





Citation: Peklar J, Kos M, O'Dwyer M, McCarron M, McCallion P, Kenny RA, et al. (2017) Medication and supplement use in older people with and without intellectual disability. An observational, cross-sectional study. PLoS ONE 12(9): e0184390. https://doi.org/10.1371/journal.pone.0184390

Editor: Gianluigi Forloni, Istituto Di Ricerche Farmacologiche Mario Negri, ITALY

Received: April 16, 2017

Accepted: August 23, 2017

Published: September 6, 2017

Copyright: © 2017 Peklar et al. This is an open access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

Data Availability Statement: All relevant data are within the paper.

Funding: The TILDA study is supported by Irish Life; the Department for Health and Children; and by The Atlantic Philanthropies. The IDS-TILDA study is funded by the Health Research Board and the Department of Health. The sponsors had no role in the design and conduct of the study; collection, management, analysis and interpretation of the data; or preparation, review and approval of the manuscript.

RESEARCH ARTICLE

Medication and supplement use in older people with and without intellectual disability: An observational, cross-sectional study

Jure Peklar¹*, Mitja Kos¹, Máire O'Dwyer^{2,3}, Mary McCarron^{3,4}, Philip McCallion⁵, Rose Anne Kenny^{6,7}, Martin C. Henman²

- 1 Faculty of Pharmacy, University of Ljubljana, Ljubljana, Slovenia, 2 School of Pharmacy and Pharmaceutical Sciences, Trinity College Dublin, Dublin, Ireland, 3 IDS-TILDA School of Nursing and Midwifery, Trinity College Dublin, Dublin, Ireland, 4 Dean of the Faculty of Health Sciences, Trinity College Dublin, Dublin, Ireland, 5 School of Social Work, Temple University, Philadelphia, Pennsylvania, United States of America, 6 The Irish Longitudinal Study on Ageing (TILDA), Trinity College Dublin, Dublin, Ireland, 7 Trinity College Institute of Neuroscience (TCIN), Trinity College Dublin, Dublin, Ireland
- * jure.peklar@gmail.com

Abstract

Introduction

Understanding the medication and supplement use of aging people is critical to ensuring that health service providers in primary care can optimise use of these agents. An increasing number of people with different levels of intellectual disability (ID) are living in the community and becoming for the first time substantial users of primary health care services. This, however, brings new challenges that need to be addressed at the primary health care level. We quantified the use of medicines and food supplements and described the associated patterns of morbidity in the two comparable cohorts of aging population with and without intellectual disability.

Method

This research aligned participants of 50 years and over who lived in the community from two nationally representative cohorts of older people; those with ID from the Intellectual Disability Supplement (n = 238) and those without ID (n = 8,081) from the Irish Longitudinal Study on Ageing.

Results

Data showed that both medication and supplement use in the two groups was prevalent but that those with ID received more of both medications and supplements (e.g. polypharmacy was 39.0% in ID vs. 18.1% in non-ID cohort). Moreover, based on an analysis of the therapeutic groups and medications used that treatment was more intense in the ID cohort (95.8 vs. 7.0 International Non-proprietary Names per 100 participants). Supplement use was almost twice as prevalent in the ID group but substantially less diverse with only 10 types of supplements reported. Morbidity was higher in the ID group and showed a higher prevalence of neurological and mental health disorders.



Competing interests: The TILDA study is supported by Irish Life; the Department for Health and Children; and by The Atlantic Philanthropies. The IDS-TILDA study is funded by the Health Research Board and the Department of Health. The sponsors had no role in the design and conduct of the study; collection, management, analysis and interpretation of the data; or preparation, review and approval of the manuscript. This does not alter our adherence to PLOS ONE policies on sharing data and materials.

Conclusion

The results highlight that the burden of therapy management and the potential risks in those ageing with ID differs substantially from those ageing without ID. Understanding the medication and supplement use of people aging with intellectual disability (ID) is critical to ensuring that health service providers in primary/ambulatory care can optimise use of these agents.

Introduction

Ageing in the general population is strongly associated with multimorbidity (2 or more chronic conditions [1]), functional impairments and geriatric syndromes (falls, delirium, frailty) [2], requiring complex clinical management [3] including polypharmacy (5–9 medications) and excessive polypharmacy (≥ 10) [4–6]. Polypharmacy is associated with low adherence [7], adverse drug reactions [6,8]. Furthermore, with aging, people become more susceptible to adverse drug events owing to diminished physiological reserve and the accumulation of chronic conditions [9]. For the treatment of minor illness and the prevention of chronic illness and the intensification of treatment of established conditions prescription and non-prescription medications and supplements may be used on differing combinations over the lifespan. However, this adds to the complexity of the person's medication regimens and may further increase the potential risks and occurrence of harm from adverse drug reactions and interactions in the elderly [10–11].

Intellectual disability (ID) is a disability characterised by significant limitations in both intellectual functioning and in adaptive behaviours, which covers many everyday social and practical skills [12]. Delayed development and long term marked deficits in intellectual functioning characterised by impaired communication, social skills and motor impairments are among the problems posed by the disorder [13]. Intellectual disability begins during the developmental period and it often co-occurs with other mental conditions such as depression, autism spectrum disorder, and attention-deficit-hyperactivity disorder [12]. Indeed, due to comorbidities and challenging behaviours people with ID have on average up to 2.5 times the number of health problems reported for the general population [14–16] and they are one of the most heavily medicated groups in society [16].

In line with general population trends, the life expectancy of people with ID has increased significantly, however, it remains lower than that of the general population [17–18]. Older adults with ID may be particularly challenged as a result of premature aging, age-related decline in health, increased dependency and a further decline in cognitive function, which may reduce an individual's ability to cope, adapt and maintain quality of life [19]. Recent research in the same Irish cohort of people with ID as presented in this paper has shown, that 90% of participants reported medication use; 31.5% polypharmacy and 20.1% excessive polypharmacy [20].

