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Gait metrics analysis utilizing single-point inertial measurement units: a systematic review

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Background: Wearable sensors, particularly accelerometers alone or combined with gyroscopes and magnetometers in an inertial measurement unit (IMU), are a logical alternative for gait analysis. While issues with intrusive and complex sensor placement limit practicality of multi-point IMU systems, single-point IMUs could potentially maximize patient compliance and allow inconspicuous monitoring in daily-living. Therefore, this review aimed to examine the validity of single-point IMUs for gait metrics analysis and identify studies employing them for clinical applications.

Methods: The Preferred Reporting Items for Systematic Reviews and Meta-Analyses Guidelines (PRISMA) were followed utilizing the following databases: PubMed; MEDLINE; EMBASE and Cochrane. Four databases were systematically searched to obtain relevant journal articles focusing on the measurement of gait metrics using single-point IMU sensors.

Results: A total of 90 articles were selected for inclusion. Critical analysis of studies was conducted, and data collected included: sensor type(s); sensor placement; study aim(s); study conclusion(s); gait metrics and methods; and clinical application. Validation research primarily focuses on lower trunk sensors in healthy cohorts. Clinical applications focus on diagnosis and severity assessment, rehabilitation and intervention efficacy and delineating pathological subjects from healthy controls.

Discussion: This review has demonstrated the validity of single-point IMUs for gait metrics analysis and their ability to assist in clinical scenarios. Further validation for continuous monitoring in daily living scenarios and performance in pathological cohorts is required before commercial and clinical uptake can be expected.

Keywords: Accelerometry; gait analysis; wearable electronic devices

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Introduction

Human gait is affected by ageing as well as numerous musculoskeletal and neurological ailments. Consequently,

gait analysis has wide-ranging clinical applications from diagnosis and severity assessment as well as evaluation of intervention and rehabilitation efficacy in neurological and

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orthopedic conditions (1-8) to identifying falls risk and frailty status (9,10). Qualitative and subjective measures generally constitute routine clinical gait analysis, with patient self-reporting and clinician observation sometimes integrated with clinical tests such as the Timed-Up-And-Go and 6-minute walking test (6MWT) (11). These approaches impose significant interobserver inaccuracies and deny appreciation of kinematic and kinetic intricacies that can be obtained from quantitative gait assessment (12).

The gold-standard for quantitative gait analysis, optoelectronic stereophotogrammetry, features infrared cameras that capture three-dimensional trajectories of reflective markers placed on the subject that are processed to accurately assess spatio-temporal and kinematic variables of gait (13). Stereophotogrammetry is often combined with force plates that measure ground-reaction forces (GRF) to determine kinetic forces and electromyography (EMG) systems to measure muscle activity during gait. However, these systems are expensive, time-consuming, and require expert operation and equipment. Furthermore, restriction of their performance to dedicated laboratory settings limits portability, access and external validity of measures obtained to free-living gait (11).

In response, wearable sensors (goniometers, EMG systems, sensing fabric etc.), particularly accelerometers alone or combined with gyroscopes and magnetometers in an inertial measurement unit (IMU), are proving to be the logical alternative for gait analysis. Cheap, small and portable, wearables could potentially enable continuous gait metrics analysis in daily living (14-17). Furthermore, fast preparation and processing negate the need for expert operation, enhancing practicality. Multi-point IMU systems have been validated against standards (18-23) and employed clinically for gait metrics analysis (24-28); however, issues with intrusiveness and consistency of complex sensor placement limit real-life adoption (29). Although not as accurate and reliable as multi-point IMU systems (30), single-point IMUs have nonetheless demonstrated enormous potential. They have clinical uses, such as with the assessment of Parkinson's Disease (PD) severity (31,32) and the evaluation of falls risk in preventative health care (33,34), and personal uses, such as with the tallying of daily steps in consumer-grade watches (4,35). Commonly used single-point IMUs and their specifications are detailed for comparison in *Table 1*.

Moreover, presuming validated accuracy is demonstrated, single-point IMUs could potentially maximize patient compliance and comfort and allow inconspicuous monitoring in daily-living. Consequently, any future mass uptake of IMUs in clinical and commercial settings is likely to be dependent upon the validation and applications of single-point IMUs and not of multi-point IMU systems. Therefore, this review will provide a synopsis of inertial sensor principles, practical considerations and gait metrics analysis capabilities before examining the validity and clinical applications of single-point devices for gait analysis. We present the article in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-Analyses Guidelines (PRISMA) checklist (available at https://dx.doi.org/10.21037/mhealth-21-17).

Inertial sensors: principles and practical considerations

Inertial sensors, accelerometers and gyroscopes, are often fabricated into microelectromechanical systems (MEMS) alone or together as an IMU and employed for gait metrics analysis (78).

Accelerometers measure acceleration along their sensitive axis, ranging from uni-axial to the commonly employed tri-axial sensitivity which allows appreciation of movement along the antero-posterior, horizontal and vertical planes. However, current devices are susceptible to drift errors due to change in mechanical or electrical properties and noise from amplified mechanical motions. Furthermore, the measured acceleration comprises both the inertial acceleration associated with changes in velocity, and gravitational acceleration superimposed along the accelerometer's sensitive axes. Removing this confounding effect of gravity can be difficult (78). To appreciate velocity and distance, numerical integration of acceleration data is required, causing noise and drift errors to accumulate, imposing significant limitations on long-term accelerometer employment. Without compensation for this drift, readings become useless. Compensation requires frequent accelerometer recalibration, achieved through zero-velocity updates (ZUPT), using an external event indicating an instantaneous null in movement such as a footstep (79).

