

Review

Gait Kinematic Parameters in Parkinson's Disease: A Systematic Review

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Abstract.

Background: Gait impairments are common and highly disabling for Parkinson's disease (PD) patients. With the development of technology-based tools, it is now possible to measure the spatiotemporal parameters of gait with a reduced margin of error, thereby enabling a more accurate characterization of impairment.

Objective: To summarize and critically appraise the characteristics of technology-based gait analysis in PD and to provide mean and standard deviation values for spatiotemporal gait parameters.

Methods: A systematic review was conducted using the databases CENTRAL, MEDLINE, Embase, and PEDro from their inception to September 2019 to identify all observational and experimental studies conducted in PD or atypical parkinsonism that included a technology-based gait assessment. Two reviewers independently screened citations and extracted data.

Results: We included 95 studies, 82.1% ($n = 78$) reporting a laboratory gait assessment and 61.1% ($n = 58$ studies) using a wearable sensor. The most frequently reported parameters were gait velocity, stride and step length, and cadence. A statistically significant difference was found when comparing the mean values of each of these parameters in PD patients versus healthy controls. No statistically significant differences were found in the mean value of the parameters when comparing wearable versus non-wearable sensors, different types of wearable sensors, and different sensor locations.

Conclusion: Our results provide useful information for performing objective technology-based gait assessment in PD, as well as mean values to better interpret the results. Further studies should explore the clinical meaningfulness of each parameter and how they behave in a free-living context and throughout disease progression.

Keywords: Parkinson's disease, gait, objective assessment, technology, wearable sensor

BACKGROUND

Parkinson's disease (PD) gait impairments increase with disease progression and are a marker of global health, cognition status, falls risk, and institutionalization [1, 2].

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The use of accurate and reliable quantitative information about the mechanics of PD gait is perhaps one of the most promising outcomes that enables early diagnosis, assessment of disease progression and evaluation of therapeutic interventions [3, 4]. In the last decades, with the appearance of technology-based objective measures (TOMs), the evaluation of different spatial and temporal parameters of gait paved the way for a more ecological (i.e., closer to patients' real-life environment performance) and efficient assessment, with a reduced margin of error. Two types of devices have been commonly used: non-wearable sensors (NWS) and wearable sensors (WS) [4]. The NWS are considered the gold standard. They require a controlled and calibrated environment, where individuals walk with skin-mounted markers whose instantaneous positions are obtained using stereophotogrammetry (motion capture) most often based on optoelectronic sensors. WS are small, lightweight sensors (e.g., inertial measurement units) that are attached to one or several body segments, enabling human motion reconstruction in both the context of a laboratory or during activities of daily living [4].

The International Society of Biomechanics has attempted to standardize reports of joint motion in the field of biomechanics for human movement [5]. However, in the PD field, there is a lack of consensus on the best type of sensors and which gait spatiotemporal parameters are clinically relevant. This limits the use of objective measurements of gait in clinical practice and research. [6–8]. Therefore, we aimed to summarize and critically appraise the characteristics of technology-based gait analysis in PD and to provide mean and standard deviation values for spatiotemporal gait parameters.

METHODS

Literature search

We searched CENTRAL, MEDLINE, and PEDro from their inception to September 2019 using “Parkinson*”, “Gait”, “Walking”, “Accelerometer”, “Algorithm” and “Body-fix sensor” as key words. Reference lists from the identified articles were cross-checked to identify any further potentially eligible studies.

Study selection

We included all observational and experimental studies, or study protocols, conducted in PD

patients or atypical parkinsonisms, that included a technology-based gait analysis focused on continuous gait disturbances and that specified which parameters had been studied. There were no restrictions regarding the type of intervention in the active and control arms.

We excluded reviews and studies written in languages other than English, French, Spanish, and Portuguese. All retrieved abstracts were independently screened by two authors. The full texts of potentially relevant articles were retrieved for further assessment. Disagreements were resolved by consensus.

Data extraction

Five pre-defined domains of items were extracted: general information (year and journal of publication, aim of the study, study design, population, intervention, time point assessments, technology development phase), gait assessment supplies (equipment, type of sensor, type of assessment), gait assessment procedures (protocol, medication status, and other outcome tools) and gait parameters values.

According to Maetzler's classification [6], we classified studies according to their technology development phase, which covered three phases: i) preclinical development and testing (those studies focused on how to measure, i.e., testing algorithms or validating a new gait assessment system), ii) clinical development and testing phase (studies focused on the parameters that can be measured and on their clinical relevance) and iii) clinical validation (experimental and observational studies that use gait analysis as an outcome).

We also used an adaptation of the conceptual model of gait presented by Del Din, 2016 [9] to present and analyze the gait parameters reported in the included studies. Parameters that were only reported in one study, and not fitting the model, were included in the “other parameters” section. Data were extracted by two independent authors. Discrepancies were resolved through discussion.