From the mid-19th century, care for people with intellectual disabilities in Ireland was mainly in hands of religious orders who maintained special residential centres. From the 1950s, the development of community-based services began to emerge and from 1980 a stronger policy towards community inclusion started [21]. An increasing number of people with different levels of ID are living in the community and becoming for the first time substantial users of Primary Care services (e.g. GP, community pharmacy). This, however, brings new challenges that need to be addressed in Primary Care so that the special needs of this population may be met.



The aim of this study was to describe the patterns of use of medications and supplements and the associated patterns of morbidity in two groups of older, community dwelling people, those with, and without, Intellectual Disability since these groups have not been compared before in Ireland.

Methods

Study design and population

The data for this study was drawn from two sources: the Irish Longitudinal Study on Ageing (TILDA) study and the Intellectual Disability Supplement to TILDA (IDS TILDA). To facilitate comparison the groups were aligned on the basis of age—those over 50 years from each cohort; and living arrangements—those living independently or in Community Group Homes from the IDS TILDA cohort and the participants of the TILDA cohort, all of whom were living in the community. Thus, two similar groups were formed that were aligned at the group level.

The same measure for depression (moderate and severe) was used in both groups—the Centre for Epidemiologic Studies Depression Scale [22].

TILDA

The Irish Longitudinal Study on Ageing (TILDA) is a validated national prospective cohort study that describes the social (social network, home-care), economic (income, employment, life standard) and health status (physical and mental health, medication and food supplement use, need of health services and its utilisation) of older Irish adults. More details about the study cohort profile have been published elsewhere [23].

Participants were selected by means of a three stage process of RANSAM sampling procedure [23]. Inclusion criteria for participation in this study were age 50 years or older and being resident in the Republic of Ireland. Institutionalized people were excluded from recruitment.

The data used here were taken from the first wave of TILDA from October 2009 to February 2011. Medication and supplement use data were collected during the in-home interviews performed by trained interviewers using Computer-Aided Personal Interview software (Quancept SPSS®). The in-home inventory of medications and food supplement was conducted by asking the question "Now I would like to record all medications that you take on a regular basis, like every day or every week. This will include prescription and non-prescription medications, over-the-counter medications, vitamins, and herbal and alternative medications." No information about dose, frequency and quantity or prescription status was obtained.

A total of 8,175 people participated in but in 94 cases data on medication and supplement use were not collected (participants didn't want to give or didn't know the details about the medications/ supplements use during the interview) which corresponds to a participation rate of 98.9%. The TILDA study was approved by the research ethics committee of Trinity College Dublin, and subjects were required to sign an informed consent document prior to participation.

IDS TILDA

Data on older adults with intellectual disabilities was drawn from the first wave of data collected between 2009 and 2010 for the Intellectual Disability Supplement to the Irish Longitudinal Study on Ageing (IDS TILDA). The sample was randomly selected from Ireland's National Intellectual Disability Database (NIDD) which collects information on 26,000 people with all levels of ID, eligible for or receiving services and in a full range of residential circumstances [24]. This study has been described in detail elsewhere [25−26]. Inclusion criteria were: age ≥40 years with intellectual disability and written consent to participate and/or family/guardian



written agreement, where required. The recruited, consented and protocols completed sample was 753 persons with an ID, aged between 41 and 90 years and represented 8.9% of the total population aged 40 and over registered on the 2009 NIDD database. A pre-interview question-naire (PIQ) was sent to each participant/ carer before a formal in-person interview. The PIQ covers demographical information including age, level of ID (Down syndrome or ID from other aetiology) physical and mental health status (including physician confirmed diagnoses of physical and mental health concerns), healthcare utilisation and medication usage taken on a regular basis ('every day or every week', as in TILDA). Additional data was gathered in a subsequent face to face interview when PIQ reports including the information on the regular use of medications and supplements was cross-checked and verified for 736 (97.7%) participants. For the purpose of this paper the cohort of participants living in the community aged 50 years and over was only used to be aligned with the TILDA cohort inclusion criteria. This group comprised 238 participants for whom data on medications and health status was available. Medication usage included drugs recorded by brand/generic name, including prescription and non-prescription herbal and alternative drugs and any supplements.

The IDS TILDA study was approved by the research ethics committee of Trinity College Dublin and all of the participating services providers (n = 138).

The aim of this study was to describe the patterns of medication and supplement use in two groups of older, community dwelling people and to facilitate this comparison the groups were aligned on the basis of age, those over 50 years from each cohort, and living arrangements, those living independently or in Community Group Homes (IDS TILDA) whereas those living in residential care facilities are excluded from the TILDA cohort and were not included from the IDS TILDA cohort records. Residents of residential facilities and nursing homes are known to receive more medications than those living in the community both in the elderly [27] and in older people with ID [20].

Medication and supplement data collection

A supplement was defined according to the Directive 2002/46/EC of the European Parliament and of the Council, of 10 June 2002: "*Food supplements* means foodstuffs the purpose of which is to supplement the normal diet and . . . other substances with a nutritional or physiological effect. . . ." [28].

Medication data was coded using the ATC (Anatomical Therapeutic Chemical) classification system [29] which uses five levels from body system to drug name (International Non-Proprietary Name) to categorise each type of preparation.

Products with ATC groups C10AX06 (omega-3-triglycerides), M01AX05 (glucosamine), A12AA (calcium), A11D, A11E and B03B (vitamins B), B03A (iron), A11A and A11B (multivitamins with minerals), A11GA01 (vitamin C) and A11CC (vitamin D and analogues) were classed as supplements.

Morbidity, medication and supplement use was compared across genders and age groups $(50-59, 60-69 \text{ and } \ge 70 \text{ years})$ both the TILDA and IDS TILDA. Due to large difference in number of participants in the cohorts, number of medications and supplements were compared using same denominator ("per 100 enrolled participants").

Statistical analysis

Descriptive statistics were used to illustrate differences in the prevalence of medication and supplement use, morbidity and clinical characteristics of both populations. Prevalences were presented as percentages including 95% confidence intervals (CI). Group differences for each cohort were assessed by Pearson's chi-square test for categorical variables and analysis of



variance (ANOVA) for continuous ones. Two-sided significance tests were used throughout and p-value <0.05 were considered statistically significant.

