Performance of Monosyllabic vs Multisyllabic Diadochokinetic Exercises in Evaluating Parkinson's Disease Hypokinetic Dysarthria from Fluency Distributions

Pedro Gómez-Vilda¹[®]^a, Andrés Gómez-Rodellar²[®]^b, Daniel Palacios-Alonso³[®]^c

and Athanasios Tsanas²^{od}

¹NeuSpeLab, Center for Biomedical Technology, Universidad Politécnica de Madrid,

Campus de Montegancedo, s/n, 28223, Pozuelo de Alarcón, Madrid, Spain

²Usher Institute, Medical School, University of Edinburgh, Old Medical School, Teviot Place, Edinburgh, EH8 9AG, U.K.

³Escuela Técnica Superior de Ingeniería Informática, Universidad Rey Juan Carlos,

Campus de Móstoles, Tulipán, s/n, 28933 Móstoles, Madrid, Spain

Keywords: Parkinson's Disease, Speech Diadochokinetics, Hypokinetic Dysarthria.

Abstract: Hypokinetic Dysarthria (HD) is a major debilitating symptom in the vast majority of people diagnosed with Parkinson's Disease (PD). It has been traditionally evaluated using diadochokinetic exercises to estimate its degree of severity, among them, the fast repetition of monosyllables as [pa], [ta], and [ka] and multisyllable sequences as [pataka], [pakata], [badaga] and others alike. However, the real efficiency of these exercises in differentiating the participant behaviour as pathological or normative has not been investigated in depth. The aim of the present work is to explore the timely responsive performance of two of these exercises (a monosyllabic [ta] vs a multisyllabic [pataka]). A method to characterize statistically syllabic and inter-syllabic interval durations in the execution of these diadochokinetic exercises, based on Kolmogorov-Smirnov approximations and Jensen-Shannon Divergence has been used to assess the efficiency of both types of exercises. The results from the evaluation of 24 gender-balanced participants (12 PD and 12 controls) show that the monosyllabic exercise does not seem to differentiate well, whereas the multisyllabic exercise has a better differentiation performance. These findings, although relatively preliminary due to the limited sample size, underline the need to carefully consider the battery of tests towards assessing HD.

1 INTRODUCTION

Parkinson's Disease (PD) is a neurodegenerative disorder second in prevalence to Alzheimer's Disease (De Lau and Breteler, 2006). Its origin is mainly caused by the lack of a specific neurotransmitter known as dopamine in midbrain (Dauer and Przedboski, 2003), resulting in relevant neuromotor deterioration affecting body movement (Duffy, 2013). Since the early work of Dr. James Parkinson (Parkinson, 1817) describing observable neuromotor alterations in patients of shaking palsy, including speech problems, most commonly known as

114

Gómez-Vilda, P., Gómez-Rodellar, A., Palacios-Alonso, D. and Tsanas, A.

Hypokinetic Dysarthria (HD). It is a well-established fact that PD causes considerable alterations in speech and phonation (Ricciardi et al., 2016, Brabenec et al., 2017). Roughly, speech alterations may be classified as dysphonia (alterations to the production of voice), dysarthria (alterations in the articulation of speech), dysprosody (alterations in the definition of the fundamental frequency) and dysfluency (alterations in the rhythm and in speech blocking). Although these terms refer to specific and different aspects of anomalous speech production, as all these effects are included in HD, this term will be used for the remainder of this study. The extraction of acoustic markers caused by HD in PD speech allows to

^a https://orcid.org/0000-0003-3283-378X

^b https://orcid.org/0000-0001-8643-9871

^c https://orcid.org/0000-0001-6063-4898

^d https://orcid.org/0000-0002-0994-8100

Performance of Monosyllabic vs Multisyllabic Diadochokinetic Exercises in Evaluating Parkinson's Disease Hypokinetic Dysarthria from Fluency Distributions DOI: 10.5220/0010380301140123

In Proceedings of the 14th International Joint Conference on Biomedical Engineering Systems and Technologies (BIOSTEC 2021) - Volume 4: BIOSIGNALS, pages 114-123 ISBN: 978-989-758-490-9

Copyright © 2021 by SCITEPRESS - Science and Technology Publications, Lda. All rights reserved

conclude that speech analysis might become a noninvasive and cost-effective tool to characterize and monitor PD. There is "compelling evidence to suggest that speech can help quantify not only motor symptoms ... but generalized diverse symptoms in PD" (Tsanas, 2012). There has been a substantial body of work aimed to characterize PD induced HD, focusing on diadochokinetic exercises to assess its degree of severity. Indicative diadochokinetic exercises include the repetition of monosyllables as [pa], [ta], and [ka] and multisyllable sequences as [pataka], [pakata], [badaga] and others of the same nature and function. Exercises consist of repetitions of the sequences as fast as possible, and this setup has been commonly used in PD speech assessment (Ziegler, 2002). The efficiency of these exercises as a way of differentiating participant behaviour as pathological or normative has not been fully evaluated (for a comprehensive review see Karlsson, et al., 2020). The aim of the present study is to explore if the timely responsive evaluation of these exercises may serve as a reliable biomarker or if different or better organized protocols would have a better performance. The main objective of the present study is to compare the performance of two classical diadochokinetic exercises as the repetition of a single syllable [...ta...] where an apical-alveolar pattern is involved, versus the repetition of a multisyllabic sequence as [...pataka...] that presents bilabial and dorsal-velar patterns. These two exercises may allow to properly differentiate between PD participants and Heal Control (HC) participants. The paper is organized as follows: Section 2 is devoted to describe the number of participants in the experimental framework, the speech recording conditions used, the biometrical characteristics of the participants, and the statistical methods used in the study. Section 3 describes the results produced by the statistical analysis of speech recordings. Section 4 focus on analysing and discussing the results. Section 5 summarises the main conclusions derived from the present work.

