

## Access Care for the patient undergoing Intensive Hemodialysis

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## Objectives

- Define intensive HD
- Benefits of intensive HD
- Review association between intensive HD and VA morbidity

### Intensive Hemodialysis

- Frequency > 3 sessions / week (2-8 hours) or extended duration > 4 hours/session
- Includes both home and in centre HD pts
- Unclear number of patients in US

Susantitaphong AJKD 2012

### Benefits of Intensive HD

- Quality of Life
- Cardiac benefits
  - BP control, reduction anti-hypertensive meds
  - Volume control
  - LVH reduction
  - LVEF improvement
- Phosphate control, reduction phosphate binders



#### In-Center Hemodialysis Six Times per Week versus Three Times per Week

The FHN Trial Group\*

#### In-Center Hemodialysis Six times per week vs Three times per week

- Frequent HD Network Daily Trial (FHN)
  - Randomized controlled trial; n =245; duration 12m
  - Coprimary composite outcome:
    - Death or increase in LVMI
    - Death or decrease in physical health composite score
  - <sup>o</sup> 5.2 sessions/wk (154 min) vs 3/wk (214min)

### FHN Daily Trial: VA Outcomes

- VA related complications
  - access failure,
  - infection requiring a procedure,
  - thrombectomy,
  - angioplasty and
  - fibrin sheath stripping or replacement of catheters

### FHN Daily Trial: Results

- 60% AVF, 20% CVC and 19% AVG
- Primary composite outcome death /reduction LVM significantly lower in frequent group:
  - □ HR 0.6 (0.46 0.82)
  - adjusted mean decrease in LV mass was 16 g vs 2.6; p<0.001 in frequent pts</li>
- No significant impact of Intensive HD on hospitalization (unrelated to vascular access)

### FHN Daily Trial: VA results

Outcome	Conventional Hemodialysis (N=120)		Frequent Hemodialysis (N=125)		Hazard Ratio (95% CI)	P Value
	no. of events	no. of patients with event	no. of events	no. of patients with event		
Death	9		5		—	—
All hospitalizations	114	47	109	58	0.88 (0.60–1.28)	0.50
Unrelated to vascular access	90	44	79	47	0.80 (0.53–1.21)	0.30
Related to vascular access	24	14	30	20	0.99 (0.54–1.82)	0.97
Cardiovascular-related	15	12	17	15	0.83 (0.44–1.59)	—
Infection related	27	20	27	23	0.83 (0.49–1.40)	<u> </u>
All interventions related to vascular access	65	29	95	47	1.35 (0.84–2.18)	0.22
Correction of access failure	23	15	19	15	0.71 (0.35–1.44)	0.35
Other procedures	42	21	76	38	1.71 (0.98–2.97)	0.06

No increased rate of access failure despite increased number of VA procedures ? Hypervigilance with monitoring Chertow NEJM 2010

### FHN Daily Trial: VA Results

- 47% FHD underwent one procedure
  51% AVF, 32% AVG, 17% CVC had a VA event
- 29% CHD underwent one procedure
  48% AVF, 38% AVG, 14% CVC had a VA event

#### ?Did $\uparrow$ HD frequency lead to $\uparrow$ VA dysfunction?

# So What is the impact of Frequent or Intensive HD on VA?

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In-Center Hemodialysis Six Times per Week versus Three Times per Week

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Kidney Int. 2011 November ; 80(10): 1080-1091. doi:10.1038/ki.2011.213.

