Key Words: psychosocial impairments, adult ADHD, social impairment, work impairment, DSM-V, unmet medication coverage

Unmet Medication Coverage Needs among Adults with Attention Deficit/Hyperactivity Disorder (ADHD)

By Thomas E. Brown, Emuella Flood, Phillip Sarocco, Norman Atkins, Alexandra Khachatryan

BACKGROUND

- Attention-deficit/hyperactivity disorder (ADHD) is a common neurobehavioral condition in childhood that often persists into adulthood.¹ Psychostimulant medication has demonstrated efficacy for managing ADHD symptoms in adults^{2,3}
- The development of formulations with varying durations of effect has greatly expanded the available treatment options for individuals with ADHD. Management of ADHD symptoms in adults may involve treatment with either short-acting (SA) medication, long-acting (LA) medication, or an adjunctive LA + SA (AU) medication regimen. These different formulation dosing regimens provide differing durations of effect and thus differing coverage of symptoms that can impair adult patients in their social, work, school, and/or family settings across the entire day (in early morning, late afternoon, and into the evening)
- Few studies, however, have investigated how LA/SA formulations and differing formulation dosing regimens relate to differing medication coverage resulting in symptom impairments in adult patients with ADHD^{5,6}

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OBJECTIVE

• The objective of this study is to assess the relationship between unmet daily symptom coverage needs and LA vs. SA vs. AU formulation dosing regimens of adults with ADHD

METHODS

- A cross-sectional online survey was conducted among adults with ADHD who had been taking prescription psychostimulant medication for treatment of ADHD symptoms
- The online survey was developed following a series of qualitative interviews with adults who have ADHD (n=31) to understand symptoms and life challenges/impairments particular to ADHD. After providing consent, the participants were interviewed by a trained interviewer who used a semistructured interview guide to elicit information regarding the patients' experiences with ADHD, focusing on identifying burdensome symptom impairments and unmet symptom treatment needs. A draft survey was then tested on a small sample of adults with ADHD (n=5) using cognitive interviewing methodology, and the content of the survey was finalized based on the feedback received

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Participant selection

- Participants were recruited in the United States through research participant panels according to the IRB- approved protocol
- Participants were eligible for inclusion in the study if all of the following criteria were present at the time of screening:

Online Survey

• Participants completed an online survey that lasted approximately 30 minutes. The survey asked questions relating to the participants'

TABLE 1

INCLUSION CRITERIA FOR ADHD PARTICIPANTS

- Patient-reported diagnosis of ADHD
- Aged 18 years or older
- Currently taking at least one prescription medication indicated for ADHD and began taking it at least 6 months prior to screening
- · Adequate written and oral fluency in English
- Willing to provide informed consent
- Access to the internet
- Able to comply with the study procedures

experiences with ADHD treatment and their everyday activities, as well as sociodemographic and clinical questions. All participants provided informed consent and were compensated for their participation

Data Analysis

- Survey respondents were stratified according to their current treatment:
 - o patients receiving short-acting (SA) medication ≤2 times a day (SA group)
 - o patients receiving long-acting (LA) medication once a day (LA group)
 - patients who augmented treatment by receiving more than one medication throughout the day, i.e., LA medication more than once a day; LA plus SA medication; or SA medication more than twice a day (AU group)
- Descriptive statistics were calculated for each survey question: mean with standard deviation (SD) for normally distributed continuous data; median with range and/or interquartile range for non-normally distributed continuous data; frequency with percentages for categorical data
- The statistical significance of differences between subgroups was assessed using independent sample t-tests or Mann-Whitney tests for comparing normally and non-normally distributed continuous data, respectively, between two groups; analysis of variance (ANOVA with post-hoc Tukey tests for pairwise comparisons) and Kruskal-Wallis tests for comparing scores between three or more groups; and chisquared tests to compare categorical data. Statistical significance was set at the 5% level (P < 0.05)

