

# Interdialytic 24-Hours Ambulatory Blood Pressure versus Dialysis Unit Blood Pressure for the Diagnosis of Electrocardiographic-Left Ventricular Hypertrophy in Chronic Hemodialysis Black Patients

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## Abstract

**Background and Aim:** In hemodialysis patients, 24-hours interdialytic ABPM better detects TOD than dialysis unit blood pressure. Therefore, the present study was aimed to assess the diagnostic performance of 24-hours ABPM vs. dialysis unit BPs for the diagnosis of ECG-LVH in steady state chronic hemodialysis black patients. **Methods:** From March 31 to September 30, 2018, interdialytic ABPM was performed after a mid-week hemodialysis session for 24 hours using a Spacelab 90207 ABPM monitor in the non-access arm in 45 stable chronic hemodialysis black patients (age  $\geq 20$  years, hemodialysis for at least 3 months and informed consent) attending 3 hemodialysis centers in Kinshasa. Ambulatory BP was recorded every 20 minutes during the day (6 AM to 10 PM) and every 30 minutes during the night (10 PM to 6 AM). ECG-LVH was defined using Cornell product criteria. ROC curve method was used to assess the performance of dialysis unit BPs vs. interdialytic 24-hours ABPM in diagnosing ECG-LVH.  $P < 0.05$  defined the level of statistical significance. **Results:** Whatever the method of BP measurement, all the SBP values were related to ECG-LVH with similar AUC and overlapping 95% CI; however, they were not significantly different from each other. 24-hours interdialytic ambulatory SBP (AUC 0.748; 95% CI 0.58 - 0.94) had the highest

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area under the curve. **Conclusion:** The present study showed that although all the two BP measurement methods equally detected ECG-LVH, 24-hours ABPM tended to have the highest diagnostic performance.

### Keywords

24-Hours ABPM, Dialysis Unit BPs, ECG-LVH, Diagnostic Performance, Chronic Hemodialysis, Black Africans

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## 1. Introduction

Hypertension, a common clinical finding among patients with chronic kidney disease (CKD), often remains poorly controlled in maintenance hemodialysis (MHD) and is associated with an increased risk for cardiovascular (CV) events [1]. Therefore, accurate and reliable measurement of blood pressure (BP) in this specific group of patients is of diagnostic, prognostic and therapeutic importance [1]. In this regard, the relationship between BP and CV outcomes in MHD has been reported to depend on the setting and the technique of BP measurement [1]. As the variability of BP within the patient at least as great as variability seen between patients [2], BP obtained routinely using oscillometric method in the dialysis unit without attention to details that differ strikingly from BP obtained using standardized methods [1]. However, BP obtained in the dialysis unit (peridialytic BP), even if obtained by standardized methods, agrees poorly with interdialytic ambulatory BP (ABP) [3] [4] [5], and cannot be used to accurately predict the presence or absence of left ventricular hypertrophy (LVH) [6]. In contrast, BP obtained outside the dialysis unit, whether obtained by ABP measurement (ABPM) or home BP, is more useful and accurate to diagnose LVH [6]. It appears thus rationale for each setting to evaluate the relationship between inside and outside BP measurement and intermediate end-point such as LVH as well final end-points such as mortality in MHD patients. Unfortunately, most studies on this topic have been conducted out-of sub-Saharan Africa highlighting the need for data from this region of the world where the access to renal replacement therapy has substantially improved. In the Democratic Republic of the Congo (DRC), a recent study has reported an inverse relationship between peridialytic BP and mortality in MHD patients with stroke [7]. However, the predictive value of peri-dialytic BP compared to 24-hour ABPM for the diagnosis of LVH in steady state MHD patients has not yet been evaluated. Therefore, the aim of the present study was to assess the predictive value of peridialytic BPs and 24-hour ABPM for the diagnosis of electrocardiographic (ECG)-LVH in steady state MHD patients in Kinshasa, the DRC.

