RESEARCH

Open Access

Check for updates

Anti-cholinergic drug burden in patients with dementia increases after hospital admission: a multicentre cross-sectional study

Annabelle Hook^{1,2*†}, Jessica L. Randall¹⁺, Carla M. Grubb^{1,3}, Natalie Ellis^{1,4}, Jack Wellington¹, Aayushi Hemmad⁵, Agisilaos Zerdelis^{6,7}, Andrew R. D. Winnett⁸, Benjamin D. W. Geers⁹, Bethany Sykes¹⁰, Charlotte N. Auty^{9,11}, Cecilia Vinchenzo¹², Christiane E. Thorburn¹³, Daniella Asogbon¹⁴, Emily Granger^{15,16}, Heather Boagey¹⁷, Juliet Raphael¹⁸, Kajal Patel⁹, Kartik Bhargava⁵, Mary-Kate M. Dolley¹⁹, Matthew J. Maden⁹, Mehdin M. Shah²⁰, Qao M. Lee²⁰, Ratnaraj Vaidya⁵, Simran Sehdev²¹, Sneha Barai^{22,23}, Sophie Roche¹⁷, Uzair Khalid^{24,25}, David A. Codling^{26†} and Judith R. Harrison^{27,28†}

Abstract

Background: Anticholinergic medications are drugs that block cholinergic transmission, either as their primary therapeutic action or as a secondary effect. Patients with dementia may be particularly sensitive to the central effects of anticholinergic drugs. Anticholinergics also antagonise the effects of the main dementia treatment, cholinesterase inhibitors. Our study aimed to investigate anticholinergic prescribing for dementia patients in UK acute hospitals before and after admission.

Methods: We included 352 patients with dementia from 17 UK hospital sites in 2019. They were all inpatients on surgical, medical or Care of the Elderly wards. Information about each patient's medications were collected using a standardised form, and the anticholinergic drug burden of each patient was calculated with an evidence-based online calculator. Wilcoxon's rank test was used to look at the correlation between two subgroups upon admission and discharge.

Results: On admission to hospital, 37.8% of patients had an anticholinergic burden score ≥ 1 and 5.68% ≥ 3 . On discharge, 43.2% of patients with an anticholinergic burden score ≥ 1 and 9.1% ≥ 3 . The increase in scores was statistically significant (p = 0.001). Psychotropics were the most common group of anticholinergic medications prescribed at discharge. Of those patients taking cholinesterase inhibitors, 44.9% were also prescribed anticholinergic medications.

Conclusions: Our cross-sectional, multicentre study found that people with dementia are commonly prescribed anticholinergic medications, even if concurrently taking cholinesterase inhibitors, and are significantly more likely to be discharged from hospital with a higher anticholinergic burden than on admission.

Keywords: Dementia, Alzheimer disease, Cholinesterase inhibitors, Muscarinic antagonists, Antidepressive agents, Antipsychotic agents, Cognitive dysfunction, Memory disorders, Cognition, Polypharmacy

[†]Annabelle Hook and Jessica L. Randall should be considered joint first authors. [†]David A. Codling and Judith R. Harrison should be considered joint last authors.

*Correspondence: annabelle.hook@nhs.net

² Great Western Hospital, Marlborough Road, Swindon SN3 6BB, UK Full list of author information is available at the end of the article

Background

Dementia is a collective term for chronic neurodegenerative illnesses characterised by progressive cognitive deficits and functional decline [1], of which Alzheimer's dementia is the most common [2]. The neurotransmitter

© The Author(s) 2022. **Open Access** This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit http://creativecommons.org/licenses/by/4.0/. The Creative Commons Public Domain Dedication waiver (http://creativecommons.org/publicdomain/zero/1.0/) applies to the data made available in this article, unless otherwise stated in a credit line to the data.

Acetylcholine is thought to play a key role in memory, and loss of cholinergic neurons correlates with memory loss in Alzheimer's and other dementias [3].

Anticholinergic medications are medications that block cholinergic transmission, either as their primary therapeutic action or as a secondary effect [4]. Older patients are particularly prone to side-effects from these drugs including impaired memory and attention [5], delirium [6], falls [5], constipation and urinary retention [7]. The effects on cognition seem to be cumulative [8], with increased anticholinergic activity associated with a marked decline in cognition. Patients with dementia may be particularly sensitive to the central effects of anticholinergic drugs. They are more likely to develop delirium and increased cognitive decline [9], potentially due to increased bloodbrain barrier permeability [10]. Anticholinergics also antagonise the effects of the main dementia treatment, cholinesterase inhibitors [11]. Examples of such anticholinergic drugs include tricyclic antidepressants, bladder antimuscarinics and antipsychotics [12].

Despite this, older people with dementia are subject to considerable polypharmacy, including drugs with anticholinergic effects [13]. Polypharmacy as a whole [14], and anticholinergic polypharmacy specifically [15], have been correlated with increased hospitalisation and mortality.

Recent studies have attempted to quantify the anticholinergic effect of different medications. Online calculators such as medichec and acbcalc have been developed using these models. Medichec, based on Bishara et al. [16], reports an anticholinergic burden (ACB) score for each drug, calculated using the following factors: i) the magnitude of anticholinergic action, ii) the extent of bloodbrain-barrier penetration, and iii) reports of association with cognitive impairment. It also provides an aggregate ACB score based on all drugs prescribed to an individual.