To control for problems associated with multiple comparisons (which increases the likelihood of rejecting the null hypothesis when it is true; Type I error), and the False Discovery Rate (FRD) we adopted the Bonferroni correction procedure to maintain the Family wise Error Rate (FWER) [30]. In relation to demographic and clinical characteristics, we tested 20 and 21 hypotheses in the TILDA and IDS TILDA, respectively, with a desired α of 0.05, the Bonferroni correction tested each individual hypothesis at $\alpha = 0.05/20 = 0.0025$ and $\alpha = 0.05/21 = 0.0024$, respectively.

All analyses were performed using SPSS 20 (SPSS for Windows Release 20.0).

Results

TILDA and IDS TILDA cohorts

The age and sex profiles of the two cohorts presented in Table 1 differ and are consistent with the shorter life expectancy of people with ID and the preponderance of men in this population. Both medication and supplement use was substantially higher in the IDS group (73.1 IDS TILDA vs 49.5% TILDA respectively) in line with greater multimorbidity. The range of reported conditions was 0 to 9 in the TILDA group and 0 to 7 in IDS TILDA. However, the pattern of chronic conditions varied with a substantially greater prevalence in the TILDA group of hypertension (36.9% vs. 21.0%), bone and joint disease (33.3% vs. 21.4%), cancer (6.3% vs 4.6%) and moderate to severe depression (27.2% vs. 14.7%) while conversely, gastro-intestinal conditions (peptic ulcer, constipation; 17.2% vs. 7.8%) and eye conditions were more prevalent in the IDS group (67.6% IDS TILDA vs. 16.4% TILDA). Information on other mental health conditions (39.5%) and on epilepsy (24.1%) was collected only in IDS TILDA.

Use of medications and supplements

Table 2 illustrates that use of medications and supplements in the IDS TILDA group was greater than in the TILDA group; however, the most notable difference was in the degree of excessive polypharmacy which was 6 times more prevalent (TILDA 1.9% vs 11.3% IDS TILDA). Furthermore, use of a supplement with at least one medication, was over twice as prevalent in the IDS group (14.6% TILDA vs 32.8% IDS TILDA). The mean number of medications in both groups grew as age increased but the increase was greater in the TILDA group and the extent of the difference between the two groups diminished with increasing age. Supplement use also increased with age in both groups but in the IDS TILDA group the increase with age was greater and the disparity between the two cohorts also grew. Women used more medications and more supplements than men within both TILDA and IDS TILDA and the differences were more marked within IDS TILDA than in TILDA.

Types of medications and supplements used

The diversity of reported medications (Table 3 and Table 4) and supplements (Table 5) differs between the cohorts with 568 different drugs and 134 different supplements in TILDA and only 228 drugs and merely 10 supplements in IDS TILDA. However, when the number of participants is taken into account (568/8,081 TILDA vs 228/238 IDS TILDA) there was much higher diversity of drug use in the IDS cohort: more than 13-times (TILDA 7.0 vs. IDS TILDA 95.8 drugs per 100 participants).

The usage was greatest in certain therapeutic groups within the IDS cohort; for conditions of the Gastro-Intestinal tract and metabolism (A)—three times as many medications and 17



Table 1. Comparison of the demographic and clinical characteristics (not all characteristics have been explored equally in both groups).

	TILDA	IDS TILDA	
N (%) 8,081		Respondents	N (%) 238
3,706 (45.9%)		Male	134 (56.3%)
4,375 (54.1%)		Female	104 (43.7%)
		Age	
3,256 (40.3)		50–59	152 (63.9)
2,563 (31.7)		60–69	67 (28.2)
2,262 (28.0)		≥70	18 (8.0)
5,622 (69.6)	Education (≥ secondary school)		
	Disability [#]	Level of ID ^{\$} [missing 26]	
275 (3.4)	IADL disability	Mild	76 (35.8)
687 (8.5)	Any ADL disability	Moderate	109 (51.4)
		Severe/Profound	27 (12.8)
		Living status	
1,782 (22.1)	Living alone	Independent/Family	69 (29.0)
		Community/Group Home	169 (71.0)
2.35 (±2.55)	Number	of medications (mean+SD)	4.55 (±3.87)
1,405 (17.4)	Any s	upplement reported (%)	80 (33.6)
0.25 (0.64)	Number	of supplements (mean+SD)	0.47 (±0.76)
1,239 (15.4)	Self-ra	ted health [Fair/Poor] (%)	31 (13.1)
[missing = 13]			[missing = 2]
4,002 (49.5)	(≥:	Multimorbidity 2 chronic conditions)	174 (73.1)
1.71 (1.45))	Number of c	chronic conditions [mean+SD]	2.51 (±1.43)
2,981 (36.9)		Hypertension	50 (21.0)
1,165 (14.4)	(angina, h	Heart disease eart attack, AF, heart failure)	27 (11.3)
286 (3.5)	Str	roke (Stroke and TIA)	6 (2.5)
621 (7.7)		Diabetes	29 (12.4) [missing = 4]
1,326 (16.4)		Eye disorders	161 (67.6)
2,691 (33.3)	Bone/joint d	isease (osteoporosis, arthritis)	51 (21.4)
512 (6.3)		Cancer	11 (4.6)
325 (4.0)	Lung	disease (COPD, asthma)	16 (6.7)
634 (7.8)	Pep	tic ulcer, constipation	41 (17.2)
2,160 (27.2) [missing = 126]	Depre	ssion (severe/moderate)	35 (14.7)
		Mental health & neurological disorders *	94 (39.5)
		(schizophrenia, psychosis, depression)	
		Epilepsy [missing = 1]	57 (24.1)

^{*}ADL and IADL (basic and instrumental activities of daily living) are measures of one's independent living skills and is measured with the questionnaire

times as many different drugs, Dermatological conditions (D) - 14 times as many medications and 16 times as many drugs, and Nervous system conditions (N) - 6 times as many medications and 20 times as many drugs. In contrast, the use of medications for cancer and

^{\$} Level of Intellectual Disability indicates both intellectual and adaptive functioning deficits in conceptual, social, and practical domains.

