1	Title: A cross-sectional feasibility study of nutrient intake patterns in people with
2	Parkinson's compared to government nutrition guidelines.
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21	Abstract
22	Diet could have implications for disease progression and management in people with
23	Parkinson's (PwP). However, the knowledge of diet intake patterns in PwP is limited.
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25	Objectives
26	We set out to assess the feasibility of collecting diet data in PwP to determine food and
27	nutrient intake, in order to compare to national nutrition guidelines and thus understand the
28	habits in this population.
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30	Methods
31	In this cross sectional feasibility study PwP were approached through local support groups
32	through out the Thames Valley and were asked to complete a Food Frequency Questionnaire
33	Eligibility criteria included a self-reported neurologist confirmed diagnosis of Parkinson's.
34	Completeness of questionnaires was reported and 80% was considered appropriate for each
35	measure including demographic information.
36	
37	Results
38	Response rate was 61% and missing data for the 121 returned questionnaires was 74%,
39	however of the 90 used for analysis there was 100% completion of the questionnaires.
40	Compared to the UK government guidelines, protein was significantly higher for both males
41	and females and fluid intake lower for both genders (p<0.001). There were several other
42	differences in nutrient intake compared to guidelines.
43	
44	Conclusion

- We observed high levels of engagement from PwP and found that assessing food and nutrient patterns in PwP was feasible. Importantly, the diet was generally healthy overall, yet there
- 47 were specific nutrients that may affect medication metabolism in PwP which were found to
- be high. Therefore further research into this emerging and important area is warranted.

50 Keywords: Protein; diet; Parkinson's

Introduction

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A healthy diet is known to benefit physical and cognitive functioning throughout the lifespan 54 [1, 2], and to protect against a number of long term conditions and associated symptoms [3, 55 4]. However there is limited evidence of the role of diet in disease progression and symptom 56 management in people with Parkinson's (PwP) [5, 6]. 57 58 Whilst diet is known to impact on the effectiveness of drug metabolism [7], dietary patterns 59 are not typically measured clinically in PwP and there is a need to consider diet in both 60 disease progression and management. Although only a modest effect of diet on risk of 61 Parkinsons's has been shown [8], the role of diet after diagnosis is more promising [9, 10]. 62 63 Evidence so far has suggested a more healthy diet is associated with lower levels of disability and symptom severity in PwP, yet good quality comprehensive research is lacking and 64 subsequently there are no specific nutritional guidelines for PwP [6, 11]. 65 66 PwP have been found to consume less fluids than the control population, partially as a result 67 of symptoms such as dysphagia [12]. High protein intake have also been noted in PwP and 68 the higher the intake the larger the dose of levodopa required, with a correlation seen between 69 higher protein intake and levodopa related motor complications [13]. A recent study looked 70 71 into the effect of food groups on Parkinson's progression and found that 'healthy' food such as fresh vegetables, fresh fruit, nuts and seeds, non fried fish, olive oil, wine, coconut oil, 72 fresh herbs, and spices were related to a reduced progression of disease, where as fruits and 73 vegetables, diet and non diet soda, fried foods, beef, ice cream, yogurt, and cheese were 74 associated with increased progression [14]. 75

The aim of the present study is to determine the feasibility of collecting information about food and nutrient intakes in PwP using a standardised comprehensive diet questionnaire, and to highlight habits and potential areas of concern for PwP considering the disease and its management.

Methods

This was a cross sectional pilot study designed to assess aspects of feasibility including the efficiency of data collection methods through completion and return of questionnaires, identification of missing data and recruitment rate. Participants were recruited throughout Buckinghamshire and Oxfordshire either via the Parkinson's UK Research Network or through Parkinson's UK support groups and questionnaires were administered in person and were returned through mail. Potential participants were provided with information about the study and if they agreed to take part, consent was implied through the completion and return of anonymised questionnaires. Eligibility criteria included a self-reported neurologist confirmed diagnosis of Parkinson's. Ethical approval for this study was granted by University Ethics Committee (150895).

The EPIC-Norfolk Food Frequency Questionnaire [15] was used to measure habitual food intake over the previous 12 months. It included questions about specific food items, such as seasonal consumption of fruit and vegetables and habitual consumption of meat, fish, dairy products, potatoes, breads, rice, fats and sugars. The FFQ was analysed using software from the European Prospective Investigation into Cancer (EPIC-Norfolk) Cohort study; from which the accuracy of the analysis was originally validated and therefore has not been

validated in those with Parkinson's. Demographic information was self reported and included weight, height, gender, and date of birth.