NICE guidelines on the assessment and management of dementia recommend reviewing and, where possible, replacing anticholinergic drugs in patients with dementia [17]. However, it is not clear how often this occurs in clinical practice. Our study aimed to characterise anticholinergic prescribing for dementia patients in UK acute hospitals before and after admission.

Methods

Study aim

To characterise anticholinergic prescribing for dementia patients in UK acute hospitals before and after admission.

Study design

A cross-sectional, multicentre study was conducted in 2019.

Study sites

Seventeen hospital sites from across the United Kingdom were involved in the study (Additional file 1: Appendix B). To be included in the audit, each hospital site was required to have a minimum of one surgical, one medical and one Care of the Elderly ward. The audit was registered at each site and approval from each trust audit committee was gained prior to data collection.

Study population

The sample comprised 352 medical records of patients with dementia. They were all inpatients on a surgical, medical or Care of the Elderly wards. Only patients with a formal diagnosis of dementia (pre-existing or made during that admission) were included. We included only patients who were ready for discharge to ensure that we captured their discharge medications.

Data collection

DC, NE and CG designed a form to capture data on each study participant. The form included: patient demographics, medications on admission and discharge, and discharge destination (Additional file 1: Appendix C). No identifying information was recorded.

Data was collected from February to May 2019. For each participant identified, their medical notes and drug charts were reviewed. The data collection tool Enketo [18] was used to record the data. To calculate the ACB for each participant, an online calculator was used (http://medichec.com [19]). This gave an overall ACB score for each patient based on their list of medications as reported by Bishara et al. [16].

Data analysis

Data was analysed by AH and DC using the statistical software package 'R' [20]. A mean overall ACB Score was calculated for admission and discharge. As the data was non-parametric, Wilcoxon's rank test was used to ascertain whether this change in score was statistically significant.

Frequencies were calculated to show the number of medications per drug class within the dataset. These were weighted by the effect of the drug and the frequency of prescription to show the contribution to the overall ACB score.

Further analysis looked at the subset of patients taking the anti-dementia cholinesterase inhibitors. The frequency of patients prescribed both cholinesterase inhibitors and ACB medications were calculated, and their change in ACB score from admission to discharge.

Data was recorded on the input patients received from old-age psychiatrists, geriatricians and dementia

specialists. Using change in ACB score as the outcome variable, we used linear regression using the ordinary least squares method against the binary input variables of 'input from each specialty', modelled as interacting variables. A separate linear regression was performed according to the type of ward the patient was on– modelled as non-interacting binary variables, with change in ACB score as the outcome variable.

Ethics

As our study was accessing routinely-collected data to evaluate service provision through the collection of anonymised and aggregated data, we registered with local audit committees but did not apply for research ethics. All methods were performed in accordance with the relevant standards for service evaluations.

Results

Patient demographics

Three hundred fifty-two patient records were included in the audit across 17 sites. A summary of patient demographics is presented in Table 1.

Patients were distributed across the hospitals with 18.5% admitted to acute wards, 9.7% admitted to a dementia ward, 52.8% admitted to a geriatrics ward, 10.2% admitted to surgical wards and 8.5% admitted to 'other' wards. 86.1% of patients were either seen by a geriatrician, dementia specialist or old age psychiatrist during their admission.

Anticholinergic burden

Figure 1 shows the distribution of ACB scores on admission and on discharge. Mean ACB score on admission was 0.61, with a mean score on discharge of 0.77.

On admission to hospital, 37.8% of patients had an ACB ≥ 1 and 5.68% had an ACB of ≥ 3 . This reflected a total sample ACB score on admission of 215. On discharge, 43.2% of patients with an ACB score ≥ 1 and 9.1% of patients had an ACB score of ≥ 3 . This corresponded to a total sample ACB of 272. As the value counts for total ACB at admission and total ACB at discharge are non-parametric, Wilcoxon analysis could then be used to compare whether the change from total admission ACB score to total discharge ACB score was significant. This analysis showed that there was in fact a statistically significant increase from total admission ACB score to the total discharge ACB score (t=693.0, p = 0.001).

Table 2 shows the results of regression with change in ACB as the outcome variable and specialist input as the input variables. Significant values included negative association between ACB scores and input from old age psychiatry alone (p = 0.024), and from all 3 specialties (p = 0.005). Input from both old age psychiatrist and dementia specialist was positively associated with ACB scores (p = 0.013).

Results of the regression of change in ACB against ward type are shown in Table 3. All wards were significantly positively associated with change in ACB scores other than dementia wards.

On discharge, 152 patients were taking anticholinergic medications. 103 (67.8%) were taking one anticholinergic medication, 34 (22.4%) patients were taking two anticholinergic medications, 12 (7.9%) patients were taking three anticholinergic medications and 3 (2.0%) patients were taking four anticholinergic medications. In total, there were 219 instances of anticholinergic medications being prescribed.

Figure 1 shows the distributions of ACB scores at the beginning and end of the admission, as well as the change across the admission.

Of the 219 anticholinergic medications prescribed at discharge, the most common drug class was psychotropics (antidepressants, antipsychotics, mood stabilisers or benzodiazepines; 74%) (see Fig. 2). The other drug classes were antihistamines (8.2%), anticholinergic antispasmodics (7.3%), non-steroidal anti-inflammatories (5.9%), opiates (2.3%), anti-arrhythmics, anti-sickness, quinine and Parkinson medications (1.4%). The most frequently prescribed medications with anticholinergic activity were Mirtazapine (n = 41), Sertraline (n = 35), Citalopram (n = 18), Cyclizine (n = 17) and Midazolam (n = 14). When adjusted for their respective ACB scores, the drugs contributing most to the total ACB were Mirtazapine, Sertraline and Amitriptyline. The breakdown drug classes and their contribution to the ACB scores is shown in Table 4.

