## EXCESS RIGHT ATRIAL CONGESTION AND PULMONARY CIRCULATION LOADING OVER THE LONG INTERDIALYTIC INTERVALIN PATIENTS RECEIVING THRICE-WEEKLY HEMODIALYSIS

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**Introduction:** Among hemodialysis patients, pulmonary hypertension is a strong and independent predictor of all-cause and cardiovascular mortality [1-3]. It remains unclear whether volume accumulation over the interdialytic period acutely influences right ventricular (RV) sizing and function [1-3]. Accordingly, this study investigated the changes RV function during the 2-day and 3-day interdialytic intervals.

Materials and Methods: Forty-one stable end-stage-renal-disease patients receiving thrice-weekly hemodialysis for at least 3 months underwent 2-dimensional echocardiographic and tissue-Doppler imaging evaluation with a standard cardiac ultrasound device (Vivid 7, GE, Horten, Norway) at start and end of the 3-day and 2-day interdialytic intervals. Participants were randomly assigned into two cross-over sequences of recordings; 3-day followed by 2-day interval or vice versa.

Results: The baseline demographic characteristics and routine predialysis biochemical parameters of study participants are presented in Table (1). Study participants had a mean age of 58.7±14.7 years and were receiving hemodialysis for a mean time-period of 57.8±57.2 months. As shown in Table 2, Inferior-vava-diameter (IVCD), a marker reflecting intravascular volume among patients on dialysis [4], increased during both interdialytic intervals. It was rather expectable that IVCD increase (0.54±0.3 vs 0.25±0.2) was higher during the 3-day versus the 2-day interval (P<0.001). RV systolic function assessed by SM RV and tei index of myocardial performance was stable during both intervals. Significant elevations in right atrial (RA) volume index, RV enddiastolic diameter, E/Em RV, tricuspid regurgitation maximum velocity (TRVmax) and tricuspid regurgitation peak gradient (PGr TRV) were noted during both 3-day and 2-day interdialytic periods. Increase in RV systolic pressure (RVSP) was also evident in both intervals (3-day: 29.3±12.5 vs 44.6±16.3 mmHg, P<0.001; 2-day: 33.1±12.4 vs 37.9±13.7 mmHg, P<0.01). Interdialytic increase in RVSP, TRVmax and PGr TRV was more prominent during the 3-day compared with the 2day interdialytic interval (Table 3). In order to identify possible factors associated with the interdialytic increase in RVloading, we have performed a univariate and multivariate linear regression analysis with change in RVSP being the dependent variable. As shown in Table 4, in multivariate analysis IDWG (β=1.645; 95% CI= 0.332 - 2.958), E/Em RV change ( $\beta$ =1.142, 95% CI= 0.293 - 1.990), and PVR change  $(\beta=7.034, 95\% \text{ CI} = 1.766 - 12.303)$  were shown to be independently associated with changes in RVSP, whereas changes BP, HR and Sm RV changes were not.

Conclusion: Excess fluid accumulation results in higher RA congestion and pulmonary circulation loading, providing one plausible pathway for the excess cardiovascular risk associated with the long interdialytic interval in thrice-weekly hemodialysis.

## REFERENCES

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Table 1: Baseline characteristics and routine predialysis biochemical parameters of study participants.

| Characteristic                | Value      |  |  |
|-------------------------------|------------|--|--|
| N                             | 41         |  |  |
| Age (years)                   | 58.7±14.7  |  |  |
| Male, n (%)                   | 24 (59%)   |  |  |
| Height (m)                    | 1.65±0.1   |  |  |
| BMI (kg/m <sup>2</sup> )      | 25.2±4.8   |  |  |
| Duration of dialysis (months) | 57.8±57.2  |  |  |
| Hemoglobin (g/dl)             | 12.7±4.4   |  |  |
| Hematocrit (%)                | 36.5±3.3   |  |  |
| Serum glucose (mg/dl)         | 98.5±36.3  |  |  |
| Serum urea (mg/dl)            | 141.7±31.8 |  |  |
| Serum creatinine (mg/dl)      | 8.3±3.0    |  |  |
| Total protein (g/dl)          | 7.4±0.8    |  |  |
| Serum albumin (g/dl)          | 4.3±0.4    |  |  |
| Serum calcium (mg/dl)         | 9.0±0.8    |  |  |
| Serum phosphate (mg/dl)       | 4.7±1.1    |  |  |