^{*} Data on these conditions were not available in TILDA cohort



Table 2. Medication and supplement use compared.

	TILDA (n = 8,081)	IDS TILDA (n = 238)
At least 1 medication	5,572 (69.2%)	212 (89.1%)
5–9 medications	1,309 (16.2%)	66 (27.7%)
≥ 10 medications	154 (1.9%)	27 (11.3%)
At least 1 supplement	1,404 (17.4%)	80 (33.6%)
Concomitant use	1,180 (14.6%)	78 (32.8%)
Medication use per chronic condition (Mean No of medications ±SD)	1.34 (±1.24)	2.06 (±1.68)
Medication use with age (Mean No of medications±SD)	2.35 (±2.55)	4.55 (±3.87)
50–59	1.43 (±2.04)	3.95 (±3.68)
60–69	2.39 (±2.47)	5.46 (±4.11)
70+	3.62 (±2.73)	6.16 (±3.53)
Supplement use with age (Mean No of supplements ±SD)		
50–59	0.2 (±0.64)	0.43 (±0.77)
60–69	0.26 (±0.65)	0.51 (±0.77)
70+	0.29 (±0.64)	0.68 (±0.67)
Medication use by sex (Mean No of medications ±SD)		
Male	2.31 (±2.60)	4.15 (±3.75)
Female	2.38 (±2.49)	4.87 (±3.94)
Supplement use by sex (Mean No of supplements±SD)		
Male	0.14 (±0.48)	0.25 (±0.52)
Female	0.34 (±0.74)	0.65 (±0.87)
No of chronic conditions	use (Mean No of c	ation and supplement ombined medications ements ±SD)
0	0.66 (±1.27)	1.00 (±1.15)
1	1.67 (±1.81)	2.96 (±2.78)
2	2.86 (±2.17)	3.58 (±3.27)
3	4.16 (±2.58)	6.57 (±4.08)
4	5.51 (±2.79)	6.26 (±3.86)
5	6.50 (±3.00)	9.14 (±3.42)
6	7.85 (±2.93)	11.8 (±5.40)
7	7.69 (±4.20)	10.5 (±7.78)
8	9.78 (±4.27)	-

immunomodulation—L was 73% lower overall in IDS TILDA but with more drugs in use compared to the TILDA group. Similar pattern was observed in the Cardiovascular conditions (C) group.

Comparing the 20 most frequently reported therapeutic groups (2nd ATC level) in IDS-TILDA showed that 15 of them were common to TILDA cohort (Table 4). Of the 5 other therapeutic groups used in the IDS cohort, 3 were used to treat gastro-intestinal conditions whereas in the TILDA cohort, 3 of the 5 other therapeutic groups were associated with cardiovascular and haematological conditions.

In contrast to the greater variety of medications per participant identified in the IDS TILDA group, supplement utilisation illustrated minimal diversity with only 10 types of supplements in use. However, 9 supplements in the IDS TILDA group and 8 in the TILDA group were used by 1% or more of the cohort and in both, calcium with or without vitamin D and vitamins, alone or in combination preparations were the most prevalent.



Table 3. Reported number of medications and their INNs (ATC 5th level).

	TILDA [n =	= 8,081]		Therapeutic class		IDS TILI	DA [n = 238]	
Medications/ 100 enrolled participants	N of medications	Different drugs/ 100 enrolled participants	Different drugs	1 st ATC level	Different drugs	Different drugs/ 100 enrolled participants	N of medications	Medications/100 enrolled participants
34.7	2,807	1.0	78	A—Alimentary	40	16.8	256	107.6
27.4	2,211	0.2	13	B- Blood	3	1.3	28	11.8
100.4	8,110	1.3	107	C- Cardiovascular	36	15.1	167	70.2
1.2	95	0.4	32	D- Dermatology	15	6.3	40	16.8
6.7	545	0.5	41	G- Genito-Urinary	11	4.6	19	8.0
9.2	744	0.2	15	H- Hormones	7	2.9	41	17.2
1.8	143	0.4	33	J-Anti-infective	7	2.9	12	5.0
3.0	244	0.3	27	L-Malignancy & Immuno- modulation	1	0.4	2	0.8
16.1	1,299	0.5	39	M- Musculo- Skeletal	11	4.6	43	18.1
32.4	2,617	1.3	105	N- Nervous system	61	25.6	486	204.2
0.1	6	0.0	3	P—Antiparasitic	1	0.4	1	0.4
15.0	1,214	0.6	45	R- Respiratory	21	8.8	72	30.3
2.9	238	0.3	27	S- Sensory	11	4.6	21	8.8
0.1	8	0.0	3	V- Various	3	1.3	10	4.2
251.0	20,281	7.0	568	All	228	95.8	1,198	503.4

Table 4. Comparison of the 20 most frequently reported therapeutic classes (2nd ATC level).

	IDS T	ILDA	TII	LDA
Most frequently reported medications by ATC (2nd level)	Rank	%	%	Rank
N05 Psycholeptics (antipsychotics, anxiolytics, hypnotics)	1	11.0	3.1	11
N06 Psychoanaleptics (antidepressants, stimulants)	2	6.1	3.0	12
R03 Drugs for obstructive airway diseases	2	6.1	4.8	6
C09 Agents acting on renin-angiotensin system	4	5.3	10.2	3
N03 Antiepileptics	5	4.8	1.3	18
A06 Drugs for constipation	6	4.4	0.5	25
S01 Ophtalmologicals	6	4.4	1.1	20
A02 Drugs for acid related disorders	8	3.9	6.2	4
A10 Drugs used in diabetes	9	3.1	3.6	9
G04 Urologicals	9	3.1	1.7	16
N02 Analgesics	9	3.1	3.0	12
C10 Lipid modifying agents	12	2.6	13.9	1
D01 Antifungals for dermatological use	12	2.6	0.1	37
J01 Antibacterials for systemic use	12	2.6	0.6	23
M01 Anti-inflammatories and antirheumatics	12	2.6	4.1	7
A03 Drugs for functional GI disorders	16	1.8	0.5	25
C07 Beta blocking agents	16	1.8	5.6	5
C03 Diuretics	16	1.8	3.6	9
C08 Calcium channel blockers	16	1.8	3.9	8
A07 Antidiarrhoeals	16	1.8	0.3	33

https://doi.org/10.1371/journal.pone.0184390.t004



Table 5. Reported supplements for TILDA and IDS TILDA.