Seventy-six medications with anticholinergic activity were started during hospital admissions. The most frequently commenced group of medications were psychotropics (63.2%). Of the started medications, 26.3% were antidepressants, 22.4% were antipsychotics, 17.1% were antihistamines, 10.5% were antispasmodics, 14.5% were benzodiazepines, 6.6% were anti-inflammatories and 2.6% were opiates.

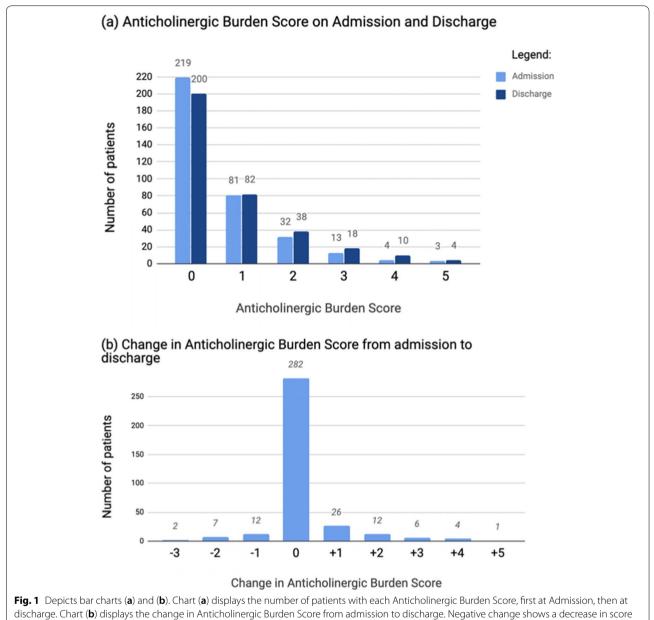
Twenty-nine anticholinergic agents were stopped. 69% of stopped anticholinergic medications were psychotropics. The most commonly stopped drug classes were antidepressants (34.5%), followed by antipsychotics (17.2%). Others included antihistamines (10.3%), anticholinergic antispasmodic (10.3%), benzodiazepines (10.3%), antiinflammatory (6.9%) and mood stabilisers, antiarrhythmic and Parkinson's medications (3.5%). Table 5 shows the changes in antipsychotic prescribing between admission and discharge.

Sixty-nine patients were taking anti-dementia cholinesterase inhibitors. Of those patients, 31 (44.9%)

Table 1 Patient demographics

Charactersitic	Ν	Percentage (%
Age (years)		
<65	6	1.7
65–74	25	7.1
75–84	137	38.9
85–94	161	45.7
>95	22	6.3
Sex		
Female	190	54
Dementia Subtype		
Alzheimer's	130	36.9
Vascular	80	22.7
Mixed	47	13.4
Lewy Body	34	9.7
Frontotemporal	5	1.4
Other (e.g. Unspecified Dementia, Dementia in Parkinson's etc)	56	15.9
Diagnoses of Dementia made during admission		
Yes	26	7.4
New diagnoses of dementia made despite delerium recorded during the ad	nission	
Yes	8	30.1
Ward		
Acute	65	18.5
Dementia	34	9.7
Geriatric	186	52.8
Surgical	36	10.2
Other (e.g. Delayed dishcarged ward, Medical rehabilitation etc)	30	8.5
Specialist Input		
Geriatrician	216	61.4
Dementia Specialist	18	5.1
Old Age Psychiatirst	4	1.1
Input from 2+ of above	65	18.5
Admitted from		
Own Home	265	75.3
Residential Home	41	11.6
Nursing Home	44	12.5
Discharged to		
Own Home	148	42.0
Residential Home	74	21.0
Nursing Home	98	27.8
Rehabilitation	24	6.8

were also taking anticholinergic medications. Eighteen patients had an ACB score of 1, eight patients scored 2, three patients scored 4 and one patient scored 5. Two patients on cholinesterase inhibitors had a decreased ACB from admission to discharge. The medications stopped during admission were antipsychotics (100%). Nine patients had an increase in score from admission to discharge. Medications added during admission were antidepressants (20%), antipsychotics (40%), benzodiazepines (20%), anti-inflammatories (10%) and antihistamines (10%). The remaining 58 patients had no change in score between admission and discharge.