RESULTS

Participant Characteristics

- A total of 616 treated ADHD participants completed the online survey (SA: 27% [n = 166]; LA: 33% [n = 201]; AU: 40% [n = 249]). The mean (SD) age of all participants was 39.0 (12.4) years and the majority were female (70%), 452 participants (88%) were currently working, and 129 (21%) were currently attending school or taking classes (Table 2)
- The distribution of self-reported ADHD severity and the number of comorbidities were similar across the current medication groups, except that participants in the AU medication group were less likely, and those in the LA group more likely, to report having "moderate" ADHD severity (P < 0.005)

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TABLE 2

PARTICIPANT CHARACTERISTICS OVERALL AND BY MEDICATION GROUP

	$\frac{AH\ ADHD}{(N=616)}$	$\frac{\text{SHORT-ACTING}}{(N = 166)}$	$\frac{\text{LONG-ACTING}}{(N = 201)}$	$\frac{AUGMENTERS}{(N = 249)}$
Age (years)				
Mean (SD)	39.0 (12.4)	38.8 (13.2)	40.0 (13.3)	38.5 (11.1)
Range	18-74	19-70	18-74	18–67
Gender, n (%)				
Male	183 (29.7)	44 (26.5)	52 (25.9)	87 (34.9)
Female	433 (70.3)	122 (73.5)	149 (74.1)	162 (65.1)
Racial background, n (%)				
White	551 (89.4)	154 (92.8)	183 (91.0)	214 (85.9)
Other	65 (10.6)	12 (7.2)	18 (9.0)	35 (14.1)
Relationship status, n (%)				
Married or living with a partner	338 (54.9)	88 (53.0)	111 (55.2)	139 (55.8)
Employment status, n (%)				
Employed, full time	349 (56.7)	96 (57.8)	107 (53.2)	146 (58.6)
Employed, part time	66 (10.7)	17 (10.2)	20 (10.0)	29 (11.6)
Self-employed	37 (6.0)	13 (7.8)	7 (3.5)	17 (6.8)
Student	32 (5.2)	8 (4.8)	12 (6.0)	12 (4.8)
Stay-at-home	33 (5.4)	2 (1.2)	18 (9.0)	13 (5.2)
parent/homemaker	,	` /	,	` '
Unemployed	24 (3.9)	6 (3.6)	14 (7.0)	4 (1.6)
Retired	27 (4.4)	9 (5.4)	12 (6.0)	6 (2.4)
Disabled	48 (7.8)	15 (9.0)	11 (5.5)	22 (8.8)
Currently attending school/ taking classes, n (%)				
Yes	129 (20.9)	35 (21.1)	36 (17.9)	58 (23.3)
No	487 (79.1)	131 (78.9)	165 (82.1)	191 (76.7)
Self-reported ADHD	107 (77.1)	101 (70.7)	103 (02.1)	171 (70.7)
severity, n (%)				
Very mild	28 (4.5)	4 (2.4)	4 (2.0)	20 (8.0)*
Mild	94 (15.3)	27 (16.3)	29 (14.4)	38 (15.3)
Moderate	338 (54.9)	94 (56.6)	126 (62.7)	118 (47.4)
Severe	128 (20.8)	33 (19.9)	35 (17.4)	60 (24.1)
Very severe	28 (4.5)	8 (4.8)	7 (3.5)	13 (5.2)
Age at diagnosis (years)	20 (110)	0 (110)	, (6.5)	10 (0.2)
Mean (SD)	27.8 (14.6)	28.9 (15.5)	28.4 (15.5)	26.7 (13.3)
Range	0–69	2–62	3–69	0–60
Number of comorbidities,	0 0)	2 02	3 0)	0 00
n (%)				
None	209 (33.9)	60 (36.1)	73 (36.3)	76 (30.5)
One	159 (25.8)	44 (26.5)	42 (20.9)	73 (29.3)
Two	138 (22.4)	37 (22.3)	50 (24.9)	51 (20.5)
Three or more	110 (17.9)	25 (15.1)	36 (17.9)	49 (19.7)
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Note: *P < 0.05, **P < 0.001 group comparison.

Abbreviations: ADHD, Attention-deficit/hyperactivity disorder; SD, standard deviation.

