



Published in final edited form as:

*Am J Kidney Dis.* 2021 November ; 78(5): 640–648. doi:10.1053/j.ajkd.2021.04.013.

## Comparison of Dialysis Unit and Home Blood Pressures: An Observational Cohort Study

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### Abstract

**Rationale & Objective:** Past studies show that, on average, predialysis BP runs higher than BP measured at home in a dialysis population. We hypothesized that in a subset of patients, BP was higher at home than in the dialysis unit and that the prevalence of left ventricular hypertrophy, a surrogate for cardiovascular events and death, was increased.

**Study Design:** Prospective cohort

**Settings and Participants:** 97 hypertensive hemodialysis patients enrolled in the Blood Pressure in Dialysis Study (BID), a randomized trial of treatment to predialysis BP 140/90 vs. 155–165 /90 mm Hg, with 6 pairs home and predialysis BP readings.

**Exposure:** Differences between predialysis and next day home systolic BP measured over one year

**Outcome:** Left ventricular mass index (LVMI) by cardiac magnetic resonance imaging

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**Authors' Contributions:** research idea and study design: DCM, AG, JGG, PGZ; data acquisition: DCM, MJ, RYK, LN, DWP, PGZ, AMH; data analysis/interpretation: DCM, HJ, AG, VSP, SSP, JGG, PGZ, SKS; statistical analysis: HJ, SSP, VSP. Each author contributed important intellectual content during manuscript drafting or revision and agrees to be personally accountable for the individual's own contributions and to ensure that questions pertaining to the accuracy or integrity of any portion of the work, even one in which the author was not directly involved, are appropriately investigated and resolved, including with documentation in the literature if appropriate.

**Analytic Approach:** A hierarchical clustering analysis divided patients into 3 clusters based on the average and variability of differences in systolic BP predialysis and at home. We compared clinical factors and LVMI across clusters.

**Results:** The predialysis-to-home systolic BP differences were least square (LS) means (95% CIs) 19.1 (17.0, 21.1) for Cluster 1 ('home lower'), 3.7 (1.6, 5.8) for Cluster 2 ('home and predialysis similar'), and -9.7 (-12.0, -7.4) mm Hg for Cluster 3 ('home higher'). Systolic BP declined during dialysis in Clusters 1 and 2 but increased in Cluster 3. Interdialytic weight gains did not differ. After adjusting for sex and treatment arm, LVMI was higher in Cluster 3 versus Clusters 1 and 2 (differences in LS means of 10.6 (SE 4.96, p=0.04) and 12.0 (SE 5.08, p=0.02) gm/m<sup>2</sup>, respectively).

**Limitations:** Limited statistical power

**Conclusions:** In Cluster 3, which accounted for 31% of participants, home BP was higher than predialysis BP. Patients in Cluster 3 had a higher LVMI than those in Clusters 1 and 2 indicating that their BP may have been undertreated.

### Keywords

masked hypertension; cardiovascular risk; left ventricular mass

## INTRODUCTION

Hypertension is common among hemodialysis (HD) patients.<sup>1</sup> However, the optimal blood pressure (BP) target, as well as the timing and location for BP measurements, remain controversial. Thrice weekly HD results in significant BP variability throughout the week. BP usually decreases significantly during dialysis and increases slowly during the interdialytic interval.<sup>2,3</sup>

The KDOQI and the Canadian Society of Nephrology recommend using pre- and post-dialysis BP measurements to guide hypertension management.<sup>4,5</sup> Results from previous studies indicate that home BP is lower than predialysis BP.<sup>6,7</sup> However, these observations were based on population averages from a two week period of intermittent home readings and a single 44-h ambulatory blood pressure monitoring (ABPM) session. Since BP is highly variable there may be significant variability in the observed differences between predialysis and home BP within individuals over repeated measurements and, across individuals in a population. Assessing population averages over a short time period may miss the detection of a pattern in which home BP is consistently higher or lower than dialysis unit readings within certain individuals. To the extent that home and dialysis unit BP measurements differ, excessive reliance on dialysis unit BP measurements may lead to over or under treatment of BP.

The present study was designed to explore the hypothesis that there is a subset of HD patients in whom BP is often higher at home than in the dialysis unit. As treatment decisions are based on dialysis unit BP these patients maybe undertreated, leading to left ventricular hypertrophy (LVH). We explored this hypothesis using data obtained in the Blood Pressure in Dialysis (BID) Pilot Study, a randomized trial of treating hypertensive HD patients to

a predialysis systolic BP (SBP) of 110–140 mm Hg versus 155–165 mm Hg.<sup>8</sup> BP was measured immediately before each dialysis and at home the day after the midweek HD over one year. In the present study we classified individuals into three clusters based on the mean and variability of the predialysis minus home BP differences over one year. We then assessed the differences in demographic and clinical factors, including left ventricular mass index (LVMI) across the clusters.

## METHODS

### Study Design

This is a prospective cohort study involving individuals who participated in the BID Study. Examining differences in home and standardized dialysis unit BP measurements was a pre-specified secondary outcome.

### Study Population and Setting

We previously described the BID methods.<sup>9</sup> Briefly, patients were recruited from 18 dialysis units in Albuquerque NM, Boston MA, Charleston SC, Pittsburgh PA, and Cleveland OH. The study was approved by the university affiliated Institutional Review Boards at each site and patients provided informed consent. Inclusion criteria for the BID Study were (1) age ≥ 18 years; (2) treated with HD for ≥ 90 days; and (3) 2-week averaged standardized predialysis systolic BP < 155 mm Hg. Exclusion criteria were (1) fluid overload evident from clinical exam; (2) frequent intradialytic hypotension; and (3) life expectancy < 1 year. Patients had to have ≥ 6 weekly pairs of midweek predialysis and next day home BP readings to be included in the present study.