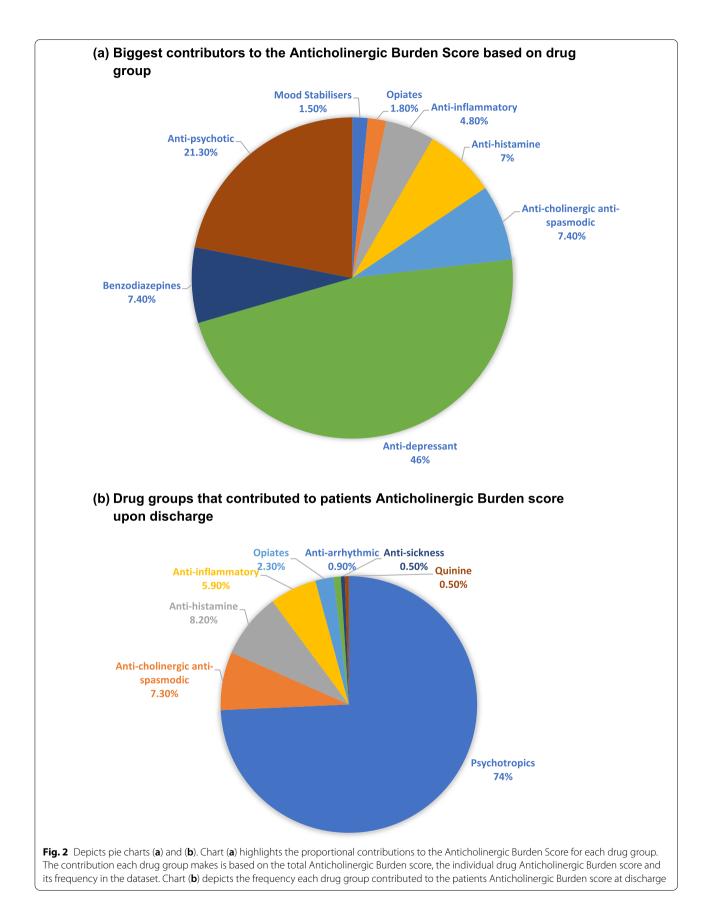
from admission to discharge. Positive change shows an increase in score from admission to discharge

Table 2 Results of regress	sion of specialist input against change
in ACB score	

Specialist input	P =	0.025	0.975
Geriatrician	0.448	-0.366	0.162
Old age Psychiatry	0.024	-1.867	-0.133
Geriatrician + Old age Psychiatry	0.060	-0.041	1.937
Dementia Specialist	0.476	-0.626	0.293
Geriatrician + Dementia Specialist	0.521	-0.743	0.377
Old Age Psychiatry + Dementia Specialist	0.013	0.311	2.594
Geriatrician + Old age Psychiatry + Dementia Specialist	0.005	-3.297	-0.601

 Table 3
 Linear regression analysis testing whether ward type influence the total ACB at discharge

Ward type	P=	0.05	0.95
Acute	0.023	0.283	3.717
Geriatric	0.038	0.104	3.52
Surgical	0.038	0.106	3.56
Dementia	0.057	-0.052	3.405
Other	0.032	0.168	3.632



Drug Class	Medicine	ACB score	N Prescribed	Percentage of total ACB for sample (%)
Antidepressants		1.4	107	55.07
	Mirtazapine	1	41	15.07
	Sertraline	1	35	12.87
	Amitriptyline	3	9	9.93
	Citalopram	1	18	6.62
	Fluoxetine	1	4	1.47
Antipsychotics		1.67	31	19
	Quetiapine	2	11	8.09
	Olanzapine	2	10	7.35
	Prochlorperazine	2	3	2.21
	Levomepromazine	2	3	2.21
	Aripiprazole	1	3	1.1
	Flupentixol Decanoate	1	1	0.37
Antihistamines	·	1.5	18	9.93
	Cyclizine	1	17	6.25
	Chlorphenamine	2	1	0.74
Antispasmodics	·	1.33	16	7.84
	Hyoscine Butyl Bromide	1	9	3.31
	Tolterodine	2	4	2.94
	Solifenacin	1	3	1.1
Benzodiazepines		1	20	7.35
·	Midazolam	1	14	5.15
	Diazepam	1	5	1.84
	Temazepam	1	1	0.37
Anti-inflammatories		1	13	4.78
	Prednisolone	1	13	4.78
Opiates		1	5	1.84
opiates	Fentanyl	1	5	1.84
Mood Stabilisers		1	4	1.47
	Carbamazepine	1	3	1.1
	Lithium	1	1	0.37
Anti-sickness	Litingini	3	1	1.1
	Hyoscine Hydrobromide	3	1	1.1
Anti-arrhythmics	, oseine rijaiosionnae	1	2	0.74
	Amiodarone	1	2	0.74
Anti-Parkinson's	, and date the	2	1	0.74
	Amantadine	2	1	0.74
Quinine	/ manaanie	2	1	0.37
Quinnic	Quinine Sulphate	1	1	0.37

Table 4 Drugs prescribed, N prescribed, ACB drug score for drug, percentage of total ACB at discharge

Discussion

Our cross-sectional, multicentre study found that people with dementia are commonly prescribed anticholinergic medications, even if concurrently taking cholinesterase inhibitors, and are significantly more likely to be discharged from hospital with a higher ACB than on admission. We found that psychotropic medications accounted for almost three quarters of anticholinergic medications taken on discharge. Psychotropics are both the most added and the most stopped medication during hospital admissions.

Whilst there is a significant literature on ACB [21, 22], particularly in dementia [23–26], our study adds a large cohort of patients from a variety of hospitals around the UK, with granular patient-level data including treatment setting and specialist input.

