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Management of primary hypothyroidism in adults: An analysis of the results of a survey in 546 primary care physicians



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KEYWORDS

Diagnosis; Hypothyroidism; Monitoring; Primary care; Survey; Treatment

Abstract

Objective: To document current practices in the management of adult patients with hypothyroidism in the setting of primary healthcare.

Methods: We designed a web-based survey to inquire information on real-life practices regarding management of hypothyroidism by primary care physicians in the region of Madrid (Spain). Results: In total, 546 out of 3897 (14%) physicians (aged 50.9 ± 8.5 yr, 404 females) completed the survey. More than 90% of respondents requested serum thyrotropin measurement in subjects with symptoms of thyroid hypofunction, family history of thyroid disease and history of autoimmune disease. A thyroid ultrasound was requested to evaluate subclinical and overt hypothyroidism by 27.1% and 69.6% of respondents, respectively. Only 22.1% of respondents stated that they do not treat subclinical hypothyroidism with thyrotropin values less than $10\,\text{mU/l}$. Most physicians use brand-name formulations of levothyroxine and advise patients on how to take the tablets. To start treatment, the gradual replacement rate was the option chosen by most of the respondents, even in young patients. The thyrotropin target preferred by most respondents was $0.5-5.0\,\text{mU/l}$, especially in older patients. In patients with persistent symptoms, 61.4% search for the causes through complementary investigations. A longer professional practice time was not always accompanied by better adherence to guidelines and expert recommendations.

Conclusion: Our results reveal a proactive attitude in the diagnosis and of therapy by most of the respondents. However, we observed a tendency to perform unnecessary diagnostic tests and an excessive propensity to treat mild subclinical hypothyroidism.

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Abbreviations: FRD, full replacement dose; FT4, free thyroxine; IoR, interval of reference; L-T4, levothyroxine; OH, overt hypothyroidism; SH, subclinical hypothyroidism; TFT, thyroid function tests; TSH, thyrotropin; US, ultrasound.

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PALABRAS CLAVE

Diagnóstico; Hipotiroidismo; Control; Atención primaria; Encuesta; Tratamiento

Manejo del hipotiroidismo primario en adultos: análisis de los resultados de una encuesta en 546 médicos de atención primaria

Resumen

Objetivo: Documentar la práctica clínica actual en el manejo del hipotiroidismo en adultos. *Métodos*: Se diseñó una encuesta basada en la web para recabar información sobre las prácticas relativas al manejo del hipotiroidismo por médicos de Atención Primaria de Madrid.

Resultados: Quinientos cuarenta y seis médicos (edad 50,9±8,5 años, 404 mujeres) de un total de 3.897 (14%) completaron la encuesta. Más del 90% solicitaba cuantificación de tirotropina en sujetos con síntomas de hipofunción tiroidea, antecedentes familiares de enfermedad tiroidea e historia de enfermedad autoinmune. El 27,1 y el 69,6% solicitaban ecografía tiroidea para evaluar el hipotiroidismo subclínico y manifiesto, respectivamente. Solo el 22,1% declaró que no trataba el hipotiroidismo subclínico con valores de tirotropina <10 mU/l. La mayoría utilizaba preparados de marca de levotiroxina y aconsejaban a los pacientes sobre cómo tomar los comprimidos. La mayoría de los encuestados comenzaba el tratamiento con dosis bajas de levotiroxina, incluso en pacientes jóvenes. El objetivo de tirotropina preferido por la mayoría fue 0,5-5,0 mU/l, especialmente en pacientes mayores. En pacientes con síntomas persistentes, el 61,4% buscaba las causas mediante pruebas complementarias. El tiempo de práctica profesional más prolongado no siempre se acompañó de una mejor adherencia a las pautas y recomendaciones de expertos.

Conclusión: Nuestros resultados revelan una actitud proactiva de la mayoría de los encuestados en el diagnóstico y el tratamiento del hipotiroidismo. Sin embargo, observamos una tendencia a realizar pruebas diagnósticas innecesarias, así como una propensión excesiva a tratar el hipotiroidismo subclínico leve.

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Introduction

Hypothyroidism is the most common hormonal deficiency worldwide and is managed mainly by primary care physicians (PCPs).¹ In a meta-analysis of European studies, the prevalence of undiagnosed hypothyroidism was 4.9%, while the prevalence of known hypothyroidism was clearly lower (3.1%).² Another study carried out specifically in primary care in Spain³ showed that 8.8% of the general population suffered from hypothyroidism. This frequency was clearly higher in women (13.3%) than in men (4.2%) and raised as the age of the subjects studied increased.