### BP Measurements

Standardized predialysis BP was measured each treatment after five minutes of rest in a sitting position with back supported, legs uncrossed, and in accord with other aspects of the American Heart Association guidelines for the duration of the study.<sup>10</sup> Three readings were taken and the latter two were averaged. Patients and dialysis unit staff were trained to obtain standardized predialysis and home measurements, respectively, using the Lifesource UA-767 oscillometric device (A&D Medical, San Jose, CA). Home measurements were taken twice daily the day after the midweek dialysis. We requested readings on only one day per week to increase patient adherence over the one year study. The day after the midweek treatment was chosen because a prior study had demonstrated that BP taken the morning after the midweek HD matched most closely with the average of twice daily home and pre- and postdialysis readings over 7 days.<sup>7</sup> Quarterly 44-hour ambulatory blood pressure monitoring (ABPM), beginning immediately after a midweek HD, was obtained in all sites except Boston. We collected intradialysis BP readings (taken every 30 minutes) from one dialysis session every other week. These were averaged starting at time 0 and excluding the standardized pre- and postdialysis readings. Intra- and postdialysis BPs were measured via the sphygmomanometer attached to the dialysis machine.

## Outcome Variables

Cardiac magnetic resonance imaging was done the day after the midweek treatment at both baseline and 12 months. Left ventricular mass, excluding papillary muscle, was estimated by manual tracing. LVH was defined as LVMI (LVM/ body surface area)  $> 84.1 \text{ g/m}^2$  in males and  $> 66.8 \text{ g/m}^2$  in females.<sup>11,12</sup>

## Statistical Analyses

Data were described as boxplots of the mean of observed values per individual for each BP type (pre-, post-, intradialytic, home, ABPM). Linear mixed models were constructed with patients as random effects to obtain least squares (LS) means with 95% confidence intervals (95% CIs) for differences between home and other BP measures after accounting for non-independence of observations and adjusting for time since randomization, treatment arm, and an interaction between time and treatment arm. Pre-, post-, and intradialysis BP were time matched to the closest home sitting the following day. The daytime (8 AM to 10 PM) average from the ABPM was matched to the average of the AM and PM home BP the following day. Dipping was defined as a decrease in average nocturnal SBP of 10% or more as compared to the average daytime SBP.

## Cluster Analysis

Using a hierarchical clustering analysis ( R version 3.6.0) we constructed a dissimilarity matrix and calculated Euclidian distances<sup>13</sup> based on the average and variability of differences between matched predialysis and home SBP per patient. We chose Ward's minimum variance method,<sup>13,14</sup> a clustering structure based on an agglomerative coefficient.

We compared characteristics across clusters using generalized linear models adjusted for treatment arm. Continuous data were log transformed if not normally distributed. Continuous data were expressed as LS means with 95% CIs. Count (IDH rate, cramps rate) and categorical data were modeled with generalized linear mixed effects modeling. To examine potential differences in LVMI and the change in LVMI over 12 months by cluster, we constructed a linear regression model with LVMI (or change in LVMI, respectively) as the dependent variable with adjustment for sex and treatment arm.

Reported p values are two-sided, and were considered statistically significant at  $p < 0.05$ . Analyses were conducted using SAS 9.4 (SAS Institute Inc., Cary, NC) and R version 3.6.0.

## RESULTS

Characteristics of the 97 participants with 6 matched pairs of home and predialysis readings are shown (Table 1). The median number (IQR) of weekly BP pairs per individual was 24 (15, 38). These participants were not significantly different from the original 126 BID participants (Table S1). The mean age of the study population was 56.7, while 48.5% were African American, 54% had diabetes as the cause of ESKD, 4.1% had a history of myocardial infarction, 8.3% of peripheral vascular disease, and 12.4% of congestive heart failure. The median time on dialysis prior to study start was 3.0 years, the median (IQR) duration of a dialysis treatment was 3.8 (3.5–4.0) hours. The median (IQR)

number of antihypertensive medications prescribed was 3 (2, 3) and 21% held at least one antihypertensive prior to starting dialysis.

### Differences between Blood Pressure Measurements

The distributions of pre-, post-, intradialytic and home SBP measurements in the 97 patients who had 6 pairs of matched midweek standardized dialysis unit and next day home readings and in the 72 patients who had all of the former readings as well as ABPM are shown (Figure 1, Table S2). Predialysis SBP was 5.91 (4.50, 7.33) mm Hg higher than home SBP (Table 2). Intradialytic and ABPM were similar to home SBP, postdialysis was lower than home SBP. Home, intradialytic, pre- and postdialysis diastolic BP (DBP) readings were similar.