2 MATERIALS AND METHODS

2.1 Materials

Twelve gender-balanced PD participants were selected from the patient associations of Alcorcón and Leganés (APARKAM). The inclusion conditions for HC participants were non-smoking for the last five years, and not presenting any known laryngeal or neurological diagnosis. The study was approved by

the Ethical Committee of Universidad Politécnica de Madrid. Each participant signed a voluntary participation informed consent. The study was fully aligned with the Declaration of Helsinki. The participants were asked to utter two different exercises, the first one consisting in the repetition of the syllable [ta] at the fastest speed possible and as long as they could sustain it, as [...tatata...] (monosyllabic repetition). The second exercise consisted of repeating the sequence [pataka] as fast and as long as possible. These two sequences were selected for being regular and monosyllabic (the former one) and for involving three different articulation points (bilabial, apical-alveolar and dorsal-velar, the latter). These exercises are especially well suited for the examination of the speaker's fluency, as they do not have any meaning per se. The first one is regular and serves as a reference both for HC and PD participants. The second one invokes the three main articulation points in Spanish (bilabial, dento-alveolar, velar), and it forces the speaker to change facial, lingual, velar and jaw positions, to extract meaningful features from the distribution of time intervals (inter-syllabic and intrasyllabic). The recordings were taken in the speech therapist service room at two different locations of the patient association, no soundproofing or any other quality-preserving measures were undertaken, except keeping a silent environment inside the room with access limited to participants and assistants. The speech recordings were originally sampled at 50 kHz with 16 bits of resolution by a phantom-fed wireless Audio Technica cardioid microphone, and digitized on a Motu Traveller board. The data were downsampled to 8 kHz (antialias filtering at 4 kHz was previously used) to comply with standard telephone channel conditions, making it compatible with remote recordings obtained from a smart phone using the protocol defined in MonParLoc (Palacios et al., 2020). The participants were divided in four data sets for the study: 6 male and 6 female HC participants, and 6 male and 6 female PD participants as shown in Table 1.

Table 1: Participants' biometrical data. MC: male control participants; MP: male PD participants; FC: female control participants; FP: female PD participants; H&Y: Hoehn and Yahr PD rating scale; State: medication state (on: under medication; -: not applicable).

Dataset	Code	Gender	Age	H&Y	State
	MC1	М	69	-	-
	MC2	М	70	-	-
MC	MC3	М	68	-	-
MC	MC4	М	67	-	-
	MC5	М	61	-	-
	MC6	М	68	-	-
	FC1	F	66	-	-
	FC2	F	62	-	-
FC	FC3	F	65	-	-
re	FC4	F	67	-	-
	FC5	F	65	-	-
	FC6	F	65	-	-
	MP1	М	71	2	on
	MP2	М	69	2	on
MP	MP3	М	73	2	on
IVIF	MP4	М	73	2	on
	MP5	М	73	2	on
	MP6	М	69	2	on
	FP1	F	73	2	on
	FP2	F	73	2	on
FP	FP3	F	66	2	on
I.L.	FP4	F	71	2	on
	FP5	F	78	2	on
	FP6	F	70	2	on

2.2 Methods

An experimental framework has been devised to test the relative effects of HD by means of the extraction of syllabic and inter-syllabic interval durations estimated from the speech signal produced by the participants. The main features considered are mean, standard deviation, skewness, and kurtosis of the duration of the syllabic and inter-syllabic (silence) interval distributions, and their normality. The methodology used in the study is based on the estimation of the following acoustic characteristics of the speech recordings:

- The energy profile estimated using the Teager-Kaiser Energy Operator (TKEO, Dimitriadis, Potamianos and Maragos, 2009).
- The glottal residual using the Iterative Adaptive Inverse Filtering (IAIF, Alku et al., 2019).
- The Voiced-Unvoiced Intervals (VUI) using the zero-crossings function of the Linear Prediction (LP) residual.

All these characteristics can be considered correlates showing relevant semantic clues present in speech, that affect the quality of phonation, the prosody and the fluency. The TKO, and the VUI, are defined as

$$E_{\text{TKO}}(n) = s^{2}(n) - s(n+1)s(n-1)$$

$$F_{\text{VIII}}(n) = \dim (z(n));$$
(1)

where

$$r(n) = \begin{cases} 1; s(n) > 0; \\ 0; \text{ otherwise} \end{cases}$$

$$q(n) = r(n) - r(n-1)$$

$$z(n) = \begin{cases} q(n); q(n) \neq 0 \\ 0; \text{ otherwise} \end{cases}$$
(2)