Various international⁴⁻⁶ and national guidelines⁷ offer recommendations for the screening, diagnosis, and treatment of hypothyroidism. Despite this, many studies have found that 40–50% of patients receiving levothyroxine (L-T4) are poorly controlled due to either under- or overtreatment.⁸⁻¹¹

The proper management of this common disease by PCPs is essential to achieve an accurate diagnosis and treatment and avoid comorbidities associated with excess or defect of thyroid hormone. In recent years, several studies dealing with the adherence of patients to replacement treatment and the opinion of patients on their disease, have been reported. 12-18 Other investigations have evaluated diagnostic and therapeutic attitudes in primary hypothyroidism by endocrinologists, 16,19,20 and two studies have included endocrinologists and general practitioners in France²¹ and the United States. ²² However, we did not find any study investigating this topic in Spain. Therefore, we aimed to

document the current practices of PCPs in the management of adult patients with primary hypothyroidism in Madrid (Spain).

Methods

Scope of the study

The scope of this study was all the PCPs (n = 3897) working at the Gerencia Asistencial de Atención Primaria (GAAP) of the Community of Madrid (Spain), a public primary care health system that serves a population of 6,784,804 inhabitants.²³

Survey design

We worked out a questionnaire to be answered anonymously online. The questionnaire collected information regarding demographic and professional data of participants and their attitude in the detection, diagnosis, treatment, and monitoring of primary hypothyroidism in non-pregnant adults (Supplementary Material, Table S1).

Our survey is an original creation of the authors. The authors designed the questions inquiring about issues commonly encountered in clinical practice, following the natural order of clinical work, that is, screening, diagnosis, treatment, and monitoring. Some questions were based on previously published surveys, 16,19–22 and others were newly created. The survey was designed in order that most questions were answered by indicating yes or no. Other questions

were multiple-choice, with the possibility of choosing one or more of the answers offered. To avoid bias in multiple-choice questions, we tried to omit phrases that induce the interviewee's answer, and we included a wide range of options ordered alphabetically or randomly. The study was designed to record, but not modify, the usual clinical practice of participants.

Dissemination of the survey

GAAP divides the territory of the Community of Madrid into 7 health areas (South, North, Northwest, East, West, Centre and Southeast). Each of these areas is directed by a Healthcare Management (Dirección Asistencial, DA). All potential interviewees were contacted through the procedures established by GAAP. Thereby, the authors sent the survey information to the directors of the seven DAs in Madrid. An initial mailing was sent immediately before starting the survey (18 September 2019), and a second reminder mailing was sent on 5 November 2019. Each DA was responsible for forwarding the information to the family doctors working at the primary care centres in their area. The survey remained hosted on the web from 18 September to 31 December 2019.

Data collection

Participants' responses were collected anonymously and stored electronically in a form hosted on an open-access form creation website (https://www.google.com/forms/).

Statistical analysis

Results are expressed as $\operatorname{mean} \pm \operatorname{SD}$ for normally distributed data and as median (interquartile range, IQR) for nonparametric data. The Kolmogorov–Smirnov test tested adjustment to normal distribution. For comparisons of means between two groups of subjects, the Student t-test or the Mann–Whitney U-test were used. Categorical variables are described as absolute values and percentages. Because not every participant answered all the questions, the percentage of respondents providing a given answer was calculated individually for each question, using the number of respondents to that question as the denominator. Chisquare tests and Fisher's exact test were used to compare proportions. Differences were considered significant when P < 0.05.

Results

Surveyed physicians

This web-based survey was responded to by 556 out of 3897 PCPs. Ten registries were excluded because of a lack of information in most questions. The final analysed data set consisted of completed questionnaires from 546 physicians (404 females). The mean age of subjects responding to the survey was 50.9 ± 8.5 years, and the mean time of professional practice was 23.8 ± 8.3 years. Males were older and had a longer time of professional activity than females.

The geographical distribution of respondents was as follows: South, 11.4%; North, 12.3%; Northwest, 14.7%; East, 18.1%; West, 11.2%; Centre 12.8%; and Southeast 19.6%. Details on demographic and professional characteristics of participants are provided in Supplementary Material (Table S2).

