# Factors responsible for increased percent recirculation in arterio-venous fistula among the haemodialysis patients

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### Abstract

Recirculation is an important issue in haemodialysis (HD) patients as increased percent recirculation causes decreased dialysis delivery of the patients. The purpose of the study was to determine the amount and factors of recirculation in those patients. The study was a cross sectional one carried in the department of Nephrology, Dhaka Medical College and Hospital during October 2010 to September 2011. A total of 118 end stage renal disease patients with arterio-venous fistula who were on HD for more than 3 months were purposively selected. The degree of recirculation was measured with urea based two needle technique method. For each patient distances between arterial and venous and distances of needles from fistula and its directions were recorded. Echocardiography and A-V fistula Colour Doppler Ultrasound were also performed. The mean A-V fistula recirculation was 8.1±5.5% with a range 0-66%. The most common factors were close proximity and improper arterial and venous needles placement. No difference was observed between diabetic and non diabetic also between hypertensive and normotensive. A-V fistula recirculation is common occurrence in HD patients and the most common factors of recirculation are misplacement and close proximity of needles therefore emphasis should be given on education and training of HD staffs.

#### Introduction

End stage renal disease (ESRD) is one of most common life threatening diseases and the number of ESRD patients are expanding. Therefore the number of Haemodialysis(HD) patients are also increasing rapidly<sup>1</sup>.World wide HD is method of renal replacement therapy for majority of ESRD patients and survival of haemodialysis patients is an important and very well discussed issue.,<sup>2</sup> Though the life expectancy of patients with ESRD has improved a lot after the introduction of dialysis but unfortunately still it is far below that of the general population. A study from Iran observed that one, three, and five-year survival of ESRD patients on HD is 89.2%, 69.2%, and 46.8%, respectively<sup>3</sup>. Another study observed that the mean life span at age 49 in the United States is 33 years in the general population but it was only approximately 7 years in patients receiving maintenance dialysis <sup>4</sup>.So more emphasis should be given for adequate dialysis. Different factors affect in the survival of ESRD patients, one of the important factors is inadequate dialysis<sup>5,6,7</sup>. It was well established that recirculation is one of the important causes of inadequate dialysis. Recirculation is defined as the volume of blood that has crossed the dialyzer and is

recirculated directly through the extracorporeal circuit again rather than returning to the systemic circulation. There by it cuts down the efficiency of dialysis.8 Many factors are responsible for recirculation especially low blood flow through fistula, low left ventricular ejection fraction ,Stenosis in vessels and improper arterial and venous needle placement and close proximity between needles and can be prevented by appropriate measures.<sup>9,10,13</sup> Another importance of recirculation study is to early detection of access dysfunction especially to predict the presence of venous stenosis and low blood flow through fiatula.<sup>12</sup> According to NFK-DOQI guideline any recirculation more than 10% by two needle urea based method and more than 5% by ultrasound dilution method should be investigated by further extensive investigation to identify the cause.<sup>11</sup> Thus screening for recirculation is an important issue in HD patients.

Due to scarcity of donor, HD is the choice of treatment in majority of patient with ESRD in our community and the treatment cost of HD is very high, especially for our poor people. More over each year Government has to take an extra burden to serve this group of patients and the demand is also increasing day by day. So we should be more careful in dealing with this group of patients and ensure adequate dialysis delivery in each session to minimize our economic burden. The measurement of A-V fistula recirculation in Haemodialysis is an important cause of inadequate dialysis as well as an indicator of access dysfunction.<sup>10,11</sup> Therefore by detecting the amount of recirculation and causes responsible it we can improve the quality of dialysis delivery to our patients. Considering recirculation as an important issue in haemodialysis and as there was no similar such study in Bangladesh, this study was directed to estimation of recirculation among our haemodialysis patients as a baseline work to improve dialysis delivery in future as well as this work will create a ground for future extensive study.

#### **Materials and Methods**

*Objectives:* Different study show that haemodialysis access recirculation is a common occurrence and is a preventable factor for underdialysis. Thereby the study was directed to estimation of recirculation among haemodialysis patients as well as to identify the factors responsible for it.

