

# The socioeconomic burden of adult attention-deficit/ hyperactivity disorder in Spain

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## **ABSTRACT**

Introduction: The symptoms of attention-deficit/hyperactivity disorder (ADHD) in adults highly interfere with function in multiple dimensions, increasing the economic burden associated with ADHD. The aim of this study was to explore the impact of ADHD in Spanish adults and estimate the associated economic burden within the healthcare, social, economic, and legal domains.

Methods: An economic model was developed from a social perspective using a bottom-up approach, based on the scientific literature and a multidisciplinary expert group.

Results: The cost incurred per diagnosed adult patient with ADHD included an annual cost of €15,652 and a one-time cost of €7,893 (3,035 M€ and 1,531 M€ for Spain, respectively). Regarding the annual cost, 50% was attributed to costs within the economic domain, of which 53% were work-absenteeism-related. Moreover, 28% was attributed to costs within the social domain, of which 74% were substance-abuse-related. Regarding the one-time cost, 52% was attributed to costs within the healthcare domain, of which approximately 50% were hospitalization-related costs. Moreover, 42% was attributed to costs within the legal domain, of which 62% were imprisonment-related costs.

Conclusions: This is the first report on the socioeconomic burden of ADHD in Spanish adults, shedding light on the large burden that adult ADHD poses on the healthcare system and society at large, as symptoms have been shown to impact almost every aspect of life. This is particularly important for undiagnosed/untreated patients with ADHD in Spain, as appropriate treatments have shown positive results in these areas and may reduce its associated socioeconomic burden.

Keywords: adult ADHD, burden of disease, healthcare system, social perspective, substance abuse, work productivity loss

# Introduction

Attention-deficit/hyperactivity disorder (ADHD) is a neurodevelopmental disorder characterized by a persistent pattern of inattention, hyperactivity, and/or impulsivity that

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interferes with functioning or development, with symptoms present before the age of 12, in two or more settings, and not exclusively related to other mental disorders (1). The main characteristics of ADHD encompass symptoms of inattention (e.g., forgetfulness, distractibility, doubtfulness, etc.), hyperactivity (e.g., restlessness, difficulty relaxing, talking too much and too loud, etc.), and impulsivity (e.g., acting without thinking, interrupting others, impatience, etc.), which pose a substantial burden on patients and society at large (2).

Typically diagnosed in early childhood, ADHD is among the most common group of mental disorders in children and adolescents with a worldwide pooled prevalence of 3.4% (3). However, considered a lifespan disorder (2), ADHD may persist



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into adulthood (57.0%-78.2%) as patients continue to meet the criteria for ADHD as adults (4). Accordingly, the worldwide pooled prevalence of adult ADHD has been estimated at 2.8% of the total population (4).

The latest update on ADHD diagnostic criteria specified the need for clear evidence that symptoms interfere with or reduce the quality of social, academic, or occupational functioning (1,2). While children with ADHD present social and academic difficulties, impairments in adults with ADHD extend beyond these areas (1). Specifically, adults with ADHD present social and interpersonal difficulties such as unstable relationships and marital problems leading to separation or divorce: work-related problems such as unemployment and underachievement; and antisocial and criminal behavior leading to arrests, convictions, and/or imprisonment, among others (2,5). Accordingly, a recent systematic review showed that adults incur most of the economic burden associated with ADHD (89.5%) compared to that incurred by children and adolescents (10.5%) (6). In this regard, most ADHD economic burden studies have typically examined only direct costs (medical and nonmedical costs), followed by indirect costs (indirect medical costs, absenteeism, presentism), and a paucity of studies have further captured educational or justice system associated costs, hence underestimating the real economic burden (7). However, particularly for adults. the multidimensional impact of ADHD on function prompts an analysis encompassing a greater number of domains (e.g., healthcare, social, academic, occupational, etc.) to provide a comprehensive description of the economic burden of adult ADHD (2,7).

Impaired functioning associated with ADHD may be prevented or mitigated by an appropriate multimodal

treatment (8). Specifically, pharmacological treatment of ADHD (i.e., methylphenidate, lisdexamfetamine, and atomoxetine) has been reported to be both effective and well-tolerated, and has shown a positive impact on important aspects of daily functioning (8). Unfortunately, adult ADHD is often underdiagnosed or misdiagnosed due to the overlap of symptoms with other psychiatric disorders and hence undertreated (9). Moreover, the difficult transition from pediatric to adult healthcare services may compromise treatment continuity (8). These patients continue to show greater healthcare utilization and economic burden compared to those without ADHD (10). Consequently, untreated adult patients with ADHD highly contribute to the overall burden associated with ADHD.

While the socioeconomic burden of children and adolescents with ADHD in Spain has been previously studied (11), we are unaware of any previous study on the socioeconomic burden of adult ADHD in Spain (7). Given that childhoodonset ADHD may persist into adulthood (4) and enhance the socioeconomic burden of ADHD (7), this should be studied in detail across relevant domains. Therefore, the aims of the present study were to identify the most relevant impacts of ADHD in the adult population of Spain and to estimate their associated burden within the healthcare, social, economic, and legal domains.

#### Methods

The burden of adult ADHD in Spain was estimated through an economic model developed from a social perspective using a bottom-up approach and divided into four domains (Fig. 1).