Serum thyrotropin request

The clinical situations in which the surveyed PCPs request a serum thyrotropin (TSH) measurement as a detection test for hypothyroidism appear in Table 1. Physicians with more than 25 years of professional practice were more likely to screen in some situations, such as women older than 60 years (P < 0.05), history of autoimmune disease (P < 0.01), dyslipidemia (P < 0.05), diabetes (P < 0.01), family history of thyroid disease (P < 0.001) and symptoms (P < 0.01) (Table 1).

Diagnostic work-up

In the diagnostic evaluation of patients with biochemical evidence of hypothyroidism, most respondents requested the second quantification of thyroid function tests (TFT), i.e., TSH and free thyroxine (FT4), and thyroid antibodies and lipid profile measurement. Thyroid ultrasound (US) was requested for the diagnostic evaluation of subclinical hypothyroidism (SH) and overt hypothyroidism (OH) by 27.1% and 69.6% of the respondents, respectively (Fig. 1A).

Again, we found few differences in the respondents classified by gender, extra-healthcare activity and the number of patients under follow-up (Supplementary Material, Table S3). However, physicians with more than 25 years of practice were more likely to request a second determination of TFT, antibodies, lipid profile and US in patients with SH and lipid profile and US in patients with OH (Fig. 1B).

Treatment

Most respondents (81.3%) stated that they treat most of their patients with SH or OH. Only 2 respondents (0.4%) do not start treatment and send their patients to the specialist (Table 2). Physicians with more than 50 patients on follow-up were more proactive in treating most of their patients. However, physicians with more than 25 years of practice were less proactive than those with fewer years of experience (78.5 vs 84.5%, P < 0.05).

22.1% of respondents stated that SH with TSH values between 5 and 10 mU/l (mild SH) should not be treated. The criteria for L-T4 treatment of patients with mild SH were symptoms (57.7% of respondents), antibodies (32.0%), and goitre (22.7%). On the contrary, only 0.7% of respondents believed SH with TSH values >10 mU/l should not be treated. The criteria for treatment of these patients were symptoms (32.7%), antibodies (25.6%) and goitre (18.4%).

Most of the respondents recommended taking L-T4 tablets on an empty stomach (90.6%) and avoiding taking other medications (78.0%) (Table 3). Approximately two-thirds of physicians preferred to use a brand-name

87 (31.2)

67 (28.2)

97 (33.7)

68 (26.4)

106 (30.1)

59 (30.4)

43 (30.3)

122 (30.2)

165 (30.2)

Lipid profile request

	Table 1 Clinical situations in which surveyed primary care physicians request serum thyrotropin determination.	which surveyed	primary care physi	cians request ser	rum thyrotropin	determination.				
	Clinical situation	All	Gender	der	Non-assistance activities	e3	Time of practice (yr)		Patients in follow-up	
		(546)	Female (404)	Male (142)	No (194)	Yes (352)	<25 (258)	>25 (288)	<50 (238)	>50 (279)
	Fatigue, weight gain and constipation	532 (97.4)	395 (97.8)	137 (96.5)	188 (96.9)	344 (97.7)	245 (95.0)	287 (99.7)**	235 (98.7)	269 (96.4)
	Familial history of thyroid disease	501 (91.8)	374 (92.6)	127 (89.4)	172 (88.7)	329 (93.5)	225 (87.2)	276 (95.8)***	218 (91.6)	256 (91.8)
	History of autoimmune disease	497 (91.0)	373 (92.3)	124 (87.3)	170 (87.6)	327 (92.9)*	224 (86.8)	273 (94.8)**	211 (88.7)	258 (92.5)
2	Diabetes	388 (71.1)	300 (74.3)	88 (62.0)**	133 (68.6)	255 (72.4)	165 (64.0)	223 (77.4)**	159 (66.8)	204 (73.1)
292	Known dyslipidemia	374 (68.5)	285 (70.5)	89 (62.7)	128 (66.0)	246 (69.9)	164 (63.6)	210 (72.9)*	166 (69.7)	187 (67.0)
<u>.</u>	Women older than 60 yr	356 (65.2)	271 (67.1)	85 (59.9)	126 (64.9)	230 (65.3)	157 (60.9)	199 (69.1)*	146 (61.3)	191 (68.5)
	Men older than 60 yr	262 (48.0)	199 (49.3)	63 (44.4)	91 (49.9)	171 (48.5)	122 (47.3)	140 (48.6)	102 (42.9)	148 (53.0)*
	Health test in heahtly	169 (31.0)	131 (32.4)	38 (26.8)	61 (31.4)	108 (30.7)	79 (30.6)	90 (31.3)	79 (33.2)	80 (28.7)

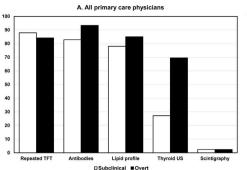
Data are the number (percentage) of affirmative answers for each of the situations in which the respondents' opinion is requested about the screening. Figures in parentheses in the headings of each column indicate the number of subjects in each group or subgroup.