#### The effects of frequent nocturnal home hemodialysis: the Frequent Hemodialysis Network Nocturnal Trial

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#### HD Frequency and VA Complications

- Combine VA data from FHN daily and FHN Nocturnal trials (6x NHD vs 3x CHD)
- Primary VA outcome time to first access event (repair, loss, access hospitalizations)
- Secondary VA outcomes: time to all repairs and time to all losses
   Suri JASN 2012

#### Table 1. Baseline characteristics

		Daily Trial		Nocturnal Trial			
Variable	Conventional ( <i>n</i> =120)	Daily ( <i>n</i> =125)	All (n=245)	Conventional (n=42)	Nocturnal (n=45)	All (n=87)	
Age (yr)	52.0±14.1	48.9±13.6	50.4±13.9	54.0±12.9	51.7±14.4	52.8±13.6	
Male sex	73 (60.8)	78 (62.4)	151 (61.6)	28 (66.7)	29 (64.4)	57 (65.5)	
Race/ethnicity							
Black	53 (44.2)	49 (39.2)	102 (41.6)	11 (26.2)	12 (26.7)	23(26.4)	
White	46 (38.3)	43 (34.4)	89 (36.3)	21 (50.0)	27 (60.0)	48 (55.2)	
Native American, aboriginal Canadian, Alaska Native, First Nation	4 (3.3)	4 (3.2)	8 (3.3)	2 (4.8)	1 (2.2)	3 (3.4)	
Asian	5 (4.2)	11 (8.8)	16 (6.5)	7 (16.7)	5 (11.1)	12 (13.8)	
Native Hawaiian/other Pacific Islander	3 (2.5)	1 (0.8)	4 (1.6)	0 (0)	0 (0)	0 (0)	
Other/mixed/unknown	9 (7.5)	17 (13.6)	26 (10.5)	1 (2.4)	0 (0)	1 (1.2)	
Hispanic/Latino	31 (26)	38 (30)	69 (28.2)	0 (0)	0 (0)	0 (0)	
ESRD vintage							
<2 yr	35 (29.2)	37 (29.6)	72 (29.4)	30 (71.4)	28 (62.2)	58 (66.7)	
2–5 yr	42 (35.0)	34 (27.2)	76 (31.0)	5 (11.9)	8 (17.8)	13 (14.9)	
>5 yr	43 (35.8)	54 (43.2)	97 (39.6)	7 (16.7)	9 (20.0)	16 (18.4)	
Comorbid conditions							
Diabetes	50 (41.7)	50 (40.0)	100 (40.8)	18 (42.9)	19 (42.2)	37 (42.5)	
Congestive heart failure	24 (20)	25 (20)	49 (20.0)	7 (16.7)	5 (11.1)	12 (13.8)	
Laboratory values							
Albumin (g/dl)	3.94±0.46	3.94±0.37	$3.94 \pm 0.42$	3.92±0.51	3.90±0.48	3.91±0.49	
Hemoglobin (g/dl)	$12.0 \pm 1.2$	$11.9 \pm 1.3$	$11.9 \pm 1.3$	11.9±1.1	11.6±1.1	11.8±1.1	
Vascular access used at randomization			$\frown$			$\frown$	
Arteriovenous fistula	71 (59)	82 (66)	153 (63)	17 (40)	22 (49)	39 (45)	
Arteriovenous graft	23 (19)	22 (18)	45 (18)	4 (10)	3 (7)	7 (8)	
Tunneled catheter	26 (22)	21 (17)	47 (19)	21 (50)	20 (44)	41 (47)	
Buttonhole technique used (n)			$\backslash - /$	14	19	33	

Unless otherwise indicated, values are expressed as number of patients (percentage). Values expressed with a plus/minus sign are the mean  $\pm$  SD.

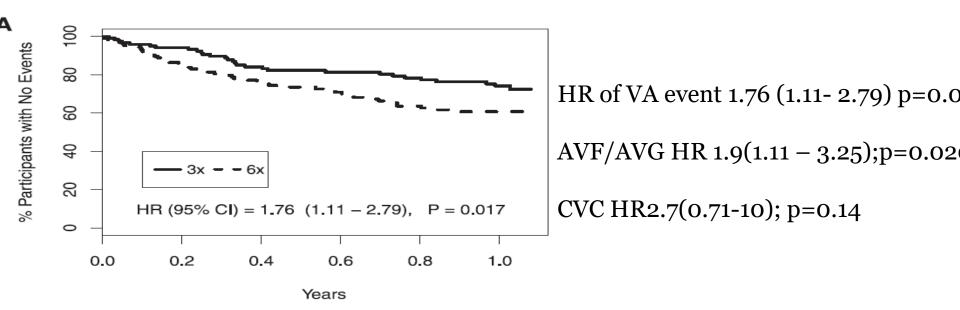
#### Suri JASN 2012

#### HD Frequency and VA Complications

Table 2. Description of all repairs and losses, considering all accesses used during follow-up