### Pattern of the Home-to-Predialysis Difference over Repeated Measures within Individuals

Ninety-seven participants were divided into three clusters based on the within-patient differences between predialysis and home SBP and the variability in these differences over one year. There were 31 (32%), 36 (37%), and 30 (31%) patients allocated to clusters 1 ('home lower'), 2 ('home and predialysis similar'), and 3 ('home higher'), respectively. The LS mean (95% CI) predialysis minus home SBP differences were 19.1 (17.0, 21.1) vs. 3.7 (1.6, 5.8) vs. -9.7 (-12.0, -7.4) mm Hg in clusters 1, 2 and 3, respectively ( $p < 0.001$ ). Home SBP was *lower* than predialysis by 10 mm Hg for 69%, 34%, and 17% of readings per patient in Clusters 1, 2 and 3, respectively, while, home was *higher* than predialysis SBP by 10 mm Hg for 9%, 24% and 48% in Clusters 1, 2 and 3, respectively. Patients in cluster 1 were older than those in clusters 2 and 3, however, these differences did not attain statistical significance (Table 3a). There were no significant differences in sex, race, vintage, cause of ESKD, body mass index, smoking status, the number of antihypertensive drugs taken or the number held prior to dialysis at baseline or in quarter 4, respectively, by cluster. There was no difference in predialysis systolic BP across clusters (Table 3b). BP declined during dialysis in Clusters 1 and 2 (LS means pre-post systolic difference (95% CI) 10.0 (5.4, 14.6), and 9.3 (5.0, 13.6) mm Hg, respectively) while there was a nonsignificant increase in Cluster 3 (-3.3 (-8.0, 1.4) mm Hg). Postdialysis BP was higher in Cluster 3 than Clusters 1 and 2 (LS means systolic BP: 151.7 vs. 142.4 vs. 142.9 mm Hg, and diastolic BP 80.4 vs. 71.7 vs. 77.9 mm Hg, respectively). The daytime ABPM ( $n=56$ ) was higher for systolic in Cluster 3 vs. Clusters 1 and 2 (LS means 149.0 vs. 135.6 and 146.6 mm Hg, respectively,  $p=0.05$ ) and for diastolic BP (86.0 vs. 72.8 and 81.1 mm Hg, respectively,  $p < 0.01$ ). Non-dipping status was highly prevalent and did not differ by cluster (100%, 85%, and 87% in Clusters 1, 2 and 3, respectively). Interdialytic weight gain (IDWG) did not differ by cluster. Intradialytic hypotension (1 or more occurrences of intradialytic SBP  $\geq 90$  mm Hg during a treatment) was less frequent in Cluster 3 vs. Clusters 1 and 2 (incidence rates per 100 patient-treatments (95% CI) 1.8 (1.0, 3.1) vs. 4.9 (2.9, 8.1) vs. 3.2 (2.3, 4.3), respectively  $p=0.04$ ).

### Left Ventricular Hypertrophy (LVH) and Mass Index (LVMI) by Cluster

The prevalence of LVH tended to be higher in Cluster 3 (61.5%) vs. Clusters 1 (36.7%) and 2 (37.0%), although these differences did not attain statistical significance. Figure 2 shows boxplots of LVMI at 12 months after study end, stratified by sex and cluster. After

adjusting for sex ( $p=0.02$ ) and treatment arm ( $p=0.53$ ), LVMI was higher in Cluster 3 versus Clusters 1 and 2 with differences in LS means of  $10.6 \text{ gm/m}^2$  (SE 4.96,  $p=0.04$ ) and  $12.0 \text{ gm/m}^2$  (SE 5.08,  $p=0.02$ ), respectively. The difference between clusters 1 and 2 was  $1.42 \text{ gm/m}^2$  (SE 4.89,  $p=0.8$ ). In a simple linear regression relating the one year averaged predialysis-to-home systolic BP difference with LVMI, there was an increase in LVMI with declining negative values of the predialysis-home BP difference (Fig S2). This is consistent with the results of the cluster analysis showing higher LVMI in Cluster 3 ('home higher').

### **Predialysis-to-Home Systolic BP Difference and Change in Left Ventricular Mass Index Over 12 months**

In a model adjusted for sex and treatment arm, there was no difference by Cluster in the change in LVMI over 12 months (predicted change (95% confidence) in Cluster 1, 2 and 3 of 1.41 (-3.33, 6.14), 2.60 (-2.38, 7.58), and 1.17 (-3.93, 6.26) g/m, respectively,  $p=0.91$ ). The predialysis-to-home BP difference treated as a continuous variable was also not significantly related to the change in LVMI over 12 months (Figure S3).

## **DISCUSSION**

Past studies indicate that on average, BP runs lower at home than predialysis and this is also seen in this study. However, when we examine the home-to-predialysis BP difference within individuals over repeated measures of each obtained over the course of a year, we find that for nearly one third of patients, BP was usually *higher* at home than predialysis. Moreover, this group of patients (Cluster 3) had higher LVMI than those in Cluster 1 ('home lower') or 2 ('home and predialysis similar'). The clusters were indistinguishable with regards to predialysis SBP and interdialytic weight gain. The 'home higher' group had a small increase in BP during dialysis, in contrast to a significant decline in BP during dialysis that was seen in the other patients. Postdialysis BP was also significantly higher in Cluster 3 than Clusters 1 and 2. The lack of a decline in BP with dialysis may be a signal identifying a patient who will exhibit a 'home higher' BP pattern. The higher LVMI among patients in Cluster 3, despite similar predialysis BPs, indicates that home and postdialysis BPs may better reflect the average BP load than the predialysis BP. The higher LVMI in this group suggests that their high blood pressure at home is under recognized and undertreated.

The present study supports the European Renal Association guidelines recommending home over dialysis unit BPs to guide management of dialysis patients.<sup>15</sup> The evidence to support such recommendations is quite limited. A study of predominantly African Americans ( $n=140$ ) who underwent a single ABPM session and two weeks of intermittent home and dialysis unit BP readings showed higher predictive accuracy for LVH of home followed by ABPM, pre- and postdialysis readings<sup>6</sup>. In largely the same population, the risk for cardiovascular and all-cause mortality increased monotonically with increasing quartile of home BP and ABPM, which was not found for pre- and postdialysis readings<sup>16</sup>. From these results, one does not appreciate that there is a group of patients, (sizable in the present study at 30%) that run higher BPs at home than in the dialysis unit and have higher LVMI than other patients. Such findings provide further evidence of the value of home readings and are

interpreted in clinical terms more readily than a series of ROC curves or the linearity of the relationship of quartiles of BP with an outcome.