TILDA [n = 8,081]			IDS TIL	DA [n = 23	8]
Supplements	N	Per 100 enrolled participants	Supplements	N	Per 100 enrolled participants
Calcium with or w/o D vitamin	619	7.7	Calcium with or w/o D vitamin	36	15.1
Vitamins (alone or multiple)	468	5.8	Vitamins (alone or multiple)	34	14.3
Omega-3-triglycerides	360	4.5	Iron	20	8.4
Glucosamine	263	3.3	Evening primrose oil	6	2.5
Iron	97	1.2	Omega 3	6	2.5
Evening primrose oil	56	0.7	Glucosamine	3	1.3
Garlic	23	0.3	Linseed	4	1.7
Carotenoids (lutein, meso- zeaxanthin)	23	0.3	Carotenoids (lutein, meso- zeaxanthin)	2	0.8
Echinacea	14	0.2	Acidophilus	2	0.8
Other non-ATC supplements*	171	2.1	Peppermint oil	1	0.4
All	2,094	25.9	All	114	47.9

^{*} aloevera, starflower, cranberry, milk thistle,...

The pattern of medication and supplement usage alone and in combination shows considerable variation across the range of reported conditions (Table 6). In both groups, 9 out of 10 participants reported use of either or both medications and supplements but in the IDS TILDA group only 5.7% (95% CI 3.6, 7.7) reported taking neither, whereas in the TILDA group this proportion was almost double 10.1% (95% CI 6.2, 13.9).

Use of supplements alone was only reported in eye disorders within the IDS group (1.2%) whereas use alone was reported in all of the conditions within the TILDA group and in particular in those reporting bone or joint disease (2.6%), depression (2.3%) or cancer (2.1%). Use of medications alone was most frequently associated with diabetes mellitus (80.2%) and hypertension (74.4%) in the TILDA group and in moderate or severe depression (77.1%) and mental health conditions (60.6%) in the IDS TILDA group.

Concomitant use was most prevalent in both cohorts in those reporting bone or joint disease (54.9%) with a calcium preparation with or without vitamin D taken with medications. By contrast the lowest concomitant usage in the TILDA group and one of the lowest in the IDS group are associated with those reporting diabetes mellitus 15.3% TILDA vs 37.9% IDS TILDA). Concomitant use was highest in the IDS group in stroke although this was the smallest subgroup (n = 6).

The use of neither medications nor supplements was frequently reported among the TILDA group most notably by those reporting moderate or severe depression (21.4%) and in 18.2% of those with peptic ulcer and 12.4% of those with bone or joint disease. Of the larger subgroups among the IDS TILDA participants, those reporting eye disorders (8.7%), hypertension (10.0%) and bone or joint disease (7.8%) also recorded neither medication nor supplement use.

Discussion

There are few comparative studies of medication use in these populations and this is the first to describe the use of both medications and supplements. This study shows that in two similar groups of community dwelling elderly patients' medication and supplement use was prevalent but in those with ID both medication and supplement use was higher in all age groups. The pattern of morbidity reflected, in part, the therapeutic classes used but for some, such as peptic ulcer and moderate to severe depression, 20% or more of the TILDA group reported no use of

Table 6. Medication (M) and supplement (S) use and chronic conditions in TILDA (T) and IDS TILDA (IDS) cohort.

Taking/ Condition (TILDA vs. IDS TILDA)	Hypert 36.9%:	Hypertension 36.9%: 21.0%		Heart disease (angina, heart attack, AF, heart failure) 14.4%: 11.3%	Stroke (Stroke and TIA) 3.5%: 2.5%	Stroke) 3.5%: %	Diabetes mellitus 7.7%: 12.2%	etes 8 7.7%: 2%	Eye disorders 16.4%: 67.6%	orders 67.6%	Bone/joints disease (osteoporosis, arthritis) 33.3%: 21.4%	oints ase prosis, tis) 21.4%	Cancer 6.3%: 4.6%	6.3%:	Lung disease 4.0%: 6.7%	sease 3.7%	Peptic ulcer 7.0%: 4.2%	ulcer 1.2%	Depression symptoms (severe/ moderate) 26.7%: 14.7%	ssion toms ere/ rate)	Mental health condition 39.5%	Epilepsy 23.9%
	T (n = 2981)	IDS (n = 50)	T (n= 1165)	IDS (n = 27)	(n = 286)	IDS (n = 6)	T (n = 621)	IDS* (n = 29)	T (n = 1326)	IDS (n = 161)	T (n = 2691)	IDS (n = 51)	T (n = 512)	IDS (n = 11)	(n = 325)	IDS (n = 16)	T (n = 567)	IDS (n = 10)	T [§] (n = 2160)	IDS (n = 35)	IDS (n = 94)	IDS* (n = 57)
No M, no S	201	2	59	-	6	0	23	-	149	14	334	4	89	-	38	0	103	2	462	0	5	-
	%2.9	10.0%	5.1%	3.7%	3.1%	%0.0	3.7%	3.4%	11.2%	8.7%	12.4%	7.8%	13.3%	9.1%	11.7%	. %0.0	18.2%	20.0%	21.4%	%0.0	5.3%	1.8%
M only, no S	2219	27	853	14	199	2	498	17	833	06	1553	19	328	9	214	6	341	14	1261	27	57	29
	74.4%	54.0%	73.2%	51.9%	%2.69	33.3%	80.2%	28.6%	62.8%	25.9%	27.7%	37.3%	64.1%	54.5%	62.9%	56.3%	60.1%	%9.69	58.4%	77.1%	%9.09	20.9%
M & S concomitantly	542	18	241	12	17	4	96	11	323	55	734	28	105	4	70	7	112	9	388	8	32	27
	18.3%	36.0%	20.7%	44.4%	26.9%	%2.99	15.3%	37.9%	24.4%	34.2%	27.3%	54.9%	20.5%	36.4%	21.5%	43.8%	19.8%	27.3%	18.0%	22.9%	34.0%	47.4%
S only	19	0	12	0	-	0	2	0	21	2	70	0	11	0	8	0	=	0	49	0	0	0
	%9.0	%0.0	1.0%	%0.0	0.3%	%0.0	%8.0	%0.0	1.6%	1.2%	2.6%	%0.0	2.1%	%0.0	%6:0	%0.0	1.9%	%0.0	2.3%	%0.0	%0:0	%0.0

* Missing = 4.