*Sample Size:* We considered every available ESRD patient on Haemodialysis through A-V fistula at the Department of Nephrology, Dhaka Medical College Hospital. There was no past record of similar study in Bangladesh, we followed a study done by Abdollahi et al<sup>16</sup> as reference. With this the sample size was determined by following formula:

 $n = \frac{Z^2 pq}{d^2}$ = 196.

[Z=1.96, the critical value of z at 95% level of significance. P=0.15 (Considering the prevalence of recirculation 15%) (Abdollahi et al. 2010) q=1-p=0.85 d=0.05 margin of error] Therefore, n=196 (Sample size).

Total 118 patients were included as per selection criterion.

*Sampling Method:* Sampling was done by purposive sampling. Inclusion criteria were Patients undergoing haemodialysis by arterio venous fistula for more than three months, More than 18 years and those who gave consent. Exclusion criteria,

patients undergoing haemodialysis by temporary access or permanent catheter was excluded.

Subjects and Methods/methodology: This was a Cross sectional study from October 2010 to

September 2011 Conducted on ESRD patients in Dept of Nephrology, Dhaka Medical college and Hospital. HD was performed for 4 hours, two to three times per week, using triacetate hollow fiber dialyzer Nipro SUREFLUX-130N, Japan with HD machine Fresenius medical care type 4008B, Germany with bicarbonate-based dialysate at a delivered bicarbonate concentration of 35m Eq/L. Blood flow was maintained at 250-350 ml/min and the dialysate flow rate at 500ml/min.A standardized questionnaire was used to collect demographic data, cause of ESRD, the date of HD onset, the type of access for HD, the date of creation and use of A-V fistula. The degree of recirculation was measured with Urea based two needle method from the following formula: Percent recirculation=([P-A] ÷ [P-V]) x 100.Where P, A, and V refer to the urea concentrations in the peripheral blood, predialyzer arterial line, and postdialyzer venous circuit, respectively. The following protocol was used for blood sampling. The ultrafiltration was turned off approximately 30 minutes after the initiation of HD and then arterial and venous line samples were obtained. Access blood flow was reduced to 50 mL/min and 15 to 30 seconds later, the systemic blood sample from the arterial blood line was obtained<sup>13</sup>. The measurements of all blood samples were done in single laboratory. The patients were divided into three groups: group 1 patients 0-.9%[most well functioning fistula] recirculation and group II patients with recirculation 1-10% [Within acceptable limit]. Group III>10% recirculation. [Above the acceptable limit of recirculation]. For each patient distances between arterial and venous needles and distances of needles from A-V fistula and its directions was recorded. Echocardiography and A-V fistula Color Doppler Ultrasound were also performed. The findings were analyzed by SPSS statistical program. Chi square test or Fisher Exact test were used for qualitative data. For quantitative data, the were compared by using ANOVA test. Association between risk factors and recirculation rate was evaluated by COX regression model. Statistical significance was assessed at the 0.05 probability level in all analyses.

## Results

Maximum of the patients were found in three groups between 41-50 years. In this current study it was observed that the age range varied from 18 to 75 years in whole study patients and majority of the patients were found in the 5<sup>th</sup> decade, which was 52.4% in group I, 30.1% in group II and 42.9% in group III. The mean age was  $43.81\pm11.95$  years in group I,  $43.58\pm12.89$  years in group II and  $44.36\pm11.45$  years in group III. Age difference was not stastially significant (p>0.05).

Male were 92 and female 26 with a male female ratio 2.5:1.male was found 85.7% in group I, 80.7% in group II and 40.0% in group III. Male was significant higher in group I and group II. Female were predominant in group III. The difference was statistically significant (p<0.023).

Among the cause of ESRD, DM was found 9(42.9%) in group I, 39(47.0%) in group II and 4(28.6%) in group III. GN was found 7(33.3%) in group I, 25(30.1%) in group II and 3(21.4%) in group II. Hypertension was found 4(19.0%) in group II. ADPKD and obstructive nephropathy was not found in group I, 3(3.6%) in group II and 2(14.3%) in group III. ADPKD and obstructive nephropathy was not found in group I, 3(3.6%) in group II and 2(14.3%) in group III. ADPKD and 1(7.1%) were unknown etiology. There difference was not statistically significant (p>0.05) among three groups in chi square test.