Table 5 Antipsychotics present at discharge (new Vs old Vs change in prescription)

Antipsychotics	Ν	Percentage (%)
Admitted and discharged on the same antipsy- chotic	13	41.9
Aripiprazole	1	
Quetiapine	4	
Olanzapine	5	
Levopromazine	2	
Prochlorperazine	0	
Flupentixol	1	
Newly prescribed an antipsychotic (no previous)	17	54.8
Aripiprazole	2	
Quetiapine	7	
Olanzapine	4	
Levopromazine	1	
Prochlorperazine	3	
Change in type of antipsychotic during admission	1	3.2
Risperidone→ Olanzapine	1	

Our findings are broadly consistent with the wider literature. Gutierrez-Valencia et al. [21] reported an increase in medications with anticholinergic effects among 200 patients discharged from an acute geriatric unit. Similarly, Wawruch et al. [22] found a statistically significant increase in elderly patients prescribed anticholinergic medications at discharge in 1636 patients admitted to long-term care facilities in Slovakia [22]. In a large retrospective study of Italian dementia patients, Reinold et al. [24] found that ACB was higher at discharge, with 46.1% of patients having a moderate to high ACB score at discharge compared to 25.4% at admission. Of prescribed medications with anticholinergic activity, they reported cardiovascular drugs were the most common, with antipsychotics second. Upon discharge, they noted that furosemide (23.4%), quetiapine (15.3%) and promazine (8.9%) were most commonly prescribed [24].

Whilst we also found an increased ACB at discharge and identified antipsychotics as the second biggest malefactor (22.4% of anticholinergic medications), in our study antidepressants were the largest contributor to ACB score (26.3%). The three most commonly prescribed medications were mirtazapine, sertraline and citalopram, and the drugs contributing most to the total ACB were mirtazapine, sertraline and amitriptyline. Psychotropic medications are frequently started in acute hospitals. In an audit of 35 acute hospitals in Ireland in 2013, Gallagher et al. [23] reported a significant increase in patients prescribed antipsychotics on discharge [23]. Indeed, 41% of those prescribed antipsychotics had doses increased [23]. Almost half of patients in our study (44.9%) who were prescribed acetylcholinesterase inhibitors as treatment for dementia were also given anticholinergic drugs. A small number of these patients (<3%) had anticholinergic medications stopped or reduced. Other studies have also reported that inappropriate co-prescription of anticholinergics and acetylcholinesterase inhibitors is common [11]. The action of acetylcholinesterase inhibitors is opposed by anticholinergic medications [27], and this may lead to a reduction in therapeutic benefit.

In our study, the majority of patients (86.1%) were seen either by a geriatrician, dementia specialist or old age psychiatrist during their admission. Our analysis shows that specialist input was not associated with increased ACB, and that input from old-age psychiatry and a combination of old age psychiatry, geriatrician and dementia specialist was associated with a reduction in ACB, although this finding must be treated with significant caution due to the small numbers (n = 38 and 14 respectively) involved. It was also notable that, of all wards, only dementia wards were not associated with increases in anticholinergic burden. This presents some evidence that specialist input may be helpful in counteracting the tendency towards increasing anticholinergic burden in inpatients. This is despite recommendations that reviews can be undertaken by any doctor or pharmacist or specialist nurse [28, 29], with the use of tools such as STOPP/ START Version 2 (Screening Tool of Older People's Prescriptions and Screening Tool to Alert to Right Treatment) [30].

Although there is consensus on many findings, it is important to note the heterogeneity in methodology. Each study uses a different method to assess ACB. For example, Wawruch et al. [22] identified anticholinergic drugs as those that scored ≥ 2 of 3 on lists published by Han et al. [8] and Rudolph et al. [31], but did not calculate an ACB score for individual patients. Gutierrez-Valencia et al. [21] used the anticholinergic risk scale (ARS) based on Rudolph et al. [31] The ARS ranks medications known to have anticholinergic effects on a scale from 0 (limited/none) to 3 (very strong). This ranking is based on the medication's dissociation constant for the cholinergic receptor and its anticholinergic adverse effects. A patient's ARS score is the sum of the ARS ranking for all of their medications [31]. However, the ARS is not based on a systematic review of the literature and instead identifies medications based on one American healthcare system. In contrast, Reinold et al. [24] used the 'Anticholinergic Cognitive Burden Scale' based on Boustani et al. [30] The developers of this scale classified medications into mild, moderate or severe anticholinergic effects based on a literature review and evaluation by an expert interdisciplinary team. These medications were then

scored from 0 to 3 (0 = no anticholinergic effect, 1 = possible anticholinergic effects, 2/3 = clinically relevant anticholinergic effects) based on their in vitro affinity for muscarinic receptors, blood-brain barrier permeability and association with development of delirium. A patient's total score is the sum of the score assigned to each medication [32]. Another possible system is the Anticholinergic Drug Scale developed by Carnahan et al. [33] This scale ranks medications based on their serum anticholinergic activity from 0 (no known activity) to 3 (marked activity). The total score for each patient is determined by the sum of the score for each medication [31]. In our study we chose to use Bishara et al.'s [16] Anticholinergic Effect on Cognition (AEC) scale, as described above. The AEC scale uses information on drug classes and medications from the British National Formulary. It also assesses penetration of the blood-brain barrier [16].

Strengths

This study has a number of strengths. We are the first to investigate anti-cholinergic prescribing in dementia patients in UK hospitals. Our study was a cross-sectional study, comprising 17 sites. Therefore, the results should be generalisable to other acute hospital settings. We have a large sample size, which minimised the chance of type II error. We used an evidence-based online calculator to ensure that we made standardised assessments of ACB score.

Limitations

Some limitations should be noted. We used a convenience sample of dementia patients who were ready for discharge. This could have introduced some selection bias. We did not enhance case ascertainment by using standardised dementia diagnostic assessments. Thus, patients that did not have a diagnosis of dementia made by their clinical team and clearly documented would have been missed. Whilst Bishara et al.'s [16] AEC scale comprehensively assesses adverse effects due to anti-cholinergic action, it does not consider the broader appropriateness of the prescription [19]. Future studies would benefit from longitudinal follow-up of patients after hospital discharge, to assess the effect of anticholinergic prescriptions on long-term outcomes.