# Healthcare domain

Medical consultations

Primary care consultations Specialized care consultations

Addiction therapy consultations

Hospital nurse consultations

Social work consultations

Emergency department visits

Hospitalizations

Medication

# Social domain

Substance abuse

Alcoholism problems

Smoking problems

Cocaine consumption problems

Interpersonal relationships

Divorce

Problems with intimate relationships

Problems with their children

Problems with social relationships

Personal safety

Suicide

## **Economic domain**

Mean wage reduction due to ADHD

Labour productivity losses

Work absenteeism

Work presenteeism

(i.e., reduced productivity)

Economic problems

## Legal domain

Driving vehicles

Traffic tickets

Driving license withdrawals

Traffic accidents

Problems with the legal system

Arrest

Imprisonment

**FIGURE 1** - Study domains and associated variables. ADHD = attention-deficit/hyperactivity disorder.

#### Data sources

Relevant data regarding the prevalence of adult ADHD in Spain and its impact on the healthcare, social, economic, and legal domains were obtained from the scientific literature using the PubMed search engine and selected based on their representation of Spain or any of its regions, or, otherwise, on study methodology. Thereafter, an Advisory Board of a multidisciplinary group of experts on ADHD, including a hospital psychiatric department chief, a prison health service physician, a hospital psychiatric department psychologist, a mental health service social worker, the director of the ADHD association of patients, and a former hospital manager and psychiatrist, was convened to agree on which of the presented data points were to be included in the socioeconomic model. Whenever experts considered that a data point was not representative of the Spanish adult population with ADHD and/or no alternative data point was available in the scientific literature, especially regarding healthcare resource consumption, assumptions were made.

## **Population**

Estimates of the burden of diagnosed and undiagnosed ADHD were based on prevalence data. Accordingly, to estimate the burden of diagnosed adult ADHD in Spain, a 0.5% prevalence was used (12). Moreover, to estimate the burden of undiagnosed adult ADHD in Spain, the difference between the estimated prevalence of adult ADHD in Spain (3.3%) (4) and the prevalence of diagnosed adult ADHD in Spain (0.5%) was used, yielding a prevalence of 2.8%. Applied to the adult population of Spain (≥18 years) on July 1, 2019 (13), it was estimated that, at the time, there were 1,279,861 adults with ADHD in Spain, of which 193,918 were diagnosed (15%) and 1,085,943 were undiagnosed (85%).

## Costs

The economic model included two types of costs, onetime and recurrent annual costs, given the characteristics of the first, which are timely and lack an annual frequency. Costs are of tangible and/or intangible nature, the latter of which were estimated from financial proxies (e.g., willingness to pay for an effective alcoholism therapy). Overall, healthcare costs were estimated by multiplying the number of annual resources consumed by their unit costs, while the remaining costs were estimated by further applying prevalence reference data (e.g., multiplying the prevalence of addictions in patients with ADHD by the number of annual addiction therapy consultations and their unit cost). Estimates of the healthcare burden associated with ADHD medication were based on the cost of treating an average patient with each one of the active agents used to treat ADHD and their market share (14), considering that between 10.0% and 27.3% of adult patients with ADHD are undergoing treatment (based on estimates by the Advisory Committee and Takeda, respectively). For each active agent, the presentations with the maximum daily dose as detailed in their summary of product characteristics were selected (15-17), and among them,

the ones with the lowest cost per mg (retail price + taxes) (18). Moreover, costs associated with labor productivity were estimated using the human-capital method, by which work time lost as a result of illness is valued at the market wage. A detailed description of the variables and unit costs included in the socioeconomic model has been provided in Tables 1 and 2.

Unit costs were obtained from the median rates for health services published in the official bulletins of the Spanish Autonomous Communities, official statistics (published by the Spanish National Institute of Statistics and the Spanish Ministry of Health, Consumer Affairs, and Social Welfare), the scientific literature, and market prices. Prices were updated to 2019 Euro according to the corresponding general or medical Consumer Price Index (19), except for medication costs, which were obtained on September 10, 2020, from the medication database of the General Council of Official Associations of Pharmacists.

Results are presented as mean costs per adult patient with ADHD and total costs of ADHD within domains, broken down into one-time costs and recurrent annual costs. Moreover, to estimate the burden of ADHD on the healthcare system, the analysis was broken down into three distinct time periods to account for differences in resource consumption: diagnosis, post-diagnosis follow-up (first year), and post-diagnosis longterm follow-up (following years). While the diagnosis time period and the first year after diagnosis incurred one-time costs, the following years were associated with recurrent annual costs. Results are also presented as total costs of adult ADHD in Spain, which were obtained by multiplying the mean cost per patient by the prevalence of diagnosed and/or undiagnosed ADHD according to the bottom-up method (4,13). Given the lack of studies on the consumption of healthcare resources by undiagnosed adults with ADHD, the one-time and annual total cost of undiagnosed ADHD was obtained by excluding healthcare costs from the one-time and annual cost per adult patient diagnosed with ADHD and applying such costs to the estimated number of undiagnosed adults with ADHD.

# Sensitivity analysis

A sensitivity analysis on assumption-based data points was carried out to account for worst and best-case scenarios, relative to a reference scenario (Tabs 1 and 2).

## Results

Overall, the total cost incurred per adult patient diagnosed with ADHD included an annual cost of €15,652 and a one-time cost of €7,893. On the one hand, the annual cost was mainly attributed to costs within the economic domain (49.80%), followed by the social domain (28.35%), the healthcare domain (21.70%), and the legal domain (0.15%). On the other hand, the one-time cost was mainly attributed to costs within the healthcare domain, which accounted for those associated with diagnosis and the first year of post-diagnosis follow-up (52.08%), followed by the legal domain (42.45%), and the social domain (5.46%). Applied to the number of

TABLE 1 - Variables included in the socioeconomic model according to domains: diagnosis and post-diagnosis (first year)