Pertaining to whether intermittent home readings vs. ABPM should be the preferred method of out-of-dialysis-unit BP measurement, this study found poor adherence with quarterly ABPM, with completion rates of a single ABPM during Q1–4 of 32%, 29%, 28% and 58%, respectively.<sup>8</sup> This was also found in the Hypertension in Hemodialysis Patients Treated with Atenolol or Lisinopril Study.<sup>17</sup> ABPM certainly poses additional logistical challenges, and, given our experience, it may not be practical. A recent US study testing the feasibility of home BP monitoring, found a high rate of adherence (97.4%) with intermittent home readings measured twice daily for one day every two weeks over 4 months<sup>34</sup>.

The pattern of home vs. predialysis unit BP seen in Cluster 3 is similar in concept to masked hypertension in the general population, in which BP is elevated at home but not in the office. Masked hypertension, estimated to affect 13% of the general population,<sup>18</sup> is associated with increased LVMI<sup>19</sup> and an increased risk of major adverse cardiovascular events as compared with normotension or white coat hypertension.<sup>20,21</sup> Given the strong and consistent relationship of LVMI with cardiovascular events, and that this risk is altered with a change in LVMI,<sup>22–26</sup> we suspect that the patients in Cluster 3 are at increased risk. Whether more aggressive BP lowering could reduce this risk remains to be proven.

Why for some patients BP is higher at home than in the office, or in this case, the dialysis unit, is not clear. Smoking and use of a greater number of antihypertensive agents have been associated with masked hypertension in the general population.<sup>27</sup> We found no difference in smoking status, the number of antihypertensive drugs used or the frequency at which drugs were withheld prior to dialysis by cluster. That BP is higher at home the next day than predialysis is particularly striking as the predialysis BP is measured at the highest degree of fluid overload in the dialytic cycle. One may postulate that Cluster 3 patients would have higher IDWGs driving the higher BP at home however, we found no difference in IDWG by cluster. Alcohol use (not measured in this study), stress, endothelial dysfunction, other factors, as discussed in the masked hypertension and intradialytic hypertension, may be operative.<sup>28–30</sup>

This study has unique strengths. Data was collected over one year as opposed to previous studies that examined only a few weeks of BP measurements.<sup>6,7</sup> This permitted characterization of direction, magnitude and variability of the dialysis unit minus home BP differences within individuals and assessment of long term adherence. Great care was taken to measure the predialysis BPs per a standardized protocol, including use of stopwatches to time the 5 minute rest period prior to the first measurement, attention to other aspects of the recommended procedure for measuring BP,<sup>10</sup> and certification of staff and patients. The BP measurements on which these analyses are based are likely to be more accurate than routine dialysis unit readings. As standardized BP readings are usually lower than routine readings, the predialysis-to-home BP difference may be greater than that reported here. LVMI was assessed with MRI, which is more accurate than echocardiogram.<sup>31</sup>

The study also has several limitations. The study population was relatively small which limited the statistical power to detect differences across clusters. The smaller sample size also restricted our choice of methods for forming the clusters. Although we recognize that latent mixture modeling methods are often preferred over a hierarchical approach, these require significantly larger sample sizes.<sup>32,33</sup> However, when we attempted to use the latent class methods, the clusters displayed a moderate level of agreement with the clusters reported in the manuscript. The present study was done within the framework of a randomized trial, raising the possibility that differences in clusters might be due to the intervention. We think this is less likely as the proportions by treatment arm did not differ across clusters (Table 3), analyses were adjusted for treatment arm and there was a non-significant trend towards a higher proportion with LVH in Cluster 3 patients at baseline, prior to randomization (Table 3). The number of paired readings per individual varied, mainly due to missing home readings. There was no difference by the quantity of paired readings ( 12 vs. >12 pairs) in the proportion of patients within Cluster 3 (Table S1B) or in the predialysis-to-home BP difference (Table S1C). Nonetheless, we cannot exclude the possibility that the missing data may have introduced bias. The study participants did not represent the general hemodialysis population since eligibility required a 2 week standardized predialysis SBP >155 mmHg and the absence of frequent intradialytic hypotension. The prevalence estimates of the home higher pattern, may be lower in a less hypertensive population. The present study did not demonstrate that treating patients based on home readings as compared with dialysis unit readings would improve outcomes or identify the optimal BP target.

In summary, this study finds that for that in close to one-third of hypertensive dialysis patients, BP runs higher at home than predialysis and these patients had a higher LVMI than the other patients. Accordingly, these patients may be at higher risk for cardiovascular events. Dialysis facilities should educate patients and staff on obtaining out-of-dialysis-unit BP readings and undertake regular programs to enhance and maintain adherence.

## Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

## Acknowledgements:

There was no direct financial support for this manuscript. The BID Study was supported by National Institute of Diabetes and Digestive and Kidney Diseases (R01DK083424) and Dialysis Clinic, Inc.

## Disclosures

Drs. Miskulin and Pankratz receive salary support and Dr. Shaffi receives research support from Dialysis Clinic, Inc. (DCI); H. Jiang, S. Paine, Drs. Gul, Harford, and Zager are employees of DCI; Drs. Jhamb, Kwong, Negrea, Gassman and Ploth have no disclosures.