Radiocephalic was found 19(90.5%) in group I, 66(79.5%) in group II and 9(64.3%) in group III. Brachiocephalic was found 2(9.5%) in group I, 17(20.5%) in group II and 5(35.7%) n group III. The difference was not statistically significant (p>0.05) among the three groups in chi square test.

Table I shows, the mean percent recirculation of study patients. Mean recirculation was found  $0.09\pm0.27\%$  with range from 0 to 0.9% in group I,  $4.15\pm2.56\%$  with range from 1 to 9% in group II,  $21.29\pm15.07\%$  with range from 11 to 66% in group III and mean recirculation  $8.1\pm5.5\%$  with range from 0 to 66% in the whole study patients. The differences was statistically significant (p<0.001) between three groups in ANOVA test.

Table II shows, distribution of improper arterial and venous needle placement in study patients. Improper needle placements were not found 21(100.0%) in group I, 78(94.0%) in group II and 4(28.6%) in group III. In approved direction was found 5(6.0%) and 10(71.4%) in group II and group III respectively but not found in group I. The differences was statistically significant (p<0.001) among the three groups in chi square test.

Table III shows distribution of distance between arterial and venous in study patients. A distance less than 4 cm was found 2(9.5%) in group I, 31(37.3%) group II and 7 (50%) in group III. The mean distance between arterial and venous needle was found  $6.28\pm2.1$  cm with range from 4.2 to 8 cm in group I,  $4.6\pm2.65$  cm and 2.5 to 14 cm in group II and  $2.5\pm1.25$  cm with range from 2.4 to 13.0 cm in group III. The difference was statistically significant (p<0.03) among the three groups in ANOVA test.

Table IV shows the distribution of ejection fraction in study patients. Ejection Fraction were found  $61.52\pm5.17$  % with range from 51 to 72% in group I, 57.28±6.95% with range from 40 to 70 % in group II and 52.71±4.95% with range from 35 to 60% in group III. Ejection fraction also had significant negative correlation with recirculation. (r=0.250; p<0.001). The difference was statistically significant (p<0.001) among the three groups in ANOVA test.

Table V shows the distribution of the mean blood flow through the fistula in study patients. Blood flow were found  $1330.9\pm622.4$ ml/min with range from 677.25 to 2279 ml/min in group I,  $1174.1\pm554.8$  ml/min with range from 456 to 2893 ml/min in group II and  $1073.3\pm363.4$  ml/min with range from 475 to 1571 ml/min in group III. Though the difference were not statistically significant (p>0.05) among the groups, there was significant negative correlation (r=0.201; p<0.046) was observed between the two variables. The study patients had no stenosis in fistula.

The mean duration of fistula was found 7.71±5.81 months with range from 3 to 36 months in group I, 9.89±6.48 months with range from 5 to 24 months in group II and 6.81±5.18 months with range from 3 to 27 months in group III. The mean difference was not statistically significant (p>0.05) among the three groups in ANOVA test. The interval between fistula constriction and first needling of the study patients. The mean interval between fistula construction and first needling was found  $51.17\pm35.92$  days with range from 27 to 150 days in group I, 40.62±35.36 days ranging from 21 to 210 days in group II and 29.5±2.36 days ranging from 21 to 33 days in group III. The difference was not statistically significant (p>0.05) among the three groups in ANOVA test

Table I: The percent recirculation of the study patients (n=118)

		Group I	Group II	Group III	
	Total	(n=21)	(n=83)	(n=14)	P value
	Mean ±SD	Mean±SD	Mean±SD	Mean±SD	
Percent of recirculation (%)	8.1±5.5	0.09±0.27	4.15±2.56	21.29±15.07	0.001s
Range(min-max)	(0 - 66)	(0-0.9)	(1-9)	(11-66)	

**Table II:** Distribution of the study patients according to Improper arterial & venous needle placement (n=118).

Improper arterial &	Group I		Group II		Group III		Р
venous needle placement	(n=21)		(n=83)		(n=14)		value
	Ν	%	Ν	%	n	%	
Absent	21	100.0	78	94.0	4	28.6	0.001 <sup>s</sup>
Present	0	0.0	5	6.0	10	71.4	0.001