	Variable	Best scenario	Reference scenario	Worst scenario	Resource/financial proxy	Unit cost (€, 2019 Euro)	Cost typology
	Diagnosis						
	Medical consultations						
	Number of specialized care consultations (27)	3.8	3.8	3.8	First specialized care consultation <sup>‡</sup>	151.8	One-time
	Post-diagnosis (first year)						
	Medical consultations						
	Number of annual primary care consultations†	6.0	7.0	8.0	Follow-up consultation with a primary care physician <sup>‡</sup>	50.1	One-time
u	Number of annual psychiatric consultations⁺	4.0	5.0	0.9	Follow-up specialized care consultation <sup>‡</sup>	88.3	One-time
iem	Number of annual addiction therapy consultations $^{\dagger \S}$	12.0	12.0	12.0	Follow-up specialized care consultation <sup>‡</sup>	88.3	One-time
op a	Number of annual hospital nurse consultations⁺¶	1.0	1.0	1.0	Hospital nurse consultation <sup>‡</sup>	42.0	One-time
csre	Number of annual social work consultations⁺	1.0	1.0	1.0	Social work consultation <sup>‡</sup>	36.3	One-time
alth	Medication						
θH	Maximum daily adult methylphenidate dose (mg) (16)	80.0	80.0	80.0	Cost per mg of methylphenidate (retail price + taxes) (16,18)	0.03403	One-time
	Maximum daily adult atomoxetine dose (mg) (17)	100.0	100.0	100.0	Cost per mg of atomoxetine (retail price + taxes) (17,18)	0.02603	One-time
	Maximum daily adult lisdexamfetamine dose (mg) (15)	70.0	70.0	70.0	Cost per mg of lisdexamfetamine (retail price + taxes) (15,18)	0.05553	One-time
	ED visits and hospitalizations						
	Patients visiting the ED in the last 6 months $^{*}(28)$	30.2%	30.2%	30.2%	Hospital ED visit*	182.2	One-time
	Patients hospitalized in the last 6 months# (28)	26.0%	26.0%	26.0%	Hospitalization (29)	3,509.7	One-time

All variables refer specifically to patients with ADHD.

ADHD = attention-deficit/hyperactivity disorder; ED = emergency department.

Advisory board.

Median rates for health services published in the official bulletins of the Spanish Autonomous Communities.

Applies to the percentage of ADHD patients with addiction.

Applies to the percentage of patients undergoing treatment with stimulant medication who present with high blood pressure.

Two annual consumptions were assumed for this resource.

 TABLE 2 - Variables included in the socioeconomic model according to domains: post-diagnosis (following years)