Conclusions

Our cross-sectional, multicentre study found that older adults with dementia are commonly prescribed anticholinergic medications, even if also prescribed cholinesterase inhibitors. Anticholinergic burden increased after hospital admission. Psychotropic medications accounted for the majority of anticholinergic burden. Increased awareness of anticholinergic effects and standardised tools for medication reviews may help to address this problem. It may be that specialist input from old age psychiatry along with other specialists or specialist dementia wards can help reduce this tendency toward increased anticholinergic burden, but this is a preliminary finding that requires further validation. Further research is needed to investigate the impact of anticholinergic burden on long-term patient outcomes.

Abbreviations

ACB: Anticholinergic burden; STOPP/START Version 2: Screening Tool of Older People's Prescriptions and Screening Tool to Alert to Right Treatment Version 2; ARS: Anticholinergic risk scale; AEC scale: Anticholinergic Effect on Cognition scale.

Supplementary Information

The online version contains supplementary material available at https://doi.org/10.1186/s12877-022-03235-9.

Additional file 1: Appendix A. SPARC collaborators. Appendix B. Table 1 - Details of the 17 hospital sites in the United Kingdom involved in the study. Appendix C. Data Collection Form (questions included on software Enketo [18]).

Acknowledgements

This project was planned and executed by the authors on behalf of SPARC (Student Psychiatry Audit and Research Collaborative). SPARC Collaborators are listed in Additional file 1: Appendix A. We thank the National Student Association of Medical Research for allowing us use of the Enketo [18] platform.

Authors' contributions

Conceptualization: JR Harrison, DA Codling, C Grubb, N Ellis. Data acquisition: A Hemmad, A Zerdelis, ARD Winnett, BDW Geers, B Sykes, CN Auty, C Vinchenzo, CE Thorburn, D Asogbon, E Granger, H Boagey, J Raphael, K Patel, K Bhargava, MKM Dolley, MJ Maden, MM Shah, QM Lee, R Vaidya, S Sehdev, S Barai, S Roche, U Khalid. Data curation: A Hook, DA Codling. Formal analysis: A Hook, DA Codling. Funding acquisition: JR Harrison. Investigation/Methodology: A Hook, JL Randall, C Grubb, N Ellis, DA Codling, JR Harrison. Project administration: A Hook, JL Randall, C Grubb, N Ellis, DA Codling, JR Harrison. Manuscript writing - original draft: A Hook, JL Randall, J Wellington; Manuscript writing - review and editing: A Hook, JL Randall, DA Codling, JR Harrison; A Hemmad, A Zerdelis, ARD Winnett, BDW Geers, B Sykes, CN Auty, C Vinchenzo, CE Thorburn, D Asogbon, E Granger, H Boagey, J Raphael, K Patel, K Bhargava, MKM Dolley, MJ Maden, MM Shah, QM Lee, R Vaidya, S Sehdev, S Barai, S Roche, U Khalid. Supervision: DA Codling, JR Harrison. The manuscript was reviewed and approved by all authors.

Funding

JR Harrison was supported by a Wellcome Trust GW4 Clinical Academic Fellowship. SPARC was supported by the National Centre for Mental Health (NCMH), Wales. A Hook was a Psych Star and is currently a Psychiatry Foundation Fellow with the RCPsych.

Availability of data and materials

The datasets used and analysed during the current study are available from the corresponding author, AH, on reasonable request.

Declarations

Ethics approval and consent to participate

Our study was accessing routinely-collected data to evaluate and feedback locally regarding service provision, which was subsequently anonymised and aggregated. As such, we consider it audit and service improvement and therefore covered by the 'public task' provision (section 6(1)(e)) of GDPR legislation and did not seek

permissions from a Research Ethics Committee nor individual consent to participate. All methods were carried out in accordance with relevant guidance and protocols and necessary permissions were gained from local Trust audit committees.

Consent for publication

Not applicable.

Competing interests

The authors declare that they have no competing interests.