Data analysis

We summarized the publication characteristics using frequencies and percentages. Review Manager software (v 5.3; Cochrane Collaboration) was used for calculating pooled mean difference (MD) and the 95% confidence interval (CI). Heterogeneity was assessed using the Q test and I^2 statistic. An I^2 value of

133 <25% was chosen to represent low heterogeneity and
 134 an I^2 value of >75% to indicate high heterogeneity.
 135 A random-effects model was used to pool all out-
 136 comes. A p -value of <0.05 was considered to be
 137 statistically significant.

138 RESULTS

139 The electronic and hand searches identified 3727
 140 citations. Full-text assessment for eligibility resulted
 141 in 95 studies being included (Fig. 1). Overall, the
 142 main reasons for exclusion were inappropriate study
 143 population ($n=2607$) and inadequately defined out-
 144 come ($n=378$) (Supplementary Material 1).

145 The most common study designs used were
 146 case-control studies (34.7%, $n=33$), cross-sectional
 147 studies (28.4%, $n=27$), and randomized controlled
 148 trials (27.4%, $n=26$). Of the 95 included studies,
 149 61.1% ($n=58$ studies) used WS, 32.6% ($n=31$ stud-
 150 ies) NWS, and 6.3% ($n=6$ studies) both types of
 151 devices. Seventy-eight studies (82.1%) reported a
 152 laboratory gait assessment, 6.3% ($n=6$) a free-living
 153 assessment, and 11.6% ($n=11$) made the assessment
 154 in both contexts (Table 1).

155 Since only two studies [9, 10] presented values for
 156 spatiotemporal gait parameters in free-living assess-
 157 ments, and patients are known to perform differently
 158 in the laboratory and free-living contexts, these values
 159 were excluded from data analysis [11].

160 Gait parameters measured with non-wearable 161 sensors

162 Table 3 lists the gait parameters using NWS
 163 reported in the included studies; the most frequently
 164 used unit of measurement and the mean and standard
 165 deviations of the reported values are also listed.

166 The most frequently reported parameters ($\geq 20\%$
 167 of the studies) were gait velocity (81.1%, $n=30$, PD
 168 mean value = 0.99 ± 0.24 m/s), stride length (56.8%,
 169 $n=21$, PD mean value = 1.06 ± 0.18 m), cadence
 170 (48.7%, $n=18$, PD mean value = 102.71 ± 10.50
 171 steps/min), step length (46.0%, $n=17$, PD mean
 172 value = 0.58 ± 0.13 m), double support phase
 173 (27.0%, $n=10$, PD mean value = $25.89 \pm 7.23\%$)
 174 and step width (24.3%, $n=9$, PD mean
 175 value = 0.13 ± 0.02 m).

176 Gait parameters measured with wearable sensors

177 Table 2 lists the gait parameters assessed with a WS
 178 reported in the included studies; the most frequently
 179 used unit of measurement and the mean and standard
 180 deviations of the reported values are also listed.

181 The more frequently reported parameters
 182 ($\geq 20\%$ of the studies) were gait velocity (60.9%,
 183 $n=39$, PD mean value = 1.01 ± 0.26 m/s),
 184 stride length (37.5%, $n=24$, PD mean
 185 value = 1.14 ± 0.25 m), stride time (28.1%,
 186 $n=18$, PD mean value = 1.18 ± 0.18 s), cadence
 187 (28.1%, $n=18$, PD mean value = 106.42 ± 19.60
 188 steps/min), step length (23.4%, $n=15$, PD mean
 189 value = 0.60 ± 0.06 m), step time (21.9%, $n=14$, PD
 190 mean value = 0.55 ± 0.03 s), stride time variability
 191 (21.9%, $n=14$, PD mean value = $4.33 \pm 2.81\%$
 192 of the coefficient of variation (%CV)) and
 193 step time variability (20.3%, $n=13$, PD mean
 194 value = 0.02 ± 0.00 s).

195 Three studies evaluated gait in a controlled envi-
 196 ronment and nine in a free-living context. Due to both
 197 the low number of studies presenting a value for this
 198 parameter and the heterogeneity of the measurement
 199 units, we did not summarize the data nor present a
 200 reference value.

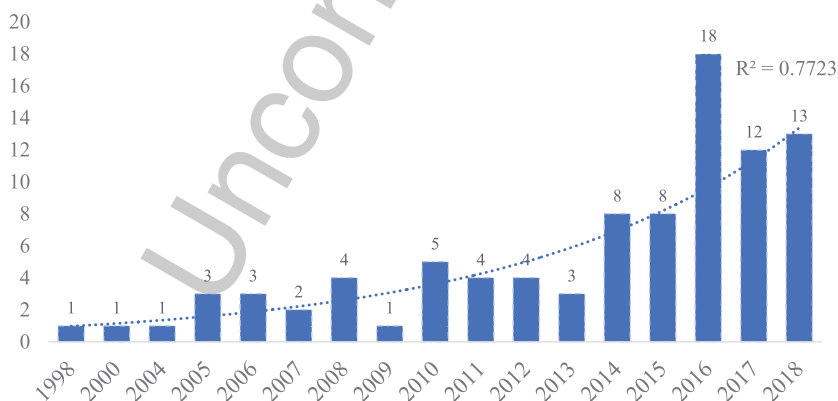


Fig. 1. Number of studies including a technology-based assessment per year in PD.