Dietary Reference Values (DRVs) are a series of estimates of the energy and nutritional requirements of different groups of healthy people and are based on the UK populations [16]. These were set by the Committee on Medical Aspects of Food and Nutrition Policy (COMA) in 1991. Therefore in the current study the nutrient intakes from the participants were compared with the optimal intake values based on the government guidelines.

Statistical analysis

A sample size of 82 PwP was calculated based on a population of 554 PwP throughout Oxfordshire (assuming 227 new diagnoses annually and 227 existing), a confidence interval of 10% and a confidence level of 95%. Demographic data was described using descriptive analysis and recruitment rate was determined. Completeness of questionnaires was reported and 80% was considered appropriate for each measure including demographic information [17].

Nutrient data was compared to the SACN guidelines to identify any obvious and major differences in nutrient intake in PwP compared to the UK population (\pm 1 SD). These nutrients were then further analysed. Therefore, the data were first descriptively analysed before performing further analysis.

Significance level was set at 5%. Multicollinearity was assessed and collinear variables were not included. Data were analysed using SPSS Statistics Version 23 (IBM SPSS Statistics for Windows, IBM Corp, Armonk, NY, USA). Independent two sided t tests for males and females were performed to compare mean values for each nutrient to the UK guideline recommendation for these nutrients.

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127	Results
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129	This was a cross-sectional feasibility study including 90 PwP (men 52; women 38) over 18
130	years of age (mean years 68 ± 9.64 SD, body mass index 25.02 ± 4.27 SD mg/k ²) recruited
131	between 2016 to 2017. 200 questionnaires were administered randomly in the support group
132	meetings and of those 121 returned (61%), 90 were used for analysis (45%). Missing data for
133	the 121 returned questionnaires was 74%, however of the 90 used for analysis there was
134	100% completion of the questionnaires (fig 1).
135	
136	[See appendix for: figure 1]
137	
138	Comparison to UK nutrition guidelines
139	
140	In the current study, the nutrient intake in PwP was compared to the SACN guidelines and
141	table 1 represents nutrients that were significantly different from the UK guidelines. Protein
142	intake was significantly higher compared to the UK guidelines and fluid intake lower in both
143	genders (p<0.001). Also, other nutrients were shown to have significance compared to SACN
144	guidelines, for example energy intake was lower for men (p=0.002), zinc was higher for
145	females (p<0.001) and fibre and vitamin D were lower for both genders (p<0.001) yet iron,
146	vitamin C, folate, phosphorus, B6, B12, calcium were higher for both genders (p<0.001).
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148	[See appendix for: table 1]
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Discussion

We assessed diet in a representative sample of PwP and notably observed that protein intake was twice that recommended by UK guidelines. Other interesting findings were lower fluid, fibre and vitamin D intake and more differences were found in men compared to SACN guidelines who generally had a poorer diet than women. Although research is limited, nutritional status may play a significant role in symptoms in PwP [17]. Our findings suggest that monitoring diet is feasible and well received and that the development of approaches to effectively support optimal and specific diets for PwP may be important for management of the disease and symptoms.

Although fluid consumption tended to be similar between the sub groups in this cohort of PwP, the intake of fluids was at the lower end of 'optimal' when compared to UK guidelines. In the current study intakes were around 800 ml whereas guidelines recommend a minimum of 1200 ml a day for health [18]. Men have been found to have fewer coping strategies when it comes to bladder management and may restrict fluid intake in order to manage urinary frequency, resulting in dehydration [19], however the reason for lower intakes in women is not known. Symptoms such as dysphagia may be responsible for the lower fluid intake in PwP as reported previously [12].

PwP in the current study were found to have double the required intake of protein which confirms findings from 2006, when a smaller study observed similar high intakes of protein [6]. Protein intake interrupts the efficiency of levodopa and therefore PwP are advised to reduce and redistribute their protein intakes. Therefore it may be detrimental for PwP to be consuming above the recommendations for this nutrient, especially if on disease modifying medication for their condition. However, when limiting protein intake, Virmani et al. [7]

found 60% of patients experienced weight loss and therefore this must be monitored closely for optimal effects especially considering the low energy intake particularly in men. This shows that dietary behaviour does not appear to have altered over time and the need for research of possible interventions to support optimal diet for symptom management in PwP.

The interaction between protein restriction for improving GABA and glutamate concentrations in the brain may show potential in neurodegenerative disorders including Parkinson's and therefore may be another incentive for reducing protein intake in those with the condition [20].

