

# Intradialytic Parenteral Nutrition in End-Stage Renal Disease

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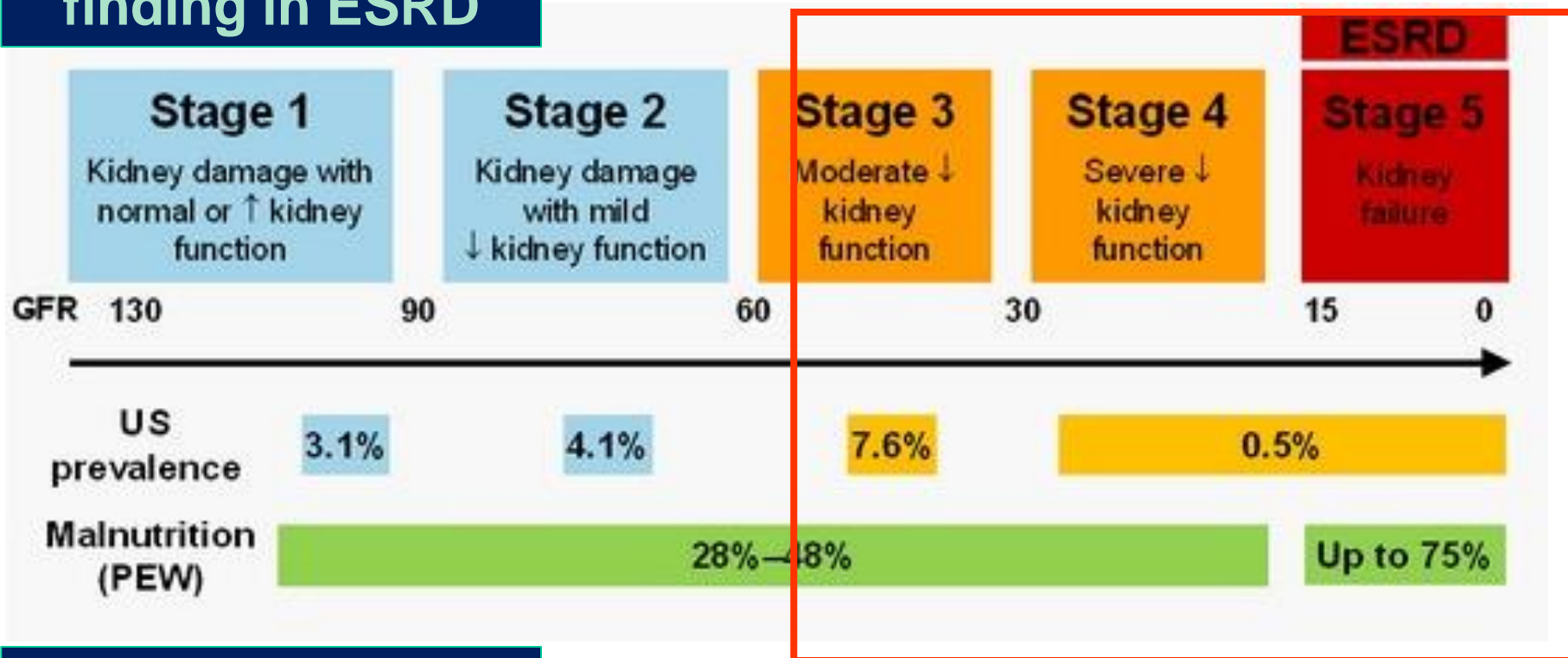


# Agenda

- Why nutritional support in End-Stage Renal Disease (ESRD) on hemodialysis (HD)
- Choice of nutritional support in ESRD on HD
- Oral Nutritional Supplementation (ONS) and Intradialytic Parenteral Nutrition (IDPN)
- The role of ONS and IDPN in ESRD on HD