Table 1
Demographic data, clinical data and mean values of gait parameters assessed with non-wearable devices.
Unk, unknown; NA, not applicable; SD, standard deviation; CV, coefficient of variation

Demographic and clinical characteristics Non-wearable devices (n = 37)						
			PD	HC		
Age (Mean, SD (n))			67.64 ± 4.76 (33)	66.72 ± 5.96 (9)		
Average % Male (Mean, SD (n))			65.25 ± 15.83 (26)	48.14 ± 13.02 (9)		
Height (Mean, SD (n))			1.68 ± 0.07 (13)	1.68 ± 0.05 (7)		
BMI (Mean, SD (n))			26.34 ± 1.92 (17)	27.23 ± 1.68 (7)		
Disease duration (Mean, SD (n))			7.71 ± 2.51 (28)	NA		
UPDRS III (Mean, SD (n))			29.31 ± 8.24 (26)	NA		
Hoehn & Yahr (Mean, SD (n))			2.46 ± 0.40 (27)	NA		
Gait Parameters Mean Values						
Domain	Variable	Studies (n)	Units	Most frequent unit (n,%)	PD mean value (mean, SD (n))	HC mean value (mean, SD (n))
Ambulatory activity	Step count	3	number or mean number of steps	NA	NA	NA
	Gait Velocity	30	km/h, m/s, cm/s	m/s (22, 73.33%)	1.00 ± 0.25 (19)	1.15 ± 0.32 (5)
	Cadence	18	strides/min, steps/min	steps/min (15, 83.33%)	104.04 ± 9.57 (15)	NA
Pace	Stride length	21	cm, m	m (11, 52.38%)	0.99 ± 0.22 (19)	1.20 ± 0.28 (4)
	Stride velocity	1	m/s	NA	NA	NA
	Step length	17	cm, m	m (13, 76.47%)	0.54 ± 0.13 (17)	0.64 ± 0.06 (6)
	Step velocity	2	m/s	m/s (2, 100.00%)	0.98 ± 0.21 (2)	1.10 ± 0.26 (2)
	Stance phase	8	% of gait cycle	% of gait cycle (8, 100.00%)	65.47 ± 3.76 (8)	NA
	Swing phase	5	% of gait cycle	% of gait cycle (4, 80.00%)	34.98 ± 1.92 (4)	NA
	Swing velocity	2	m/s	m/s (2, 100%)	1.73 ± 0.08 (2)	NA
	Double support phase	10	% of gait cycle	% of gait cycle (8, 80.00%)	22.71 ± 8.94 (8)	NA
	Rhythm	Stride time	6	msec, seconds, strides/second	seconds (3, 50.00%)	1.22 ± 0.12 (3)
Step time		6	msec, seconds	seconds (3, 50.00%)	0.60 ± 0.05 (3)	NA
Stance time		4	seconds	seconds (3, 75%)	0.74 ± 0.11 (3)	NA
Swing time		4	msec, seconds	seconds (3, 75.00%)	0.43 ± 0.07 (3)	NA
Double support time		4	msec, seconds	seconds (2, 50.00%)	0.34 ± 0.19 (2)	NA
Variability		Stride time variability	2	SD, % CV	NA	NA
	Stride length variability	2	SD, % CV	NA	NA	NA
	Step length variability	2	m	m (2, 100%)	0.020 ± 0.000 (2)	0.019 ± 0.001 (2)
	Step time variability	3	msec, %CV	NA	NA	NA
	Step velocity variability	1	m/s	NA	NA	NA
	Stance time variability	1	Unk	NA	NA	NA
	Swing time variability	0	NA	NA	NA	NA
	Double support variability	1	%	NA	NA	NA
	Asymmetry	Step time asymmetry	1	Unk	NA	NA
Stance time asymmetry		1	Unk	NA	NA	NA
Swing time asymmetry		1	Unk	NA	NA	NA
Postural control	Step length asymmetry	2	cm, m	NA	0.030 ± 0.014 (2)	NA
	Step width	9	m	m (8, 88.89%)	0.129 ± 0.027 (9)	0.100 ± 0.014 (2)
Other parameters						
Range of motion of shoulder, trunk, hip, pelvis, knee, ankle						
Support base (cm), Latency of postural response to backward translation of center of mass						
Maximal voluntary contraction, rate and peak rate of force development						
Peak heel clearance (mm), Landing (heel) gradient, Take-off toe (gradient), Max and Min toe clearance (mm)						
Magnitude, Smoothness, Attenuation, Regularity, Symmetry, Harmonic ratio						
Fractal index						
Phase Coordination Index (PCI, %), Asymmetry Index						

Table 2
Demographic data, clinical data and mean values of gait parameters assessed with wearable devices.
Unk, unknown; NA, not applicable; SD, standard deviation; CV, coefficient of variation