## 2. Patients and Methods

- **Participants**

From March 31 to September 30, 2018, patients 20 years or older who had

been on maintenance hemodialysis (MHD) for more than 3 months and were dialyzed 2 or 3 times a week in three dialysis units in Kinshasa, DR Congo (University of Kinshasa Hospital, Ngaliema Medical Center and Medical Center of Kinshasa) were consecutively enrolled in a cross-sectional on the predictive value of dialysis unit BPs (predialysis, intradialysis and postdialysis BPs) and interdialytic 24-hours ambulatory BP monitoring (ABPM) for the diagnosis of electrocardiographic-left ventricular hypertrophy (ECG-LVH). Inclusion criteria were age  $\geq 20$  years, regular hemodialysis for at least 3 months and informed consent. Patients with chronic atrial fibrillation (AF) or body mass index (BMI)  $\geq 40$  Kg/m<sup>2</sup> were excluded. Patient's medical records were used to collect data on past medical history, sociodemographic (age, gender, profession, marital status, financial support), dialysis (type of vascular access, type of dialysis, current month's mean interdialytic weight gain and KT/V, number of sessions per week, dialysate sodium profile) and biological parameters (last month's blood urea nitrogen, serum creatinine, serum albumin, total and ionized calcium, phosphorous) as well as current treatment. The study was approved by the Ethical Committee of Kinshasa School of Public Health and all patients gave written informed consent.

- **Measurements**

Peridialysis (pre- and postdialysis BPs) BP measurement was obtained by trained dialysis unit staff using a validated automated sphygmomanometer OMRON MIT5 Connect with patients at a sitting position, within 30 minutes prior to and following the dialysis session on the non-fistula arm or the non-dominant arm for patients using catheters. Three readings 1 to 3 minutes apart were recorded after the patient had been resting in a quiet room for at least 5 minutes; the average of the two last readings was used as the standard BP value for the present study. Intradialysis BP recordings were obtained by the dialysis unit staff using the sphygmomanometer equipped with hemodialysis machines without a specified technique. All BP recordings were averaged over two weeks surrounding the ambulatory BP measurement. Thus, depending on the number of session a week (2 or 3 sessions a week), each patient had 4 or 6 predialysis, postdialysis BP recordings, respectively, to provide routine dialysis unit.

Interdialysis 24-hours ambulatory BP measurement (ABPM) was performed after a mid-week hemodialysis session for 24 hours using a GIMA ABPM PULSE RATE monitor (Gima Spa, Milano, Italia) in the non-access arm. Ambulatory BP was recorded every 20 minutes during the day (6 AM to 10 PM) and every 30 minutes during the night (10 PM to 6 AM). Patients were instructed to keep their arm immobile during measurement and follow their daily activity. Awake and sleep readings were calculated for each patient by self-reported sleep and wake times by means of a diary. Patients with <70% recording were excluded for the analysis.

- **Outcome**

Resting ECG records lasting less than 3 months were retrieved from patients' medical files whereas it was performed in patients with ECG records lasting

more than 3 months. Left ventricular mass (LVM) was estimated using Cornell product index; ECG-LVH was defined  $LMV > 2.440 \text{ mm}\cdot\text{sec}$  [8].

- **Statistical analyses**

Baseline characteristics were summarized as mean (standard deviation) or median (interquartile range) for continuous variables and as absolute (n) and relative (in %) frequencies for categorical variables. The comparison of means of dialysis unit BP and 24-hours ABP was performed using one way analysis of variance (ANOVA). Receiver operating characteristic (ROC) analysis was conducted for evaluating the predictive performance (Area Under the Curve, AUC) of 24-hours ABP vs dialysis unit BP and for the diagnosis of ECG-LVH. P value  $< 0.05$  defined the level of statistical significance.

### 3. Results

- **General and hemodialysis characteristics of the study population**

Of the 62 patients eligible for the study, 17 of them were excluded [non-consent: 2, dialysis arrest: 1, hemodynamic instability: 5, death during the study: 4, travel abroad: 3 and non-valid ABP recordings (arrhythmia: 1, intolerance: 1)]. Finally, 45 patients (31 men and 14 women) constituted the sample population of the present study (**Table 1**), their mean age and body mass index (BMI) was  $59.1 \pm 12.3$  years and  $24.4 \pm 3.9 \text{ Kg/m}^2$ , respectively. Hypertension and diabetes as the initial underlying kidney disease were reported by 25 (55.6%) and 13 (28.9) patients, respectively. Hypertension with a mean duration of  $12.1 \pm 3.1$  years was present in 44 (97.8%) patients. Calcium channel blockers (CCB) and diuretics were the antihypertensive drug classes reported by 39 (88.6%) and 29 (65.9%) patients, respectively. ECG-LVH using Cornell product index was present in 11 (25%) patients.