§ Missing = 126. \neq Missing = 1.

https://doi.org/10.1371/journal.pone.0184390.t006



either a medication or a supplement; whereas the use of a medication and/or a supplement was almost universal in the IDS group. In the IDS group there was extensive use of medications in epilepsy, mental health conditions and depression and particularly the high level of excessive polypharmacy, may indicate more intensive drug treatment [31]. Although in the TILDA group a disparate group of medications was used, when this was standardised per 100 participants, it was clear that the IDS group was exposed to a proportionately wider range of medications. In contrast, the types of supplements used in the IDS group were extremely limited in comparison to the TILDA group suggesting that TILDA participants were prescribed and/or chose from a wider range of alternatives than those in the IDS group, although the two most frequently used classes were the same. Use of supplements alone was only significant in the TILDA group but the level of usage was low at only just over 2%.

Medications may pose some risk to most groups but some types of medication are particularly risky for the elderly. Long term use of sedative and anticholinergic medications have been associated with cognitive impairment, increased risk of falls, of hospitalisation and of hospital admission. Increased sedative load has been reported for the TILDA population with those most exposed having poorer health and sedative load was associated with being prefrail and frail [32] while in the IDS cohort (including those aged 40–49 years and those in nursing homes) a high anticholinergic burden was found and it was associated with daytime dozing and constipation [33]. This evidence strongly suggest that the groups studied in this work are exposed to risks and vulnerable to harm from both the types and the high levels of medications and supplements that they use. This reinforces the need for a systematic and structured approach to managing the level of exposure, to high risk medications and to reducing their vulnerability. Since these groups live in the community, it is most appropriate that they continue to be cared for in the community. Those Primary Care providers who share responsibility for the provision of medications and supplements, general practitioners and community pharmacists, should be supported and empowered to collaborate in the care of these diverse populations. Our findings highlight the scale and complexity of the challenge of the future management of these agents in both these populations.

For the TILDA group there is also a substantial but different composition of sources of risk from medications and supplement use. The diversity of medications and supplements used in this large group reduces the likelihood that Primary Care providers will have extensive experience in using the less frequently encountered drugs and supplements and the even rarer permutations of combined use that may occur. Previous detailed research in the TILDA population showed that 4.5% of concomitant users could suffer some type of major drug-supplement interaction [11]. Recently, considerable interest has been shown in approaches to reducing the numbers of medications used, deprescribing, in older people, particularly those receiving palliative care [34]. Nevertheless, this requires a mechanism for reviewing all of the medications and supplements and although pilot work has been carried out [35], no initiative has been brought forward by the health service in Ireland. Similarly, while providers of nursing home care are required to ensure that medication reviews are performed every 6 months, medication reviews would also benefit people with ID living in the community but the requirement for this has not been acknowledged [36].

In the IDS group the burden is the most marked because of the high use of both medications and supplements at all ages and because their capacity to take part in the management of their conditions, symptoms and treatments is limited and may diminish with ageing. Consequently, this burden is mainly carried by non-family carers because as people with ID live longer they are more likely to outlive their parents. The present and future consequences of this have not been evaluated and in Ireland, few carers receive training about medication and supplement use and unless the care providers address medications information and management



issues, it is unlikely that anyone else does [37]. Furthermore, the complexity of the clinical conditions and the combination of medications and supplements used would be difficult for anyone, health professional or carer, to manage, and in many instances, such as people with epilepsy, care is provided both by specialists in Secondary Care clinics and generalists in Primary Care which adds potential communication difficulties to the care process. Thus the potential for adverse drug reactions and interactions as well as drug-supplement interactions and administration errors must be considerably greater in the IDS than in the TILDA group. Therefore as people with ID are enabled to live in the community in greater numbers suitable mechanisms for the regular review of medication and supplement use, the education of carers about the use of medications and supplements and the co-ordination of care are increasingly needed to mitigate the associated risk and to prevent harm [38].

Strengths and limitations

Both the TILDA and IDS TILDA cohorts used robust sampling techniques to establish two nationally representative cohorts. The data collection periods were contemporaneous and almost all the participants in both cohorts recorded medical data (98.9% and 99.6% respectively), with validated response quality data capture that allowed for detailed analysis of medication and food supplement usage patterns, morbidity and socio-economical and lifestyle characteristics. Data on medication and food supplement use was collected by in-home interview (TILDA) or with a preinterview questionnaire and then validated within the face-to-face interview (IDS TILDA) which improves reliability compared to self-report recall alone [39]. The use of the Bonferroni correction addressed the problem of multiplicity. Both populations were matched regarding the age and living status.

There are also certain limitations: Only regularly taken medications and supplements were recorded and not those taken "when required" (prn). There was no data on the dose of medications or supplements or the way they were obtained (prescribed or purchased). The reporting of chronic conditions was based on participant or proxy self-report in both cohorts and thus may be prone to a misclassification bias [28] and the severity of the condition was not captured, except in the instance of depression. Matching at the level of the individual and adjustment using a technique such as propensity scoring was not undertaken because it would have required creation of a specific dataset which was not possible within the constraints of the Ethics approvals, the range of variables included, and the time available for this study.