Author details

¹Cardiff University School of Medicine, Neuadd Meirionnydd, Cardiff CF14 4YS, UK. ²Great Western Hospital, Marlborough Road, Swindon SN3 6BB, UK. ³Betsi Cadwaladr University Health Board, Bangor LL57 2PW, UK. ⁴Withybush Hospital, Fishguard Road, Haverfordwest SA61 2PZ, UK. ⁵The Medical School, Newcastle University, Framlington Place, Newcastle upon Tyne NE2 4HH, UK. ⁶The James Cook University Hospital, Marton Road, Middlesbrough TS4 3BW, UK. ⁷University Hospital of Wales, Heath Park Way, Cardiff CF14 4XW, UK. ⁸Whipps Cross University Hospital, Whipps Cross Road, Leytonstone, London EN11 1NR, UK. 9School of Medicine, University of Manchester, Manchester M13 9PL, UK. ¹⁰University of Exeter Medical School, Heavitree Road, Exeter EX1 2LU, UK. ¹¹Queen's Medical Centre Nottingham, Clifton Boulevard, Derby Road, Nottingham NG7 2UH, UK. ¹²Lancaster Medical School, Faculty of Health and Medicine, Lancaster University, Furness Building, Lancaster LA1 4YG, UK. ¹³Bristol Medical School, University of Bristol, First Floor, 5 Tyndall Avenue, Bristol BS8 1UD, UK.¹⁴Birmingham Medical School, College of Medical and Dental Sciences, University of Birmingham, Edgbaston, Birmingham B15 2TT, UK. ¹⁵University Hospitals of Morecambe Bay NHS Foundation Trust, Burton Road, Kendal LA9 7RG, UK. ¹⁶Lancashire Teaching Hospitals NHS Foundation Trust, Sharoe Green Lane, Fulwood, Preston, Lancashire PR2 9HT, UK. ¹⁷Medical Sciences Division, University of Oxford, John Radcliffe Hospital, Headley Way, Oxford OX3 9DU, UK. ¹⁸Norwich Medical School, University of East Anglia, Norwich Research Park, Norwich, Norfolk NR4 7TJ, UK. ¹⁹Peninsula Medical School, The Faculty of Medicine and Dentistry, The John Bull Building, Plymouth Science Park, Research Way, Plymouth PL6 8BU, UK. ²⁰Bart's and The London School of Medicine and Dentistry, Queen Mary's University of London, Garrod Building, Turner Street, Whitechapel, London E1 2AD, UK.²¹Faculty of Medicine, University of Southampton, Building 85, Life Sciences Building, Highfield Campus, Southampton SO17 1BJ, UK.²²School of Clinical Medicine, University of Cambridge, Box 111, Cambridge Biomedical Campus, Cambridge CB2 0SP, UK. ²³Peterborough City Hospital, Edith Cavell Campus, Bretton Gate, Peterborough PE3 9GZ, UK. ²⁴University College London Medical School, 74 Huntley St, Bloomsbury, London WC1E 6DE, UK.²⁵St George's Hospital, Blackshaw Road, Tooting, London SW17 0QT, UK. ²⁶King's College London, Institute of Psychiatry, Psychology and Neuroscience, 16 De Crespigny Park, London SE5 8AF, UK. ²⁷Biomedical Research Building Campus for Ageing and Vitality, Newcastle University, Newcastle upon Tyne NE4 5PL, UK. ²⁸Cardiff University Brain Research Imaging Centre (CUBRIC), Cardiff University, Maindy Road, Cardiff CF24 4HQ, UK.

Received: 9 September 2021 Accepted: 9 May 2022 Published online: 06 October 2022

References

- Alzheimer Society of Canada. Mild cognitive impairment. 2019. https:// alzheimer.ca/en/Home/About-dementia/Dementias/Mild-Cognitive-Impairment. Accessed 24 Apr 2019.
- Brunnstrom H, Gustafson L, Passant U, Englund E. Prevalence of dementia subtypes: a 30-year retrospective survey of neuropathological reports. Arch Gerontol Geriatr. 2009;49(1):146–9. https://doi.org/10. 1016/j.archger.2008.06.005.
- 3. Perry E. The cholinergic hypothesis ten years on. Br Med Bull. 1986;42:63-9.
- Gerretsen P, Pollock B. Drugs with anticholinergic properties: a current perspective on use and safety. Expert Opin Drug Saf. 2011;10(5):751–65. https://doi.org/10.1517/14740338.2011.579899.
- Aizenberg D, Sigler M, Weizman A, Barak Y. Anticholinergic burden and the risk of falls among elderly psychiatric inpatients: a 4-year case-control study. Int Psychogeriatr. 2002;14(3):307–10. https://doi.org/10.1017/s1041610202008505.
- Flacker J, Cummings V, Mach J Jr, Bettin K, Kiely D, Wei J. The association of serum anticholinergic activity with delirium in elderly medical patients. Am J Geriatr Psychiatry. 1998;6(1):31–41.