Gyroscopes measure angular velocity, demonstrating greater accuracy than accelerometers as measurement is absolute with no external information considered (80). However, gyroscopes only return rate of change of angular position; to detect relative orientation, integration of the signal is required. This leads to accumulation of drift errors and noise, similar to accelerometers (80). Furthermore, lack of an initial reference compared to accelerometers means gyroscopes cannot be recalibrated, resulting in accumulation of errors and limited long-term

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References	Sensor	Placement	Gait metrics demonstrated to be captured	Component MEMS sensors	Other known specifications
(31,36-40)	Locometrix	Lower back, L3–4	GV and stride frequency, length, symmetry and regularity	Tri-axial accelerometer	100 Hz frequency of data capture, resolution 0.001 g
(41-45)	MT product line IMUs (Xsens, Enschede, The Netherlands)	Lower back, L4	MTx was able to capture GV, cadence, step regularity, stride regularity, gait symmetry, step time, step time variability, RMS and HR	Tri-axial accelerometer, gyroscope, and magnetometer	According to MTi (newer than MTx) specifications: Output frequency up to 2 kHz. Accelerometer range 16 g. Gyroscope range 2,000 deg/s. 12.1×12.1×2.55 mm³ (without encasing). 0.6 grams (without encasing)
(46,47)	Pi-node (Philips, The Netherlands)	Lower back, L4	GV, step time, step time variability, cadence, stride length, stride length variability and non-linear measures (gait variability and symmetry)	Tri-axial accelerometer	100 Hz frequency of data capture
(48-59)	DynaPort IMUs (McRoberts, The Hague, The Netherlands)	Lower back	Step count, GV, stride time variability, stride regularity and cadence	Tri-axial accelerometer, and gyroscope	100 Hz frequency of data capture. 106.6x58x11.5 mm³. 55 grams. 14-day maximum measurement duration
(60-64)	G-walk (BTS Bioengineering Milan, Italy)	Lower back, L5	GV, SL, stride length and cadence. Gait cycle duration, stance duration, swing duration and double support duration	Tri-axial accelerometer, gyroscope, and magnetometer	Accelerometer range 2, 4, 8, or 16 g (configurable). Gyroscope range 250, 500, 1,000, or 2,000 deg/s (configurable). Magnetometer range ±1,200 uT. Accelerometer sampling rate 4–1,000 Hz (configurable). Gyroscope sampling rate 4 to 8,000 Hz (configurable). 37 grams
(65,66)	Physilog product line (Gaitup, Lausanne, Switzerland)	Chest, 5 cm below sternal notch	GV, non-linear measures (gait stability), RMS and walk-ratio	Tri-axial accelerometer and gyroscope	According to Physilog5 specifications: Accelerometer range 16 g. Gyroscope up to 2,000 deg/s. 26.5×10×47.5 mm³
(67,68)	GENEActiv (Activinsights, Kimbolton, England)	Lower back or wrist	Step count, stride time, stride time variability, step time, step time variability and non-linear measures (gait variability)	Tri-axial accelerometer	100 Hz sampling frequency
(34,69-71)	Axivity AX3 (Axivity, York, England)	Lower back, L5	Step count, mean bout length, step time, swing time, stance time, step time asymmetry, swing time asymmetry, SL asymmetry, step velocity, SL, swing time variability, stance time variability, GV and SI variability	Tri-axial accelerometer	Up to 8 g. 100 Hz. Waterproof. 23×32.5×7.6 mm³, 11 g weight. Accelerometer range 2, 4, 8, or 16 g (configurable). Accelerometer sampling rate 12.5–3,200 Hz (configurable)

Table 1 (continued)

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References Sensor	Sensor	Placement	Gait metrics demonstrated to be captured	Component MEMS sensors	Other known specifications
(72)	MIMAMORI-Gait system (LSI Medience Corp, Tokyo, Japan)	Back of waist	GV, SL, cadence, step time, step time Tri-axial accelerometer variability and double support time	Tri-axial accelerometer	100 Hz frequency of data capture 75×50×20 mm³, 120 grams
(73)	activPAL (PAL Technologies, Glasgow, Scotland)	Upper thigh	Step count, cadence	Uni-axial accelerometer	20 Hz frequency of data capture. 53.0×35.0×7.0 mm³. 20 grams
(74,75)	Actigraph wGT3X-BT activity Lateral waist monitor (Actigraph, Pensacola, FL, USA)	Lateral waist	Cadence, stride regularity and non- linear measures (intensity, dynamic stability)	Tri-axial accelerometer	50 Hz frequency of data capture. Acceleration range 8 g
(76,77)	e-AR sensor	Behind ear	Stride duration and step time asymmetry. Gait asymmetry	Tri-axial accelerometer	
(4)	Apple Watch (Apple, San Francisco, CA, USA)	Wrist	Daily step count, gait velocity, estimated caloric expenditure, distance travelled	Tri-axial accelerometer and gyroscope	

MU, inertial measurement unit; g, magnitude of acceleration due to gravity, 9.8 m/s²; MEMS, microelectromechanical systems; GV, gait velocity; SL, step length; RMS, oot-mean square; HR, harmonic ratio. precision (80). This limitation is often minimized by incorporating a magnetometer in the IMU, able to calibrate sensor orientation with reference to the Earth's magnetic field. However, these devices are prone to interference by magnetic fields created by other devices (80).

Inertial sensor-based gait analysis

While single-point inertial sensors are unable to appreciate kinetic and many kinematic variables of gait, they can determine spatio-temporal parameters. Spatiotemporal parameters are of importance clinically, as they objectively characterize key gait events (GE) and common gait abnormalities (81). A plethora of spatio-temporal parameters are employed in the literature, with some [such as gait velocity (GV) and gait regularity in predicting the staging of PD severity being more relevant than others in different clinical scenarios (31,32). This review focusses on spatio-temporal parameters based on a validated model (82,83) and clinical guidelines from The Biomathics and Canadian Gait Consortiums Initiative (84). These parameters encompass the mean, variability and asymmetry of temporal (cadence, step time, stride time, stance duration, swing duration, single-support duration, double-support duration) and spatial [step length (SL), stride length, GV step width] components of gait.

As acceleration data retains a time-series nature when extracted, by determining GEs such as heel-strike (HS) and toe-off (TO) within the gait cycle, mean temporal parameters can be quantified. Methods for GE detection are based on signal feature extraction of peaks, valleys or zerocrossings from raw accelerometric or gyroscopic data that may indicate a HS or TO (78). This can be complemented by applying hidden Markov models or Gaussian continuous wavelet-transformation (CWT) to increase GE detection accuracy (29). These methods have been implemented for single sensors placed on the trunk (85-87), waist (88,89), shank (90), ear (76) and foot (91,92). A thigh-based singlepoint IMU, the Activpal (PAL Technologies, Glasgow, Scotland), has also been used to measure mobility (73,93). However, this sensor has, to our knowledge, not analysed gait metrics beyond step identification in a single-point

Spatial parameter estimation proves more difficult due to the aforementioned technical limitations of inertial sensors (29). Current methods are based on abstraction models (e.g., machine-learning, linear regressive models), locomotion models [e.g., inverted pendulum (IP), double-