Statistically significant differences (chi-square test) are highlighted in bold.

^{*} P < 0.05. * P < 0.01. *** P < 0.001.

ے ا		Ğ	Gender	Non-assistance activities	Ce	Time of practice (yr)		Patients in follow-up	
ıts with		Female	Male	o N	Yes	<25	≥ 25	<50	>50
tients with	L					i i	000		
	444/546	330/404	114/142	150/194	294/352	218/258	78 57	181/238	241/2/9
hypothyroidism. I only refer to the specialist cases with	(c.10)	():10	(60.3)	(5:77)	(c.co)	(c.+.o)	(C.07)	(78.1)	(00.4)
t of	63/546	48/404	15/147	73/194	40/352	19/258	44/288	37/738	20/279
ral but	(11.5)	(11.9)	(10.6)	(11.9)	(11.4)	(7.4)	(15.3)	(15.5)	(7.2)
ubclinical	37/546	26/404	11/142	20/194	17/352	21/258	16/288	20/238	16/279
hypothyroidism and refer to the specialist overt hypothyroidism	(6.8)	(6.4)	(7.7)	(10.3)	(4.8)	(8.1)	(5.6)	(8.4)	(8.4)
	2/546	0/404	2/142	1/194	1/352	0/258	2/288	0/238	2/279
patients with hypothyroidism, I always consult the specialist	(0.4)	<u>(</u>)	(1.4)	(0.5)	(0.3)	(0)	(0.7)	(0)	(0.7)
	446 (60.4	000,000	,	706/1406	900	730,000	266 (242)	700,007	740,742
on an empty stomach.	47.27,74 (90.6)	(93.1)	(83.6)	(89.2)	(91,4)	(87.6)	(93.4)*	(88.8)	(91.5)
kfast	(2)			(1)					
I recommend not taking other	400/513	303/382	97/131	135/181	265/332	188/248	212/265	172/222	208/267
	(78.0)	(79.3)	(74.0)	(74.6)	(79.8)	(75.8)	(80.0)	(77.5)	(77.9)
					!		!		
l use a brand-name preparation	343/530	253/392	90/138	130/187	213/343	158/253	185/277	137/230	188/272
	(64.7)	(64.5)	(65.2)	(69.5)	(62.1)	(62.5)	(66.8)	(59.6)	(69.1)*
l use a generic pharmaceutical	187/530	139/392	48/138	57/187	130/343	95/253	92/277	93/230	84/272

Data are the number of affirmative responses for each item/number of valid responses (percentage). $^{\circ}$ P < 0.05 $^{\circ}$ P < 0.01 (chi-square test).



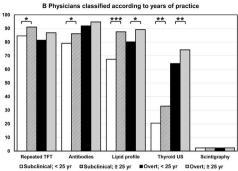


Figure 1 Affirmative responses (percentage) about the request for complementary investigations after the biochemical diagnosis of subclinical (open columns) and overt hypothyroidism (closed columns) in adult patients. (A) All surveyed physicians. (B) Physicians classified according to years of professional practice. Abscissa scale: percentage of affirmative responses. Abbreviations: TFT, thyroid function tests; US, ultrasound. *P < 0.05; **P < 0.01; ***P < 0.001.