Medical consultations         4.0         5.0         6.0         Follow-up consultation with a primary care consultation         8.3         Recurrent           Mumber of amnual posthator consultations*         4.0         5.0         6.0         Follow-up specialized care consultation*         88.3         Recurrent           Mumber of amnual posthator consultations*         1.2         1.2         1.2         Follow-up specialized care consultation*         88.3         Recurrent           Maximum daily adult action wether dose (mg) (13)         10.0 <th></th> <th>Variable</th> <th>Best scenario</th> <th>Reference scenario</th> <th>Worst</th> <th>Resource/financial proxy</th> <th>Unit cost (€, 2019 Euro)</th> <th>Cost typology</th>		Variable	Best scenario	Reference scenario	Worst	Resource/financial proxy	Unit cost (€, 2019 Euro)	Cost typology
Number of annual primary care consultations¹  Number of annual addiction therapy consultations²  Number of annual addiction therapy consultations²  Number of annual addiction therapy consultations²  Navinum adily adult inselessment dose (mg) [15]  Number of annual addiction therapy consultations²  Naxinum adily adult inselessment dose (mg) [15]  Number of annual adult and the text of a solution and the solution and solut		Medical consultations						
Number of annual psychiatric consultations³ (a. b.		Number of annual primary care consultations†	4.0	5.0	0.9	Follow-up consultation with a primary care physician*	50.1	Recurrent
Medication         12.0         12.0         12.0         12.0         12.0         12.0         Follow-up specialized care consultation¹         88.3           Medication         Medication         Medication         12.0 <th></th> <td></td> <td>4.0</td> <td>5.0</td> <td>0.9</td> <td>Follow-up specialized care consultation*</td> <td>88.3</td> <td>Recurrent</td>			4.0	5.0	0.9	Follow-up specialized care consultation*	88.3	Recurrent
Meximum daily adult methyliphenidate dose (mg) (15)         8.0.0         8.0.0         8.0.0         Cost per mg of methylphenidate (retail price + taxes) (15,18)         0.03403           Maximum daily adult methyliphenidate dose (mg) (15)         100.0         100.0         100.0         100.0         100.0         0.03 teer mg of isdexamfetamine (retail price + taxes) (15,18)         0.03533           ED visits and hospitalizations         ED visits and hospitalizations         26.0%         26.0%         100.0	nisı		12.0	12.0	12.0	Follow-up specialized care consultation <sup>‡</sup>	88.3	Recurrent
Maximum daily adult methylphenidate dose (mg) (15) 80.0 80.0 80.0 Cost per mg of methylphenidate (retail price + taxes) (15,18) 0.03403 Maximum daily adult methylphenidate dose (mg) (17) 100.0 100.0 100.0 Cost per mg of atomoxetine (retail price + taxes) (15,18) 0.03533 Maximum daily adult atomoxetine dose (mg) (15) 70.0 70.0 70.0 70.0 Cost per mg of isodexamfetamine (retail price + taxes) (15,18) 0.03533 Maximum daily adult isodexamfetamine dose (mg) (15) 70.0 70.0 70.0 70.0 Cost per mg of isodexamfetamine (retail price + taxes) (15,18) 0.03533 Patients visiting the ED in the last 6 months <sup>2</sup> (28) 26.0% 26.0% Hospitalization (29) 40.05533 Patients biophialized in the last 6 months <sup>2</sup> (28) 26.0% 26.0% Hospitalization (29) 40.00564 Patients who were ever alcoholics (28) 19.6% 19.6% Annual WTP for a 100% effective alcoholism treatment* (4,800.0 Patients who sere ever alcoholics (28) 48.3% 48.3% Annual cost for treating a patient with COPD*(31) 2,200.0 Patients who sere divorced (32) 8.9% 20.8% Annual cost for treating a patient with COPD*(31) 2,200.0 Patients who are divorced (32) 8.9% 8.9% Cost of a contentious divorce for the patient* (33) 1.456.2 Patients who have problems with their children (35) 15.0% 15.0% Cost of a contentious divorce for the patient* (34) 2.23.0 Patients with social relationship problems (38) 63.2% 63.2% Cost of six psychotherapy sessions (family therapy)*** 5.28 Patients with social relationship problems (38) 63.2% 63.2% Cost of six psychotherapy sessions (family therapy)*** 5.98 Patients with social relationship problems (38) 63.2% 63.2% 63.2% Cost of six psychotherapy sessions (family therapy)*** 5.98 Patients with social relationship problems (38) 63.2% 63.2% 63.2% Cost of six psychotherapy sessions (family therapy)*** 5.98 Patients with social relationship problems (38) 63.2%	шор							
Maximum daily adult atomoxetine dose (mg) (17)         100.0         100.0         100.0         Cost per mg of atomoxetine (retail price + taxes) (17.18)         0.05633           Raximum daily adult lisdexamfetamine dose (mg) (18)         70.0         70.0         70.0         70.0         6.0         70.0 <th>are</th> <td></td> <td>80.0</td> <td>80.0</td> <td>80.0</td> <td>Cost per mg of methylphenidate (retail price + taxes) (16,18)</td> <td>0.03403</td> <td>Recurrent</td>	are		80.0	80.0	80.0	Cost per mg of methylphenidate (retail price + taxes) (16,18)	0.03403	Recurrent
Expuisits and hospitalizations         Pool of the position of the last of months* (28)         70.0	odtl		100.0	100.0	100.0	Cost per mg of atomoxetine (retail price + taxes) (17,18)	0.02603	Recurrent
Patients visiting the ED in the last 6 months*(128) 30.2% 30.2% 30.2% Hospitalization (29) 3,5097  Patients hospitalized in the last 6 months*(128) 26.0% 26.0% Hospitalization (29) 3,5097  Substance abuse Patients who were ever alcoholics (28) 19.6% 19.6% Annual WTP for a 100% effective alcoholism treatment* (4,800.0) 1,200.0  Patients who smoke (28) As 3% 48.3% Annual wtrp*(30) 1,200.0  Patients who smoke (28) As 3% Annual cost for treating a patient with COPD*(31) 4,245.2  Patients who consume cocaine (32) 20.8% 20.8% Annual cost for treating a patient with COPD*(31) 4,245.2  Patients who are divorced (32) 8.9% 8.9% Cost of a contentious divorce for the patient* (34) 2,420.0  Patients who have problems (35) 8.9% 8.9% Cost of a contentious divorce for the patient* (34) 2,420.0  Patients with social relationship problems (35) 8.9% 8.9% Cost of six psychotherapy sessions (family therapy)*** (35) 8.99% Patients with social relationship problems (38) 63.2% Cost of six psychotherapy sessions (individual therapy)*** (59.9% Personal safety  Personal safety  Odds ratio of suicide vs. non-ADHD (37) 2.5 5.9 14.3 Social and work-related cost of suicide verson (38) 618.1	Hea		70.0	70.0	70.0	Cost per mg of lisdexamfetamine (retail price + taxes) (15,18)	0.05553	Recurrent
Patients visiting the ED in the last 6 months (28) 26.0% 26.0% 10.0% Hospitalization (29) 3.02%		ED visits and hospitalizations						
Patients hospitalized in the last 6 months* (28)         26.0%         26.0%         Hospitalization (29)         3.509.7           Substance abuse         Substance abuse         19.6%         19.6%         19.6%         Annual WTP for a 100% effective alcoholism treatment*         4.800.0           Patients who were ever alcoholics (28)         19.6%         19.6%         19.6%         Annual WTP* (30)         1,200.0           Patients who smoke (28)         48.3%         48.3%         Annual cost for treating a patient with COPD*(31)         4,245.2           Patients who smoke (28)         20.8%         20.8%         Annual cost for treating a patient with COPD*(31)         4,245.2           Patients who smoke (28)         8.9%         Annual cost for treating a patient with COPD*(31)         4,245.2           Patients who are divorced (32)         8.9%         8.9%         Cost of a contentious divorce*         4,840.0           Patients who have problems (32)         8.9%         8.9%         Cost of 14 psychotherapy sessions (couple's therapy)*** (34)         2,20.0           Patients with intimate relationship problems (38)         59.0%         59.0%         Cost of 14 psychotherapy sessions (family therapy)*** (34)         2,20.0           Patients with social relationship problems (28)         63.2%         63.2%         Cost of is psychotherapy sessions (family therapy)*** (35)		Patients visiting the ED in the last 6 months¶(28)	30.2%	30.2%	30.2%	Hospital ED visit*	182.2	Recurrent
Substance abuse       Substance abuse         Patients who were ever alcoholics (28)       19.6%       19.6%       19.6%       4mual WTP for a 100% effective alcoholism treatment*       4,800.0         Patients who smoke (28)       48.3%       48.43.3       48.45.2       48.45.2       48.45.2       48.45.2       48.45.2       48.45.2       48.45.2       48.45.2       48.45.2       48.45.2       48.45.2       48.45.2       48.45.2       48.45.2       48.45.2       48.40.3       48.45.2       48.40.0       48.40.0       48.40.0       48.40.0       48.40.0       48.40.0       48.40.0       48.40.0       48.40.0       48.40.0       48.40.0       48.40.0		Patients hospitalized in the last 6 months" (28)	26.0%	26.0%	26.0%	Hospitalization (29)	3,509.7	Recurrent
Patients who were ever alcoholics (28) 19.6% 19.6% 19.6% Annual WTP for a 100% effective alcoholism treatment* 4,800.0 $\rightarrow$ Patients who smoke (28) $\rightarrow$ 48.3% 48.3% Annual cost for treating a patient with COPD*(31) 3,500.0 $\rightarrow$ Patients who consume cocaine (32) 20.8% 20.8% 20.8% Annual cost for readiction rehabilitation treatment*(33) 1,456.2 $\rightarrow$ Patients who are divorced (32) 8.9% 8.9% Cost of a contentious divorce for the patient*(34) 2,420.0 Patients with intimate relationship problems (35) 8.9% 8.9% Cost of a contentious divorce for the patient*(34) 2,420.0 Patients with social relationship problems (28) 63.2% 63.2% 63.2% Cost of six psychotherapy sessions (couple's therapy)** 529.8 Patients with social relationship problems (28) 63.2% 63.2% Cost of six psychotherapy sessions (individual therapy)** 529.8 Personal safety  Personal safety  Personal safety  Annual cost for a contentious divorce for the patient** (34) 2,420.0 12.36.0 12.3		Substance abuse						
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Patients who smoke (28) $+ 8.3\%$ $+ 8.3\%$ $+ 8.3\%$ $+ 8.3\%$ Annual cost for treating a patient with COPD*(31) $+ 3.60.0$ Patients who consume cocaine (32) $+ 2.0.8\%$ $+ 2.0.8\%$ $+ 2.0.8\%$ Annual cost for addiction rehabilitation treatment*(33) $+ 1.456.2$ Interpersonal relationship problems (35) $+ 2.0.8\%$ $+ 2.0.8\%$ $+ 2.0.8\%$ Annual cost for addiction rehabilitation treatment*(33) $+ 2.0.8\%$ $+ 2.0.9\%$ Patients with intimate relationship problems (35) $+ 2.0.9\%$ $+ $						→ Patients' annual WTP# (30)	1,200.0	
Patients who smoke (28) 48.3% 48.3% 48.3% Annual cost for treating a patient with COPD*(3.1) 4.245.2 Patients who consume cocaine (32) 20.8% 20.8% Annual cost for addiction rehabilitation treatment**(33) 1,456.2 Interpersonal relationships  Patients who are divorced (32) 8.9% 8.9% 8.9% Cost of a contentious divorce for the patient**(34) 2,420.0 Patients with intimate relationship problems (35) 59.0% 59.0% Cost of 14 psychotherapy sessions (couple's therapy)***(36) 15.0% Patients with social relationship problems (28) 63.2% 63.2% 63.2% Cost of six psychotherapy sessions (individual therapy)*** 529.8 Personal safety  Personal safety  Odds ratio of suicide vs. non-ADHD (37) 2.5 5.9 14.3 Social and work-related cost of suicide per person (38) 618.1						→ Caregivers′ annual WTP# (30)	3,600.0	
Patients who consume cocaine (32) 20.8% 20.8% Annual cost for addiction rehabilitation treatment**(33) 1,456.2 Interpersonal relationships  Patients who are divorced (32) 8.9% 8.9% 8.9% Cost of a contentious divorce for the patient** (34) 2,420.0 Patients with intimate relationship problems (35) 59.0% 59.0% Cost of 15.0% Cost of six psychotherapy sessions (couple's therapy)***(36) 1,236.2 Patients with social relationship problems (28) 63.2% 63.2% Cost of six psychotherapy sessions (individual therapy)*** 529.8 Patients with social relationship problems (28) 63.2% 63.2% Cost of six psychotherapy sessions (individual therapy)*** 529.8 Personal safety  Odds ratio of suicide vs. non-ADHD (37) 2.5 5.9 14.3 Social and work-related cost of suicide per person (38) 618.1		Patients who smoke (28)	48.3%	48.3%	48.3%	Annual cost for treating a patient with $COPD^\#(31)$	4,245.2	Recurrent
Interpersonal relationships8.9%8.9%8.9%Cost of a contentious divorce# $4,840.0$ Patients who are divorced (32) $\Rightarrow$ 8.9% $\Rightarrow$ 8.9% $\Rightarrow$ Cost of a contentious divorce for the patient# (34) $\Rightarrow$ 7.420.0Patients who have problems with their children (35) $\Rightarrow$ 59.0% $\Rightarrow$ 59			20.8%	20.8%	20.8%	Annual cost for addiction rehabilitation treatment# (33)	1,456.2	Recurrent
Patients who are divorced (32) 8.9% 8.9% (cost of a contentious divorce $f$ and work-related cost of a contentious divorced (32) 4.840.0 9.9% (a.9.%) (cost of a contentious divorce for the patient** (34) 2.420.0 9.9% (cost of 14 psychotherapy sessions (couple's therapy)*** (3.2.% (cost of six psychotherapy sessions (family therapy)*** (5.2.% Personal safety)  Personal safety  Odds ratio of suicide vs. non-ADHD (37) 2.5 5.9 14.3 Social and work-related cost of suicide per person (38) 6.18.1	nisr							
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59.0% 59.0% 59.0% Cost of a contentious divorce for the partner# (34) 2,420.0 (12.0% 59.0% Cost of 14 psychotherapy sessions (couple's therapy)## (36) 1,236.2 (15.0% 15.0% Cost of six psychotherapy sessions (family therapy)## 529.8 (14.3 Social and work-related cost of suicide per person (38) (18.1)	Socia					$\rightarrow$ Cost of a contentious divorce for the patient" (34)	2,420.0	
59.0%       59.0%       Cost of 14 psychotherapy sessions (couple's therapy)*** (36)       1,236.2         15.0%       15.0%       Cost of six psychotherapy sessions (family therapy)***       529.8         63.2%       63.2%       Cost of six psychotherapy sessions (individual therapy)***       529.8         2.5       5.9       14.3       Social and work-related cost of suicide per person (38)       618.1						$\rightarrow$ Cost of a contentious divorce for the partner* (34)	2,420.0	
15.0% 15.0% Cost of six psychotherapy sessions (family therapy)** 529.8 63.2% 63.2% Cost of six psychotherapy sessions (individual therapy)** 529.8 2.5 5.9 14.3 Social and work-related cost of suicide per person (38) 618.1		Patients with intimate relationship problems (35)	29.0%	29.0%	29.0%	Cost of 14 psychotherapy sessions (couple's therapy)***(36)	1,236.2	Recurrent
63.2% 63.2% Cost of six psychotherapy sessions (individual therapy)** 529.8  2.5 5.9 14.3 Social and work-related cost of suicide per person (38) 618.1		Patients who have problems with their children (35)	15.0%	15.0%	15.0%	Cost of six psychotherapy sessions (family therapy)**	529.8	Recurrent
icide vs. non-ADHD (37) 2.5 5.9 14.3 Social and work-related cost of suicide per person (38) 618.1		Patients with social relationship problems (28)	63.2%	63.2%	63.2%	Cost of six psychotherapy sessions (individual therapy)***	529.8	Recurrent
2.5 5.9 14.3 Social and work-related cost of suicide per person (38) 618.1		Personal safety						
		Odds ratio of suicide vs. non-ADHD (37)	2.5	5.9	14.3	Social and work-related cost of suicide per person (38)	618.1	One-time