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IP] and numerical integration. Those employed for singlepoint sensors include: direction integration (88), linear regression models (89), IP model with double-integration of antero-posterior (86,87) or vertical acceleration from trunk sensors (85); IP model with double integration of AP-acceleration from a foot sensor (91); double-IP model with integration of angular velocity of the shank (90); and autocorrelation procedures also able to determine temporal parameters, and a measure of regularity and symmetry (94). However, the requirement of numerical integration in these models causes accumulation of drift errors (78). Drift compensation is performed using kinematical reset through ZUPT by assuming foot velocity as zero (91) and shank inclination as vertical during midstance (90); however, these methods only prevent growth of drift error without minimizing the already accumulated error (29). As zero-velocity reset is not possible with trunk sensors due to continuous pelvic motion, drift correction is achieved by applying a highpass (86) or Kalman filter (88) to retrospectively correct errors, occasionally in combination with direct and inverse integration at every step (88). However, correction efficacy may be limited in pathological gait where vertical trunk acceleration amplitude is lower and variability higher (78).

Linear measures of spatio-temporal variability and asymmetry are subsequently determined from mean spatio-temporal values, commonly expressed as the standard deviation or coefficient of variation and the absolute difference between left and right mean values respectively (95).

In addition to these traditional spatio-temporal parameters, accelerometer-based systems and non-linear calculations introduce new measures (11). Although these complex non-linear, autocorrelation and acceleration-based measures are dimensionless and unable to be validated against a standard, they are employed extensively clinically. Non-linear measures derived from the theory of stochastic dynamics (e.g., phase plot analysis, fractal-scaling index, sample entropy, Lyapunov exponents) allow appreciation of dynamic fluctuations and patterns between gait cycles throughout a walking bout, contrary to traditional linear measures that treat each as independent to the last (11,49,83). These measures represent the smoothness, regularity, stability, variability, complexity and symmetry of gait, showing sensitivity delineating between pathological and healthy subjects (96-98) equal, or superior to the sensitivity of linear measures (99). Similarly, autocorrelation measures of regularity and symmetry represent a clinically relevant (31,44,67), dynamic substitute for spatio-temporal variability and symmetry respectively (36,94). Other measures, including harmonic ratio (HR) based on Fourier analysis and the root-mean square (RMS) of acceleration magnitudes, are clinically relevant indicators of the smoothness, rhythmicity and symmetry of gait (33,100-102).

Methods

Literature Search

The PRISMA guidelines were followed for this systematic review (103) utilizing the following databases: PubMed; MEDLINE; EMBASE; Cochrane. Firstly, key search terms "gait" AND "accelerometer or inertial" were used to locate studies using inertial sensors to monitor gait. Next, "spatiotemporal" or "temporal" or "phase" or "stride" or "length" or "velocity" were used to locate publications that measure clinically relevant gait metrics beyond stepcount and activity. Finally, the terms "clinical or valid or validity or test or reference or standard" were included to reflect studies that had tested the validity of these wearable technologies or applied them clinically. Relevant MeSH (Medical Subject Heading) terms, variations and synonyms were adjusted for each database.

Study selection

Studies from the above databases were collated and duplicate studies removed. Primary screening by an independent reviewer (JP) was performed based on the title and abstract of the remaining studies following the developed inclusion and exclusion criteria detailed below. Subsequent eligibility assessment was performed based on the full texts of remaining articles by an independent reviewer (JP) following the inclusion and exclusion criteria.

Inclusion criteria

- (I) Articles involving wearable technology/ies.
- (II) The wearable technology features an inertial sensor (accelerometer, gyroscope) or is an IMU.
- (III) The wearable technology is a single-point sensor.
- (IV) Articles written in English.
- (V) Journal papers.

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Exclusion criteria

- (I) Wearable technology/ies only capable of identifying activity or step count.
- (II) Wearable technology/ies features multiple sensor points.
- (III) Wearable technology/ies classed as robotic or exoskeletons.
- (IV) Systematic reviews, books, or conference papers.

Data collection

Following final article selection, results were classified as validation, clinical application or both. Data for validation studies was collected including sensor type(s); sensor placement; study aim(s); conclusions of study; primary gait metrics and methods. Critical analysis of validation studies was also included. Data collected for clinical applications studies included: sensor type(s); sensor placement; primary gait metrics and methods; and clinical application.

Bias analysis

Three different tools were used for risk of bias assessment based on the nature of the studies. Validation and reliability studies were assessed using the Scottish Intercollegiate Guidelines Network (SIGN) methodology checklist for diagnostic tests (104,105). Of the clinical application studies, observational studies were assessed using the Newcastle-Ottawa Scale (106) while randomized-controlled trials (RCTs) were assessed using the SIGN checklist for RCTs (105,107). Case series and case reports were excluded from the use of these bias assessment scales as questions pertaining to comparability no longer apply. Studies were assessed by 3 independent reviewers (WJC), (SMR) and (MM) with at least 2 different reviewers for each study. Discrepancies in assessment were resolved by discussion and reaching a consensus.

Results

From the 1,068 articles retrieved after duplicate removal, 90 articles were selected for inclusion (*Figure 1*). Thirty-two articles assessed the validity of single-point sensors and 48 used single-point sensor gait metrics analysis for clinical applications. Ten articles concurrently assessed validity and employed the device in a clinical application.

Validity of single-point sensors

Among the 42 articles (*Figure 2*), 30 studies utilized trunk-based sensor methods; while 12 studies used alternate locations for the sensor placement which include four studies at the subjects' ear; two at the subjects' shank, two at the subjects' foot, one at the waist; while three other studies utilized smart devices with inertial sensors. The parameters used for validating the sensors include: HS, stride length/duration/regularity, SL/count/duration/length variability/variability/time asymmetry/cycle time/regularity/frequency, GV, cadence, traversed distance, walking time, stance duration, swing duration, single/double support duration, HR, TO and time averaged acceleration.

Validation studies by subject cohort

Among the 42 studies, 59 different cohort of patient population were studied (*Figure 3*). Twenty-five studies involved healthy adults, 11 studies involved elderly subjects, seven studies involved patients with PD, four studies in post-stroke patients with ataxia and three studies in Huntington's Disease (HD). There were two studies each for patients with lower limb amputees and diabetic patients. The remainders were single studies in subjects comprising healthy children, multiple sclerosis (MS) patients, orthopedic patients, muscular dystrophy patients, and patients suffering from motor neuron disease (Table S1).