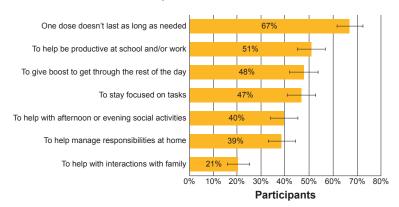
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Medication Experience Related to Duration of Effect

- Duration of effect was among the top 3 most commonly reported reasons for choosing current treatment. Only doctor's recommendation differed between the medication groups:
 - Doctor's recommendation: n = 367,60% (LA 68% [n = 136]: SA 59% [n = 98]; AU 53% [n = 133]; P < 0.01)
 - Insurance coverage: 48% [n = 298]
 - Duration of effect: 44% [n = 268]
 - \circ Treatment regimen (e.g., dosing frequency or regimen difficulty, etc.): 42% [n = 258]
- "Duration of effect" was a key factor driving the AU group's choice to take more than one medication. The most commonly reported reasons for augmenting (Figure 1) were:
 - One dose doesn't last as long as needed: 67% [n = 50]
 - Helps being productive at work/school: 51% [n = 38]
 - Gives the boost needed to get through the rest of the day: 48% [n = 36]
- Duration of effect was an important medication characteristic for many patients, influencing their likes and dislikes of their current medication and their ideal medication
 - A majority of respondents (56%) indicated that an ideal medication would last all day, with 27% of respondents reporting that their current medication does not last long enough, and 42% that they have to plan activities around their medication wearing off (Table 3). The need to plan activities was most commonly related to social (53%) and work-related activities (53%)
 - Respondents in the LA group were most likely to indicate wanting a medication that lasts all day and to report liking how long their

FIGURE 1

Reasons for Participants (N = 75) Augmenting Medication



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medication lasts. However, 34% reported having to plan activities around their medication wearing off

Medication Wearing Off

• When asked how they noticed that their medication was wearing off, the participants indicated that they experienced a number of symptoms, with 45% [n = 278] of participants indicating they experienced four or more impacts (Figure 2). A minority of participants (7% [n = 42]) could not tell when their medication was wearing off

Impact of Medication Wearing Off

• Participants commonly noted that the wearing off of their medication had negative effects on various aspects of everyday life, including their relationships (at work/school and socially), their ability to carry out responsibilities (at home or work/school), and their emotional responses and mood, with the largest effect being reported for ability to manage household and work responsibilities, school work/homework, and emotional responses or mood (Table 4)

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Burden on Everyday Life

- Participants who report being treated for their ADHD still experienced considerable burden. Participants indicated they experienced at least one impact on various aspects of life, including home life (83%, n = 510/616), social life (76%, n = 471/616), relationships (68%, n = 417/616), work (79%, n = 359/452), and school (81%, n = 104/129)
- A greater percentage of LA and SA participants versus AU participants reported issues finishing tasks (LA 70% [n = 140]; SA 72% [n = 120]; AU 54% [n = 135]; P = 0.0001), a lack of motivation (LA 59% [n = 118]; SA 49% [n = 81]; AU 41% [n = 102]; P < 0.001), and being organized in daily life (LA 56% [n = 113]; SA 50% [n = 83]; AU 41% [n = 103]; P < 0.01). LA participants were more likely to report forgetfulness (LA 61% [n = 122]; SA 43% [n = 72]; AU 47% [n = 118]; P < 0.01)

Difficult Time of Day

• Close to half of participants reported that the afternoon was the most difficult time of the day (44%, n = 269), with there being no