Acknowledgments

The TILDA study is supported by Irish Life; the Department for Health and Children; and by The Atlantic Philanthropies. The IDS-TILDA study is funded by the Health Research Board and the Department of Health. The sponsors had no role in the design and conduct of the study; collection, management, analysis and interpretation of the data; or preparation, review and approval of the manuscript. This does not alter our adherence to PLOS ONE policies on sharing data and materials.

The authors would like to thank the people with ID who participated in this study, their families, the services involved, the IDS-TILDA Scientific Advisory Committee, and the Intellectual Disability Consultative Groups for their support.

Author Contributions

Conceptualization: Jure Peklar, Mitja Kos, Máire O'Dwyer, Mary McCarron, Martin C. Henman.



Data curation: Jure Peklar, Máire O'Dwyer. Formal analysis: Jure Peklar, Máire O'Dwyer.

Funding acquisition: Mary McCarron, Rose Anne Kenny.

Investigation: Jure Peklar, Mary McCarron, Philip McCallion, Rose Anne Kenny.

Methodology: Jure Peklar, Mitja Kos, Máire O'Dwyer, Mary McCarron, Philip McCallion,

Rose Anne Kenny, Martin C. Henman. **Project administration:** Rose Anne Kenny.

Resources: Philip McCallion.

Supervision: Mitja Kos, Mary McCarron, Martin C. Henman.

Validation: Mitja Kos.

Writing – original draft: Jure Peklar.

Writing - review & editing: Jure Peklar, Máire O'Dwyer, Martin C. Henman.

References

- 1. Uijen AA, van de Lisdonk EH. Multimorbidity in primary care: prevalence and trend over the past 20 years. Eur J Gen Pract. 2008; 14(Suppl 1): 28–32.
- Marengoni A, Angleman S, Melis R, Mangialasche F, Karp A, Garmen A, et al. Aging with multimorbidity: a systematic review of the literature. Ageing Res Rev. 2011 Sep; 10(4):430–9. https://doi.org/10.1016/j.arr.2011.03.003 PMID: 21402176
- 3. Koné Pefoyo AJ, Bronskill SE, Gruneir A, Calzavara A, Thavorn K, Petrosyan Y, et al. The increasing burden and complexity of multimorbidity. BMC Public Health. 2015 Apr 23; 15(1):415.
- Hilmer SN, Gnjidic D. The effects of polypharmacy in older adults. Clin Pharmacol Ther. 2009 Jan; 85 (1):86–8. https://doi.org/10.1038/clpt.2008.224 PMID: 19037203
- Peklar J, Henman MC, Richardson K, Kos M, Kenny RA. Food supplement use in the community dwelling population aged 50 and over in the Republic of Ireland. Complement Ther Med. 2013 Aug; 21 (4):333–41. https://doi.org/10.1016/j.ctim.2013.06.004 PMID: 23876564
- Gnjidic D, Le Couteur DG, Pearson SA, McLachlan AJ, Viney R, Hilmer SN, et al. High risk prescribing in older adults: prevalence, clinical and economic implications and potential for intervention at the population level. BMC Public Health. 2013 Feb 7; 13:115. https://doi.org/10.1186/1471-2458-13-115 PMID: 23388494
- Tsai KT, Chen JH, Wen CJ, Kuo HK, Lu IS, Chiu LS, Wu SC, Chan DC. Medication adherence among geriatric outpatients prescribed multiple medications. Am J Geriatr Pharmacother. 2012 Feb; 10(1):61– 8. https://doi.org/10.1016/j.amjopharm.2011.11.005 PMID: 22264853
- Gnjidic D, Hilmer SN, Blyth FM, Naganathan V, Waite L, Seibel MJ, et al. Polypharmacy cutoff and outcomes: five or more medications were used to identify community-dwelling older men at risk of different adverse outcomes. J Clin Epidemiol. 2012 Sep; 65(9):989–95. https://doi.org/10.1016/j.jclinepi.2012.02.018 PMID: 22742913
- Mallet L, Spinewine A, Huang A. The challenge of managing drug interactions in elderly people. Lancet. 2007 Jul 14; 370(9582):185–91. https://doi.org/10.1016/S0140-6736(07)61092-7 PMID: 17630042
- Nahin RL, Pecha M, Welmerink DB, Sink K, DeKosky ST, Fitzpatrick AL. Concomitant use of prescription drugs and dietary supplements in ambulatory elderly people. J Am Geriatr Soc. 2009 Jul; 57 (7):1197–205. https://doi.org/10.1111/j.1532-5415.2009.02329.x PMID: 19515113
- Peklar J, Henman MC, Kos M, Richardson K, Kenny RA. Concurrent Use of Drugs and Supplements in a Community-Dwelling Population Aged 50 Years or More: Potential Benefits and Risks. Drugs Aging. 2014 Jul; 31(7):527–40. https://doi.org/10.1007/s40266-014-0180-6 PMID: 24890574
- American Psychiatric Association (2013). Diagnostic and Statistical Manual of Mental Disorders (5th ed.) Arlington, Virginia: American Psychiatric Publishing. Available from: http://www.dsm5.org/documents/intellectual%20disability%20fact%20sheet.pdf (Accessed on 18.07.2016)
- Matson L, Cervantes PE. Comorbidity among persons with intellectual disabilities. Res Autism Spectr Disord. 2013 Jul 7; 1:1318–1322.