The TKO and the VUI may be used to determine the inferior and superior syllabic interval limits as

$$G_{\text{VAD}}(n) = E_{\text{TKO}}(n)F_{\text{VUI}}(n)$$

$$n_{i}^{\text{lh}} = n \begin{cases} G_{\text{VAD}}(n-1) < \vartheta \\ and \ G_{\text{VAD}}(n) \ge \vartheta \end{cases}$$
(3)
$$n_{i}^{\text{hl}} = n \begin{cases} G_{\text{VAD}}(n-1) \ge \vartheta \\ and \ G_{\text{VAD}}(n) < \vartheta \end{cases}$$

to divide the speech signal produced by diadochokinetic exercise into syllabic (Sy) and silence (Si) intervals, containing the interval duration of syllabic segments $d_{sv}(i)$ and silence segments $d_{si}(i)$

$$d_{Sy}(i) = n_i^{hl} - n_i^{lh}$$

$$d_{Si}(i) = n_{i+1}^{lh} - n_i^{hl}$$

$$\forall d_{Sy}(i) > 20 ms$$
(4)

The normalized distributions of the syllable and silence interval durations might be considered good candidates to establish a differentiation protocol between the behaviour of PD and HC participants in mutual information terms(Cover and Thomas, 2006), using the Jensen-Shannon Divergence (JSD). The resulting sequences of syllable and silence interval durations as $d_{sy}(i)$ and $d_{si}(i)$ are approximated as Kolmogorov-Smirnov distribution densities $p_{sy}(i)$ and $p_{si}(i)$ following Simard & L'Ecuyer, 2011. For each probability density a distance to the average HC subsets ($p_{msy}(i)$ and $p_{msi}(i)$ for males, and $p_{fsy}(i)$ and $p_{fsi}(i)$ for females) was obtained using the JSD (see Gómez et al., 2019 for a detailed description of the JSD estimation).

3 RESULTS

The speech signal produced by each participant are split into the corresponding diadochokinetic exercises [...tatata...] and [...pataka...]. Then they are segmented into intervals with speech activity (syllables) and with no speech activity (silences) using the TKO and VUI indexes. One example of a segmented speech sequence from a diadochokinetic exercise can be seen in Figure 1.

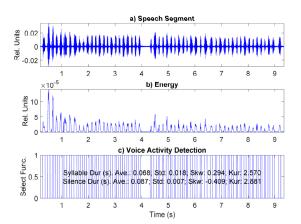


Figure 1: Segmentation of the speech signal corresponding to the diadochokinetic exercise [...pataka...] from a male HC participant (MC6): a) Speech segment under analysis; b) Results of TKO and VUI; c) Voice activity detection (segmentation into syllable and silence durations).

The distributions from two diadochokinetic exercises (...tatata... and ...pataka...) uttered by the male and female HC and PD participants are shown in Figure 2 to Figure 9.

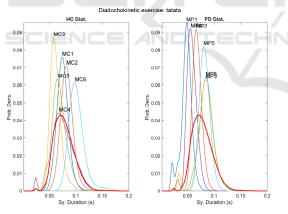


Figure 2: Male Syllable sequence distributions from [...tatata...]. HC (left) and PD (right).

The distribution from the average HC densities is shown in red in both plots for an easy comparison. For the following figures the representation conditions are the same as in figure 2.

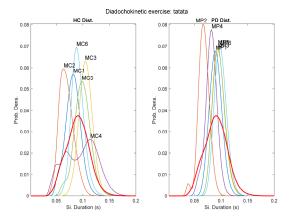


Figure 3: Male Silence sequence distributions from [...tatata...]. HC (left) and PD (right).

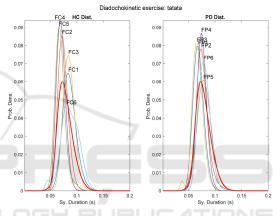


Figure 4: Female Syllable sequence distributions from [...tatata...]. HC (left) and PD (right).

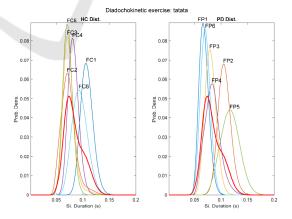


Figure 5: Female Silence sequence distributions from [...tatata...]. HC (left) and PD (right).

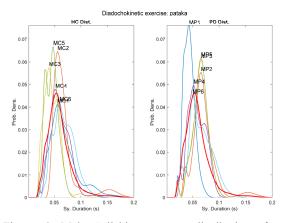


Figure 6: Male Syllable sequence distributions from [...pataka...]. HC (left) and PD (right).

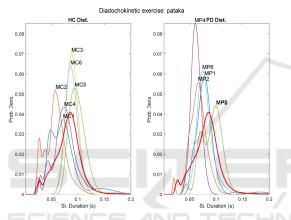


Figure 7: Male Silence sequence distributions from [...pataka...]. HC (left) and PD (right).

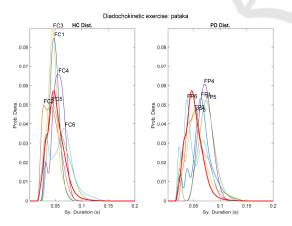


Figure 8: Female Syllable sequence distributions from [...pataka...]. HC (left) and PD (right).

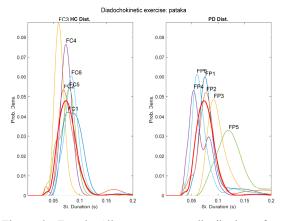


Figure 9: Female Silence sequence distributions from [...pataka...]. HC (left) and PD (right).