Veriable		Daily Trial		Nocturnal Trial		
Variable	Conventional	Daily	HR (95% CI)	Conventional	Nocturnal	HR (95% CI)
AVF/AVG <sup>a</sup>						
Patients (n)	106 (79/28) <sup>b</sup>	114 (90/28) <sup>b</sup>		28 (23/6) <sup>b</sup>	32 (29/3) <sup>b</sup>	
Total follow-up (yr)	87.9 (67.1/20.8)	95.8 (76.9/18.9)		24.2 (18.8/5.4)	24.3 (22.0/2.3)	
Repairs						
Angioplasty	21 (11/10)	28 (14/14)		5 (3/2)	14 (13/1)	
Stent placement	2 (2/0)	2 (1/1)		0 (0/0)	0 (0/0)	
Thrombectomy	10 (3/7)	22 (5/17)		1 (0/1)	2 (1/1)	
Surgical revision	5 (3/2)	14 (7/7)		1 (1/0)	0 (0/0)	
Overall rate (per 100 patient-yr)	43	69	1.68 (1.13–2.51) <i>P</i> =0.011	29	66	2.29 (0.94–5.59) <i>P</i> =0.069
Losses						
Stenosis/thrombosis	8 (5/3)	13 (5/8)		2 (1/1)	2 (2/0)	
SItenosis/thrombosis						
Infection	2 (0/2)	2 (0/2)		1 (0/1)	0 (0/0)	
Other <sup>c</sup>	1 (1/0)	3 (2/1)		0 (0/0)	0 (0/0)	
Unknown	4 (2/2)	2 (1/1)		1 (0/1)	2 (2/0)	
Overall rate (per 100 patient-yr)	17	21	1.21 (0.61–2.39) P=0.58	—	—	—
Catheters						
Patients (n)	34	37		24	23	
Total follow-up (yr)	20.8	18.9		19.0	18.1	
Repairs (n)						
Fibrin sheath stripping	1	4		0	1	
Repair broken component	0	0		1	0	
Losses (n)						
Poor flows/thrombosis	7	2		8	8	
Infection	2	7		4	5	

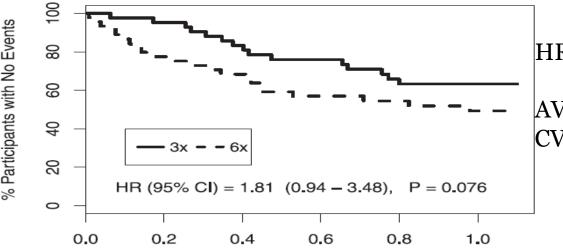
#### Time to VA Complication: Daily FHN



Time to first access loss, first access repair or access hospitalization in Daily FHN trial

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#### Time to VA Complication: Nocturnal FHN



HR of VA event 1.81(0.94-3.48) p=0.07

AVF/AVG HR 3.2(1.07 – 10.3);p=0.038 CVC HR 1,45(0.59-3.58); p=0.42

Time to first access loss, first access repair or access hospitalization in Nocturnal FHN trial