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	Variable	Best scenario	Reference scenario	Worst scenario	Resource/financial proxy	Unit cost (€, 2019 Euro)	Unit cost Cost (€, 2019 typology Euro)
	Mean wage reduction compared with non-ADHD** (39)	20.0%	20.0%	20.0%	Annual earnings per employee in Spain (40)	24,197.6	Recurrent
ain	Labor productivity losses						
mor	Absenteeism as annual days on work leave⁺⁺ (41)	33.6	33.6	33.6	Earnings per day of work (42)	123.8	Recurrent
o oin	Percentage of patients with work presenteeism <sup>++</sup> (28)	45.8%	45.8%	45.8%	Annual cost of presenteeism <sup>‡‡</sup> (43)	1,950.5	Recurrent
ouo	→ Daily effective work minutes lost (43)	15.0	30.0	45.0			
DOE	<b>Economic problems</b>						
	Patients with economic problems (35)	49.0%	49.0%	49.0%	Interest cost linked to personal loans (9% interest rate)* $(44)$	545.9	Recurrent
	Driving vehicles						
	Patients with >1 traffic ticket in the last 12 months (27)	17.6%	17.6%	17.6%	Cost of a speeding ticket (45)	132.0	Recurrent
u	Mean number of driving license withdrawals (46)	1.1	1.1	1.1	Cost of a driving license withdrawal	998.5	One-time
iism					ightarrow Very serious speeding ticket + 6-point withdrawal (47)	0.009	
op I					→ Driving license recovery course (48)	398.5	
ь <b>3</b> э.	Patients involved in traffic accidents while driving (27)	18.5%	18.5%	18.5%	Annual insurance cost increase for traffic accidents# (49,50)	110.2	One-time
1	Problems with the legal system						
	Patients who have ever been arrested	37.0% (51)	42.0%*	47.0% (52)	47.0% (52) Cost of legal counsel for criminal detainee assistance# (53)	350.0	One-time
	Patients who have ever been imprisoned	3.5% (41)	9.2%*	15.0% (52)	15.0% (52) Cost per day per inmate (54)	61.8	One-time