Complementary descriptions of the fluency sequence of syllables and silences by the male and female datasets (HC vs PD participants) are given in Table 2 to Table 9. The provided normality tests and the Jensen-Shannon divergence are with respect to the HC pool set. The subset gender, sequence and exercise are highlighted in bold.

Table 2: Parametric description of the **male** Syllabic PD and HC distributions, **sequence** [...tatata...].

Code	Ints	Mean	StdDev	Skew	Kurt	p-vLil	JSD
MP1	37	0.047	0.009	-1.018	4.430	0.029	0.597
MP2	38	0.067	0.008	-2.190	11.664	0.002	0.335
MP3	23	0.083	0.014	-1.571	7.584	0.103	0.277
MP4	26	0.058	0.006	0.405	2.183	0.244	0.462
MP5	22	0.085	0.011	-0.046	3.755	0.288	0.268
MP6	38	0.082	0.005	-0.039	2.404	0.500	0.326
MC1	42	0.075	0.008	0.115	4.243	0.276	0.242
MC2	38	0.083	0.009	0.343	4.953	0.427	0.277
MC3	54	0.062	0.009	1.142	5.317	0.079	0.406
MC4	43	0.076	0.018	-0.085	3.235	0.500	0.040
MC5	45	0.068	0.012	0.066	2.820	0.500	0.223
MC6	36	0.101	0.012	1.991	9.594	0.030	0.489

The first column from the left (code) gives each participant's code according to its gender (M: males, F: females), health condition (C: HC, P: PD), and a consecutive number from 1 to 6. The second column (Ints) give the number of syllable or silence intervals detected in each sample utterance. The third column gives the value of the mean interval in seconds. The fourth column gives its standard deviation in seconds. The fifth column (Skew) gives the skewness distribution, and the sixth (Kurt) one gives its kurtosis. The seventh column (p-vLil) gives the p-value of Lilliefors' hypothesis test of the distribution being normal (H0) on the confidence value of 0.05 (p-

value<0 means rejecting the normality hypothesis). The eighth column (JDS) gives the Jensen-Shannon distance of the sample distribution with respect to the average pool of HC distributions. The data in bold refer to the minimum and maximum of columns 2-4 and 8, and to the distributions rejecting H0 (5-7).

Table 3: Parametric description of the **male** Silence PD and HC distributions, **sequence** [...tatata...].

Code	Ints	Mean	StdDev	Skew	Kurt	p-vLil	JSD
MP1	36	0.089	0.007	-0.463	2.543	0.147	0.267
MP2	37	0.067	0.008	-1.009	5.262	0.500	0.493
MP3	22	0.096	0.007	0.278	2.641	0.500	0.272
MP4	25	0.081	0.008	-1.430	5.102	0.104	0.331
MP5	21	0.095	0.007	0.342	2.937	0.500	0.278
MP6	37	0.099	0.006	-0.126	2.086	0.463	0.304
MC1	41	0.081	0.012	-0.376	3.006	0.352	0.241
MC2	37	0.069	0.012	0.253	2.516	0.500	0.432
MC3	53	0.105	0.007	-1.080	4.170	0.181	0.357
MC4	42	0.096	0.028	-0.246	1.932	0.168	0.247
MC5	44	0.098	0.012	-0.626	3.995	0.245	0.223
MC6	35	0.090	0.008	0.331	4.166	0.420	0.266

Table 4: Parametric description of the **female** Syllabic PD and HC distributions, **sequence** [...tatata...].

Code	Ints	Mean	StdDev	Skew	Kurt	p-vLil	JSD
FP1	34	0.069	0.007	0.017	2.620	0.500	0.277
FP2	41	0.076	0.009	0.740	5.701	0.018	0.118
FP3	- 39	0.067	0.008	-0.598	5.224	0.221	0.325
FP4	28	0.074	0.006	-1.222	4.904	0.053	0.167
FP5	29	0.080	0.011	0.132	2.983	0.500	0.090
FP6	44	0.079	0.007	0.007	2.345	0.406	0.140
FC1	29	0.086	0.010	0.203	3.100	0.500	0.253
FC2	30	0.074	0.006	0.449	2.928	0.221	0.175
FC3	43	0.085	0.006	-0.265	2.399	0.500	0.267
FC4	37	0.070	0.005	0.002	2.302	0.500	0.275
FC5	45	0.069	0.007	1.352	6.635	0.002	0.316
FC6	30	0.086	0.015	-0.592	3.137	0.108	0.253

Table 5: Parametric description of the **female** Silence PD and HC distributions, **sequence** [...tatata...].

Code	Ints	Mean	StdDev	Skew	Kurt	p-vLil	JSD
FP1	33	0.068	0.006	0.298	2.318	0.500	0.379
FP2	40	0.106	0.006	-0.242	6.016	0.473	0.515
FP3	38	0.080	0.007	-0.202	2.476	0.500	0.229
FP4	27	0.083	0.012	0.319	3.131	0.417	0.153
FP5	28	0.116	0.013	-0.343	2.339	0.364	0.525
FP6	43	0.070	0.006	-0.460	2.824	0.500	0.322
FC1	28	0.107	0.005	0.076	2.583	0.380	0.527
FC2	29	0.073	0.013	1.028	5.076	0.222	0.239
FC3	42	0.073	0.009	1.589	8.805	0.028	0.269
FC4	36	0.082	0.006	0.289	3.022	0.500	0.279
FC5	44	0.071	0.005	-0.196	2.552	0.269	0.310
FC6	29	0.094	0.011	-0.029	2.357	0.500	0.309

Table 6: Parametric description of the **male** Syllabic PD and HC distributions, **sequence** [...**pataka...**].