- Lieberman J. Managing anticholinergic side effects. Prim Care Companion J Clin Psychiatry. 2004;6(2):20–3.
- Han L, Agostini J, Allore H. Cumulative anticholinergic exposure is associated with poor memory and executive function in older men. J Am Geriatr Soc. 2008;56(12):2203–10. https://doi.org/10.1111/j.1532-5415.2008.02009.x.
- Fox C, Richardson K, Maidment I, et al. Anticholingeric medication use and cognitive impairment in the older population: the medical research council cognitive impairment in the older population: the medical research council cognitive function and ageing study. J Am Geriatr Soc. 2011;59(8):1477–83. https://doi.org/10.1111/j.1532-5415.2011.03491.x.
- Campbell N, Boustani M, Limbil T, et al. The cognitive impact of anticholinergics: a clinical review. Clin Interv Aging. 2009;4:225–33. https://doi. org/10.2147/cia.s5358.
- Johnell K, Fastborn J. Concurrent use of anticholinergic drugs and cholinesterase inhibitors: register-based study of over 700,000 elderly patients. Drugs Aging. 2008;25(10):871–7. https://doi.org/10.2165/00002512-200825100-00006.
- 12. Gray S, Hanlon J. Anticholinergic medication use and dementia: latest evidence and clinical implications. Ther Adv Drug Saf. 2016;7(5):217–24. https://doi.org/10.1177/2042098616658399.
- Sergi G, De Rui M, Sarti S, Manzato E. Polypharmacy in the elderly: can comprehensive geriatric assessment reduce inappropriate medication use? Drugs Aging. 2011;28(7):509–18. https://doi.org/10.2165/11592010-000000000-00000.
- 14. Mueller C, Molokhia M, Perera G, et al. Polypharmacy in people with dementia: associations with adverse health outcomes. Exp Gerontol. 2018;106:240–5. https://doi.org/10.1016/j.exger.2018.02.011.
- Bishara D, Perera G, Harwood D, et al. The anticholinergic effect on cognition (AEC) scale - associations with mortality, hospitalisation and cognitive decline following dementia diagnosis. Int J Geriatr Psychiatry. 2020:1–9. https://doi.org/10.1002/gps.5330.
- Bishara D, Harwood D, Sauer J, Taylor D. Anticholinergic effect on cognition (AEC) of drugs commonly used in older people. Int J Geriatr Psychiatry. 2017;32(6):650–6. https://doi.org/10.1002/gps.4507.
- National Institute for Health and Care Excellence. Dementia: assessment, management and support for people living with dementia and their carer NICE guideline [NG97]. 2020. https://www.nice.org.uk/guidance/ ng97/chapter/Recommendations#medicines-that-may-cause-cognitiveimpairment. Accessed 29 June 2020.
- Sharif B, Lundin R, Morgan P, et al. Developing a digital data collection platform to measure the prevalence of sepsis in Wales. J Am Med Inform Assoc. 2016;23(6):1185–9. https://doi.org/10.1093/jamia/ocv208.
- South London and Maudsley NHS Foundation Trust. The anticholinergic effect on cognition tool. 2020. https://medichec.com/. Accessed 11 Aug 2020.
- 20. R Core Team. R: A language and environment for statistical computing. Vienna: R Foundation for Statistical Computing; 2022. https://www.R-project.org/.
- Gutierrez-Valencia M, Izquierdo M, Malafarina V, et al. Impact of hospitalization in an acute geriatric unit on polypharmacy and potentially inappropriate prescriptions: a retrospective study. Geriatr Gerontol Int. 2017;17:2354–60. https://doi.org/10.1111/ggi.13073.
- Wawruch M, Macugova A, Kostkova L, et al. The use of medications with anticholinergic properties and risk factors for their use in hospitalised elderly patients. Pharmacoepidemiol Drug Saf. 2012;21:170–6. https://doi.org/10.1002/pds.2169.
- Gallagher P, Curtin D, de Siun A, et al. Antipsychotic prescription amongst hospitalized patients with dementia. QJM. 2016;109(9):589–93 https:// doi-org.abc.cardiff.ac.uk/10.1093/qjmed/hcw023.
- Reinold J, Palese F, Romanese F, Logroscino G, Riedel O, Pisa FE. Anticholinergic burden before and after hospitalization in older adults with dementia: increase due to antipsychotic medications. Int J Geriatr Psychiatry. 2019;34:868–80. https://doi.org/10.1002/gps.5084.
- Weichert I, Romero-Ortuno R, Tolonen J, Soe T, Lebus C, Choudhury S, et al. Anticholinergic medications in patients admitted with cognitive impairment or falls (AMiCI). The impact of hospital admission on anticholinergic cognitive medication burden. Results of a multicentre observational study. J Clin Pharm Ther. 2018;43(5):682–94. https://doi.org/10.1111/jcpt.12694 Epub 2018 May 4. PMID: 29729025.
- Kable A, Fullerton A, Fraser S, et al. Comparison of potentially inappropriate medications for people with dementia at admission and discharge during an unplanned admission to hospital: results from the SMS dementia study. Healthcare (Basel). 2019;7(1):8. https://doi.org/10.3390/healthcare7010008.
- Nair V, Hunter J. Anticholinesterases and anticholinergic drugs. Contin Educ Anaesth Crit Care Pain. 2004;4(5):164–8. https://doi.org/10.1093/ bjaceaccp/mkh045.

- National Institute for Health and Care Excellence. Medicines optimisations Quality Standard [QS120]. 2020. https://www.nice.org.uk/guida nce/qs120/chapter/quality-statement-6-structured-medication-review. Accessed 4 Aug 2020.
- National Institute for Health and Care Excellence. Medicines management in care homes Quality standard [QS85]. 2020. https://www.nice. org.uk/guidance/qs85/chapter/quality-statement-5-medication-reviews. Accessed 4 Aug 2020.
- O'Mahony D, O'Sullivan D, Byrne S, O'Connor M, Ryan C, Gallagher P. STOPP/START criteria for potentially inappropriate prescribing in older people: version 2. Age Ageing. 2015;44(2):213–8. https://doi.org/10.1093/ ageing/afu145.
- Rudolph J, Salow M, Angelini M, McGlinchey R. The anticholinergic risk scale and anticholinergic adverse effects in older persons. Arch Intern Med. 2008;168(5):508–13. https://doi.org/10.1001/archinternmed.2007. 106.
- Boustani M, Campbell N, Munger S, Maidment I, Fox C. Impact of anticholingerics on the aging brain: a review and practical application. Aging Health. 2008;4(3):311–20.
- Carnahan R, Lund B, Perry P, Pollock B, Culp K. The anticholinergic drug scale as a measure of drug-related anticholinergic burden: associations with serum anticholinergic activity. J Clin Pharamacol. 2006;46:1481–6. https://doi.org/10.1177/0091270006292126.

Publisher's Note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

Ready to submit your research? Choose BMC and benefit from:

- fast, convenient online submission
- thorough peer review by experienced researchers in your field
- rapid publication on acceptance
- support for research data, including large and complex data types
- gold Open Access which fosters wider collaboration and increased citations
- maximum visibility for your research: over 100M website views per year

At BMC, research is always in progress.

Learn more biomedcentral.com/submissions