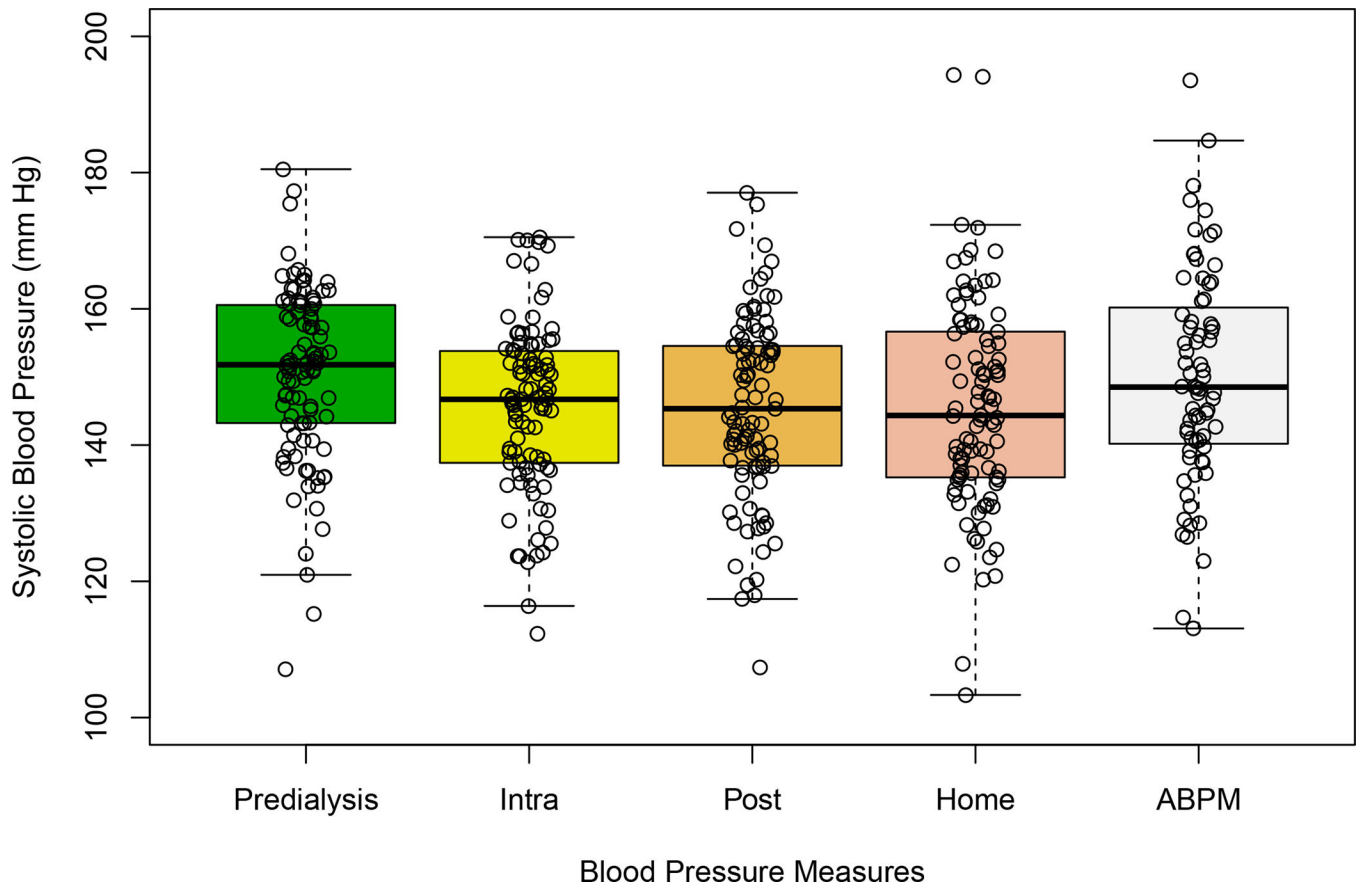
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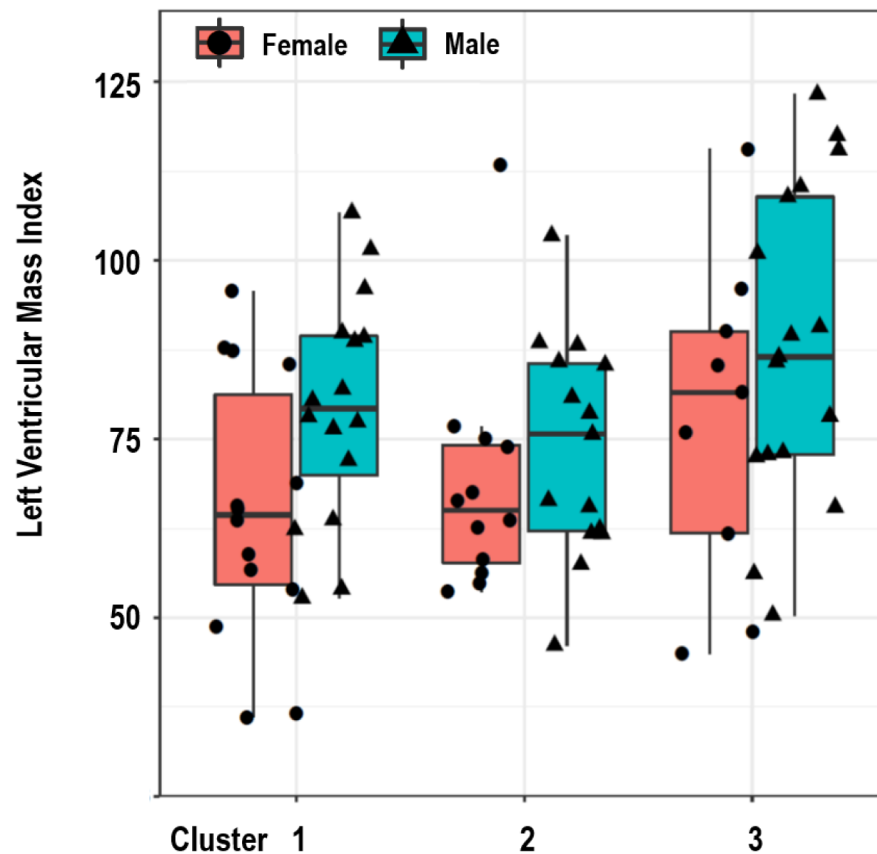
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**Figure 1. Distributions of Standardized Predialysis, Home, Postdialysis, and Average Intradialysis Systolic Blood Pressures**