Code	Ints	Mean	StdDev	Skew	Kurt	p-vLil	JSD
MP1	37	0.044	0.010	0.299	3.017	0.500	0.350
MP2	47	0.070	0.024	1.666	7.982	0.002	0.300
MP3	35	0.065	0.013	-0.270	2.748	0.277	0.286
MP4	52	0.058	0.015	0.014	2.034	0.275	0.117
MP5	35	0.065	0.012	-0.207	2.779	0.500	0.288
MP6	58	0.066	0.021	0.325	2.570	0.074	0.169
MC1	52	0.067	0.027	1.233	4.706	0.041	0.123
MC2	46	0.069	0.026	2.367	8.017	0.001	0.232
MC3	52	0.046	0.014	0.900	4.940	0.200	0.253
MC4	78	0.060	0.017	0.561	3.981	0.384	0.096
MC5	36	0.045	0.011	0.090	2.287	0.374	0.283
MC6	55	0.068	0.018	0.269	2.485	0.166	0.224

Table 7: Parametric description of the **male** Silence PD and HC distributions, **sequence** [...**pataka...**].

	Code	Ints	Mean	StdDev	Skew	Kurt	p-vLil	JSD
	MP1	36	0.080	0.021	3.330	17.749	0.001	0.247
	MP2	46	0.071	0.028	3.214	17.040	0.001	0.368
	MP3	34	0.090	0.017	-0.705	2.644	0.001	0.233
	MP4	51	0.062	0.007	-0.334	3.315	0.500	0.503
	MP5	34	0.090	0.017	-0.703	2.645	0.001	0.235
	MP6	57	0.083	0.028	4.117	23.219	0.001	0.259
	MC1	51	0.073	0.029	1.388	5.587	0.007	0.278
	MC2	43	0.058	0.018	0.184	3.022	0.233	0.440
_	MC3	51	0.091	0.018	5.351	35.660	0.001	0.280
1	MC4	76	0.074	0.021	0.133	3.488	0.114	0.177
	MC5	35	0.098	0.011	0.423	2.487	0.420	0.335
	MC6	54	0.096	0.043	5.962	40.379	0.001	0.223
		99		JBC		АТ		15

Table 8: Parametric description of the **female** Syllabic PD and HC distributions, **sequence** [...pataka...].

Code	Ints	Mean	StdDev	Skew	Kurt	p-vLil	JSD
FP1	45	0.065	0.020	0.134	3.594	0.266	0.371
FP2	68	0.056	0.017	0.454	2.661	0.277	0.129
FP3	37	0.056	0.021	0.912	3.981	0.278	0.138
FP4	33	0.072	0.011	-0.043	2.404	0.500	0.498
FP5	30	0.072	0.018	-0.919	4.466	0.247	0.489
FP6	61	0.053	0.020	0.228	1.881	0.004	0.245
FC1	38	0.047	0.008	-0.145	2.794	0.500	0.191
FC2	49	0.043	0.014	0.313	2.208	0.500	0.187
FC3	38	0.044	0.009	0.141	3.405	0.378	0.212
FC4	24	0.055	0.010	-0.622	2.704	0.342	0.231
FC5	52	0.057	0.016	0.569	2.527	0.088	0.132
FC6	35	0.058	0.022	0.357	2.846	0.500	0.263

An important estimate to be considered in analysing the data presented in Table 2 to Table 9 is the average of each sequence interval mean accordingly to gender, condition, sequence, and exercise type, as given in Table 10. The smaller and larger interval duration averages in seconds are given in bold.

Code	Ints	Mean	StdDev	Skew	Kurt	p-vLil	JSD
FP1	44	0.088	0.035	2.854	12.121	0.001	0.146
FP2	67	0.095	0.052	2.774	11.722	0.001	0.120
FP3	37	0.101	0.027	1.682	8.820	0.002	0.424
FP4	32	0.065	0.017	0.370	2.048	0.046	0.300
FP5	29	0.127	0.031	1.217	4.653	0.056	0.588
FP6	60	0.075	0.040	3.923	21.516	0.001	0.309
FC1	37	0.086	0.014	-0.060	1.823	0.151	0.231
FC2	47	0.081	0.026	2.125	7.966	0.001	0.094
FC3	37	0.061	0.008	-0.142	4.521	0.500	0.419
FC4	23	0.074	0.008	-0.536	3.211	0.500	0.204
FC5	51	0.077	0.022	3.490	21.763	0.001	0.116
FC6	34	0.087	0.020	3.084	15.917	0.001	0.235

Table 9: Parametric description of the **female** Silence PD and HC distributions, **sequence** [...pataka...].

Table 10: Interval averages in ms by category (out of 1964 syllabic and 1913 silence intervals).