Interestingly, intake of vitamin B6 was greater in this sample of PwP compared to the UK recommendations. Low vitamin B6 intake has been shown to increase risk of developing Parkinson's may slow disease progression [21]. Vitamin B6 is an essential cofactor in the conversion of homocysteine to cysteine and alpha-ketobutyrate, and homocysteine is thought to be linked to neurological pathogenesis in Parkinson's. However the vitamin could interact with certain Parkinson's medications and therefore high levels may be harmful to some PwP.

Limitations

In order to provide more comprehensive results a larger sample size would be beneficial from other geographical locations and the collection of more demographic information and possible confounders (smoking, exercise habits etc) would have provided a stronger statistical analysis of the results. Data was self-reported, including diagnosis of PD (no clinical confirmation obtained). The sample may have been biased as those who attend support groups are more likely to be physically able and also to be retired.

The EPIC FFQ is a good indicator of nutritional intake as significant correlations have been found between nutrients derived from the FFQ in comparison to biological analysis markers [22]. Despite it being comprehensive and a detailed assessment of food intake over a year time span, it does have several limitations such as it may not include all foods consumed and/or all cooking methods. Also, intake is not necessarily an indicator of nutritional status due to factors such as bioavailability, nutrient absorption and combination of food consumed [23].

Conclusion

This study provides a current snapshot of comprehensive assessment of the food and nutrient patterns in PwP compared to nutritional guidelines. Research into the diet patterns in this group was shown to be feasible and well received. This study is the first to show that PwP generally have healthy diets compared to the UK guidelines, yet may be unhealthy on some critical dietary components for their condition. The development of clear specific dietary guidance and pathways for implementation may be required to maintain the health of PwP and to help them understand the importance of optimal dietary manipulation for their condition.

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Author contributions

All authors contributed to the writing, data analysis, project design and data collection.

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References 235 236 [1] de Souto Barreto P, Delrieu J, Andrieu S, et al. Physical Activity and Cognitive Function 237 238 in Middle-Aged and Older Adults, Mayo Clin Proceed. 2016;91(11): 1515-1524. 239 [2] Lourida I, Soni M, Thompson-Coon J, et al. Mediterranean Diet, Cognitive Function, and 240 Dementia: A Systematic Review, Epid. 2013; 24(4): 479-489. 241 242 243 [3] Coe S, Axelsson E, Murphy V, et al. Flavonoid rich dark cocoa may improve fatigue in people with Multiple Sclerosis, yet has no effect on glycaemic response: an exploratory trial. 244 245 Clin Nutr ESPN. 2017; doi.org/10.1016/j.clnesp.2017.07.002 246 [4] Erro, R., Brigo, F., Tamburin, S. et al. Nutritional habits, risk, and progression of 247 Parkinson disease. J Neuro. 2017; https://doi.org/10.1007/s00415-017-8639-0 248 249 [5] Seidl SE, Santiago JA, Bilyk H et al. The emerging role of nutrition in Parkinson's 250 disease, Fron Ag Neuro. 2014;6(36). 251 252 [6] Marczewska A, De Notaris R, Sieri S et al. Protein intake in Parkinsonian patients using 253 the EPIC food frequency questionnaire, Mov Disor. 2006;21: 1229-1231. 254 255 [7] Virmani T, Tazan S, Mazzoni P et al. Motor fluctuations due to interaction between 256 dietary protein and levodopa in Parkinson's disease, J Clin MovDisor. 2016;3(8). 257 258

- 259 [8] Sääksjärvi K, Knekt P, Lundqvist A, Männistö S, Heliövaara M, Rissanen H & Järvinen
- 260 R. A cohort study on diet and the risk of Parkinson's disease: The role of food groups and diet
- 261 quality. BJN, 2013: 109(2): 329-337.

- [9] Mischley LK, Lau RC, Bennett RD. Role of Diet and Nutritional Supplements in
- Parkinson's Disease Progression. Ox Med & Cell Long, 2017: 1-9.

265

- 266 [10] Sherzai AZ, Tagliati M, Park K, Pezeghkian S, & Sherzai D. Micronutrients and Risk of
- Parkinson's Disease: A Systematic Review. Ger & Ger Med, 2016: 22(2): 1-12.

268

269 [11] Galland L. Diet and Inflammation, Nutr Clin Pract. 2010;25(6): 634-640.

270

- [12] Cassani E, Barichella M, Ferri V, et al. Dietary habits in Parkinson's disease: Adherence
- to Mediterranean diet. Park & Rela Disor. 2017; 42: 40-46.

273

- 274 [13] Barichella M, Cereda E, Cassani E, et al. Dietary habits and neurological features of
- 275 Parkinson's disease patients: Implications for practice. Clin Nutr. 2017; 36(4):1054-1061.