Hemodialysis parameters of the study population are depicted in **Table 1**. The mean HD vintage was  $34.3 \pm 10.8$  months with patients having twice and thrice a week dialysis session, respectively. The majority of patients ( $n = 28$ ) had a catheter as vascular access and were receiving either HDF ( $n = 22$ ) or the combination of HDF and HD ( $n = 17$ ). Average levels of KT/V, IDWG and RD were  $1.2 \pm 0.2$ ,  $1.9 \pm 0.4 \text{ Kg}$  and  $338.8 \pm 36.6 \text{ mL/day}$ , respectively. Nearly all patients were receiving EPO ( $n = 100$ ) and IV iron ( $n = 41$ ). The majority of patients ( $n = 35$ ) had a secured financial support. Average levels of Hb, BUN, creatinine, and albumin were  $11.1 \pm 1.9 \text{ g/dL}$ ,  $111.4 \pm 46.5 \text{ mg/dL}$  and  $9.2 \pm 3.6 \text{ mg/dL}$ , respectively (**Table 2**).

- **Dialysis unit and interdialytic 24-h ambulatory BP measurements**

**Table 3** summarizes average dialysis unit and 24-h ABP levels of the study population. Interdialytic 24-h ABP components tended to be in average lower than that of dialysis unit BPs; however, the difference was statistically significant ( $p = 0.014$ ) only for SBP.

- **Dialysis unit BPs vs. 24-h ABPM for the diagnosis of ECG-LVH**

**Figure 1** compares ROC curves of dialysis unit SBPs vs 24-h interdialytic

ambulatory SBP for the diagnosis of ECG-LVH. Whatever the method of BP measurement, all the SBP values were related to ECG-LVH with overlapping 95% CI; however, they were not significantly different from each other. Twenty four-hours interdialytic ambulatory SBP (AUC 0.748; 95% CI 0.58 - 0.94) had

**Table 1.** Demographic, clinical and biological characteristics of the study population.

Variables	All (n = 45)
Age, years	59.1 ± 12.3
Gender, n (%)	
M	31 (68.8)
F	14 (31.2)
Initial Kidney disease, n (%)	
Hypertension	25 (55.6)
Diabetes	13 (28.9)
CGN	8 (17.8)
Others	4 (8.9)
BMI, Kg/m <sup>2</sup>	24.4 ± 3.9
Hypertension, n (%)	44 (97.8)
DHT, years	12.1 ± 3.1
Antihypertensive drugs, n (%)	
CCB	39 (86.7)
Diuretic	29 (64.4)
ACEIs	14 (31.8)
Betablockers	12 (26.6)
ARBs	9 (20.0)
Others	9 (20.0)
ECG-LVH, n (%)	11 (24.4)
Hemoglobin, g/dL	11.1 ± 1.9
Hematocrit, %	33.2 ± 5.9
BUN, mg/dL	111.4 ± 46.5
Creatinine, mg/dL	9.2 ± 3.6
Albumin, g/dL	36.8 ± 13.9
Sodium, mmol/L	128.9 ± 29.6
Potassium, mmol/L	5.4 ± 1.6
Calcium, mmol/L	2.13 ± 0.31

Data are expressed as mean ± standard deviation, absolute (n) and relative (in percent) frequencies. Abbreviations: M, male F, female CGN, chronic glomerulonephritis BMI, body mass index DHT, duration of hypertension CCB, calcium channel blocker ACEIs, angiotensin converting enzyme inhibitors ARBs, angiotensin type 1 receptor blockers ECG-LVH, electrocardiographic-left ventricular hypertrophy BUN, blood urea nitrogen.