Demographic and clinical characteristics Wearable devices (n = 64)			PD		HC	
Age (Mean, SD (n))			66.98 ± 6.89 (56)		63.40 ± 13.04 (27)	
Average % Male (Mean, SD (n))			60.69 ± 15.60 (53)		50.42 ± 18.02 (25)	
Height (Mean, SD (n))			1.69 ± 0.04 (27)		1.69 ± 0.06 (14)	
BMI (Mean, SD (n))			25.76 ± 1.42 (35)		25.49 ± 1.77 (17)	
Disease duration (Mean, SD (n))			6.78 ± 5.38 (33)		NA	
UPDRS III (Mean, SD (n))			29.46 ± 12.88 (35)		NA	
Hoehn & Yard (Mean, SD (n))			2.28 ± 0.44 (39)		NA	

Domain	Variable	Studies (n)	Gait Parameters		PD mean value (mean, SD (n))	HC mean value (mean, SD (n))
			Units	Most frequent unit (n,%)		
Ambulatory activity	Step count	12	number of steps, steps/day	number of steps (7, 53.85%)	NA	NA
	Gait Velocity	39	cm/sec, m/sec	m/sec (34, 87.18%)	1.01 ± 0.26 (32); 1.04 ± 0.19 (DT, 8)	1.19 ± 0.31 (17); 1.22 ± 0.1 (DT, 8)
	Cadence	18	Hz, steps/min, steps/sec	steps/min (12, 66.67%)	106.68 ± 20.57 (11)	113.34 ± 7.55 (6)
Pace	Stride length	24	m, cm, % of the stature	meters (17, 70.83%)	1.14 ± 0.28 (18)	1.37 ± 0.08 (8)
	Stride velocity	2	seconds	NA	NA	NA
	Step length	15	cm, m	m (12, 80.00%)	0.55 ± 0.13 (13)	0.61 ± 0.21 (8)
	Step velocity	8	m/sec	m/sec (6, 75.00%)	1.18 ± 0.06 (6)	1.31 ± 0.07 (3)
	Stance phase	2	%	% (2, 100.00%)	60.25 ± 1.76 (2)	57.45 ± 2.75 (2)
	Swing phase	7	% gait cycle	% gait cycle (7, 100%)	36.95 ± 5.11 (7)	39.21 ± 3.62 (4)
	Double support phase	8	% gait cycle	% gait cycle (8, 100%)	29.03 ± 5.00 (8)	23.40 ± 5.83 (6)
	Rhythm	Stride time	18	%, msec, seconds	seconds (14, 77.78%)	1.18 ± 0.18 (12)
	Step time	14	msec, seconds	seconds (7, 50.00%)	0.55 ± 0.03 (7)	0.54 ± 0.02 (4)
	Stance time	9	seconds	seconds (5, 55.56%)	0.74 ± 0.07 (5)	0.71 ± 0.03 (3)
	Swing time	12	msec, seconds	seconds (6, 50.00%)	0.39 ± 0.03 (6)	0.39 ± 0.02 (4)
	Double support time	1	msec	NA	NA	NA
Variability	Stride time variability	14	%CV	% CV (12, 85.71%)	3.84 ± 2.94 (12)	2.18 ± 0.59 (9)
	Step length variability	6	m	m (4, 66.67%)	0.032 ± 0.012 (4)	NA
	Step time variability	13	%CV, msec, seconds	seconds (5, 38.46%)	0.030 ± 0.005 (5)	0.022 ± 0.004 (2)
	Step velocity variability	7	m/sec	m/sec (5, 71.43%)	0.057 ± 0.021 (5)	0.055 ± 0.015 (3)
	Stance time variability	8	%CV, seconds	seconds (4, 50.00%)	0.036 ± 0.015 (4)	0.024 ± 0.003 (2)
	Swing time variability	13	%CV, seconds	% CV (7, 53.85%)	4.714 ± 3.388 (7)	2.481 ± 0.624 (5)
	Double support variability	3	% CV	% CV (3, 100.00%)	9.803 ± 4.617 (3)	6.552 ± 2.224 (3)
Asymmetry	Stride time asymmetry	1	% of stature	NA	NA	NA
	Step time asymmetry	10	msec, sec	seconds (4, 40.00%)	0.021 ± 0.010 (4)	0.011 ± 0.010 (2)
	Stance time asymmetry	7	seconds	seconds (4, 57.1%)	0.021 ± 0.010 (4)	0.011 ± 0.005 (2)
	Swing time asymmetry	9	msec, seconds	seconds (4, 44.44%)	0.020 ± 0.009 (4)	0.012 ± 0.002 (2)
Postural control	Step length asymmetry	8	m	m (6, 75.00%)	0.024 ± 0.011 (6)	0.010 ± 0.004 (3)
	Step width	2	m	m (2, 100.00%)	0.080 ± 0.014 (2)	NA

Other parameters

Ambulatory activity (walking bouts, total time, activity counts/day)

Arm swing amplitude, variability, asymmetry, jerk

Angular velocity of shanks, thighs, trunk and head

Range of head, trunk, shank, thigh and knee rotation

Entropy (measure of variability)

Energy, Power

Magnitude, Smoothness, Attenuation, Regularity, Symmetry, Harmonic ratio, Jerk

SPARC (measure of smoothness)

PD patients versus healthy controls

We were able to perform a forest plot analysis comparing the mean values of PD patients versus healthy controls (HC) for the following gait parameters: gait velocity, cadence, stride length, stride time, stride time variability, step length, step time, swing time, and double support time. All, except step time using WS, presented a statistically significant difference between groups. For gait velocity and stride length, a statistically significant difference between groups was found in WS assessment, but not in the assessment using NWS (Supplementary Material 2).