#Col	Gen.	Cond.	Seq.	Exer.	Averages (ms)
1	Μ	HC	Syl.	tatata	0.077
2	Μ	HC	Syl.	pataka	0.059
3	Μ	HC	Sil.	tatata	0.090
4	Μ	HC	Sil.	pataka	0.082
5	Μ	PD	Syl.	tatata	0.070
6	Μ	PD	Syl.	pataka	0.061
7	Μ	PD	Sil.	tatata	0.088
8	Μ	PD	Sil.	pataka	0.079
9	F	HC	Syl.	tatata	0.078
10	F	HC	Syl.	pataka	0.051
11	F	HC	Sil.	tatata	0.083
12	F	HC	Sil.	pataka	0.078
13	F	PD	Syl.	tatata	0.074
14	F	PD	Syl.	pataka	0.062
15	F	PD	Sil.	tatata	0.087
16	F	PD	Sil.	pataka	0.092

Table 11: Number of samples not rejecting Lilliefors' normality hypothesis test (out of 48).

Category	Туре	#Samples	Percent (%)
By Exercise	tatata	42	87.5
	pataka	28	58.3
By Sequence	Sil.	31	64.6
	Syl.	39	81.3
By Gender	F	36	75.0
	М	34	70.8
By Condition	HC	37	77.1
	PD	33	68.8

The number of sample utterances not rejecting H0 according to the categories of exercise, sequence, gender, and condition is given in Table 11.

A summary of the regularity of each subset in terms of number of distributions not rejecting the null hypothesis, and their comparisons with respect to the HC averages is given in Table 12. Table 12: Results of comparing the JSD and StdDev with the HC subset averages. #Norm: number of samples not rejecting the normality hypothesis. #>AvJSD: number of samples over the average JSD of the HC subset. #>AvStd: Idem over the average StdDev of the HC subset.

#	Exer.	Seq.	Gen.	Cond.	#Norm	#>AvJSD	#>AvStd
1	tatata	Sil.	F	PD	6	3	2
2	tatata	Sil.	F	HC	5	1	3
3	tatata	Syl.	F	PD	5	2	3
4	tatata	Syl.	F	HC	5	3	2
5	tatata	Sil.	М	PD	6	3	0
6	tatata	Sil.	М	HC	6	2	1
7	tatata	Syl.	М	PD	4	4	1
8	tatata	Syl.	М	HC	5	2	3
9	pataka	Sil.	F	PD	1	4	6
10	pataka	Sil.	F	HC	3	3	3
11	pataka	Syl.	F	PD	5	4	5
12	pataka	Syl.	F	HC	6	3	3
13	pataka	Sil.	М	PD	1	2	2
14	pataka	Sil.	М	HC	3	2	2
15	pataka	Syl.	М	PD	5	4	2
16	pataka	Syl.	Μ	HC	4	4	2

4 **DISCUSSION**

The review by exercise, condition, gender and sequence is explained in what follows. It may be seen in Figure 2, corresponding to the monosyllabic repetition [...tatata...], that contrary to expectations, the HC distributions are more spread and over their average distribution (in red) than the PD counterparts, which are slender (concentrated). This observation may be related with the effects of repetitive regular cue rates in stabilizing the movements in PD patients (Harrison, Horin and Earhart, 2019). The sequence distribution of silence intervals in Figure 3, shows a similar behaviour, the HC distributions being more widespread than that of PD participants. Interestingly, the sequence distributions of syllable intervals in Figure 4 by female participants shows little dispersion and good alignment with the average of HC distributions. The situation is completely different regarding the sequence distributions of silence intervals shown in Figure 5, which shows a much less organized pattern of more widespread distributions. When examining the results of the multisyllabic exercise [...pataka...] the distributions become more widespread, and many of them exhibit multimodal behaviour, something not observed in the monosyllabic exercise. This behaviour may be seen in Figure 6 and Figure 7, corresponding to syllable and silence sequence distributions from the male HC

and PD datasets, although in this last figure the PD distributions are more regularly aligned than the HC counterpart. Figure 8 shows the syllable sequence distributions from females, and in this case the HC subset is more regular than the PD subset. This behaviour is also evident in Figure 9, where the PD subset is much less organized than the HC. This could be a consequence of the less regularity observed in the repetitive pattern when multisyllabic repetition is required.

The visual information provided by figures is complemented with the tabulated parameters from each distribution given in Table 2 to Table 9. Additional relevant information is provided, as the number of intervals produced by each speaker, which depends on different factors, respiratory capacity among them. Table 2 shows that the smallest and largest number of intervals, means and dispersion correspond to different members of the PD and HC subsets, respectively. The largest JSD corresponds to a PD participant, and the smallest to an HC one. Two distributions reject the null hypothesis in the PD subset (MP1, which shows the smallest syllabic interval mean, and the largest JSD to the HC average). The examination of Table 3 shows that the smallest and largest number of intervals and means correspond to a member of the PD and the HC, respectively. The largest JSD is from a PD participant, and the smallest one from an HC participant. In this case, none of the distributions reject the null hypothesis. Table 4 shows a similar behaviour regarding the number of intervals and their means, but the situation is quite different as far as JSD is concerned. In this last case, the smallest and largest distances are found in the PD dataset. The number of distributions rejecting the null hypothesis is two, one in the PD dataset and one in the HC dataset. The situation reported in Table 5 shows that the smallest number of intervals is produced by a member of the PD subset, and the largest by an HC member, but the smallest and largest interval means correspond to members from the PD subset. Contrary to what may have been expected, the largest JSD corresponds to a member of the HC subset, and the shortest JSD to a member of the PD subset (FC1). The reason has to be found in the separation of the silence sequence distribution of FC1 with respect to the HC subset average. The number of distributions rejecting H0 is two, one from the PD subset, and one from the HC subset. The multisyllabic exercises reflect a more irregular situation. Table 6 shows that the smallest and largest number of intervals correspond to the PD and HC subsets, respectively. The smallest and