Question	Options	Number	%
First analytical checkup after	2 weeks	8	1.5
starting therapy	1 month	148	27.1
	2 months	216	39.6
	3 months	171	31.3
	6 months	3	0.5
Request for analytical tests at	TSH	545	99.8
first analytical checkup	FT4	426	78.0
	FT3	138	25.3
	Antibodies	219	40.1
Preanalytical recommendations	You must go on an empty stomach	398	72.9
on fasting	You can have breakfast	63	11.5
	None	85	15.6
Preanalytical recommendations	You can take L-T4 tablet	177	32.4
on taking tablets	You cannot take L-T4 tablet	152	27.8
_	None	217	39.7
Periodicity of monitoring in a	Every 6 months	169	31.0
stable patient	Yearly	375	68.7
	Every 2 years	2	0.4
Attitude towards a patient	I refer to the specialist	279	48.9
with persistent symptoms	I request additional examinations to look for causes	335	61.4
	I change the trademark of L-T4	15	2.7
	I try combined treatment with L-T4 and L-T3	1	0.2
	I do a therapeutic trial by raising the dose of L-T4	65	11.9

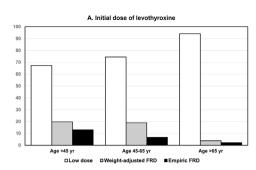
preparation. We found few significant differences in the recommendations for taking tablets and in the preferences of pharmaceutical formulations in the practitioners classified in the groups that appear in Table 3.

The most preferred option to start treating hypothyroidism was to use low doses of L-T4 with progressive increase according to TSH response, even in patients <45 years. This option was more used as the patient's age increased. Weight-adjusted or empirical full replacement dose (FRD) was used in patients under 45 years of age by 19.7% and 13.0% of the respondents, respectively. In patients aged 45 to 60, these percentages were 18.9% and 6.7%, respectively (Fig. 2A). The TSH target preferred by most respondents was 0.5–5.0 mU/l, that is, the interval of

reference (IoR). This opinion became more prevalent as the patient's age increased (Fig. 2B).

Monitoring

Table 3 shows the preferences of the respondents on monitoring and follow-up. After starting therapy, the first reevaluation of TFT was carried out between 1 and 3 months by 98.0% of the respondents. Most physicians (68.7%) perform an annual TSH measurement in stable patients. In patients with persistent symptoms, 61.4% search for the causes through complementary investigations, but 48.9% refer them to the specialist in Endocrinology.



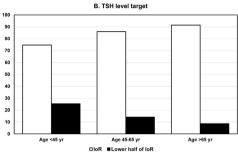


Figure 2 (A) Initial dose for starting therapy of overt hypothyroidism according to the age of patients: low dose with progressive rise according to TSH response (open columns), weight-adjusted full replacement dose (grey columns), and empiric full replacement dose (closed columns). (B) Target of serum TSH in patients classified in the same age groups: interval of reference (0.5–5.0 mU/l) (open columns), and lower half of interval of reference (0.5–2.5 mU/ml) (closed columns). Abscissa scale: percentage of affirmative responses. Abbreviations: FRD, full replacement dose; IoR, interval of reference.

Influence of professional experience and volume of patients

Table 4 shows the differences in some selected criteria for detection, diagnosis, treatment, and monitoring related to professional experience, evaluated by years of practice and number of patients under follow-up. The number of patients under follow-up was higher in physicians who decided to treat most of their hypothyroid patients, compared to those who did not (50 [30–90] vs 30 [20–60]; P < 0.01).

Time of professional practice was related to some of these criteria. Physicians who requested neck US to evaluate patients with SH or OH had significantly longer professional practice than those who did not. Furthermore, more extended time of practice was also observed in the options of choosing FRD in patients >65 years and referring patients with persistent symptoms to the specialist. On the contrary, doctors with more extended professional experience were more proactive in treating most of their patients and were more likely to select a TSH target in the lower half of the IoR in patients younger than 45 years.

Discussion

Thyroid dysfunction screening by measurement of serum TSH is a matter of international debate. ^{24–26} Our survey results show that Madrid's PCPs show a high level of awareness of hypothyroidism and agree with these general recommendations. Interestingly, some societies recommend screening in subjects over 50–60 years of age, especially in women. ^{26,27} However, only 48.0% and 65.2% of the respondents agreed with screening men and women, respectively, over 60 years of age. It is noteworthy that 71.1% of PCPs were favourable to detecting hypothyroidism in diabetic patients, as has been proposed by some authors. ^{28–30}

Most of the respondents requested a second TFT determination, antibody testing, and lipid panel in both SH and OH diagnostic evaluation. Thyroid US is not recommended routinely in hypothyroid patients in current guidelines.^{5,31} Nonetheless, it is striking that 27.1% of respondents requested this study in patients with SH and 69.6% in patients with OH. In addition, those with more than 25 years of practice are more likely to request thyroid

US. This may reflect an increased presence of US devices in primary care centres, or a greater interest of the PCPs to evaluate the presence of thyroid nodules or Hashimoto's thyroiditis in these patients.