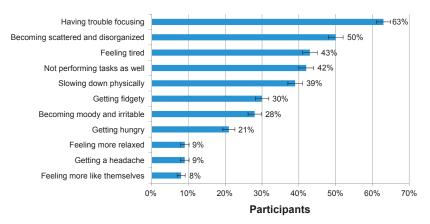
INEDICATION CHARACTERISTICS RELATED TO DURATION OF EFFECT, OVERALL AND BY INEDICATION GROUP PUBLICATION CHOICE ALLADHD SHORT-ACTING LONG-ACTING AUGMENTERS SHORT-ACTING $[N=56]$ $[N=20]$ $[N=20]$ $[N=249]$ VS.LONG-ACTING medication lasts, n (%) Dislike that current $164 (26.6) 34 (20.4) 19 (9.5) 111 (44.6)^{\dagger} < 0.001$ medication doesn't last long enough n (%)	S KELATED TO J ALL ADHD (N = 616) 277 (45.0) 164 (26.6)	SHORT-ACTING [N = 166] 27 (16.2) 34 (20.4)	LONG-ACTING (N = 201) 122 (60.7) 19 (9.5)	AUGMENTERS (N = 249) 128 (51.4) [†] 111 (44.6) [†]	P.VALUE SHORT-ACTING VS.LONG-ACTING <0.0001 <0.001	P VALUE SHORT-ACTING VS. AUGMENTERS <0.0001	P VALUE LONG-ACTING VS. AUGMENTERS <0.05 <0.001
Ideal medication would last all day, n (%)	344 (55.8)	77 (46.4)	127 (63.2)	140 (56.2)	<0.01	<0.05	NS
Have to plan activities around meds wearing off, n (%)	257 (41.7)	64 (38.6)	69 (34.3)	124 (49.8)	NS	NS	<0.01

Note: †Asked separately for each current medication. Abbreviation: NS, non-significant.

TABLE 3

FIGURE 2

IMPACTS NOTICED BY PARTICIPANTS WHEN THEIR MEDICATION WAS WEARING OFF



significant difference between the SA, LA, or AU medication groups (44% [n = 73]; 49% [n = 99]; and 39% [n = 97], respectively; Figure 2)

• Significantly more participants in the LA (47%, n = 95) than in the AU (28%, n = 70; P < 0.0001) or SA (32% [n = 53]; P < 0.01) groups reported that the evening was a particularly challenging time of the day (Figure 1). Significantly more participants in the AU (31%, n = 77) than in the LA group (21% [n = 42]; P < 0.0001) reported that midmorning was a difficult time of the day

Impairment in Home, Social, and School/Work Settings

- Despite receiving medication, ADHD impacted the participants' home lives. Difficulties were reported in terms of completing tasks (59%, n = 361), keeping the household organized (53%, n = 325), managing household chores (48%, n = 29), sleeping at night (38%, n = 233), and paying bills on time (28%, n=175). LA participants experienced greater difficulties than SA (p<0.05) and AU (P<0.05) participants
- The participants reported challenges with various aspects of their social life, including feeling awkward (38%, n = 231), tending to forget names (38%, n = 231), holding conversations (32%, n = 195), and not feeling social (31%, n = 190). The frequency of these challenges did not differ significantly between the treatment groups
- The 129 student participants indicated that they experienced a number of difficulties at school, including focusing in class (50%, n = 65) or while doing school work (46%, n = 59), prioritizing tasks (32%, n = 41), and taking notes in class (31%, n = 40). AU participants reported