**Table III:** Mean distance between arterial and venous needle of the study patients (n=118).

Distance between arterial and venous needle (in cm)	Group I (n=21)		Group II (n=83)		Group III (n=14)		P value
	Ν	%	n	%	n	%	_
<4	2	9.5	31	37.3	7	50	
≥4	19	90.5	52	62.7	7	50	
Mean±SD	6.28±2.1		4.6±2.65		2.5±1.25		0.037 <sup>s</sup>
Range (min-max)	(4.2-8)		(2.5-14)		(2.4-13)		0.037

**Table IV:** The mean left ventricular ejection fraction distribution of the study patients (n=118).

	Group I	Group II	Group III	Р
	(n=21)	(n=83)	(n=14)	value
		Mean ±SD		
Ejection fraction (%)	$61.52\pm\!\!5.17$	$57.28 \pm 6.95$	52.71±4.95	0.0018
Range (min-max)	(51-72)	(40-70)	(35-60)	0.001

**Table V:** The mean blood flow through the fistula of the study patients (n=118).

	Group I	Group II	Group III	
	(n=21)	(n=83)	(n=14)	P value
	Mean±SD	Mean ±SD	Mean ±SD	
Blood flow through	$1330.9 \pm\! 622.4$	1174.1 ±554.8	$\pm 1073.3 \pm 363.4$	
the fistula (ml/min)				$0.526^{ns}$
Range	(677.5 -2279)	(456 - 2893)	(475-1571)	

# Discussion

In this current study it was observed that 82.2% patients had recirculation, which is consistent with Salami et al, where the investigators showed 80.4% had patient had recirculation<sup>14</sup>. On the other hand, Beladi Mousavi et al <sup>13</sup>showed 17.0% and Abdollahi et al<sup>16</sup> found 15.0%, which may be due to their health professional are well trained as well as their availability of modern facilities in their hospital. The present study findings were discussed and compared with previously published relevant studies.

Recirculation was not age dependent according to this study. Similarly, Beladi Mousavi et al <sup>13</sup>showed almost similar age range in their study patients, where the authors found the age range was 16 to 75 years. On the other hand, Abdollahi et al <sup>16</sup>has observed higher mean age in their study patients, which was 50.93±17.23 years in comparision to us. This increase mean age may be due to increase life expectancy by improved quality of dialysis. Moreover due to financial burden our patient could not afford more than two session of dialysis per week rather than required three sessions. This reduced dose of dialysis affect the overall survival rate in HD patients<sup>9</sup>.

Male was more common in the whole study subjects, the male female ratio was 2.5:1. Male

was significant higher in group I and group II. Similarly, Beladi Mousavi et al <sup>13</sup> found overall, 58.69% and 41.30% patients were male and female respectively, which support the current study. This discrepancy among male female ratio occurred probably due to the fact that males are most privileged than females in the society to receive advanced medical facilities. females and diabetics have relative low blood flow through fistula than their male counterpart. <sup>11</sup> The increased amount of recirculation in females may be due to relative low blood flow through fistula .

Regarding the cause of ESRD it was observed in this current study that DM and GN were the more frequent in all three groups According to Beladi Mousavi et al <sup>13</sup> recirculation does not vary among diabetic and non diabetic and also hypertensive and normotensive. In another study Salimi et al.<sup>14</sup> mentioned in their study that history of hypertension and diabetes mellitus were not significantly correlated with recirculation, which are comparable with the current study.

Majority of the patients had radiocephalic which were 90.5%, which was not significant (p>0.05). Similar findings obtained by Salimi et  $al^{14}$ .

In this current series it was observe that the mean percent recirculation was  $8.1\pm5.5\%$  with range from 0 to 66 % in the whole study patients. range from 11 to 66 %, which was significantly higher in group III. Similarly, Vanstone et al<sup>17</sup>, Salimi et al<sup>14</sup> and Rafique et al<sup>15</sup> found the mean percent recirculation were 6.4%, 8.75% and 9.55±5.5% respectively,which are closely resembled with the current study.

Among the causes, improper arterial & venous needle placement significantly associated with increase recirculation, Similarly, a significant (p<0.001) association between recirculation and improper arterial and venous needle placement was also seen by Beladi Mousave et al <sup>13</sup>. Similar significant association also observed by Abdollahi et al<sup>16</sup>, Salimi et al<sup>14</sup>.