Clinical application of single-point IMUs

Fifty-eight articles discussed the use of single-point IMUs in clinical setting. Of these, 12 articles discussed the application of sensors in diagnosis and assessing severity of diseases (PD, MS, PN, Alzheimer's disease, age-related changes, frailty and foot & ankle health); 12 studies applied the sensors in monitoring rehabilitation and intervention efficacy (orthopedic, neurosurgical and oncological patients, foot orthoses, medical and physical intervention in neurodegenerative diseases); and 31 studies used sensors to characterize patients with different conditions from healthy subjects. Three articles described both diagnosis and severity of assessment as well as delineating healthy and participants with pathologies (*Table 2*, with more detailed findings of these articles in Tables S1,S2.

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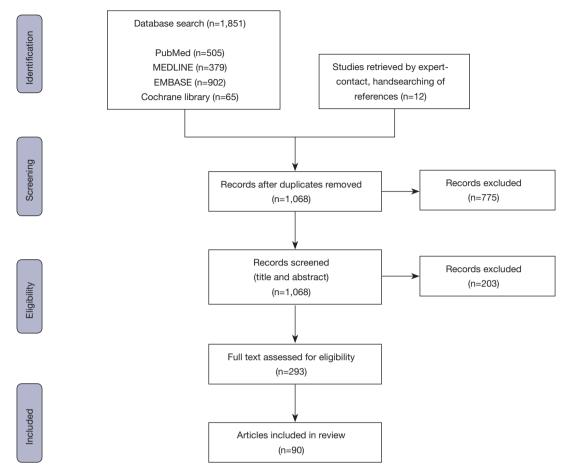


Figure 1 PRISMA methodology.

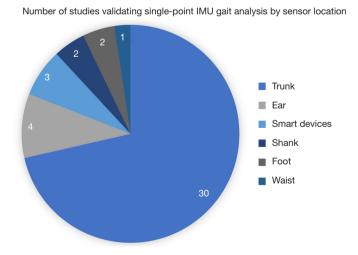


Figure 2 Number of studies validating single-point IMU gait metrics analysis by sensor location. Smart devices have been attached on various locations, some separate to where they were designed to be worn – the iPod touch G4 (iOS 6, Apple Inc.) over L3 (108), the Apple Watch (Apple, San Francisco, CA, USA) on the wrist (4), and the Apple iPhone 5 (iPhone 5, Apple Inc., Cupertino, CA, USA) on the lateral waist (109). IMU, inertial measurement unit.

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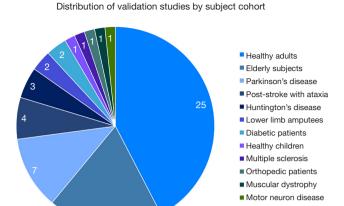


Figure 3 Distribution of validation studies by subject cohort.

Bias assessment

Risk of bias assessment of validity studies did not reveal studies with an unacceptable level of bias. 31 studies were of high quality while nine were of acceptable quality in minimizing bias. The breakdown of the assessment and interpretation is included in Table S3. Areas in which many studies had 'unclear' levels of bias were in the patient selection domain criteria of selecting a consecutive sequence or random selection of participants. Understandably, many of these studies had small sample sizes and recruited volunteers or practiced convenience sampling of patients to achieve this. Many studies were able to reduce bias by conducting simultaneous testing of the IMU and reference standard. Criteria related to the use of a reference standard were not applied to reliability studies that did not use a reference test. The criteria related to pre-specified thresholds of the index test were also largely not applicable.

Of the clinical applicability studies, 30 were scored as "good" quality, eight as "fair" and 12 as "poor". Two RCTs were deemed having acceptable quality and one RCT as high quality in minimizing bias. The breakdown of the bias assessment results and interpretation is included in Appendix 1). Most studies generally missed out a score on ascertainment of outcome using a blind investigation. This may be attributed to a lack of investigators needed to separate carrying out the test and interpreting data. However, this was regarded as not having a large influence on overall bias assessment as measurements using IMUs are automatically recorded to software and not requiring direct human measurement. It would also be difficult to

blind assessors to diseased patients with an obvious gait pathology to healthy controls. The RCTs also had unclear blinding of subjects and investigators to treatment groups. The strengths of these studies were the randomization process and standardization of testing and analysis between treatment and control groups.

Discussion

In our review, single-point IMUs have been reasonably validated in the measurement of spatial and temporal gait parameters (85,86,124). However, IMUs have shown difficulty in estimating variability and asymmetry metrics (48,70,118,125-127). Alongside this, whilst IMUs have shown promise in their clinical applications, such as in the diagnosis of disease (31,32,47,53) and the assessment of treatment efficacy (56,113,128), these studies have predominantly relied on straight-line gait metrics. This critically limits external validity to free-living analysis where day-to-day movements typically represent more complex patterns of acceleration and deceleration. Moreover, studies focusing on clinical application have predominantly described obvious gait changes and have not necessarily demonstrated IMUs to be useful in evaluating subtle differences in gait patterns. Therefore, additional studies focusing on validation and clinical application are required before any mass clinical and commercial uptake of singlepoint wearable sensors can occur.

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Table 2 Clinical applications of single-point IMUs

Reference	Application
Diagnosis and severity assessment	
Demonceau et al., 2015 (31)	Determine PD severity
Herman et al., 2014 (57)	Classify PD subtypes
Dalton et al., 2013 Collett et al., 2014 Pau et al., 2016 (47,62,110)	Determine MS severity
Esser et al., 2018 (111)	Detect PN
De Bruin et al., 2012 (53)	Determine PN severity
Gillain et al., 2016 (39)	Predict risk of Alzheimer's development
Kosse <i>et al.</i> , 2016 Terrier <i>et al.</i> , 2015 (65,108)	Predict age-related gait change
Soangra <i>et al.</i> , 2018 Martinez-Ramirez <i>et al.</i> , 2015 (43,109)	Predict frailty status/determine severity
Van Schooten et al., 2016 Del Din et al., 2017 (33,34)	Predict falls/determine falls risk
Angthong et al., 2018 (112)	Assessment of foot/ankle conditions
Rehabilitation and intervention efficacy	
Atallah <i>et al.</i> , 2014 Rapp <i>et al.</i> , 2015 (76,113)	Total hip and knee replacement efficacy and recovery
Jarchi et al., 2016 (77)	Anterior cruciate ligament repair recovery
Mobbs et al., 2018 (4)	Lumbar microdiscectomy recovery
Hojan et al., 2014 (54)	Effect of breast prostheses after mastectomy
Mutoh <i>et al.</i> , 2016 Manikowska <i>et al.</i> , 2013 (5,56)	Hippotherapy efficacy in cerebral palsy
Henderson et al., 2016 (114)	Rivastigmine efficacy in PD
Terrier et al., 2009 (115)	Prescription footwear efficacy in foot/ankle fractures
Doi et al., 2013 Pau et al., 2014 Perrochon et al., 2015 (38,60,101)	Improvement in gait after physical activity in elderly and cognitive impairment
Delineating pathological subjects from healthy controls	
Barden et al., 2016 Clermont et al., 2016 Bolink et al., 2012 (67,68,116)	Knee osteoarthritis subjects