Figure 1. Boxplots with jitter show the distributions of the midweek pre-(standardized), post-, and intradialysis blood pressure, next day home blood pressure readings averaged over all dialysis sessions over the course of the year in each of 97 patients, and all possible ABPM over the year in each of 72 patients. Each data point represents one participant.



n =	14	16	12	15	9	17
Observed mean	65.0	79.5	68.5	73.9	77.7	88.1
(SD)	(18.7)	(15.9)	(16.2)	(15.2)	(22.9)	(22.0)
Adjusted Mean	65.0	79.3	68.5	73.8	77.8	87.9
(SE)	(5.0)	(4.7)	(5.4)	(5.0)	(6.2)	(4.5)

**Figure 2. Observed and Adjusted Mean Left Ventricular Mass Index in Quarter 4 by Sex and Cluster**

Figure 2. Boxplots with jitter of left ventricular mass index by sex and cluster. The observed and adjusted LS means (SE) are shown in the table. LS means (SE) were calculated from a general linear regression model, adjusting for sex, cluster, treatment arms, and sex X cluster interaction. LVMI was higher in cluster 3 versus clusters 1 and 2 with differences in LS means of 10.6 gm/m<sup>2</sup> (SE 4.96, p=0.04) and 12.0 gm/m<sup>2</sup> (SE 5.08, p=0.02), respectively.

**Table 1.**

Baseline characteristics (N = 97).

<b>Randomized to Intensive study arm</b> <sup>a</sup>	46 (47.4)
<b>Male</b> <sup>a</sup>	56 (57.7)
<b>Race</b> <sup>a</sup>	
White	35 (36.1)
Black	47 (48.5)
Others	15 (15.5)
<b>Age (years)</b> <sup>b</sup>	56.7 (13.4)
<b>Vintage (years)</b> <sup>c</sup>	3.0 (1.1, 4.7)
<b>Cause of end stage kidney disease</b> <sup>a</sup>	
Diabetes mellitus	52 (53.6)
Hypertension	28 (28.9)
Glomerulonephritis	8 (8.3)
Other	9 (9.3)
<b>Comorbid conditions</b> <sup>a</sup>	
Myocardial infarction	4 (4.1)
Congestive heart failure	12 (12.4)
Peripheral vascular disease	8 (8.3)
Current or Former Smoking history	45 (46.4)
<b>Body mass index (kg/m<sup>2</sup>)</b> <sup>c</sup>	26.2 (22.9, 30.2)
<b>Interdialytic weight gain (kg)</b> <sup>d</sup>	2.7 (1.2)
<b>Interdialytic weight gain (% of estimated dry weight)</b> <sup>d</sup>	3.5 (1.3)
<b>Prescribed treatment time (hours)</b> <sup>e</sup>	3.8 (3.5, 4.0)
<b>Ultrafiltration volume (L)</b> <sup>e</sup>	2.9 (2.3, 3.5)
<b>Predialysis systolic blood pressure (mm Hg)</b> <sup>d</sup>	159.4 (9.2)
<b>Predialysis diastolic blood pressure (mm Hg)</b> <sup>d</sup>	80.0 (11.6)
<b>Number of antihypertensive medications taken</b> <sup>d</sup>	2.6 (1.3)
<b>Patients who had antihypertensive medication held predialysis</b> <sup>a</sup>	
1 medication held	14 (14.4)
2 medications held	2 (2.1)
<b>Percent of patients receiving specific classes of antihypertensive medications</b>	
Angiotensin converting enzyme inhibitor	42
Angiotensin receptor blocker	18
Beta-adrenergic blocking agent	75
Calcium channel blocker	60
Central alpha agonist	19

<sup>a</sup>  
n(%)

<sup>b</sup>  
mean (standard deviation)

<sup>c</sup>  
median (interquartile range)

<sup>d</sup>  
mean (standard deviation) for the month prior to randomization

<sup>e</sup>  
median (interquartile range) for the month prior to randomization

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**Table 2.**

## LS Means Differences between Home and Other Blood Pressure Measurements

<b>BP Difference</b>	<b>No. of patients</b>	<b>No. of BP pairs per patient</b>	<b>Systolic BP (mm Hg)</b>	<b>P value</b>	<b>Diastolic BP (mm Hg)</b>	<b>P value</b>
Predialysis – Home		24(15, 38)	5.91 (4.50, 7.33)	<0.001	0.16 (–0.54, 0.85)	0.9
Intradialysis – Home	97	24(16,33)	–0.74 (–2.18, 0.69)	0.9	0.56 (–0.15, 1.27)	0.2
Postdialysis – Home		24(13,38)	–1.25 (–2.62, 0.13)	0.1	0.54 (–0.14, 1.22)	0.2
ABPM – Home	72	2 (2,3)	0.57 (–2.09, 3.23)	0.7	3.99 (2.77, 5.21)	< 0.001

Predialysis, average intradialytic BP, and postdialysis BP were time matched to the closest home BP the following day. The daytime average from the ABPM was matched to the average of the AM and PM home BP. LS mean differences were estimated in linear mixed effects models adjusted for treatment arm, time between BP measurements and randomization.