The study revealed that the mean distance between arterial and venous needle was significantly (p<0.05) decline with increased percent recirculation.

In the current study the left ventricular ejection fraction was 35 to 60%. significantly (p<0.05) decline of LVEF occur with increased recirculation. Beladi Mousavi et al <sup>13</sup> on their study on patients having heart failure and showed that the average LVEF in without recirculation was  $46.76 \pm$ 

5.57% and with recirculation was  $32.67 \pm 7.52$ , which also significantly (p=0.002) associated with recirculation, which is consistent with the current study.

In this present study it was observed that the mean blood flow through the fistula was found 1330.9±622.4ml/min.In current study blood flow decline with increased percent recirculation (r=0.201) but the difference was not significant (p>0.05). Similarly, Beladi Mousavi et al<sup>13</sup> showed the mean A-V fistula flow rate in both groups was more than 400 ml/min in without recirculation group 1229.41 ±753.96 mill/min and with recirculation, 653.33 ± 209.98 mill/min) and the mean A-V fistula flow rate was significantly higher in without recirculation (p = 0.001), which is consistent with the current study.

In this present series it was observed that all patients had no stenosis in fistula, which is consistent with the study done by Beladi Mousavi et al <sup>13</sup>.Probably because of the fact that recirculation is a late indicator of Stenosis and almost >50% Stenosis is required for recirculation.<sup>11</sup>We could not found any association between the two variables as fortunately we did not find any patient with stenosis and obstruction.

The mean difference of duration of fistula used was not statistically significant (p>0.05) among the three groups. There was no correlation r=0.123; (p=0.186) was found between recirculation with duration of fistula. Similarly, Salimi et al<sup>14</sup> also found no significant (p>0.05) association between recirculation with duration of fistula, which support the current study. However, Beladi Mousavi et al<sup>13</sup> showed the length of time of A-V fistula use was 26.59± 9.37 months in without recirculation and  $33.20\pm$  7.35 months in with recirculation group. There also was a significant association (p=0.001). In our study we could not find the relationship between duration of fistula use and recirculation. As stated before In our country due to cost strain majority of the patients receive inadequate dose of HD rather than required total 12 hours dialysis per week that affect on their net survival thereby indirectly they have reduced duration of fistula use in comparison to others due to inadequate dialysis. Probably to establish such relationship we need a study group with prolong duration of fistula use.

In this current study it was observed that the interval between fistula construction and first needling was not associated with recirculation(p=0.137). Beladi Mousavi et al<sup>13</sup> observed in their study that the average time between creation and use of A-V fistula in without recirculation were  $90\pm22$ 

days and  $40 \pm 15$  days respectively. In our study we could not get the relationship cause most of the patients denied to create their fistula in stage VI of CKD and majority of them created it when they were in emergency phase of dialysis and receiving dialysis through catheter. That is they usually don't give enough time for fistula maturation after creation of AV fistula. As catheters are not suitable for dialysis for prolong period due to its complications. A newly created fistula needs almost 4 to 6 weeks for maturation<sup>8</sup>. Thereby we could not get this relationship between this two variable from this study.

Conclusion: In this observational study recirculation was a common findings but according to this study it does not dependent on age, type of fistula, cause of ESRD, but more frequent in female. Improper arterial and venous needle placement and close proximity between needles were identified as most important causes of recirculation. Left ventricular ejection fraction had significant association with recirculation. A negative correlation was observed between blood flow and recirculation. Senosis and obstruction in fistula was not to be a contributing factor in present study. No association was observed between recirculation and duration of fistula use and wait after fistula creation.

*Limitation:* The study was conducted over a short period and included only one centre which may not represent whole dialysis population. Stenosis was not excluded as gold standard method to detect stenosis is venography.

*Recommendation:* A large multicenter study is demanded to verify the existence of such high percentage of recirculation found in the present study, to evaluate the risk factors associated with recirculation found in this study as well as other risk factors ascertain in large study group. Specific training program dedicated to dialysis care personnel should be launched nationwide to adopt attitude which can eliminate this correctable cause of inadequate dialysis.

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