- Haveman M, Heller T, Lee L, Maaskant M, Shooshtari S and Strydom A. Major Health Risks in Aging Persons With Intellectual Disabilities J Policy Pract Intellect Disabil, 2010; 7: 59

 –69.
- van Schrojenstein Lantman-de Valk HM, Walsh PN. Managing health problems in people with intellectual disabilities. BMJ. 2008 Dec 8; 337.
- Straetmans JM, van Schrojenstein Lantman-de Valk HM, Schellevis FG, Dinant GJ. Health problems of people with intellectual disabilities: the impact for general practice. Br J Gen Pract. 2007; 57(534):64–6. PMID: 17244427
- Coppus AMW. People with intellectual disability: What do we know about adulthood and life expectancy? Dev Disabil Res Rev. 2013; 18(1):6–16. https://doi.org/10.1002/ddrr.1123 PMID: 23949824
- McCarron M, Carroll R, Kelly C, McCallion P. Mortality Rates in the General Irish Population Compared to those with an Intellectual Disability from 2003 to 2012. J Appl Res Intellect Disabil. 2015 Sep; 28 (5):406–13. https://doi.org/10.1111/jar.12194 PMID: 26256275
- 19. Sinai A, Bohnen I, Strydom A. Older adults with intellectual disability. Curr Opin Psychiatry. 2012; 25 (5):359–64. https://doi.org/10.1097/YCO.0b013e328355ab26 PMID: 22744404
- O'Dwyer M, Peklar J, McCallion P, McCarron M, Henman MC. Factors associated with polypharmacy and excessive polypharmacy in older people with intellectual disability differ from the general population: a cross-sectional observational nationwide study. BMJ Open, 2016; 6: e010505. https://doi.org/10.1136/bmjopen-2015-010505 PMID: 27044582
- Health Service Executive. Report of the Working Group on Congregated Settings HSE. Time to Move on from Congregated Settings: A strategy for Community Inclusion (2011). Available from: http://www.hse.ie/eng/services/list/4/disability/congregatedsettings/ (Accessed on 19.07.2016)
- Kearney PM, Cronin H, O'Regan C, Kamiya Y, Savva GM, Whelan B, et al. Cohort profile: The Irish Longitudinal study on Ageing. Int. J. Epidemiol. (2011) 40 (4): 877–884. https://doi.org/10.1093/ije/dyr116 PMID: 21810894
- 23. Radloff LS. The CES-D scale a self-report depression scale for research in the general population. Appl Psychol Measur(1977): 1(3):385–401.
- 24. Kelly CS, Kelly F. Annual Report of the National Intellectual Disability Database Committee 2009. Health Research Board; 2010.
- 25. McCarron M, Swinburne J, Burke E, McGlinchey E, Mulryan N, Andrews V, et al. Growing Older with an Intellectual Disability in Ireland 2011: First Results from the Intellectual Disability Supplement of The Irish Longitudinal Study on Ageing. School of Nursing, Trinity College Dublin, 2011.
- 26. McCarron M, Swinburne J, Burke E, McGlinchey E, Carroll R, McCallion P. Patterns of multimorbidity in an older population of persons with an intellectual disability: Results from the intellectual disability supplement to the Irish longitudinal study on aging (IDS-TILDA). Res Develop Disabil. 2013; 34(1):521–7.
- Shah SM, Carey IM, Harris T, DeWilde S, Cook DG. Quality of prescribing in care homes and the community in England and Wales. Br J Gen Pract. 2012 May; 62(598):e329–36. https://doi.org/10.3399/bjgp12X641447 PMID: 22546592
- Directive 2002/46/EC of the European Parliament and of the Council of 10 June 2002 on the approximation of the laws of the Member States relating to food. Official Journal L 183, 12/07/2002 P.0051–0057 http://eurlex.europa.eu/LexUriServ/LexUriServ.do?uri=CELEX:32002L0046:EN:HTML (accessed on 02.07.2016)
- 29. World Health Organization. Anatomical Therapeutic Classification: Structure and Principles [Internet]. Oslo: World Health Organization; 2011 [updated 2011 Mar 25; cited 2014]. Anatomical Therapeutic Classification: Structure and Principles. Available from: http://www.whocc.no/atc/structure_and_principles/.
- Benjamini Y, Hochberg Y. Controlling the false discovery rate: a practical and powerful approach to multiple testing. J R Stat Soc Series B Stat Methodol. 1995:289–300.
- O'Dwyer M., Peklar J., Mulryan N., McCallion P., McCarron M., and Henman M. C. Prevalence, patterns and factors associated with psychotropic use in older adults with intellectual disabilities in Ireland. J Intell Disabil Res. 2017 https://doi.org/10.1111/jir.12391 PMID: 28639298
- Peklar J, O'Halloran AM, Maidment ID, Henman MC, Kenny RA, Kos M. Sedative Load and frailty among community-dwelling population aged ≥65 years. J Am Med Dir Assoc. 2015 Apr; 16(4):282–9. https://doi.org/10.1016/j.jamda.2014.10.010 PMID: 25434581
- O'Dwyer M, Maidment ID, Bennett K Peklar J, Mulryan N, McCallion P, et al. Association of Anticholinergic Burden with Adverse Effects in Older People with Intellectual Disabilities in Ireland: an observational cross-sectional study Br J Psychiatry. 2016 Dec; 209(6):504–510. https://doi.org/10.1192/bjp.bp.115. 173971 PMID: 27660331
- Reeve E, Gnjidic D, Long J, Hilmer S. A systematic review of the emerging definition of 'deprescribing' with network analysis: implications for future research and clinical practice. Br J Clin Pharmacol. 2015 80:6; 1254–1268. https://doi.org/10.1111/bcp.12732 PMID: 27006985



- Henman MC. Medications use review pilot project. 2012. Available from: http://www.lenus.ie/hse/handle/10147/285066 (Accessed; 21.07.2016)
- Health Information and Quality Authority. Medications Management Guidance (2015). Available from: https://www.hiqa.ie/system/files/Medications-Management-Guidance.pdf (Accessed on 19.07.2016)
- Flood B, Henman MC. Case study: hidden complexity of medications use: information provided by a
 person with intellectual disability and diabetes to a pharmacist. Br J Learn Disabil 2015: 43(3); 234
 242.
- **38.** Flood B, Henman M. The Intellectually Disabled: a neglected group who need pharmacists' help. Int J Pharm. 2011; 27(3): 41–44.
- Qato DM, Alexander GC, Conti RM, Johnson M, Schumm P, Lindau ST. Use of prescription and overthe-counter medications and dietary supplements among older adults in the United States. JAMA 2008; 300:2867–78. https://doi.org/10.1001/jama.2008.892 PMID: 19109115