. BP, blood pressure; ABPM, ambulatory blood pressure monitoring.

**Table 3a:**

## Patient Characteristics Across Clusters

	Total	Cluster 1 “Home Lower”	Cluster 2 “Home and Predialysis Similar”	Cluster 3 “Home Higher”	<i>p</i> value <sup>e</sup>
<b>No. of patients</b>	<b>97</b>	<b>31</b>	<b>36</b>	<b>30</b>	
<b>Age (years)</b> <sup>a</sup>	56.7 (13.4)	61.0 (12.9)	54.3 (13.4)	55.1 (13.2)	0.09
<b>Male</b> <sup>b</sup>	56 (57.7)	16 (51.6)	19 (52.8)	21 (70.0)	0.2
<b>Vintage (years)</b> <sup>c,d</sup>	3.0 (1.1, 4.7)	3.3 (0.9, 5.3)	2.6 (0.7, 3.8)	2.9 (1.4, 4.6)	0.6
<b>Race</b> <sup>b</sup>					
<b>White</b>	35 (36.1)	11 (35.5)	14 (38.9)	10 (33.3)	
<b>Black</b>	47 (48.5)	14 (45.2)	17 (47.2)	16 (53.3)	0.9
<b>Other</b>	15 (15.5)	6 (19.4)	5 (13.9)	4 (13.3)	
<b>Body mass index (kg/m<sup>2</sup>)</b> <sup>c,d</sup>	26.2 (22.9, 30.2)	25.7 (22.6, 31.6)	26.8 (24.3, 31.9)	26.0 (22.5, 29.6)	0.7
<b>Cause of ESKD</b> <sup>b</sup>					
<b>Diabetes</b>	52 (53.6)	17 (54.8)	17 (47.2)	18 (60.0)	
<b>Hypertension</b>	28 (28.9)	9 (29.0)	13 (36.1)	6 (20.0)	0.9
<b>Glomerulonephritis</b>	8 (8.3)	2 (6.5)	3 (8.3)	3 (10.0)	
<b>Other</b>	9 (9.3)	3 (9.7)	3 (8.3)	3 (10.0)	
<b>Smoking status</b> <sup>b</sup>					
<b>Never</b>	52 (53.6)	15 (48.4)	24 (66.7)	13 (43.3)	0.2
<b>Current</b>	12 (12.4)	4 (12.9)	5 (13.9)	3 (10.0)	
<b>Former</b>	33 (34.0)	12 (38.7)	7 (19.4)	14 (46.7)	
<b>Number of antihypertensive medications</b> <sup>c</sup>					
<b>Baseline</b>	3 (2, 3)	2 (2, 3)	2 (1, 3)	3 (2, 4)	0.2
<b>Quarter 4</b>	3 (2, 4)	3 (2, 4)	3 (2, 4)	3 (3, 4)	0.3
<b>Percent of patients who had 1 or more antihypertensive medications held predialysis</b> <sup>b</sup>					
<b>Baseline</b>	20 (20.8)	6 (19.4)	8 (22.9)	6 (20.0)	0.8
<b>Quarter 4</b>	26 (27.1)	6 (19.4)	13 (37.1)	7 (23.3)	0.3
<b>BID randomized arm</b> <sup>b</sup>					
<b>Intensive Arm</b>	46 (47.4)	15 (48.4)	18 (50.0)	13 (43.3)	0.8
<b>Left ventricular hypertrophy</b> <sup>b</sup>					
<b>Baseline</b>	40 (48.2)	14 (46.7)	11 (40.7)	15 (57.7)	0.4
<b>Quarter 4</b>	37 (44.6)	11 (36.7)	10 (37.0)	16 (61.5)	0.1

<sup>a</sup> mean (standard deviation)<sup>b</sup> n(%)<sup>c</sup> median (interquartile range)<sup>d</sup> Log transformed for statistical testing



<sup>e</sup>The p-value represents results of comparing factor by cluster using generalized linear regressions adjusted for treatment arm

Abbreviations: BID, Blood Pressure in Dialysis Pilot Study.