Table 2 (continued)

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Table 2 (continued)

Reference	Application
Lamoth <i>et al.</i> , 2010 Houdijk <i>et al.</i> , 2008 (48,49)	Amputee subjects
Terrier et al., 2017 (74)	Chronic lower limb pain subjects
Arvin et al., 2015 (58)	Hip abductor fatigue Subjects
Esser <i>et al.</i> , 2011 Mizuike <i>et al.</i> , 2009 Meijer <i>et al.</i> , 2011 (41,52,102)	Muscular dystrophy/Motor neuron disease/Stroke subjects
Hickey <i>et al.</i> , 2016 Matsushima <i>et al.</i> , 2015 (70, 117)	Ataxia disorder subjects
Demonceau <i>et al.</i> , 2015 Del Din <i>et al.</i> , 2016 Esser <i>et al.</i> , 2013 Hatanaka <i>et al.</i> , 2016 Yang <i>et al.</i> , 2011 (31,46,69,72,118)	PD/Progressive supranuclear palsy subjects
Pau et al., 2017 Pau et al., 2018 Storm et al., 2018 (63,119,120)	MS subjects
Dalton <i>et al.</i> , 2013 Collett <i>et al.</i> , 2014 (47,110)	HD subjects
Manikowska et al., 2013 (55)	Menopausal women
Tanigawa et al., 2018 (121)	Pregnant patients with lumbopelvic pain
losa <i>et al.</i> , 2013 Saether <i>et al.</i> , 2014 (42,78)	Cerebral palsy subjects
Moe-Nilssen et al., 2003 (122)	Dyslexia subjects
Awotidebe et al., 2016 (61)	Type 2 diabetes subjects
Chung et al., 2012 Maquet et al., 2010 Martinez-Ramirez et al., 2016 Lamoth et al., 2011 (37,44,51,123)	Cognitively impaired subjects
Auvinet <i>et al.</i> , 2003 Bautmans <i>et al.</i> , 2011 (36,50)	Falls risk/fallers

IMU, inertial measurement unit; PD, Parkinson's Disease; MS, multiple sclerosis; PN, peripheral neuropathy; HD, Huntington's Disease. Detailed findings of these articles can be found in Tables S1,S2.

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Validating IMU gait metrics analysis

Proposed methods and sensor locations for determination of gait metrics from single-point IMU acceleration data are generally validated against a standard or by test-retest reliability (129). The large portion of validation research employing lower trunk sensors compared with alternative sensor locations is reflected in *Figure 2*.

Trunk-based sensor methods

Association of lower trunk accelerations with HS and TO, and the ability to predict these accelerations with an IP model of the body's center of mass (COM) trajectory, has prompted the proposal of several methods for GE and spatio-temporal parameter estimation from these signals.

Methods by Zijlstra & Hof (86)

Peak detection and IP methods proposed by Zijlstra & Hof (86) utilizing a tri-axial accelerometer over the lower trunk demonstrated accurate detection of GEs and limited mean spatio-temporal parameters when compared to GRF from a treadmill. However, the straight-walking protocol employed limits external validity to daily-living analysis. Similarly, 100% of GEs were detected in only nine subjects of a small sample (n=15) of healthy adults, with 12% of GEs identified falsely in the remaining subjects. Furthermore, SL calculation which requires input of individual leglength was consistently underestimated before application of a 1.25 correction factor, revealing limitations in the model itself. Further validation in healthy adults has demonstrated accurate estimation of mean spatio-temporal parameters when compared to stereophotogrammetry and dynamometry, however significant differences were found for gait phase durations that rely on determining TO which isn't explicitly detected by the model (124). SL estimation was again detected with less accuracy due to errors implied by double-integration in this method (124). Further model limitations were identified in neurological populations (41) as neither a generic (1.25) or pathology-specific correction factor could be applied for accurate SL estimation, alluding to the need for individual corrections in pathological cohorts. This finding was reflected when validated in child (130) and older cohorts (131). The model has been further validated in small samples of PD subjects against motion capture (132), MS subjects against an instrumented walkway (110) and through test-retest reliability (47) for mean and variability spatio-temporal measures. These studies also assessed reliability of non-linear variability measures (47) and demonstrated feasibility of anterior

trunk sensor placement, representing an attenuated version of COM accelerations (110). Methods from Zijlstra and Hof (86) have been incorporated into commercially available IMUs, G-Walk (BTS, Milan, Italy) and DynaPort (McRoberts, The Hague, Netherlands). DynaPort accuracy has been assessed in children (130), lower-limb prosthesis (48), diabetes (53), healthy elderly (125,126) and falls risk subjects (50), while G-walk accuracy has been assessed in healthy adult (133-135), PD (136) and MS (137) subjects. Despite determining mean spatiotemporal parameters accurately in most studies, caution is recommended for interpretation of linear variability and asymmetry parameters (48,125,126). The interpretation of gait phase durations reliant on TO (stance, swing, single, double support time) is also uncertain since this event is not explicitly detected by the algorithm (133,136,137). Furthermore, testing limited to small samples in controlled straight-walking conditions limits power of inference to gait metrics analysis in pathological cohorts or scenarios of daily-living.

Autocorrelation methods utilizing lower trunk accelerations from a tri-axial accelerometer (94) have demonstrated test-retest reliability in a small sample (n=20) of healthy subjects for RMS and mean spatio-temporal parameter assessment (138) in straight-walking protocols. Despite lacking further validation for traditional spatio-temporal parameters, the test-retest reliability of autocorrelation measures of regularity and symmetry have been demonstrated in small samples of elderly, falls-risk and HD subjects in straight-walking protocols (47,50,139).