Patients with SH and TSH values in the 5–10 mU/l range represent a subset in which treatment with thyroid hormone is controversial.⁵ Remarkably, more than 50% of respondents treat mild SH (TSH levels 5–10 mU/l) because of the presence of symptoms with the expectation that symptoms may improve. Many physicians also use the presence of antibodies to decide the treatment of SH, but guidelines only recommend using thyroid antibodies for investigating the cause of hypothyroidism and not for the decision to start treatment.⁵

In general, we observed a tendency to overtreatment of SH since only 22.1% of respondents stated that they do not treat SH with TSH of 5–10 mU/l. This agrees with a trend toward a lower threshold for treating SH reported in United Kingdom³² and Denmark.³³ However, this overtreatment might cause a significant proportion of patients treated with thyroid hormone to be in a situation of subclinical thyrotoxicosis, as has been shown by some studies.^{8–10,34,35}

Most PCPs offer appropriate advice to their patients on how to take the tablets. Brand name preparations of L-T4 was preferred by 64.7% of PCPs, in contrast to 49.9% of endocrinologists in the survey by Burch et al. 19 and to 83% of ATA members in the survey by McDermott et al. 22 Starting therapy with FRD is safe in otherwise healthy individuals. 22,36 However, the gradual rate of replacement, starting with low doses, was the option chosen by 67.3-94.2% of PCPs, according to the age of the patients. This gradual rate of replacement has been reported in half of the patients studied by Delemer et al., 21 and was the option chosen by 38.5% of endocrinologists in the survey of Burch et al. 19 Starting therapy with FRD in young patients was more common among specialists than among PCPs in the survey of McDermott et al.²² Our survey did not include any questions about the use of different L-T4 formulations, such as liquid solution or soft-gel capsules, as the only L-T4 available in Spain is in tablet form.

Contrary to the opinion of endocrinologists in previous studies, ^{19,22} a target of TSH in the lower half of IoR was an option scarcely chosen by our respondents, even in young patients (25.3% in patients <45 years). As previously shown, ²² PCPs often chose a broader TSH goal in patients of

Table 4 Relationships between professional experience (years of practice and the number of hypothyroid patients under follow-up) with some criteria for detection, diagnosis, treatment and follow-up of hypothyroidism.

		Tim	e of practice (yr)		r of hypothyroid ts under follow-up
		n	Mean \pm SD	n	Median (IQR)
Detection					
Women older than 60 yr	No	190	$\textbf{22.9} \pm \textbf{7.7}$	180	45 (30-74)
	Yes	356	$\textbf{24.2}\pm\textbf{8.6}$	337	50 (30-100)
Diagnosis					
Neck US in SH	No	398	23.1 ± 8.3	375	50 (30-90)
	Yes	148	$25.4 \pm 4.8^{**}$	142	50 (30–100)
Neck US in OH	No	166	22.3 - 8.8	157	50 (30-90)
	Yes	380	24.4 \pm 8.1*	360	50 (30-90)
Treatment					
I treat most of my patients	No	102	25.6 ± 8.6	95	30 (20-60)
· ·	Yes	444	$\textbf{23.3} \pm \textbf{8.2*}$	422	50 (30-90)**
Treatment of SH with 5< TSH< 10	No	424	23.7 ± 8.5	403	50 (30-90)
	Yes	120	23.7 ± 7.7	112	50 (30-60)
Starting dose in patients <45 yr	Low	356	$\textbf{23.8} \pm \textbf{8.7}$	337	50 (30-80)
	FRD	173	$\textbf{23.6} \pm \textbf{7.7}$	163	50 (30-100)
Starting dose in patients >65 yr	Low	493	$\textbf{23.5} \pm \textbf{8.4}$	465	50(30-90)
	FRD	31	$26.8\pm7.8^{\boldsymbol{*}}$	30	50 (25-73)
Empty stomach	No	49	19.9 ± 9.6	48	42 (30-50)
	Yes	475	$24.0 \pm 8.1^{**}$	448	50 (30-100)
Not taking other medications	No	113	$\textbf{22.8} \pm \textbf{8.3}$	109	50 (30-79)
	Yes	400	23.8 ± 8.3	380	50 (30-100)
Control and follow-up					
TSH target in patients <45 yr	0.5-5.0	408	23.0 ± 8.5	385	50 (30-90)
	0.5-2.5	138	$25.8 \pm 7.5^{***}$	132	48 (30-80)
TSH target in patients >65 yr	0.5-5.0	499	$\textbf{23.8} \pm \textbf{8.3}$	471	50 (30-90)
	0.5-2.5	47	$\textbf{23.8} \pm \textbf{8.6}$	46	40 (30-68)
I refer to the specialist patients	No	267	$\textbf{23.0} \pm \textbf{8.4}$	251	50 (30-100)
with persistent symptoms	Yes	279	$24.4\pm8.3^{*}$	256	48 (30-80)