All variables refer specifically to patients with ADHD.

ADHD = attention-deficit/hyperactivity disorder; COPD = chronic obstructive pulmonary disease; ED = emergency department; WTP = willingness to pay.

\*Average of best- and worst-case scenarios.

\*Advisory board.

\*Median rates for health services published in the official bulletins of the Spanish Autonomous Communities.

\*Applies to the percentage of ADHD patients with addiction.

\*Two annual consumptions were assumed for this resource.

\*Financial proxy.

\*Applies to the percentage of patients with ADHD who are currently employed.

\*\*The percentage of patients with ADHD who are currently employed.

\*\*The percentage of patients with ADHD who are currently employed.

\*\*The percentage of patients with ADHD who are currently employed.

TABLE 3 - Average cost per patient (€ 2019), total cost (M€ 2019), and relative cost (%) according to domains and type of cost

Domains/type of cost	Cost per patient (€, 2019 Euro)	Total cost <sup>†</sup> (M€, 2019 Euro)	Relative cost (%)
Healthcare burden			
Recurrent annual cost	3,395.83	658.51	21.70
One-time cost <sup>‡</sup>	4,110.88	797.17	52.08
Social burden			
Recurrent annual cost	4,437.80	860.57	28.35
One-time cost	430.99	83.58	5.46
Economic burden			
Recurrent annual cost	7,794.79	1,511.55	49.80
One-time cost	0.00	0.00	0.00
Legal burden			
Recurrent annual cost	23.23	4.51	0.15
One-time cost	3,350.84	649.79	42.45
Total cost			
Recurrent annual cost	15,651.66	3,035.14	100
One-time cost	7,892.71	1,530.54	100

ADHD = attention-deficit/hyperactivity disorder; M€ = million Euros.

adult patients diagnosed with ADHD in Spain, the estimated costs would yield a total annual cost of €3,035 million and a one-time cost of €1,531 million (Tab. 3).

Moreover, undiagnosed patients with ADHD were estimated to incur at least a total annual cost of €13,309 million and a one-time cost of €4,107 million.

# Healthcare burden

The estimated cost per adult patient with ADHD to establish a diagnosis amounted to €577, increased up to €3,534 in the first year following diagnosis, and slightly decreased to €3,396 per year during the following years. Costs associated with hospitalizations accounted for approximately 50% of the burden on the healthcare system for both the first and the following post-diagnosis years (Fig. 2A).