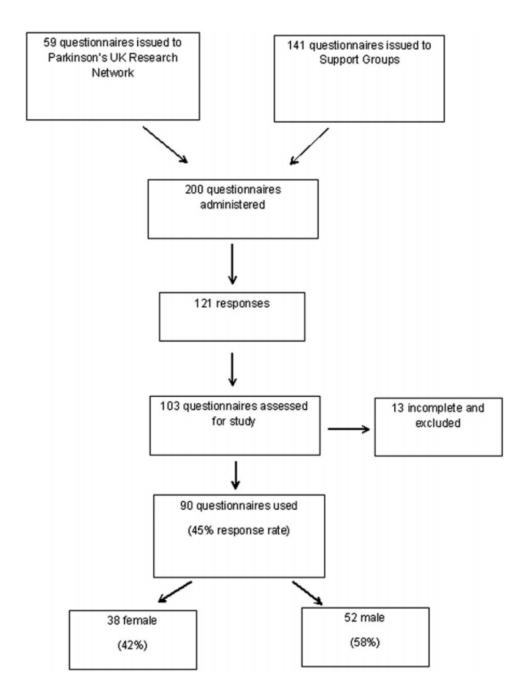
276

- 277 [14] Michley LK, Lau RC, Bennett RD. Role of Diet and Nutritional Supplements in
- 278 Parkinson's Disease Progression. Oxid Med Cell Long. 2017;
- 279 https://doi.org/10.1155/2017/6405278.

280

- 281 [15] Bingham S, Welch A, McTaggart A, et al. Nutritional methods in the European
- 282 Prospective Investigation of Cancer in Norfolk, Pub Health Nutr. 2001;4(3): 847-858.

[16] British Nutrition Foundations (2018) doi: https://www.nutrition.org.uk/ 284 285 [17] Fereshtehnejad SM, Ghazi L, Shafieesabet M, et al. Motor, Psychiatric and Fatigue 286 Features Associated with Nutritional Status and Its Effects on Quality of Life in Parkinson's 287 Disease Patients, Plos One. 2014; https://doi.org/10.1371/journal.pone.0091153. 288 289 290 [18] Benelam B, Wyness L. Hydration and health: a review, Nutr Bull. 2010;35(1): 3-25. 291 292 [19] Collett J, Dawes H, Cavey A, et al. Hydration and independence in activities of daily living in people with multiple sclerosis: a pilot investigation, Disab Rehab. 2011; 33: 1822-293 1825. 294 295 [20] Nayak P, Chatterjee AK. Dietary protein restriction causes modification in aluminum-296 induced alteration in glutamate and GABA system of rat brain. BMC Neurosci. 2003; 297 24(4):4. 298 299 [21] Shen L. Associations between B vitamin sand Parkinson's Disease. Nutrients. 2015; 7, 300 7197-7208. 301 302 303 [22] Sauvageot N, Alkerwi A, Albert A, et al. Use of food frequency questionnaire to assess relationships between dietary habits and cardiovascular risk factors in NESCAV study: 304 validation with biomarkers, Nutr J. 2013;143(12). 305 306 [23] Potischman N. Biologic and Methodological Issues for Nutritional Biomarkers, J Nutr 307 308 2003;133(3): 875S-880S.



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Nutrient	Energy (kcal)	Protein (g) Fiber (g)	Iron (mg)	Zinc (mg)	Caldium (mg) Vit C (mg) Vit D (µg) 86 (mg)	Vit C (mg)	Vit D (µg)	B6 (mg)	B12 (µg)	Phosphorus (mg	Folate (µg)	Chloride (mg)	Non alcoholic beverages (g)	Alcohol (g)
PWP															
Female	1916	83	8	12	9	362	143	m	2	7	1453	327	4315	816	36
Male	2034	88	19	13	9	866	136	m	7	60	1504	327	4190	118	961
SACN															
Female	1912	465	g	8.7	7	8	ę	01	1.2	1.5	250	200	2500	1200	
Male	2342	535	8	8.7	9.5	200	9	9	1.4	1.5	550	300	2500	1200	ı
Mote. Int Whole	ike in PwP ref food groups w	ers to mear ere also cor	data from nsidered. Vit	the 90 Fox t C: vitamin	d Frequency C; Vit D: vita	Questionnaires amin D.	. Nutrients v	vere chosen	from 60 m	utrients an	m 60 nutrients and were those ab	ove or below o	me standard de	viation from the SACN recom	nmendations.