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**Table 3b:**

## Blood Pressure Measurements and Dialysis Treatment Characteristics Across Clusters

	Total	Cluster 1 “Home Lower”	Cluster 2 “Home and Predialysis Similar”	Cluster 3 “Home Higher”	<i>p</i> value <sup>c</sup>
No. of patients	97	31	36	30	
Total No. of BP pairs per cluster	2479	951	852	676	
<b>Cluster Definition</b>					
Predialysis minus home systolic BP (mm Hg) <sup>a</sup>	4.8 (2.2, 7.4)	19.1 (17.0, 21.1)	3.7 (1.6, 5.8)	-9.7 (-12.0, -7.4)	<.001
<b>Blood Pressure (BP) Measurements</b>					
<b>Percent of observations per patient in which predialysis minus home systolic BP<sup>b</sup></b>					
10 mm Hg	36.1 (21.4, 60.0)	68.8 (57.1, 75.0)	33.7 (27.4, 43.2)	17.2 (9.1, 23.5)	<.001
± <10 mm Hg	29.6 (21.9, 44.4)	21.9 (16.7, 31.6)	42.4 (28.6, 50.0)	29.6 (24.0, 42.9)	<.001
-10 mm Hg	23.5 (12.5, 41.9)	8.7 (2.6, 12.5)	23.5 (21.1, 30.0)	48.3 (41.9, 57.1)	<.001
<b>Predialysis BP (mm Hg)<sup>a</sup></b>					
Systolic	150.9 (148.7, 153.1)	152.3 (148.5, 156.1)	151.8 (148.2, 155.3)	148.3 (144.4, 152.2)	0.2
Diastolic	76.6 (74.5, 78.8)	72.2 (68.5, 75.9)	80.6 (77.2, 84.0)	76.4 (72.7, 80.2)	<.001
<b>Postdialysis BP (mm Hg)<sup>a</sup></b>					
Systolic	145.4 (142.7, 148.2)	142.4 (137.7, 147.0)	142.9 (138.5, 147.2)	151.7 (146.9, 156.4)	<.001
Diastolic	76.7 (74.7, 78.7)	71.7 (68.5, 75.0)	77.9 (74.8, 80.9)	80.4 (77.1, 83.8)	<.001
<b>Pre- minus postdialysis BP change (mm Hg)<sup>a</sup></b>					
Systolic	7.3 (4.7, 9.9)	10.0 (5.4, 14.6)	9.3 (5.0, 13.6)	-3.3 (-8.0, 1.4)	<.001
Diastolic	0.7 (-0.6, 2.0)	0.3 (-2.1, 2.7)	2.3 (0.0, 4.5)	-4.2 (-6.7, -1.7)	<.001
<b>Home BP* (mm Hg)<sup>a</sup></b>					
Systolic	145.3 (142.2, 148.3)	132.8 (128.7, 136.9)	146.8 (143.0, 150.7)	156.6 (152.3, 160.9)	<.001
Diastolic	76.5 (74.1, 78.9)	68.2 (64.4, 71.9)	80.3 (76.8, 83.8)	80.4 (76.6, 84.2)	<.001
<b>ABPM daytime BP<sup>#</sup> (mm Hg)<sup>a</sup></b>					
Systolic	143.0 (137.9, 148.1)	135.6 (127.9, 143.3)	146.6 (138.2, 154.9)	149.0 (140.3, 157.8)	0.05
Diastolic	79.3 (75.9, 82.7)	72.8 (68.0, 77.7)	81.1 (75.9, 86.4)	86.0 (80.4, 91.5)	<.001
<b>ABPM nocturnal BP<sup>#</sup> (mm Hg)<sup>a</sup></b>					
Systolic	142.3 (136.8, 147.7)	137.7 (129.1, 146.3)	145.3 (136.0, 154.6)	144.9 (135.0, 154.8)	0.4
Diastolic	76.2 (72.8, 79.6)	71.0 (66.0, 76.0)	77.6 (72.2, 83.1)	81.5 (75.8, 87.3)	0.02
<b>Dialysis Treatment Characteristics<sup>e</sup></b>					
Length of dialysis session (minutes) <sup>a</sup>	219.8 (214.7, 224.8)	217.1 (208.0, 226.2)	220.4 (211.9, 228.8)	222.1 (212.8, 231.5)	0.7

	<b>Total</b>	<b>Cluster 1 “Home Lower”</b>	<b>Cluster 2 “Home and Predialysis Similar”</b>	<b>Cluster 3 “Home Higher”</b>	
<b>No. of patients</b>	<b>97</b>	<b>31</b>	<b>36</b>	<b>30</b>	
<b>Total No. of BP pairs per cluster</b>	<b>2479</b>	<b>951</b>	<b>852</b>	<b>676</b>	<i>p</i> value <sup>c</sup>
<b>Intradialytic hypotension<sup>d</sup> (rate per 100 treatments)</b>	3.5 (2.6, 4.7)	4.9 (2.9, 8.1)	3.2 (2.3, 4.3)	1.8 (1.0, 3.1)	0.04
<b>Intradialytic Cramps (rate per 100 treatments)</b>	7.2 (5.8, 8.9)	7.5 (5.1, 11.0)	7.2 (5.8, 9.0)	7.0 (4.7, 10.3)	0.9
<b>Interdialytic weight gain (kg)<sup>a</sup></b>	2.7 (2.5, 2.8)	2.5 (2.2, 2.8)	2.9 (2.6, 3.1)	2.6 (2.3, 2.9)	0.2

<sup>#</sup>Missing ABPM data n=21, n=24, and n=18, for clusters 1, 2, and 3, respectively.

<sup>a</sup>LS Mean (95% confidence interval)

<sup>b</sup>median (interquartile range)

<sup>c</sup>The p-value represents the results of comparing factor by cluster using generalized linear mixed effects models adjusted for treatment arm

<sup>d</sup>Defined as the occurrence of 1 or more episode of intradialytic systolic BP <90 mm Hg. Derived from n=126,158 dialysis treatments.

<sup>e</sup>The median (IQR) number of dialysis treatments from which these variables were derived was as follows: 42(33,46) for treatment duration, 144(126,158) for intradialytic hypotension and cramps, and 152(138,157) for interdialytic weight gain.

Abbreviations: ABPM, ambulatory blood pressure monitoring