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#### HD Frequency and VA

	3 TIMES PER WEEK		6 TIMES PER WEEK						
	Ν	Follow-up (yrs)	Event rate*	Ν	Follow-up (yrs)	Event rate*	HR (95% CI)	Plot of HR, 95% CI	p-value
Daily Trial									
All patients	120	127.8	23	125	121.2	40	1.76 (1.11-2.79)	I₿	0.017
AV access	94	101.3	21	104	99.5	37	1.90 (1.11-3.25)	II	0.020
Catheters	26	18.9	16	21	17.0	47	2.70 (0.71-10.2)	ŀ <b> →</b> ->	0.14
Nocturnal Tr	ial								
All patients	42	47.1	32	45	39.5	58	1.81 (0.94-3.48)	ı <b>•</b> ı	0.076
AV access	21	24.2	17	25	21.8	55	3.23 (1.07-10.35)	ŀ <b>◆</b> ≻	0.038
Catheters	21	19.9	45	20	14.1	71	1.45 (0.59-3.58)	ıı	0.42
								0.25 0.5 1 2 4	
								6 times/wk better 3 times/wk better	

Figure 2. Forest plot of time to first access repair, access loss, or access hospitalization by trial and access subgroup. \*Event rates expressed as number of events per 100 patient-years. AV access, arteriovenous fistulae and arteriovenous graft Suri JASN 2012

# Summary: HD Frequency and VA

- Daily FHN
  - More total AV access repairs occurred in daily group (HR 1.68 (1.13-2.51); p=0.011
  - HR higher for repairs in AVG than AVF: 2.2(1.26-3.87);
     p=0.0059
  - No significant difference in AV losses between groups
- Nocturnal FHN
  - More total AV access repairs occurred in nocturnal but not significant; HR 2.29; p=0.069
  - Trend to higher AVF repairs: HR 2.87;p=0.063

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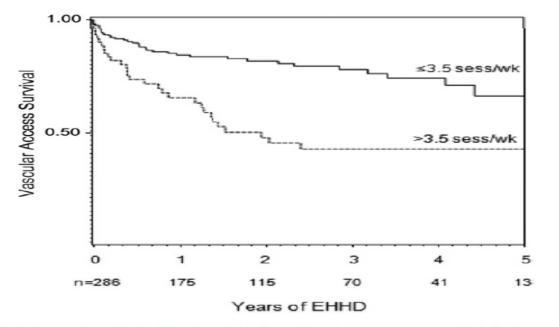
#### Increased VA Events in Intensive HD: What About Other Studies?

#### Frequent Home N-HD and VA Complications

- Australian observational cohort of extended hours N-HD (≥24h/week) 1999 - 2009
- Outcomes: all cause mortality, technique failure and access related events
- Access related event: any type of VA related intervention

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#### Frequent N-HD and VA Dysfunction



Access events predicted by dialysis frequency and age

Access events predicted death: HR 2.85(1.14 – 7.15)

Figure 2. Unadjusted Kaplan-Meier access survival curves comparing patients dialyzing for 3.5 or fewer sessions per week (solid line) with patients dialyzing for more than 3.5 sessions per

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#### Frequent Home N-HD and VA Dysfunction

Table 2. Outcomes Analyzed

Outcome	No. (%)			
All-cause mortality				
Cardiac death	7 (29.2)			
Sepsis	5 (20.8)			
Hemorrhage	1 (4.2)			
Renal carcinoma	1 (4.2)			
Unknown	10 (41.7)			
Technique failure				
Failing health	17 (26.9)			
Dialysis facility issues	8 (12.7)			
Not coping with the modality	4 (6.4)			
Not sleeping properly	4 (6.4)			
Psychosocial issues	1 (1.6)			
Access-related events	2 (3.2)			
Patient choice	1 (1.6)			
Other	4 (6.4)			
Unknown	22 (34.9)			
Access-related adverse events <sup>a</sup>				
Infection	47 (59.5)			
Bacteremia	14 (29.8)			
Local AVF infection	22 (46.8)			
Unspecified infection	11 (23.4)			
Thrombosis/occlusion	12 (15.2)			
AVF aneurysm	5 (6.3)			
Stenosis	8 (10.1)			
AVF revision requirement	3 (3.8)			
Other	4 (5.1)			

*Note:* Based on unadjusted Kaplan-Meier survival rates.