**Table 2.** Hemodialysis parameters of the study population.

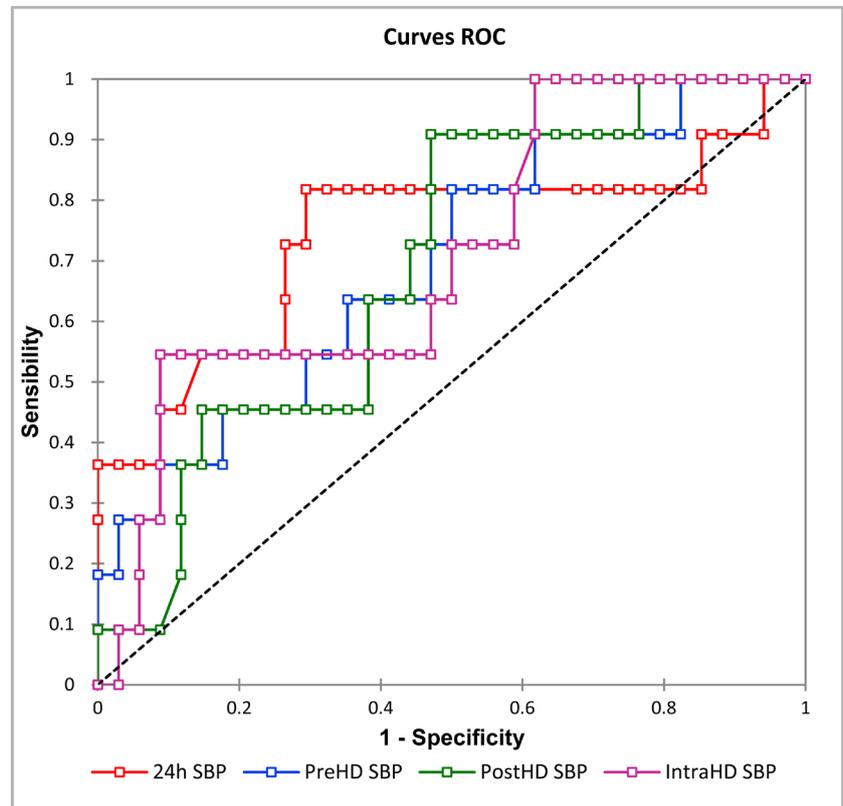
Variables	All (n = 45)
Type, n (%)	
HD	6 (13.3)
HDF	22 (48.9)
HD/HDF	17 (37.8)
Dialysis duration, mo	34.3 ± 10.8
Vascular access, n (%)	
Catheter	28 (62.2)
AVF	17 (37.8)
KT/V	1.2 ± 0.2
IDWG, Kg	1.9 ± 0.4
RD, mL/day	338.8 ± 36.6
>500	6 (20.7)
200 - 400	12 (41.4)
<200	11 (37.9)
EPO, n (%)	45 (100)
EPO dosing, IU/Kg/week	11,187.8 ± 2450.5
IV iron therapy, n (%)	41 (91.1)
IV Iron dosing, mg/week	130.6 ± 46.7
Financial support, n (%)	
Patient/family	8 (17.8)
Private and public enterprises	35 (77.8)
Government	2 (4.4)

Data are expressed as mean ± standard deviation, absolute (n) and relative (in percent) frequencies. Abbreviations: HD, hemodialysis HDF, hemodiafiltration AVF, arteriovenous fistulae KT/V, dialysis efficacy EPO, erythropoietin IU, international unit IV intravenous.

**Table 3.** Average levels of dialysis unit blood pressure and interdialytic 24-hours ambulatory blood pressure of the study population (n = 45).

Variable	PreHD	PostHD	IntraHD	24 h-ABPM	p
SBP, mmHg	152.3 ± 19.4	149.2 ± 18.6	153.4 ± 22.0	141.1 ± 17.8	<b>0.014</b>
DBP, mmHg	88.5 ± 13.6	86.5 ± 10.9	95.1 ± 10.4	82.1 ± 12.5	0.705
PP, mmHg	63.8 ± 11.9	62.7 ± 14.5	58.3 ± 11.0	59.0 ± 11.8	0.957
MAP, mmHg	109.8 ± 14.7	107.4 ± 12.1	114.5 ± 16.0	101.8 ± 13.4	0.405

Data are expressed as mean ± standard deviation Abbreviations: PreHD, prehemodialysis PostHD, Post-hemodialysis IntraHD, intrahemodialysis SBP, systolic blood pressure DBP, diastolic blood pressure PP, pulse pressure MAP, mean arterial blood pressure.



	p value	AUC	95%CI
SBP			
PreHD	0.041	0.708	0.530 - 0.886
PostHD	0.069	0.685	0.515 - 0.854
IntraHD	0.035	0.715	0.543 - 0.887
24 HABP	0.015	0.748	0.548 - 0.947

**Figure 1.** Receiver operated characteristic (ROC) curves and area under the curve (AUC) of interdialytic 24-hour ambulatory systolic BP vs dialysis unit systolic blood pressure (BP) vs for the diagnosis of ECG-LVH. (Abbreviations: PreHD, pre-hemodialysis, post-HD, post-hemodialysis, intra-HD, intra-hemodialysis, 24 HABPM, 24-hours ambulatory blood pressure).

the highest area under the curve followed by IntraHD (AUC 0.715; 95% CI 0.58 - 0.94), PreHD (AUC 0.708; 95% CI 0.53 - 0.88) and PostHD (AUC 0.685; 95% CI 0.51 - 0.85) SBPs, respectively.

#### 4. Discussion

The main findings of the present study are as follows. First, average levels of 24-h interdialytic ambulatory BP levels were lower compared with those obtained with dialysis unit measurements. Second, whatever the BP measurement method, all SBP predicted equally ECG-LVH without significant difference from each other; 24-h interdialytic ambulatory SBP had the highest area under the

curve followed by intradialytic SBP.

