the data cannot be extrapolated to the entire Guinean population.

#### 4.1. Socio-Demographic Aspects

In this study, Graves' disease was the most frequent etiology of hyperthyroidism (64%). A similar frequency was found by Baldé *et al.* [7] in 2007 in Conakry.

In this study, there was a clear predominance of women (93%) with a sex ratio of 0.07. Graves' disease is essentially a female disease. In the literature, a frequency of 7 to 10 women for one man is reported [7] [8]. The medium age of our patients was  $39.4 \pm 13$  years with a peak between 31 and 40 years. This confirms that Graves' disease remains a disease of the young adult. Sixty-five percent of the patients lived in the capital city, which could be explained by the choice of the urban site of the

study, but also by the inaccessibility of patients living within the country to care, especially to specialized care.

## 4.2. Clinical and Paraclinical Aspects

The more or less complete thyrotoxicosis syndrome was intense with a predominance of general, cardiovascular, and neuropsychological signs. The frequency of associated signs such as goiter, exophthalmos, and myxedema are in agreement with the data of the literature [4] [7] [8].

In our series, Graves' disease was associated with diabetes in 10%, with one case of vitiligo and one case of rheumatoid arthritis. This could suggest the possibility of a link between autoimmune pathologies. The prevalence of association with arterial hypertension (30% of cases) reflects the frequency of this pathology in the general population [9].

Several authors have reported a frequent observation of cardiothyreosis during hyperthyroidism: 6% in our series, 13% in Conakry [7] and Dakar [10], and 26.57% in Lomé [11]. This cardiothyreosis would occur more frequently in elderly subjects or with a delay in diagnosis and management. It is said to be more serious when the subject has pre-existing heart disease [7] [8].

Hormonal assessment, in particular TSHus, was performed in 91% of patients. This frequency is higher than the 46.66% found by Akoussou *et al.* [11] in Togo in 2001 and the 70% found by Baldé *et al.* [7] in 2017 in Guinea.

The immunological workup, anti-TSH receptor antibodies (TRAK) were performed in 38 patients and the result was positive in 89% of cases, *i.e.*, a mean level of 17.93 mIU/l with extremes of 1.7 to 44 IU/l. This immunological test was performed abroad for patients who had the means to do so. In Sub-Saharan Africa, the high cost of investigations, particularly immunological ones, and the inadequacy of the local technical facilities are limiting factors for their prescription. However, according to the literature, TSH receptor antibodies offer more than 90% of diagnostic sensitivity, especially in the absence of exophthalmos [8].

#### 4.3. Therapeutic and Evolutionary Aspects

The therapeutic strategy of Graves' disease is different in different institutions and continents. In developing countries, the poor accessibility and the lack of availability of thyroid exploration means increase the difficulty in establishing therapeutic indications for hyperthyroidism. Radioactive iodine treatment is rarely available in Africa, which limits treatment options to synthetic antithyroid drugs and surgery. In contrast, in the United States of America, 69% of patients receive radioactive iodine therapy as first-line treatment [8].

Carbimazole was rapidly effective in correcting the symptoms of hyperthyroidism with a favorable short-term course. However, compliance with current treatment regimens (12 to 18 months) is poor in Africa, and treatment discontinuation is frequent. The number of patients lost to follow-up is high: 27% in our study, 65% in the series reported in Togo [11], 23.7% in Senegal [4]. Moreover, the quality of therapeutic follow-up is made difficult by the absence or high cost of hormonal

monitoring.

Only 4% of our patients underwent surgery after stabilization of thyroid function. In Senegal, Dieudiou D *et al.* [8] in their study reported a frequency of thyroidectomy of 14.5%. Madieng *et al.* [12] in a series of 402 thyroidectomies in general surgery reported a frequency of 25.3% of Graves' disease. None of our patients had received radioactive iodine treatment because it was not available. Therapy for thyrotoxicosis in African countries appears mainly limited to surgery and antithyroid drugs.

#### 5. Conclusion

Graves' disease is the most common form of hyperthyroidism in Conakry with a clear predominance of women, especially young women. Its clinical manifestations are classic. Prolonged medical treatment with Carbimazole is the preferred therapeutic option. Nevertheless, compliance with this treatment is poor, treatment breaks are frequent, and the number of patients lost to follow-up is high. A comprehensive thyroid management program, including training for general practitioners and the access to medications, is needed to improve diagnosis, treatment, and patient compliance.

#### **Conflicts of Interest**

The authors declare no conflict of interest.

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