#### 28% (79/286) had adverse VA event

#### ~60% VA infection

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### VA outcomes and intensive HD

- Meta analysis of studies including home and intensive HD patients and VA outcomes
- 3 RCTS (Chertow NEJM 2010, Rocco KI 2011, Culleton JAMA 2006)
- 11 prospective & 5 retrospective cohorts
- Most studies >4 x per week, > 4 hours

Cornelis Blood Purif 2014

#### More VA events in Intensive HD

	Access-	Relative risk (intensive/conventional)	Event rate			
	years		difference	ratio	р	
Access admissions	0					
Access dysfunction	332	22 <u>2</u> 2	0.116	1.545	0.137	
Access infection	185		0.000			
Permanent access failure	228		0.008	1.077	0.872	
AVF (all)	744		0.027	1.391	0.162	
Access admissions	0					
Access dysfunction	117		0.429	1.456	0.171	
Access infection	47		-0.019	0.835	0.874	
Permanent access failure	47		0.176	1.449	0.547	
AVG (all)	212		0.376	1.478	0.139	
Access admissions	26		0.153			
Access dysfunction	91	· · · · · · · · · · · · · · · · · · ·	-0.024	0.696	0.516	
Access infection	103		0.193	1.683	0.264	
Permanent access failure	103		0.341	1.483	0.217	
Catheter (all)	322		0.169	1.419	0.133	
Access admissions	985		0.073	1.324	0.311	
Access dysfunction	924		-0.209	0.811	0.112	
Access infection	254		0.098	1.975	0.307	
Permanent access failure	383	· · · · · · · · · · · · · · · · · · ·	0.015	1.213	0.673	
Access unknown (all)	2,546		0.009	0.965	0.702	
Access admissions	1,011		0.073	1.350	0.237	
Access dysfunction	1,464		-0.049	1.144	0.134	
Access infection	589		0.065	1.633	0.106	
Permanent access failure	761		0.103	1.307	0.150	
All	3,824		0.067	1.224	0.009	
		0 1 2 3 4 6.	7 more VA	events	s per 100 p	ot year
		Ratio				•
		110/05/03/04	Cr	rnelis	Blood Pu	rıf 201



T Vachharajani Atlas of Dialysis Access

# So Why Might Intensive HD promote VA complications?

### VA dysfunction with Intensive HD

• Vascular access used twice as often

• Increased surveillance

# VA dysfunction with Intensive HD

- Infection
  - Increased exposure to inflammatory stimuli on HD
  - Potential exposure to bacterial pathogens (skin,water)
- Trauma
  - Frequent venipuncture
  - Increased shear stress on endothelium → promotes remodeling
  - Impact of blood flow rate and needle size?

#### How to Mitigate risk in Intensive HD

- Reduce frequency of HD (alternating days)
- Use of lower blood flow rates eg 200 vs 400
  - Nocturnal HD had fewer thrombectomy and surgical revisions than Daily HD
- Maintain vigilance for impending VA dysfunction (home)
- Train (home) patients regarding signs of VA dysfunction
- Regular audits in home patients
- Reduce buttonhole cannulation to lower infection risk

# **Questions?** Thank You

### VA Resources

International Society of HD (ISHD) Manual for Home HD Care Module 7 (VA) http://www.ishd.org/7-the-care-and-keeping-of-vascular-access-for-home-hemodialysis-patients/ Canadian Journal of Kidney Health and Disease VA Core Curriculum 1. Arteriovenous access selection and evaluation http://journals.sagepub.com/doi/full/10.1177/2054358116669125 2. Arteriovenous access failure, stenosis and thrombosis http://journals.sagepub.com/doi/full/10.1177/2054358116669126 3. Arteriovenous access infection, neuropathy and other complications http://journals.sagepub.com/doi/full/10.1177/2054358116669127 4. Practical aspects of tunneled and nontunneled HD catheters http://journals.sagepub.com/doi/full/10.1177/2054358116669128 5. Hemodialysis tunneled catheter related infections http://journals.sagepub.com/doi/full/10.1177/2054358116669129 6. Hemodialysis tunneled catheter non-infectious complications http://journals.sagepub.com/doi/full/10.1177/2054358116669130