Abbreviations: SH, subclinical hypothyroidism; OH, overt hypothyroidism; US, ultrasound; FRD, full replacement dose.

all ages, and it seems that they are not comfortable with a TSH target in the lower half of the IoR.

Rechecking TFT after starting therapy was performed between 1 and 3 months by 98.0% of respondents. It is striking that, apart from serum TSH, many respondents also request determinations not recommended by guidelines (FT4) or not helpful in monitoring (FT3 and antibodies).⁵ After achieving stable TSH values, 68.7% of respondents obtained laboratory studies yearly and 31.0% at 6-months intervals. In contrast, 34.0% and 55.5% of endocrinologists performed monitoring every 12 and 6 months, respectively.¹⁹ Although some studies have shown a transient suppression of serum TSH levels after L-T4 dosing,³⁷ only 27.8% of respondents recommended blood sampling for TSH before L-T4 ingestion.

Symptoms of hypothyroidism are not specific, and each of the symptoms generally associated with hypothyroidism may also have non-thyroid causes.³⁸ In our survey, 61.4% of respondents performed testing for other

sources of the patient's symptoms, in contrast to 84.3% of endocrinologists. ¹⁹ It is noticeable that 48.9% of family doctors refer these patients to the specialist. As recommended by guidelines ^{4,6} the use of T3-containing therapies was anecdotic.

Our data suggest that a longer professional practice time is not always accompanied by better adherence to guideline recommendations. This could be the case for the use of US in diagnosis and the preference for FRD in patients older than 65 years. Surprisingly, PCPs less prone to treat their patients and more favourable to refer to the specialist patients with persistent symptoms have more professional practice time.

We believe our results are likely to represent current practices in primary care in Madrid since the survey was answered by 14% of PCPs. This percentage is in line with that obtained in other studies on hypothyroidism. In the survey by McDermott et al.,²² the response rate was 24% of the invited PCPs. In the study by Delemer et al.,²¹ 9.5% of the PCPs agreed to participate. In the report by

^{*} P < 0.05

^{**} P < 0.01

^{***} *P* < 0.001.

Burch et al., ¹⁹ 10.9% of the members of the Endocrine Society responded to the survey. Moreover, the distribution by geographic areas of the responding PCPs from Madrid was regular, and there was no health area with less than 10% of respondents.

Furthermore, most of the respondents answered all the questions on the survey, and there were very few unanswered questions. The study was limited to Madrid; therefore, results cannot be extrapolated to other geographical regions. It is possible that PCPs who are more interested in hypothyroidism may have been more willing to respond to the survey than other physicians. In addition, results may be biased by this fact, that is, the management of hypothyroidism by such a group of dedicated doctors could differ significantly from that of the average physicians. To settle this point, a new study may be needed to eliminate any doubt about the existence of such bias. The relationship and communication of PCPs with the Endocrinology departments of Madrid's different health areas was not analysed in this study. It is possible that these relationships are different in the various areas and that, therefore, this may condition differences in the management of hypothyroidism.

In summary, the present report shows that the management of hypothyroidism by the Madrid PCPs broadly follow the recommendations of the international guidelines. However, PCPs tend to deviate from the recommendations due to the widespread use of thyroid US and thyroid antibodies, the criteria for starting therapy of SH and the definition of TSH targets according to age.

Data availability

The data are available from the corresponding author upon request.

Ethical approval

The study received favourable reports from the Research Ethics Committee of the Hospital Universitario Puerta de Hierro Majadahonda and the Central Research Committee of the GAAP (code 37/19).

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Conflict of interest

None.

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Appendix A. Supplementary data

Supplementary data associated with this article can be found, in the online version, at doi:10.1016/j.endinu.2021.04.010.

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