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TABLE 4

Effects of Medication Wearing Off, Overall and by Medication Group

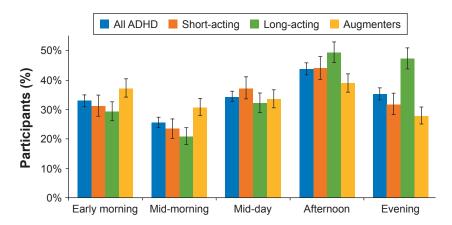
PARTICIPANTS, N (%)	ALL ADHD (N = 616)	<u>SHORT-</u> <u>ACTING</u> (N = 166)	<u>LONG-</u> <u>ACTING</u> (N = 201)	AUGMENTERS (N = 249)
Relationship with spouse/partner**	(14 - 010)	<u>[11 - 100]</u>	<u>[11 - 201]</u>	<u>(14 – 243)</u>
All negative	98 (29.4)	20 (23.3)	27 (24.3)	51 (37.5)
Not negative/some negative	235 (70.6)	66 (76.7)		85 (62.5)
N/A	283	80	90	113
Relationship with children	203	00	70	113
All negative	66 (21.9)	20 (24.1)	20 (23.3)	26 (19.7)
Not negative/some negative	235 (78.1)	63 (75.9)		106 (80.3)
N/A	315	83	115	117
Ability to be a responsible parent	313	03	113	117
All negative	66 (21.9)	22 (25.6)	19 (22.6)	25 (18.9)
Not negative/some negative	236 (78.1)	64 (74.4)		107 (81.1)
N/A	314	80	117	117
Relationship with parents	314	80	117	117
All negative	79 (17.1)	17 (147)	23 (15.9)	39 (19.4)
Not negative/some negative	383 (82.9)			162 (80.6)
N/A	154	50	56	48
Friendships	134	30	30	40
All negative	122 (20.9)	29 (17 0)	38 (19.6)	56 (23.9)
Not negative/some negative	462 (79.1)	128 (82.1)	, ,	178 (76.1)
N/A	32	10	7	178 (70.1)
Relationship with colleagues	32	10	/	13
All negative	106 (24.4)	28 (23.0)	28 (22.4)	50 (26.6)
Not negative/some negative	329 (75.6)	94 (77.0)	97 (77.6)	50 (26.6) 138 (73.4)
N/A	181	44	76	61
Ability to manage household	101	44	70	01
responsibilities				
All negative	214 (47.8)	57 (45.6)	73 (55.3)	84 (44.0)
Not negative/some negative	234 (52.5)	68 (54.4)		107 (56.0)
N/A	168	41	69	58
Ability to manage responsibilities	100	41	07	36
at work				
All negative	250 (49.0)	63 (43.8)	78 (49.4)	109 (52.4)
Not negative/some negative	260 (51.0)	81 (56.3)		99 (47.6)
N/A	106	22	43	41
Ability to do schoolwork/homework	100	44	43	41
All negative	15 (62 1)	11 (55.0)	16 (55 5)	18 (81.8)
		9 (45.0)	13 (44.8)	
Not negative/some negative N/A	26 (36.6) 545	9 (43.0) 146	172	4 (18.2) 227
	343	140	172	221
Emotional response or mood	235 (39.0)	50 (25 6)	70 (40.2)	00 (40 2)
All negative	, ,		79 (40.3)	98 (40.2)
Not negative/some negative		105 (64.4)		146 (59.8) 5
N/A	13	3	5	3

Note: *P < 0.05, **P < 0.001 group comparison.

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FIGURE 3

Most Challenging Time of Day



significantly greater difficulty taking notes than SA participants (P < 0.05)

- Similarly, the 452 participants who were employed indicated that they experienced a number of difficulties at work, including focusing on tasks (50%, n=228), being organized (42%, n=191), being productive (34%, n=155), forgetting to do simple tasks (31%, n=141), and completing tasks on time (28%, n=128)
- There is a potential for selection bias as participants were recruited through research participant panels. In addition, a majority of the participants were female, which may limit the generalizability of the results
- Participants self-reported their ADHD diagnosis and treatment in our survey without clinician confirmation

CONCLUSIONS

- Even with medication, ADHD places considerable burden on adults with the condition
- Adults with ADHD reported experiencing difficulties across the entire day, in particular during afternoon and evening hours, when medication has worn off and when work, familial, and household responsibilities are present
- Duration of effect was a key driver of treatment choice and satisfaction with treatment, with many participants indicating a desire for more adequate coverage, possibly a longer-acting medication with augmentation or a single medication that lasts all day

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• Optimizing treatment for adults with ADHD involves accounting for specific patient needs for coverage at multiple times of day and appropriate dose optimization. Differences between medications with regard to time of onset and duration of effective action should be considered in order to provide adequate coverage for various activities that may extend over substantial portions of the day, especially into the afternoon and evening, in line with individual needs of each patient *

DISCLOSURE

Drs. Brown, Flood, Sarocco, Atkins, and Khachatryan have no conflict of interest to disclose in the preparation of this manuscript.

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