Methods by McCamley et al. (85)

CWT for GE detection and IP methods for spatial parameter estimation proposed by McCamley et al. (85) using a lower trunk IMU have been validated against stereophotogrammetry and force plates. It has also been compared against previous methods from Zijlstra & Hof (86) and González et al. (87). Despite limitations of a small sample (n=18) of healthy subjects and a straightwalking protocol of only 3.6 m, the method by McCamley et al. (85) provided an improved estimate of SL and GE detection compared to previous methods. Validity for temporal parameter means and variability has been subsequently demonstrated in both controlled and freeliving conditions against pressure insoles, despite observed accuracy decreases in free-living conditions (140). Similar protocols in MS patients validated measurement of mean temporal parameters (120). However, variability measures were highly overestimated, with inaccuracies increasing

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with length of walking bout, detracting from applicability to continuous monitoring. The lower accuracy for which this method detects TO compared to HS, may account for difficulty estimating variability measures, shown to be highly sensitive to incorrect GE identification compared to mean parameters (120). Method accuracy was also shown to be speed-dependent and decrease with increasing disability, hindering reliability in pathological cohorts (120).

Method by Godfrey et al. (95,141) combining those by Zijlstra & Hof (86) and McCamley et al. (85)

Combining methods by McCamley et al. (85) for temporal parameter and Zijlstra & Hof (86) for SL estimation, Godfrey et al. (95,141) demonstrated validity against an instrumented walkway and through test-retest reliability in large populations of young and older adults (cumulative n=92) in protocols reflecting daily-living. Despite acceptable agreement for mean spatio-temporal parameters, SL underestimation was again attributed to limitations with generic correction factors, straight-walking dependent IP model and mathematical integration errors. Both methods' measures of spatio-temporal variability and asymmetry were poor, concurring with findings of other studies using an instrumented walkway as a control (70,125-127). In defense of IMUs, Godfrey et al. (95) demonstrated that discrepancies in variability and asymmetry were due to inherent differences between IMUs and instrumented walkways used in these studies, rather than IMU inaccuracy. This highlights the importance of caution when choosing a standard for validation purposes. Demonstrated discriminatory power between pathological and healthy cohorts based on IMU asymmetry and variability measures, despite poor agreement with an instrumented walkway, reinforces these conclusions (70).

Other methods of GE and spatio-temporal parameter estimation

Waist-placed sensor and algorithm development has also been validated. Direct integration methods for SL estimation based on a waist placed IMU (88) were validated against stereophotogrammetry. However, limited gait parameters, a small sample (n=9) of healthy patients and accuracy discrepancies between left and right steps due to anatomically asymmetrical sensor placement limit clinical applicability (88).

To combat the plethora of COM methods, a comparison of five methods (85,86,88,89,142) against stereophotogrammetry and force plates for determination of temporal parameters was conducted by Trojaniello *et al.* (143). The different GE identification methods

incorporated largely either the zero-crossing or waveletbased method. Zjilstra & Hof (86) used a zero-crossing method where foot contact was taken as peak forward acceleration preceding the change of sign of acceleration from positive to negative. González et al. also used a zerocrossing method to approximate a search window prior to applying certain heuristic rules to determine the peak associated with the contact event (142). In conjunction with the zero-crossing method, Shin et al. used a sliding window summing technique to reduce noise (89). The method by McCamley et al. involved integrating and differentiating the acceleration signal using Gaussian continuous wavelet transforms prior to identifying initial and final contact events from the minima and maxima of the smoothed signal (85). Köse et al. used a wavelet-based method to identify windows of interest prior to decomposition and reconstruction of the original signal based on certain threshold application. Heel strike was then detected as the timepoint between signals of the different local frame axes (88). No statistically significant difference was found between methods for stride and step duration and the standard. However, methods that detect TO in addition to HS to allow determination of gait phase durations (85,87,88) showed a statistically significant difference, due to difficulties detecting the smoother acceleration signals indicating TO (143). Omitting assessment of spatial parameter methods limits completeness of the study. While despite examining a large sample, comparison of methods in healthy controls limits external validity to pathological cohorts. In response, assessment of the three best-performing methods (85,86,89) in 10 hemiparetic, 10 PD and 10 HD subjects against an instrumented walkway was undertaken (144). This revealed a universal decrease in GE detection and temporal parameter accuracy compared to healthy subjects. However, no statistically significant differences were revealed regarding accuracy between IMU methods in any cohort, apart from PD subjects for which methods from Zijlstra & Hof (86) outperformed.

New methods are continuously being formulated for lower trunk IMU analysis of spatial and temporal parameters. Oyake *et al.* (145) recently proposed a new algorithm for SL symmetry determination, validated in stroke subjects, while Sejdić *et al.* (146) validated novel methods against motion capture data in PD and peripheral neuropathy (PN) subjects.

Alternative sensor placements

Despite trunk IMUs maintaining the lion's-share of

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research, methods based on alternative sensor positions have also garnered validation attention.

An ear-worn tri-axial accelerometer has been validated in healthy and lower-limb orthopedic subjects against an instrumented treadmill for estimation of mean step time and symmetry (76). A similar sensor has also been validated for the detection of GE's and limited gait parameters in small samples of healthy (147), PD (148) and orthopedic (77) subjects in laboratory conditions.

With placement closer to the ground allowing better GE detection, sensors on the lower limb have also been proposed and validated. GV estimation from a single shank IMU was validated on a treadmill across numerous speeds and slope gradients (90). However, decrease in effectiveness of vertical shank inclination as a ZUPT re-calibration and subsequent GV accuracy with changing incline limits dailyliving application. Magbool et al. (149) further validated an algorithm for detection of GEs across walking speeds and slopes in eight healthy and two amputee subjects using a shank gyroscope. Straight-walking protocols and need for the instrumented shank to take the first step limit these studies' clinical applicability. IP methods using one foot IMU have been validated against a treadmill for GV estimation in five healthy subjects (91). However, difficult attachment of the IMU to the shoe, straight-walking protocols and an accuracy decrease with increasing incline limit applicability. Temporal parameter detection has also been validated using a foot IMU in eight healthy adults (92), showing strong correlation with motion capture. Limited validation of comprehensive gait metrics and inability to assess asymmetry and complex trunk-accelerometer measures with lower-limb sensor placement limits clinical applicability (88).