Wearable versus non-wearable sensors assessment

Comparison between the two types of devices was possible for gait velocity, stride, and step length. While gait velocity presented a statistically significant difference ($p=0.04$, $I^2=76.7\%$), there was no difference between WS and NWS in stride ($p=0.35$, $I^2=0\%$) or step length ($p=0.14$, $I^2=55\%$) (Supplementary Material 2).

Type of wearable sensor

The use of an accelerometer was compared with the use of other types of sensors for gait velocity. The subgroup analysis was not statistically significant ($p=0.18$ and $I^2=44.7\%$). Both groups showed a statistically significant difference between PD and HC ($p \leq 0.05$). The available data did not allow other comparisons for this topic (Supplementary Material 2).

Sensor location

The impact of sensor location (lower back versus feet versus other locations) was studied for gait velocity, stride time, and stride time variability. No differences between groups were registered. Heterogeneity (I^2) ranged between 0–52.9%. All the parameters, except for stride time variability, using the sensor in the lower back, showed a statistically significant difference between PD and HC ($p \leq 0.05$) (Supplementary Material 2).

Sample characteristics

Studies using non-wearable sensors

Eleven studies used a healthy control group. The mean age of PD patients was 67.1 ± 4.8 years

($n=29$ studies) and of 66.3 ± 5.7 years ($n=7$ studies) in HC. The mean percentage of male patients was $63.5 \pm 16.0\%$ for PD ($n=22$ studies) and of 49.0 ± 11.2 for HC ($n=7$ studies). The mean disease duration of PD patients was 7.9 ± 2.3 years ($n=25$ studies). The mean Hoehn and Yahr (HY) score was 2.5 ± 0.4 (77.1%, $n=27$ studies), and the mean motor score for the Unified Parkinson's Disease Rating Scale (UPDRS III) was 28.9 ± 7.9 points (71.4%, $n=25$ studies) (Table 1).

Studies using wearable sensors

Twenty-nine studies used a healthy control group. The mean age of PD patients was 66.8 ± 6.8 years (82.3%, $n=51$ studies) and of 65.1 ± 11.3 in HC (35.5%, $n=22$ studies). The mean percentage of male patients was $60.4 \pm 15.9\%$ for PD (77.4%, $n=48$ studies) and of 47.4 ± 16.2 for HC (30.6%, $n=19$ studies). The mean disease duration of PD patients was 6.7 ± 5.4 years (51.6%, $n=32$ studies). The mean HY score was 2.3 ± 0.4 (61.3%, $n=38$ studies), and the mean motor score for the UPDRS III was 30.0 ± 13.9 points (53.2%, $n=33$ studies) (Table 2).

General characteristics of technology-based gait analysis in PD

From the 95 included studies, according to the technology development phase classification: 24.2% of the studies ($n=23$) were in the preclinical development and testing phase, 31.6% ($n=30$) were in the clinical development and testing phase and 44.2% ($n=42$) belong to the clinical validation phase.

Preclinical development and testing phase

In 56.5% ($n=13$) of the 23 studies, gait assessment was performed in the laboratory, in 17.4% ($n=4$) it was performed in a free-living context, and in 26.1% ($n=6$) it was performed in both contexts.

In 87.0% ($n=20$) WS was used, while 13.0% ($n=3$) used both type of devices. The most common types of sensors were accelerometers (56.5%, $n=13$), accelerometers and gyroscopes (17.4%, $n=4$), only gyroscopes (8.7%, $n=2$) and smartphones (using an accelerometer and gyroscope, 8.7%, $n=2$).

The most common position for the sensor was on the lower back, between the second and fifth lumbar vertebrae (43.5%, $n=10$ of the studies) (Table 3).

Clinical development and testing phase

In 83.3% ($n=25$) of the 30 studies, gait assessment was performed in the laboratory, while in 6.7%

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Table 3
General characteristics of technology-based gait analysis in PD