largest means are both from the PD subset. The smallest and largest standard deviations are from the PD and HC subsets, respectively. The largest and smallest JSDs are from the PD and HC subsets, respectively. The number of distributions rejecting the null hypothesis is three, one from the PD and two from the HC subsets. But the situation becomes much more irregular when examining the distributions of the silence intervals given in Table 7. Whereas the smallest and largest number of intervals correspond to the PD and HC subsets, respectively, the smallest and largest means are both from the HC subset. The largest and smallest JSDs are from the PD and HC subsets, as it could be expected, but the number of distributions rejecting the null hypothesis is eight, five from the PD subset and three from the HC subset. Would this mean that males have more problems in separating syllables in a regular way when facing a multisyllabic exercise? Table 8 shows a less irregular situation, although the largest and smallest number of intervals are produced by members of the PD subset, whereas the largest and smallest means correspond to two members of the PD subset and a member of the HC subset, respectively. The largest and smallest JSDs are produced by two distributions of the PD subset. Only one distribution from the PD dataset rejects the null hypothesis. Finally, the situation described in Figure 9 shows again an irregular behaviour as far as the separation of syllables by females facing multisyllabic repetitions is concerned. Contrary to what could be expected, the largest and smallest numbers of intervals were produced by a PD subset member and by an HC one, respectively. The largest and smallest silence interval mean duration were produced by two members of the PD subset. The largest and smallest JSDs corresponded to a PD subset member and to an HC member, respectively. But the most remarkable observation is that the number of distributions rejecting the null hypothesis is again eight, five from the PD subset, and three from the HC subset, showing a striking resemblance with the male cases described in Table 7.

The averages of all speakers' mean interval duration are summarized Table 10 by gender, condition, duration sequence, and exercise. It may be observed that in all cases but one, the average duration of syllables and silences from the monosyllabic exercise are longer than those from the multisyllabic exercise, with the exception of the silence sequences produced by female PD participants (rows 15 vs 16). The duration of syllables is shorter than the duration of silences comparing by

gender and condition. Compare for instance, row 1 against row 3 (0.077 ms vs 0.090 ms), and row 2 against row 4 (0.059 ms vs 0.082 ms). The results given in 0 help in explaining the regular behaviour of the data derived from the two diadochokinetic exercises, in terms of the statistical characteristics of their interval distributions. The largest number of sequences with distributions not rejecting the normality hypothesis is attributed to the sequence [...tatata...], whereas [...pataka...] produced less distributions fulfilling the same condition. This fact explains the difficulty of HC and PD participants in producing regular intervals when facing a relatively more complicate exercise. Interpreting the normality behaviour of distributions as a hallmark of regularity, the most regular subsets correspond to silence intervals from the monosyllabic exercise. The multisyllabic exercise produced more normal-like distributions for syllable than for silence intervals. Therefore normality tests to differentiate PD from HC behaviour might work better with the multisyllabic exercise than with the monosyllabic one. In this same respect, assuming that the JSD would be used as a feature, the most efficient exercises would correspond to the ones marked in bold in column #>AvJSD (number of PD participants producing a JSD larger than the average of the HC subset, compared to HC participants, as given by rows 7 vs 8, 9 vs 10 and 11 vs 12, where the number of participants from the PD subset showing larger JSDs are well over the number of HC participants in the same case. Proceeding similarly with respect to standard deviations, as given in column #>AvStd, the most differentiating conditions are the ones given by row 9 vs 10 and 11 vs 12. Therefore, the best candidates to be checked in a further study would be the syllabic interval durations from the monosyllabic exercise in the case of males, and the syllabic and silence interval durations from the multisyllabic exercise in the case of females.

It might be inferred from what has been exposed that fast purely repetitive exercises provide a timely cadence to PD participants which help them in successful fast repetition. On the contrary, mixedsyllable exercises require a conscious control of syllable sequence repetition (sequence planning and executing), presenting an added difficulty for HC and PD participants, but HC participants seem to utter the multisyllabic sequence faster. The added difficulty found in the multisyllabic exercise may be related to the extra difficulty found in executing the neuromotor changes implied in the articulation from bilabial [pa] (orofacial) to apical-alveolar [ta] (lingual) and dorsalvelar [ka] (lingual-pharyngeal), involving quite different muscular systems. These facts which would explain the complete differential behaviour in exercise planning and execution, from the purely repetitive to the alternating planning, and the control of complete different neuromotor pathways and units.

The main weakness of this study is its nonconclusive character, as the data sample examined is quite low, but it has an exploratory value to initiate a larger scale study. The slightly difference in the age range of HC and PD participants might introduce some bias in the comparisons as well, this fact needing a further study as highlighted by Gómez et al., 2019.