Smart device gait analysis

Smart devices embedded with inertial sensors have become ubiquitous in everyday life, making them an obvious solution for maximizing patient compliance and allowing inconspicuous, portable gait analysis.

Initial proof of concept using an iPhone attached to the lateral malleolus demonstrated test-retest reliability for quantification of time averaged acceleration and step duration (150). However, unrealistic device placement, limited gait parameters and a sample of only one healthy patient limit validity and clinical applicability. Following this, utilizing trunk-based methods for GE detection (85-87) and SL estimation (86), high correlations were found for an iPhone against stereophotogrammetry for the identification

of GE and mean spatio-temporal parameters in eleven healthy subjects (151). However, similar unrealistic device placement over the lumbar spine and waist has limited applicability to daily living. Addressing this, the reliability of smartphone locations: body, belt, bag, pocket and hand and validity against an instrumented walkway has been tested (152). Hand positions demonstrated poor reliability and agreement with the standard at slow speeds which only marginally improved at higher velocity, while high validity and excellent reliability were demonstrated in body, bag or belt positions at fast/comfortable speeds, lending traction to their incorporation into everyday gait monitoring. However, universal inaccuracies assessing gait at slow speeds limits application to pathological groups. Furthermore, limited emulation of free-living scenarios was employed with only five to nine steps investigated per trial (152).

Validation status of single-point IMUs

Despite lacking validation for a comprehensive set of gait metrics in alternative positions, current COM systems are a proven alternative for calculation of a range of traditional spatio-temporal measures. However, these algorithms still need development, with caution recommended with interpretation of spatio-temporal variability and symmetry parameters, measures of gait phases and estimation of spatial parameters. Further validation is required in larger samples of pathological groups; Furthermore, their accuracy for continuous monitoring in scenarios of daily living needs to be assessed. Incorporation of these algorithms into commercial devices and smart devices is promising for clinical practicality and uptake.

Clinical applications

Although the vision of single-point IMUs for gait metrics analysis in daily-living is in its infancy (4,34,69,120), these devices have been employed extensively in clinical environments to aid diagnosis and severity assessment, determine rehabilitation and intervention efficacy, and delineate pathological groups from healthy controls (*Table 2*). Of these applications, trunk-based IMUs are uniformly employed with the exception of a limited number of studies utilizing ear (76,77), foot (112,123) and smart-device (4,108,109) analysis.

Diagnosis and severity assessment

Single-point IMU gait metrics analysis has been employed as a method of assessment of ageing, orthopedic and Page 14 of 27 mHealth, 2022

neurological conditions.

Gait metrics analysis with single-point devices has aided diagnosis and severity assessment in numerous neurological diseases. Lower trunk sensor gait metrics analysis has been used to determine PD severity, demonstrating significantly reduced gait regularity and GV with increasing Hoeh and Yahr stage severity (31,32). Utilizing the commercially available DynaPort, Herman et al. (57) have also classified PD subtypes based on increased gait impairment and demonstrated classification superiority based on objective gait measures compared to conventional schemes. In MS, mean spatio-temporal parameter changes have been correlated with increasing disease severity using a G-Walk sensor, while also demonstrating high correlations between gait characteristics and patient-reported outcomes, reinforcing the applicability of gait metrics analysis as a clinical measure (62). Dalton et al. (110) also demonstrated significant differences between MS severity groups based on spatio-temporal mean and variability parameters as well as autocorrelation regularity and symmetry. Determining HD severity from trunk-based gait metrics analysis has also shown to correlate with clinical scales using both linear and non-linear measures (47). In diabetes subjects, gait parameters from a trunk IMU have shown good discriminatory power in detecting those with PN in a pilot study with a small sample (111), while De Bruin et al. (53) demonstrated the discriminatory power of SL to discern PN severity in type 2 diabetes patients in free-living gait conditions.

Similarly, gait metrics analysis has aided assessment in orthopedic conditions. Using a foot IMU, Angthong & Veljkovic (112) demonstrated significant correlations between obtained spatio-temporal parameters and subjective validated patient-reported outcomes and quality of life scores in patients with foot and ankle conditions such as arthritis, injury and tendinopathy. This is suggestive of the validity of gait assessment in clinical practice as an objective outcome measure.

In cognitive impairment, analysis of mean spatiotemporal and autocorrelation measures using a trunk IMU has been shown to predict risk of decline from mild cognitive impairment (MCI) to Alzheimer's Disease in small sample sizes (n=23) (39). Furthermore, single-point gait quantification has been used to delineate between dementia subtypes (153).

Decline in physical and cognitive capacity with age is associated with frailty and disability, with consequences including falls, hospitalization and death. Numerous studies using single-point sensors and smart-devices have attempted to predict age-related gait changes (65,108), predict frailty status and determine severity (43,109) and predict falls and determine risk (33,34) to allow early-intervention to reduce adverse outcomes.

Rehabilitation and intervention efficacy

Gait metrics analysis using single-point IMUs has been used as an objective measure of rehabilitation and intervention efficacy in a range of conditions.

Single-point IMUs have been employed in the assessment of surgical outcomes and rehabilitation. A range of objective gait measures from trunk IMUs have been obtained to determine operation efficacy and rehabilitation progress after total hip replacement (113) and decompressive laminectomy for lumbar spine stenosis (128). An ear-worn sensor has been used to evaluate recovery from anterior cruciate ligament repair based on gait symmetry (147) as well as total hip and knee replacements based on stride duration and gait symmetry in small samples (76). Similarly, employment of the consumer available Apple Watch (Apple, San Francisco, CA, USA) for detection of GV was claimed invaluable in monitoring recovery through continuous gait monitoring in daily life following lumbar microdiscectomy in a single patient (4).

Assessment of non-operable intervention efficacy has also been determined using trunk IMUs. In cerebral palsy subjects, improvement in spatio-temporal mean and symmetry parameters after one hippotherapy session (56) and increase in mean parameters over a two-year intervention course (5) has been demonstrated. While linear measures of gait variability from trunk accelerations have been used to determine the effectiveness of Rivastigmine in in PD subjects over 32 weeks (114). In orthopedics, the effectiveness of prescription footwear has been assessed through trunk-based autocorrelation gait metrics analysis in severe foot and ankle fractures (115). While in women following single-breast mastectomy, significant influence of external breast prosthesis on spatiotemporal parameters of gait has been demonstrated (54).