	Preclinical development and testing	Clinical development and testing	Clinical validation	Total
N	23	30	42	95
Type of assessment				
Lab	13	25	40	78
FL	4	2	0	6
Both	6	3	2	11
Type of device				
Wearable	20	23	15	58
Non wearable	0	5	26	31
Both	3	2	1	6
Type of sensor				
Accelerometer	13	17	9	39
Accelerometer and gyroscope	4	2	3	9
Force-sensitive insoles	0	4	3	7
Accelerometer, gyroscope and magnetometer	1	2	0	3
Gyroscopes	2	0	0	2
Smartphone – Accelerometer and gyroscope	2	0	0	2
Pressure sensor	1	0	0	1
Magnetometers	0	0	1	1
Location of the sensor				
Lower back (L2–L5)	10	18	2	30
Ankles/Feet	3	4	3	10
Lower back and ankles/feet	2	2	5	9
4–6 sensors	3	0	1	4
Other	3	1	0	4
Lower back and wrists	0	0	1	1
Unknown	2	0	4	6
Medication state				
ON-phase medication	5	15	28	48
OFF-phase medication	1	1	5	7
ON- and OFF-phase medication	1	2	1	4
Not described	12	10	8	30
Not applicable (Free-living)	4	2	0	6

($n = 2$) it was performed in a free-living context, and in 10.0% ($n = 3$) it was performed in both contexts.

In 76.7% of the studies ($n = 23$) a WS was used, 16.7% ($n = 5$) used NWS and 6.7% ($n = 2$) used both type of devices. Accelerometer (68.0%, $n = 17$) and force-sensitive insoles (16.0%, $n = 4$) were the most frequently used type of sensor. The most common position for the sensor was in the lower back, between the second and fifth lumbar vertebrae (72.0%, $n = 18$) (Table 3).

Clinical validation phase

The majority of the assessments were performed in the laboratory (95.2%, $n = 40$). NWS was used in 61.9% ($n = 26$) of the studies, a WS in 35.7% ($n = 15$) and both devices in one study. Accelerometers (60.0%, $n = 9$) were the most frequently used type of sensor. The most common position for the sensor was on the lower back and the feet/ankles (33.3%, $n = 5$). (Table 3)

Protocol details

Table 4 shows the characteristics of the gait assessment protocol. The most frequently used distance in laboratory assessments was 10 meters ($n = 23$), the shortest distance reported was 3 meters and the longest 500 meters. Table 5 compares PD patients' gait velocity using a gait assessment protocol with less than 10 meters, 10 meters and more than 10 meters. Due to the heterogeneity of the data, this comparison was only performed for gait velocity and a forest plot analysis was not possible.

The mean number of trials was 4.3 ± 2.9 . In 46.1% of the studies ($n = 41$), gait assessment was performed at a self-selected comfortable speed. In free-living assessments, the most common duration of data collection was 7 days (58.8%, $n = 10$).

In 58.5% of studies ($n = 48$), patients were in an "ON-state" during the assessment, in 7.4% ($n = 7$) in an "OFF-state" and in 4.2% of the studies ($n = 4$) the assessment was performed in both conditions

Table 4

Protocol details of laboratory and free-living gait assessments		
Protocol details		
Laboratory assessment		
Distance	Median [Min, Max in meters]	10 [3,500]
	Mode (n, %)	10 (23, 24.2%)
Trials	Mean, SD	4,52 ± 2,98
	Protocol	
	Self-selected comfortable speed	44
	Self-selected comfortable and dual task	8
	Self-selected comfortable, fast speed and dual task	6
	Self-selected comfortable and fast speed	5
	Self-selected comfortable and cueing	4
	Fast speed	2
	Fast, normal, and slow speed	2
	Other	7
	Unknown	11
Free-living assessment		
Duration	7 days	10
	3 days	3
	10 days	2

Table 5

Analysis of gait speed according to the distance covered in the gait protocol		
	Wearable	Non-Wearable
Less than 10 meters (mean, SD (n))	0.9 ± 0.2 (5)	0.9 ± 0.3 (7)
10 meters (mean, SD (n))	1.0 ± 0.1 (7)	0.9 ± 0.4 (8)
More than 10 meters (mean, SD (n))	1.1 ± 0.3 (18)	NA

(Table 4). Table 6 compares the PD mean values with and without having into account the “ON/OFF”

medication state. Due to the low number of studies assessing gait in “OFF” state medication ($n = 11$, 11.6%) and the heterogeneity of the data, this analysis was only possible to perform for some gait parameters and did not allow for a forest plot analysis. Except for stride time variability, all the mean values of the studies only including an “On” state medication assessment, were closer to those from the HC group.

DISCUSSION

The number of studies including a technology-based gait assessment is increasing (Fig. 1). Of the 95 studies included, the majority performed a laboratory assessment (82.1%, $n = 78$) and used WS (61.1%, $n = 58$). Accelerometers were the most frequently used type of sensor (67.2%, $n = 39$), usually on the lower back (51.7%, $n = 30$). The sample characteristics of the included studies were very similar, not allowing for subgroup analysis.

1) What should be measured?

The most frequently reported parameters in the included studies were gait velocity, stride and step length, and cadence. Compared to HC, PD patients had decreased velocity, reduced stride and step length, decreased swing time, increased stride time, stride time variability and dual support time ($p < 0.05$). These differences are in line with the usual description of PD gait impairments, i.e., a slow, short-stepped, shuffling, with a forward-stooped posture and asymmetrical arm swing [7, 12, 13].