5 CONCLUSIONS

The study of the potential capabilities of repetitive spoken diadochokinetic exercises might derive important benefits to plan speech databases and machine learning methods to characterize PD. The main findings in this study are the following:

- Multisyllabic exercises appear to have more discriminatory power compared to monosyllabic exercises when it comes to assessing PD vs HC..
- PD participants would produce good and regular syllable and inter-syllable intervals, and at a faster repetition rate when monosyllabic exercises are concerned. This fact should be carefully considered when analysing the differentiation capability of monosyllabic exercises.
- On the contrary, PD participants could face more difficulties when multisyllabic exercises are used, therefore these exercises should be prioritized when used combined with other diadochokinetic exercises in test design and analysis.
- The statistical behaviour of syllabic and intersyllabic interval sequence durations of multisyllabic exercises deviates from normality, therefore the statistical evaluation of these tests must stand on non-parametric methods.
- JSDs seem to be sufficiently sensitive to be used in establishing standard syllable and silence interval durations from distance estimations among duration distributions.

This last aspect was not included *per se* as an objective of the study, however the methodology we have used does not rely on expensive high quality equipment that most studies in the field rely on. This concept is very much aligned with the spirit of the Parkinson's Voice Initiative (PVI), see Arora, Bahai-Ravary and Tsanas, 2019, and the exploration of work

trying to facilitate low-cost, robust assessment of PD using readily available means. In this sense we are working on extending these findings on the PVI database.

ACKNOWLEDGEMENTS

This work is being funded by grants TEC2016-77791-C4-4-R from the Government of Spain, and CENIE_TECA-PARK_55_02 INTERREG V-A Spain – Portugal (POCTEP).

REFERENCES

- Alku, P., Murtola, T., Malinen, J., Kuortti, J., Story, B., Airaksinen, M., Salmi, M., Vilkman, E., Geneid, A., 2019. OPENGLOT-An open environment for the evaluation of glottal inverse filtering, Speech Communication 107 38-47, doi: 10.1016/j.specom.2019.01.005
- Arora, S., Baghai-Rivary, L., and Tsanas, A., 2019. Developing a large scale population screening tool for the assessment of Parkinson's disease using telephonequality voice. Journal of the Acoustical Society of America, Vol. 145, pp. 2871-2884.
- Brabenec, L., Mekyska, J., Galaz, Z., and Rektorova, I., 2017. Speech disorders in Parkinson's disease: early diagnostics and effects of medication and brain stimulation. J. Neural Transm., vol. 124:3, pp. 303– 334.
- Cover, T. M. and Thomas, J. A., 2006. Elements of information theory, Wiley, New York.
- Dauer, W. and Przedborski S., 2003. Parkinson's disease: Mechanisms and models. Neuron, vol. 39, pp. 889–909
- Dimitriadis, D., Potamianos, A., and Maragos, P., 2009. A comparison of the Squared Energy and Teager-Kaiser Operators for Short-Term Energy Estimation in Additive Noise. IEEE Trans. on Sig. Proc., vol. 57, No. 7, pp. 2569-2581.
- De Lau, L. M. and Breteler, M. M., 2006. Epidemiology of Parkinson's disease. The Lancet Neurology 5, pp. 525– 535.
- Duffy, J. R., 2013. Motor Speech Disorders, Elsevier, River Lane, St. Louis, Missouri, US.
- Gómez, A., Palacios, D., Ferrández, J. M., Mekyska, J., Álvarez, A., and Gómez, P., 2019. A Methodology to Differentiate Parkinson's Disease and Aging Speech Based on Glottal Flow Acoustic Analysis. Int. Journal of Neural Systems, Vol. 30, 205558.
- Harrison, E. C., Horin, A. P., and Earhart, G. M., 2019. Mental Singing Reduces Gait Variability More than Music Listening for Healthy Older Adults and People With Parkinson Disease. JNPT, Vol. 43, 2019, pp. 204-211.

- Karlsson, F., Schalling, E., Laakso, K., Johansson, K. and Hartelius, L., Assessment of speech impairment in patients with Parkinson's disease from acoustic quantifications of oral diadochokinetic sequences. Journal of the Acoustical Society of America, vol. 147, pp. 839-851.
- Palacios, D., Meléndez, G., López, A., Lázaro, C., Gómez, A., and Gómez, P., 2020. MonParLoc: A Speech-Based System for Parkinson's Disease Analysis and Monitoring. IEEE Access, vol. 8, pp. 188243-188255 doi: 10.1109/ACCESS.2020.3031646.
- Parkinson, J., 1817. An Essay on the Shaking Palsy. J. Neuropsychiatry Clin. Neurosci, Vol. 14:2 pp. 223-236. Ricciardi et al., 2016 (Re-edited in Neuropsychiatry Classics from the 1817 monograph, by Sherwood, Neely and Jones).
- Ricciardi, L., Ebreo, M., Graziosi, A., Barbuto, M., Sorbera, C., Morgante, L., and Morgante, F., 2016. Speech and gait in Parkinson's disease: When rhythm matters. Park. Relat. Disord., vol. 32, pp. 42–47.
- Simard, R. and L'Ecuyer, P., 2011. Computing the Two-Sided Kolmogorov-Smirnov Distribution. Journal of Statistical Software, Vol. 39:11, pp.
- Tsanas, A., 2012. Accurate telemonitoring of Parkinson's disease symptom severity using nonlinear speech signal processing and statistical machine leaning. PhD. Thesis, U. of Oxford, U.K., June 2012.
- Ziegler, W., 2002. Task-Related Factors in Oral Motor Control: Speech and Oral Diadochokinesis in Dysarthria and Apraxia of Speech. Brain and Language, vol. 80, pp. 556-575.