#### Social burden

The estimated cost per adult patient with ADHD within the social domain amounted to an annual cost of €4,438 and a one-time cost of €431. Costs associated with substance abuse accounted for 74.23% of the annual social burden, specifically those associated with smoking (46.20%) and alcoholism (21.20%). In addition, problems with intimate relationships accounted for 16.44% of the social burden. Moreover, the cost of a divorce at any given time in the patients' life further contributed to the overall social burden (Fig. 2B).

# Economic burden

The estimated cost per adult patient with ADHD within the economic domain amounted to an annual cost of €7,795, of which 53.35% were costs associated with work absenteeism, followed by wage reductions with respect to those without ADHD (31.76%) and work presenteeism (11.46%). One-time costs were not identified within this domain (Fig. 2C).

# Legal burden

The estimated cost per adult patient with ADHD within the legal domain amounted to an annual cost of €23 associated with traffic tickets. Most importantly, one-time costs amounted to €3,351, of which 62.23% were associated with imprisonment and 32.78% with driving license withdrawals at any given time in the patients' life (Fig. 2D).

## Sensitivity analysis

Overall, the results of the sensitivity analysis showed that recurrent annual costs associated with diagnosed adults with ADHD could vary up to 6%, while one-time costs could vary up to 21% with respect to the reference scenario. More specifically, one-time costs associated with the legal domain could vary up to 39% with respect to the reference scenario (Tab. 4). This large variation within the legal system was primarily associated with variations in the percentage of patients with ADHD who have ever been imprisoned (3.5%)

<sup>&</sup>lt;sup>†</sup>Cost per patient applied to the prevalence of diagnosed adult patients with ADHD.

<sup>&</sup>lt;sup>‡</sup>Includes costs associated with the diagnosis and the first-year post-diagnosis.

# (a) Healthcare burden

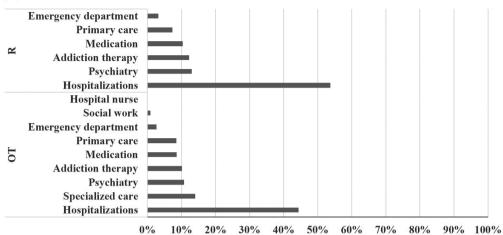
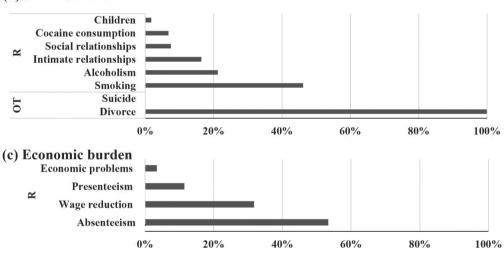
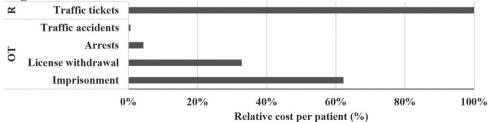


FIGURE 2 - Relative costs per patient (%) with respect to total one-time (OT) or recurrent (R) total cost within the healthcare (A), social (B), economic (C), and legal (D) domains.

# (b) Social burden







to 15%). Moreover, the mean cost per day of imprisonment depends on the number of years that the patient spends in prison, which could further increase variability.

This is the first report on the socioeconomic burden of adult ADHD in Spain (7). The results highlight a large socioeconomic impact, which was estimated to incur an annual cost of €15,652 and a one-time cost of €7,893 per adult

patient diagnosed with ADHD (€3,035 million and €1,531 million for Spain, respectively). Given the large rates of adult ADHD underdiagnosis, estimated at 2.8% (4), the socioeconomic impact is expected to be much higher (at least €13,309 million and €4,107 million, respectively). These results are not easily comparable to previous studies on the economic burden of ADHD. A recent systematic review on the global

**TABLE 4** - Best- and worst-case scenarios for the average cost per patient (€ 2019) and total cost (M€ 2019) according to domains and type of cost

Domains/type of cost	•	Cost per patient (€, 2019 Euro)		Total cost <sup>†</sup> (M€, 2019 Euro)	
	Best-case Scenario	Worst-case Scenario	Best-case Scenario	Worst-case Scenario	
Healthcare burden					
Recurrent annual cost	3,093.32	3,698.35	599.85	717.18	
One-time cost <sup>‡</sup>	3,806.94	4,415.68	738.24	856.28	
Social burden					
Recurrent annual cost	4,437.80	4,437.80	860.57	860.57	
One-time cost	430.81	431.62	83.54	83.70	
Economic burden					
Recurrent annual cost	7,147.29	8,442.29	1,385.99	1,637.12	
One-time cost	0.00	0.00	0.00	0.00	
Legal burden					
Recurrent annual cost	23.23	23.23	4.51	4.51	
One-time cost	2,033.50	4,668.18	394.33	905.25	
Total cost					
Recurrent annual cost	14,701.64	16,601.68	2,850.92	3,219.37	
One-time cost	6,271.25	9,515.48	1,216.11	1,845.23	

ADHD = attention-deficit/hyperactivity disorder; M€ = million Euros.

economic burden of ADHD reported a large variability across countries possibly associated with differences in healthcare systems, resource utilization, cost components, populations, and data sources (7). Therefore, any comparisons with previous studies should be interpreted with caution.

Overall, the estimated total annual cost per adult patient diagnosed with ADHD in Spain was above previous reports ranging between \$US 831 (€536) to \$US 20,538 (€13,247) per person (7,20). Moreover, the present study included one-time costs that increased the overall burden of adult ADHD in Spain, providing a more holistic estimate of costs. However, while previous studies on the economic burden of ADHD have typically focused on only one aspect of cost (e.g., healthcare costs) (7), the present study examined the burden of adult ADHD from a multidimensional perspective, estimating costs incurred within the healthcare, social, economic, and legal domains. Particularly important for adults with ADHD, this multidimensional analysis provides a more comprehensive description of the burden and may prevent the underestimation of costs (2,7). However, it should be noted that overestimation of costs was also prevented by taking a conservative approach within the analysis (e.g., costs that could possibly be incurred by undiagnosed adults with ADHD were not quantified, such as healthcare costs derived from traffic accidents, fights, or drug consumption).