Beyond this, a large number of different, or differently measured gait parameters, were found in the included studies. From a clinical point of view,

Table 6

Analysis of PD gait parameters according to the “ON/OFF” medication state during the gait assessment

	Wearable devices		
	All	“ON” State Medication	Healthy controls
Gait velocity	1.01 ± 0.26 (32)	1.06 ± 0.20 (29)	1.19 ± 0.31 (17)
Cadence	106.68 ± 20.57 (11)	112.33 ± 8.89 (10)	113.34 ± 7.55 (6)
Stride Length	1.14 ± 0.28 (18)	1.15 ± 0.26 (15)	1.37 ± 0.08 (8)
Stride Time	1.18 ± 0.17 (13)	1.18 ± 0.18 (12)	1.09 ± 0.07 (9)
Stride Time Var	3.84 ± 2.94 (12)	4.01 ± 3.02 (11)	2.18 ± 0.59 (9)
Double support phase	29.03 ± 5.00 (8)	29.22 ± 5.37 (7)	23.40 ± 5.83 (6)
	Non-wearable devices		
	All	“ON” State Medication	Healthy controls
Gait velocity	1.00 ± 0.25 (19)	1.01 ± 0.25 (18)	1.15 ± 0.32 (5)
Cadence	104.04 ± 9.57 (15)	105.75 ± 7.15 (14)	NA
Stride Length	0.77 ± 0.40 (19)	0.77 ± 0.43 (17)	1.20 ± 0.28 (4)
Step Length	0.54 ± 0.13 (17)	0.55 ± 0.13 (16)	0.64 ± 0.06 (6)

not every parameter that can be measured should be measured [6, 8]. The collection and interpretation of the data must lead to justified outcomes, i.e., those with an impact on activities of daily living, displayed in a visually intuitive format that covers the clinical information needs of the stakeholders (health professionals, patients, and caregivers) [6, 8]. For this, gait parameters should be correlated with robust measures of clinical meaningfulness, such as the MDS-UPDRS motor score or the Timed Up and Go Test (TUG). Once the most suitable parameters to measure PD gait impairments in different contexts are established, then the minimal clinically important differences should be addressed for each [6, 8]. Other measures emerging from the nonlinear analysis of human variability (e.g., entropy, fractals, and others) can give us a more accurate angle of patients' gait dynamics in a real-life environment. However, work is needed to make them more intuitive and clinically informative [6, 8].

Although currently, sensor-based gait analysis has demonstrated feasibility and applicability for objectively assess PD gait impairments, differences still exist measuring the same parameter, with different devices or devices from different manufacturers [3, 14, 15]. This highlights the difficulty of accurately measuring the spatiotemporal gait parameters and the need to continue developing valid and reliable mathematical algorithms. Despite the major technological advances and the current possibility of capturing and store extremely high amounts of data with TOMs, the ability to algorithmically analyze (eliminating the noise) and summarize the clinically relevant data to stakeholders remains limited. [3]

2) Which devices should be used?

The comparison between assessments using WS and NWS was investigated for gait velocity, stride and step length parameters. A statistically significant difference between groups was found in gait velocity ($p=0.04$). Although it was the analysis with the highest number of studies ($n=18$), due to the level of heterogeneity ($I^2=76.7\%$), the results should be interpreted with caution. We believe that the differences in the type of devices and in the assessment protocols of the included studies might have contributed to this result.

No statistically significant difference was found in the two other parameters (stride length – $p=0.35$, step length – $p=0.14$). Taking into account the low value for heterogeneity ($I^2=0\%$, $p<0.001$), we believe that

wearable sensors can be used in place of NWS (the gold standard of gait analysis).

WS have the added value of enabling the assessment of gait during activities of daily living in the patients' actual environment. However, more studies exploring how gait parameters behave in a real-world context are needed [4].

It was only possible to explore the impact of the type of WS for gait velocity. This was undertaken by comparing the use of accelerometer (used in 67.2% of the WS) with all other types of sensors.

Accelerometers allow the measurement of dynamic accelerations of a body, when submitted to an external force, and provide information about the device orientation related to gravity [3, 14, 15]. They are frequently combined with a gyroscope, which allow for the measurement of angular velocities [3, 14, 15]. In some devices, a 3D-magnetometer is also added for orientation purposes.

Since no difference was found in this subgroup analysis (accelerometer versus all other types of sensors) and both groups were able to detect a statistically significant difference between PD and HC, we believe that for an accurate assessment and monitoring of PD patients' gait impairments, the use of a single accelerometer is feasible. However, for the assessment of turns or of a more complex movement that requires the information captured by angular velocity, wearable devices including at least a gyroscope, seem more suitable.

In the included studies, only one study used an isolated magnetometer for gait analysis. Since magnetometers are very sensitive to magnetic changes (e.g., those produced by proximity with ferromagnetic objects) and therefore to many external interferences, they are more frequently used as a complement to accelerometers and gyroscopes, than as a single sensor [3, 14, 15].

3) Where to place the sensor?