24-h interdialytic ambulatory BP levels were in average lower than that obtained by dialysis unit BP measurements. This finding is consistent with that of Argawal *et al.* [6] who reported in a meta-analysis that pre- and postdialysis blood pressures are imprecise estimates of interdialytic blood pressure. They found that predialysis BP overestimated ambulatory BP whereas postdialysis BP underestimated it. They linked the higher predialysis measurements to increased intravascular volume, withholding antihypertensive medications just before treatment, white coat effect, and lack of standardized measurements.

Whatever the BP measurement method used, all SBP equally predicted ECG-LVH with 24-h interdialytic ambulatory SBP having the highest AUC. Consistent with our finding, Agarwal *et al.* [9] already reported that using all BP values pre- and postdialysis measured a mid-week dialysis may serve as a more useful tool to estimate interdialytic ambulatory BP. Although all BP measurement methods have already been reported to predict hypertension and related target organ damage (TOD) such as LVH in chronic hemodialysis patients, their sensitivity and specificity condition their diagnostic performance [9]. In this regard, ABPM is considered as the gold standard for the diagnosis and management of hypertension in chronic hemodialysis patients [9] [10]. The superiority of ABPM over dialysis unit BP measurement is thought to rely upon the provision of a greater number of measurements and a more accurate reflection of the patient's BP burden overtime [9]. In addition, whereas dialysis unit BP is influenced by the white-coat effect, the latter is eliminated with the use of ABPM [11]. Agarwal *et al.* [6] found that dialysis unit BP measurements were weak correlates of LVH and, whatever the BP measurement technique used, DBP was not associated with LVH. Of note, SBP outside the dialysis unit was a stronger correlate of echocardiographic [6] [12] as well as ECG-LVH [13] compared with dialysis unit BP. In an attempt to improve the diagnostic performance of dialysis unit BP measurements, Rahman *et al.* [13] found that standardizing the technique of blood pressure measurement was associated with statistically lower pre- and postdialysis BPs. The authors stress out that the magnitude of the measurement errors should attenuate the difference between peridialysis BP and ambulatory BP. However, this improvement remains still a matter of controversy and debate since it was not found in some studies, like ours, that used standardized BP [6] [14].

Intradialytic BP had the second highest area under the curve for diagnosing ECG-LVH in the present study. Another way to improve diagnostic accuracy of peridialysis BP measurements is to consider intradialytic BP recordings obtained via an automatic cuff attached to the HD machine [9] [15] [16]. Indeed, the median intradialytic BP has been suggested to serve as useful tool to assess hypertension and related TOD and may therefore be used a bedside tool to predict interdialytic ambulatory BP. A mid-week intradialytic BP  $\geq 140/90$  mmHg has sensitivity and specificity that exceeds pre- or postdialysis BP measurements.

However, this method is of last resort because better methods are available to evaluate hypertension and related TOD in hemodialysis patients. In addition, when diagnosing hypertension using ABPM as a gold standard, average intradialytic BP considered together with peridialytic BP has greater diagnostic value compared to peridialytic BP recordings alone [11]. It is possible that intradialytic BP correlates better with ambulatory BP over peridialytic BP because the latter, similar to interdialytic BP, samples the patient over a range of extracellular fluid volume and uremic states during the condensed time span of the HD session [11].

The interpretation of the results of the present study should take into account some limitations. First, the cross-sectional nature of the study precludes the establishment of any temporal relationship between the variables of interest. Second, the relatively small study sample size did not allow much power to statistical tests to detect potential association between the variables of interest. Third, single peridialytic BP and 24-h ambulatory BP measurements could have led to under- or overestimation of average BP levels. Fourth, electrocardiogram a less sensitive method was used to assess left ventricular mass and left ventricular hypertrophy.

## 5. Conclusion

Although all BP measurement methods equally diagnosed ECG-LVH among the present case series, 24-h interdialytic ambulatory BP measurement had the highest diagnostic performance compared to dialysis unit BP measurements.

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## Authors Contribution

CKI collected data, participated in data analysis and reviewed the manuscript.

FBBL conceived the study, participated in data analysis and drafted the manuscript.

YL participated in data collection and reviewed the manuscript.

TM participated in data collection and reviewed the manuscript.

NU participated in data collection and reviewed the manuscript

AN conducted statistical analysis of data and reviewed the manuscript.

VMM reviewed the manuscript.

EKS participated in statistical analysis of data and reviewed the manuscript.

NMN reviewed the manuscript.

EVK reviewed the manuscript.

## Conflicts of Interest

The authors declare no conflicts of interest regarding the publication of this paper.

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