**Table 2:** Echocardiographic indices of RV function and pressure during the 3-day and 2-day interdialytic intervals.

|                                 | 3-day Interdialytic Interval                |                                      |         | 2-day Interdialytic Interval                |                                      |            |
|---------------------------------|---|--------------------------------------|---------|---|--------------------------------------|------------|
| Parameter                       | Visit 1<br>(post-<br>HD,<br>3-day<br>start) | Visit 2<br>(pre-HD,<br>3-day<br>end) | P Value | Visit 3<br>(post-<br>HD,<br>2-day<br>start) | Visit 4<br>(pre-HD,<br>2-day<br>end) | P<br>Value |
| Sm RV (m/sec)                   | 0.14±0.04                                   | 0.15±0.04                            | 0.4     | 0.15±0.04                                   | 0.14±0.03                            | 0.2        |
| E/Em RV                         | 7.0±2.3                                     | 8.5±3.3                              | < 0.001 | 7.1±2.6                                     | 8.0±2.3                              | 0.05       |
| TRV max (m/sec)                 | 2.41±0.5                                    | 2.87±0.5                             | < 0.001 | 2.54±0.5                                    | 2.68±0.5                             | 0.009      |
| PGr TRV (mmHg)                  | 24.1±10.9                                   | 33.5±12.7                            | < 0.001 | 26.8±12.4                                   | 29.7±12.9                            | 0.008      |
| RVSP (mmHg)                     | 29.3±12.5                                   | 44.6±16.3                            | < 0.001 | 33.1±12.4                                   | 37.9±13.7                            | < 0.001    |
| IVC diameter (cm)               | 1.40±0.4                                    | 1.93±0.4                             | < 0.001 | 1.54±0.4                                    | 1.79±0.4                             | < 0.001    |
| RVEDD (cm)                      | 3.2±0.8                                     | 3.6±0.9                              | < 0.001 | 3.1±0.8                                     | 3.4±0.8                              | < 0.001    |
| RAVi (ml/m²)                    | 20.9±9.6                                    | 27.3±10.6                            | < 0.001 | 21.3±10.1                                   | 25.0±11.5                            | 0.002      |
| Tei index RV                    | 0.56±0.2                                    | 0.52±0.20                            | 0.2     | 0.59±0.20                                   | 0.50±0.13                            | 0.5        |
| PVR (dyn.sec.cm <sup>-5</sup> ) | 1.58±0.4                                    | 1.63±0.4                             | 0.5     | 1.72±1.02                                   | 1.59±0.4                             | 0.3        |

**Table 3:** Comparison of the net changes in echocardiographic indices of RV sizing and function between the 3-day and 2-day interdialytic intervals

|                                 | 3-day vs 2-day Interval                     |              |               |  |
|---------------------------------|---|--------------|---------------|--|
| Parameter                       | Net Changes Net Change 3-day 2-day interval |              | es<br>P value |  |
| Sm RV (m/sec)                   | 0.006±0.042                                 | -0.007±0.038 | 0.1           |  |
| E/Em RV                         | 1.6±2.2                                     | 0.9±2.7      | 0.2           |  |
| TRV max (m/sec)                 | 0.46±0.45                                   | 0.14±0.33    | 0.001         |  |
| PGr TRV (mmHg)                  | 9.4±10.2                                    | 2.9±6.6      | 0.001         |  |
| RVSP (mmHg)                     | 15.3±10.2                                   | 4.7±5.2      | < 0.001       |  |
| IVC diameter (cm)               | 0.54±0.3                                    | 0.25±0.3     | < 0.001       |  |
| RVEDD (cm)                      | 0.4±0.8                                     | 0.3±0.5      | 0.4           |  |
| RAVi (ml/m²)                    | 6.7±6,4                                     | 3.7±7.1      | 0.05          |  |
| Tei index RV                    | -0.04±0.23                                  | -0.09±0.18   | 0.3           |  |
| PVR (dyn.sec.cm <sup>-5</sup> ) | -0.04±0.41                                  | -0.02±0.35   | 0.5           |  |

Table 4: Univariate and multivariate linear regression analysis of parameters possibly affecting the change in RVSP during interdialytic intervals

|                | Univariate analysis |                   |            | Multivariate analysis |                 |            |
|----------------|---------------------|-------------------|------------|-----------------------|-----------------|------------|
| Parameter      | Estimate            | 95% CI            | P<br>value | Estimate              | 95% CI          | P<br>value |
| IDWG           | 1.512               | 0.104 to 2.920    | 0.04       | 1.645                 | 0.332 to 2.958  | 0.02       |
| SBP change     | 0.171               | -0.114 to 0.457   | 0.2        |                       |                 |            |
| HR change      | -0.028              | -0.283 to 0.227   | 0.8        |                       |                 |            |
| Sm RV change   | -2.246              | -55.740 to 51.248 | 0.9        |                       |                 |            |
| E/Em RV change | 0.820               | -0.012 to 1.653   | 0.05       | 1.142                 | 0.293 to 1.990  | 0.009      |
| PVR change     | 6.177               | 0.633 to 11.721   | 0.03       | 7.034                 | 1.766 to 12.303 | 0.01       |