Our results showed that in 46.9% ($n=30$) of the studies using WS, the sensor was used on the lower back, between the second and the fifth lumbar vertebra. Although it was only possible to investigate the impact of sensor location for three parameters, it was limited to the comparison between lower back, feet and all other locations, the results consistently show no statistically significant difference between groups. Stride time variability measured with the sensor in the lower back was the only parameter that did not show a statistically significant difference between PD and

466 HC. However, a heterogeneity (I^2) of 82% was found,
467 whereby these results should be interpreted carefully.

468 Several gait analyses protocols have been used.
469 However, an optimal and standardized method
470 remains for establishing [15]. The number and loca-
471 tion of the sensors are key aspects for the success of
472 assessments with TOMs, especially in a free-living
473 context [8, 16]. To increase wearing compliance with-
474 out hindering the precision of data collection the
475 number of sensors should be kept to a minimum, and
476 the least obtrusive devices preferred [8, 16]. Today,
477 although the lower back is not considered the most
478 comfortable and unobtrusive location, it has been
479 shown that a single sensor (accelerometer) in this
480 location is able to capture with precision, physical
481 activity and gait parameters in a laboratory and free-
482 living context [16, 17]. Recently, there has been a
483 move toward using sensors on the wrist or embedded
484 in smartphones. However, problems still exist when
485 collecting data. Kim et al., 2019 [16] report that sen-
486 sors used on the wrist tend to overestimate the number
487 of steps and the time spent at different intensities of
488 activity. Höchsmann et al., 2018 [18] compared the
489 accuracy of step detection of a smartphone (placed in
490 a trouser pocket, shoulder bag, and backpack) with
491 a WS used on the wrist and waist. At a gait veloci-
492 ty of 4.8 km/h (shoulder bag and backpack) and 6.0
493 km/h (all positions), smartphones did not exceed a
494 1% error deviation from the gold standard (threshold
495 to be considered an accurate measurement). How-
496 ever, for a gait velocity of 1.6 km/h, a 3% error was
497 found. In a free-living context, smartphones underes-
498 timate the number of steps [18]. Another limitation of
499 free-living assessment with smartphones is the place
500 where it is used. While for men a trouser pocket is a
501 commonly preferred position, for women it is more
502 likely to be the purse or backpack [18]. In the search
503 for a solution for a smartphone-based body location
504 the magnetometer sensor will most certainly be a cru-
505 cial sensor to consider when dealing with the device's
506 orientation.

507 4) Which gait assessment protocol

508 The comparison between all the included stud-
509 ies and those that only used an assessment in "ON"
510 state medication, revealed that PD gait parameters
511 under the effect of the medication are closer to the
512 HC. Only stride time variability did not follow this
513 pattern. According to the literature [12], stride time
514 variability is increased in PD patients and diminishes
515 in response to dopaminergic medication. In our anal-

516 ysis, we found that the difference between PD on
517 and HC increased when only studies assessing gait
518 in "ON" state medication, were taking into account.
519 However, this result should be interpreted with cau-
520 tion, since this was only a basic comparison of means
521 and gait protocols differentiated substantially in the
522 included studies.

523 The distance covered during gait analysis varied
524 in the included studies. According to the analysis
525 performed, the distance doesn't seem to have a high
526 impact on gait velocity tested in a controlled envi-
527 ronment. However, the data from the included studies
528 doesn't allow us to conclude on this topic. More stud-
529 ies are needed to understand the implications of gait
530 protocol length in PD gait parameters.

531 Almost half of the included studies (43.2%, $n = 41$)
532 used only a self-selected comfortable speed, during
533 gait assessment. Since some of the gait param-
534 eters, like stride length and cadence, are sensitive to
535 velocity and to the presence of concurrent attention
536 demands, gait assessment protocols should include
537 different velocities and both single- and dual-task
538 activities [19]. The most common duration of free-
539 living assessment data collection was seven days,
540 varying between three and ten days. Based on our
541 results, we cannot conclude if this is the best option.
542 These are challenging assessments due to the het-
543 erogeneity of ambulatory activity within habitual
544 environments. We believe that the duration of data
545 collection during free-living assessments should be
546 a balance between not performing a burdensome
547 assessment and the ability to collect enough and
548 precise data to obtain a pattern of patients' perfor-
549 mance during the day [8]. As a fluctuating disease, the
550 duration applied in other research fields, may not be
551 appropriate. This topic should be addressed in future
552 studies.

553 Conclusion

554 Our results support previous descriptions of PD
555 gait impairments when compared with HC. No sta-
556 tistically significant differences were found for the
557 impact of different types of devices (WS vs NWS),
558 or different types or locations of wearable sensors
559 during assessments. Future studies should test the
560 reported gait parameters against validated clinical
561 meaningful outcome measures in PD to select those
562 most suitable for evaluating and monitoring the pro-
563 gression of gait impairments in PD. More studies are
564 also needed to explore gait parameter behavior in a
565 free-living context, with more complex movements

(e.g., including turns, sequences of movements and others).

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CONFLICT OF INTEREST

The authors declare that there are no conflicts of interest relevant to this work.

SUPPLEMENTARY MATERIAL

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