Prescribed physical activity programs have been shown to improve physical functioning and reduce risk of falls and adverse outcomes (154). Pau *et al.* (60) utilized a trunk accelerometer to determine the increased effectiveness of vigorous compared to light physical activity on mean spatiotemporal parameters in elderly over 36 weeks. Similarly, the positive effect of PA on a range of spatio-temporal and accelerometer-based measures was demonstrated in MCI (101) and dementia subjects (38) using trunk-based

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accelerometry.

The efficacy of neurorehabilitation has also been assessed using trunk-based gait analysis. Santoyo *et al.* (155) demonstrated an increase in mean spatio-temporal parameters following a five-month neurorehabilitation program in 45 MS patients. Furthermore, Zanetta et al. (156) demonstrated similar improvements in cadence and GV after a four-week program, and significant correlation of gait parameters with validated clinical assessments (Berg Balance Scale, 6MWT), validating the usefulness of gait metrics analysis as an objective measure of outcomes.

Delineating pathological and healthy subjects

Most single-point IMU clinical applications focus on delineating between pathological groups with gait impairments and healthy controls. This application enables clinical validation of gait metrics and appreciation of metrics relevant to different pathologies.

Quantification of gait through single-point sensors in musculoskeletal disorders has been assessed. Knee osteoarthritis patients have been delineated from healthy controls based on numerous mean, autocorrelation and accelerometer-based spatio-temporal measures (67,68,116). In amputee gait, measures of stability, regularity and variability (49) as well as mean spatio-temporal parameters (48) have shown significant difference from healthy controls. Hip adductor fatigue has been demonstrated to significantly impact variability and symmetry of gait (58), while patients with chronic lower limb pain have demonstrated significant differences in cadence, variability and symmetry measures compared to controls (74). Using a trunk IMU in pregnant patients, significant differences have also been noted in trunk movement asymmetries between those with and without lumbopelvic pain. (121)

Numerous neurological conditions show altered gait quality using single-point IMUs. In stroke, RMS, autocorrelation measures of regularity and symmetry (102) and GV (52) have shown significant differences compared to controls. The gait impairments of various ataxia disorders have been assessed, demonstrating significant differences in mean, variability and asymmetry of spatio-temporal characteristics as well as autocorrelation and accelerometer-based measures (70,117). In PD, Esser et al. (46) demonstrated superior sensitivity of GV and non-linear variability measures compared to mean parameters for delineation from healthy controls in a small sample (n=24). Further works have demonstrated the ability of

mean and symmetry spatio-temporal parameters as well as autocorrelation measures in both laboratory and freeliving gait to delineate between PD subjects and healthy controls (31,69,72,118). Accelerometer-based measures of gait smoothness (HR) have also been shown to delineate between MS and healthy subjects prior to any measurable changes in mean spatio-temporal parameters (119), while dual-task gait in these patients has shown to result in a significant difference in mean spatio-temporal parameters compared to controls (63). In HD, significant differences have been demonstrated for autocorrelation measures, mean spatio-temporal parameters and both linear and nonlinear variability measures against healthy subjects (47,110). Furthermore, in type 2 diabetes (61) and normal pressure hydrocephalus (157), altered spatio-temporal parameters have been demonstrated compared to healthy controls. In child pathological cohorts, differences in gait parameters have also been demonstrated in cerebral palsy (42,78) and dyslexia subjects (122) compared to healthy controls.

The effect of cognitive impairment on gait has also been quantified compared to healthy controls using single-point sensors (37,52,123,153). Gait differences have been quantified in those at risk or with a history of falls, with autocorrelation measures of regularity and symmetry, GV, stride length and step-time asymmetry showing significant discriminative capacity against healthy controls (36,50).

Limitation of single-point IMUs in delineating diseases

Clinical use of IMUs discussed above were mainly describing an obvious gait change (e.g., MS or Parkinson disease) rather than minute structural damages such as a torn hip labrum. Therefore, current use of a single-point IMU has to be considered within the appropriate clinical context. Additionally, the clinical application studies included are largely limited to straight line gait metrics assessment which does not fully reflect real-life movements. Along with the maturation of single-point IMUs in terms of validity, future studies should attempt to assess validity of these sensors in picking up complex movements that reflect real life movements such as falls.

Future prospects

The ability for wearable devices in detecting gait and posture is maturing and undergoing continuous development. Multiple studies have demonstrated the use of wearable technologies in aiding postural analysis as well as serving as a tool for the general population in everyday Page 16 of 27 mHealth, 2022

postural/activity tracking (158,159). A novel scoring algorithm incorporating gait and postural scores has also been proposed to report patients' outcome in a manner which is simple and conducive to a continuous stream of data that can be remotely monitored by clinicians (160). This ability to remotely measure and record continuous data gives wearable devices an upper hand compared to lab-based instruments which are geographically sparse and perform gait analysis at discrete time points, though validity and standardization remains a drawback currently (4,23,158-160).

Future studies are required before the implementation of IMUs can be recommended to clinicians. In particular, there is an urgent need to validate IMU accuracy in free-living home environments, with most current validation studies instead measuring gait metrics on straight-line pathways. Other parameters such as U-turns, complex acceleration and deceleration that mimics day-to-day movements such as slowing down when approaching a door or chair could be studied. In addition, single-point IMUs have not been consistently shown to have high accuracy when measuring variability and asymmetry metrics. Future studies may assess other models for GE detection, which are continuously being developed, that may more accurately capture these metrics. Moreover, many of the studies focusing on clinical application have described obvious gait changes. While IMUs are still useful in objectively quantifying these changes, more evidence is required to demonstrate the clinical applications of IMUs in the measurement of subtle gait differences. Future studies to compare user acceptability and compliance between single-point IMUs and multipoint IMUs should also be conducted. Studies to determine ideal placement location of single-point IMUs at various body parts could also be conducted. Further developments and validation may one day bridge the gap for incorporating wearable technologies into actual clinical setting in aiding diagnosis and monitoring patient progression.

Conclusions

This review has demonstrated the validity of single-point IMUs as an alternative to current quantitative methods and their ability to assist in clinical scenarios. The accuracy of these systems for detection of traditional metrics as well as the demonstrated clinical relevance of novel, accelerometer-based measures is promising for practicality and efficacy in the clinical context. Further validation for long-term, continuous monitoring in daily living scenarios is required

as is performance in larger samples of pathological cohorts before mass commercial and clinical uptake can be expected.

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