The results of previous studies that considered costs within four major domains (direct costs, indirect costs,

education system costs, and justice system costs) closely resembled those reported in the present study. Accordingly, an annual cost per person of \$US 20,538.95, of which 72% was attributed to work-related productivity costs, has been previously reported (7). Moreover, Sciberras et al (21) reported an annual cost per person of \$US 15,664 of which 81% was attributed to work-related productivity costs. Though the latter estimated costs across the lifespan, the majority of costs were associated with adulthood (21). As the results of these studies, the estimated annual cost per patient in the present study was mainly attributed to workrelated productivity costs within the economic domain (i.e., costs associated with mean wage reductions compared to non-ADHD, absenteeism, and presenteeism). However, only 45% of published studies on the economic burden of ADHD have captured indirect costs (i.e., work-related productivity) (7).

On the other hand, healthcare costs have been widely studied within economic burden studies (7). In this domain, the present study estimated an annual cost of €3,395.83 (21% of the total annual costs) and a one-time cost of €4,110.88 (52% of the total one-time costs) per patient diagnosed with ADHD. The latter included all medical consultations leading to diagnosis and those of the first-year post-diagnosis follow-up, contributing to the large one-time healthcare costs. However, these costs may be compensated with an appropriate treatment, which may have an impact on costs within

<sup>&</sup>lt;sup>†</sup>Cost per patient applied to the prevalence of diagnosed adult patients with ADHD.

<sup>†</sup>Includes costs associated with the diagnosis and the first-year post-diagnosis.

other domains. Moreover, medication was estimated to be only 10% of the annual healthcare costs. This is particularly important for undiagnosed (i.e., untreated) adults, which accounts for 85% of adult patients with ADHD in Spain (4,13). In this regard, previous studies have shown that appropriate medication may reduce the risk of substance abuse (22,23), criminality (24), traffic accidents (25), and suicide attempts (26) in patients with ADHD, which may positively impact work-related productivity and other economic and/or legal issues.

However, the results of this study should be interpreted with caution given its limitations. First of all, the estimated burden of adult ADHD applies specifically to Spain and is therefore not representative of other countries. Within the healthcare domain, the large variability reported for the management of ADHD across regions of Spain may infer a large variability of costs as well. However, the present study did not include an interregion analysis of costs, which should be considered in future studies. Notwithstanding, a reasonable range for the consumption of healthcare resources was included in the sensitivity analysis to account for this variability. Moreover, healthcare costs incurred were mainly associated with ADHD follow-up and treatment and as such were only applied to adult patients with an ADHD diagnosis (0.5%) (12) for the estimation of total costs. Therefore, total costs incurred by adults with ADHD who are yet to be diagnosed and treated were underestimated, as the healthcare domain was not accounted for in these patients. Accordingly, future studies are encouraged to estimate the burden of adult ADHD before and after diagnosis, as it was not possible in the present study given the lack of scientific literature on the matter. Moreover, future studies should consider ADHD as a lifespan disorder in which any action taken during initial stages may drastically affect the course of the disorder and its burden (8).

## **Conclusions**

The present study highlights the large burden that adult ADHD poses on the healthcare system and society at large. In contrast to childhood ADHD, symptoms in adulthood result in very serious consequences for the patients and society at large. The impact of ADHD encompasses almost every aspect of adult life, including social (e.g., substance abuse), work-related (e.g., productivity loss), and legal aspects (e.g., criminality). This is particularly important for undiagnosed patients with ADHD as appropriate treatments have shown positive results in these areas and may therefore reduce its associated socioeconomic burden. Accordingly, strategies that promote an early diagnosis and treatment should be implemented. The results of the present study can be used to inform policy and practice to help reduce the large burden of adult ADHD.

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Conflicts of interest: FG declares no conflicts of interest, JAA, JB. MC, M Morales, and CM received consulting fees from Takeda as members of the Advisory Board. M Merino, PMH, and TML declare that they are employees of Vivactis Weber, an independent company that has received fees from Takeda for carrying out this study. JB received research grants and served as consultant, advisor, or speaker within the last 3 years for: AB-Biotics, Acadia Pharmaceuticals, Alkermes, Angelini, Ambrosetti-Angelini, Biogen, Casen Recordati, D&A Pharma, Exeltis, Gilead, Indivior, GW Pharmaceuticals, Janssen-Cilag, Jazz Pharmaceuticals, Lundbeck, Otsuka, Pfizer, Roche, Sage Therapeutics, Servier, Shire, Takeda. In addition, JB received research funding from the Spanish Ministry of Economy and Competitiveness—Centro de Investigación Biomedica en Red area de Salud Mental (CIBERSAM), and Instituto de Salud Carlos III, Spanish Ministry of Health. MC declares that the ADHD Program of the Department of Psychiatry, Hospital Universitari Vall d'Hebron, received unrestricted educational and research support from Eli Lilly and Co., Janssen-Cilag, Shire/Takeda, Rovi, Psious, and Laboratorios Rubió in the past two years. Moreover, MC received travel awards from Shire for participating in psychiatric meetings and acted as a consultant for Takeda.

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**Data availability:** The datasets generated and/or analyzed during the current study are available from the corresponding author on reasonable request.

Ethics approval: Not applicable.

Consent: Not applicable.

**Author contributions:** M Merino, PMH, and TML contributed to the study design, data collection, analysis, and interpretation. JAA, JB, MC, FG, M Morales, and CM contributed as members of the Advisory Board that provided expert appraisal for the study, including the economic model design, data collection and interpretation, and a critical review of the manuscript draft for important intellectual content. TML developed the original draft of the manuscript. M Merino and PMH critically revised the original draft. All authors approved the final submitted version of the manuscript.

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