# Epidemiology of Protein-Energy Wasting (PEW) in CKD/ESRD

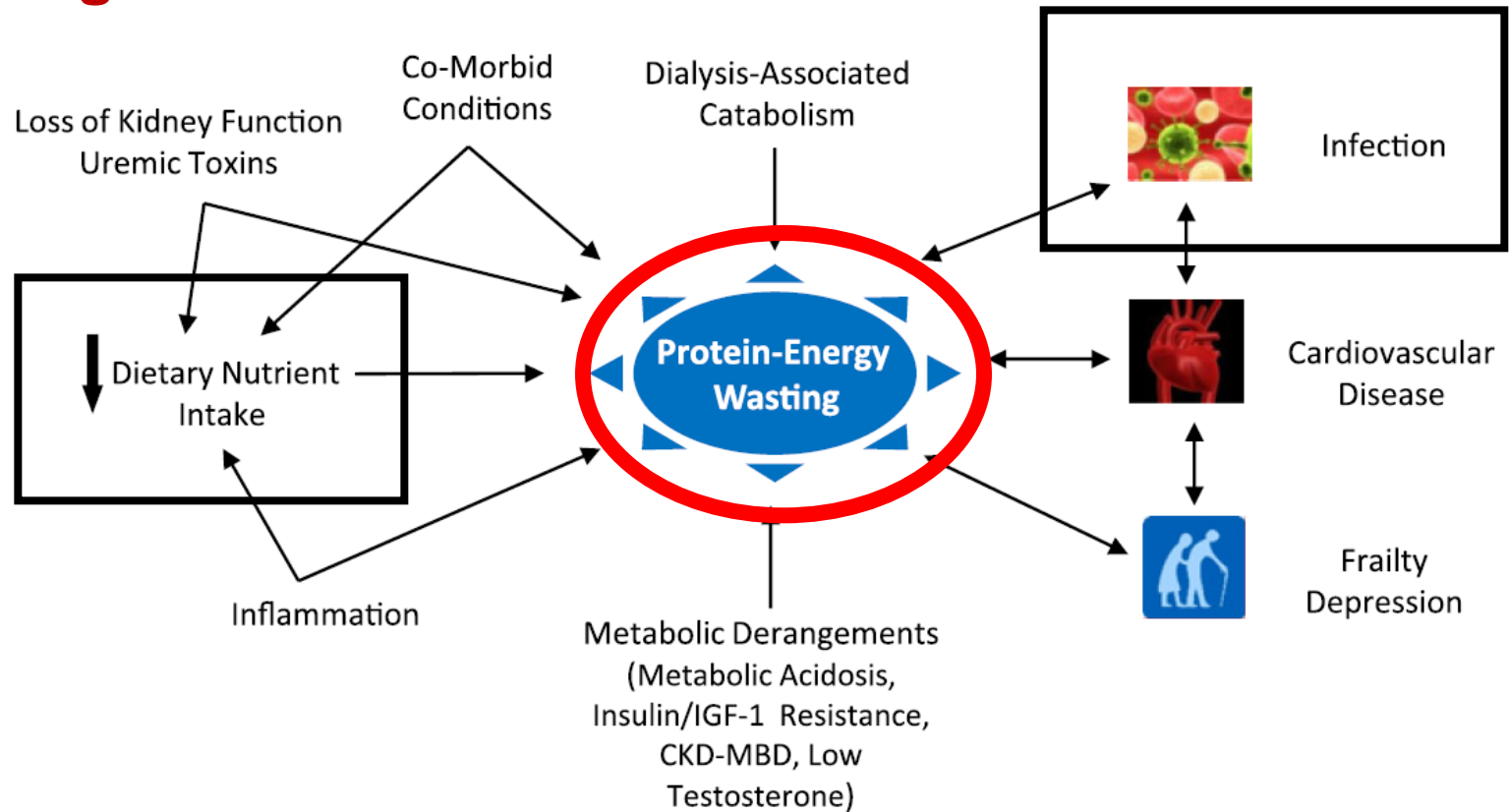
**PEW is a frequent finding in ESRD**



**PEW prevalence increases from stages 3 to 5**

USRDS 2009 Annual Data Report  
 Fouque D et al., *Kidney Int* 2008; 73:391-398  
 Kovesdy C et al. *Am J Clin Nutr* 2009; 29:3-14  
 Kovesdy CP et al. *Am J Clin Nutr* 2013; 97: 1163-1177

# Pathogenesis of PEW in CKD/ESRD



**PEW is the result of multiple mechanisms inherent to CKD/ESRD, including loss of appetite and reduced dietary intake, systemic inflammation, comorbidities, hormonal derangements, dialysis procedure, and uremic toxicity**

# Inflammation and wasting in CKD/ESRD

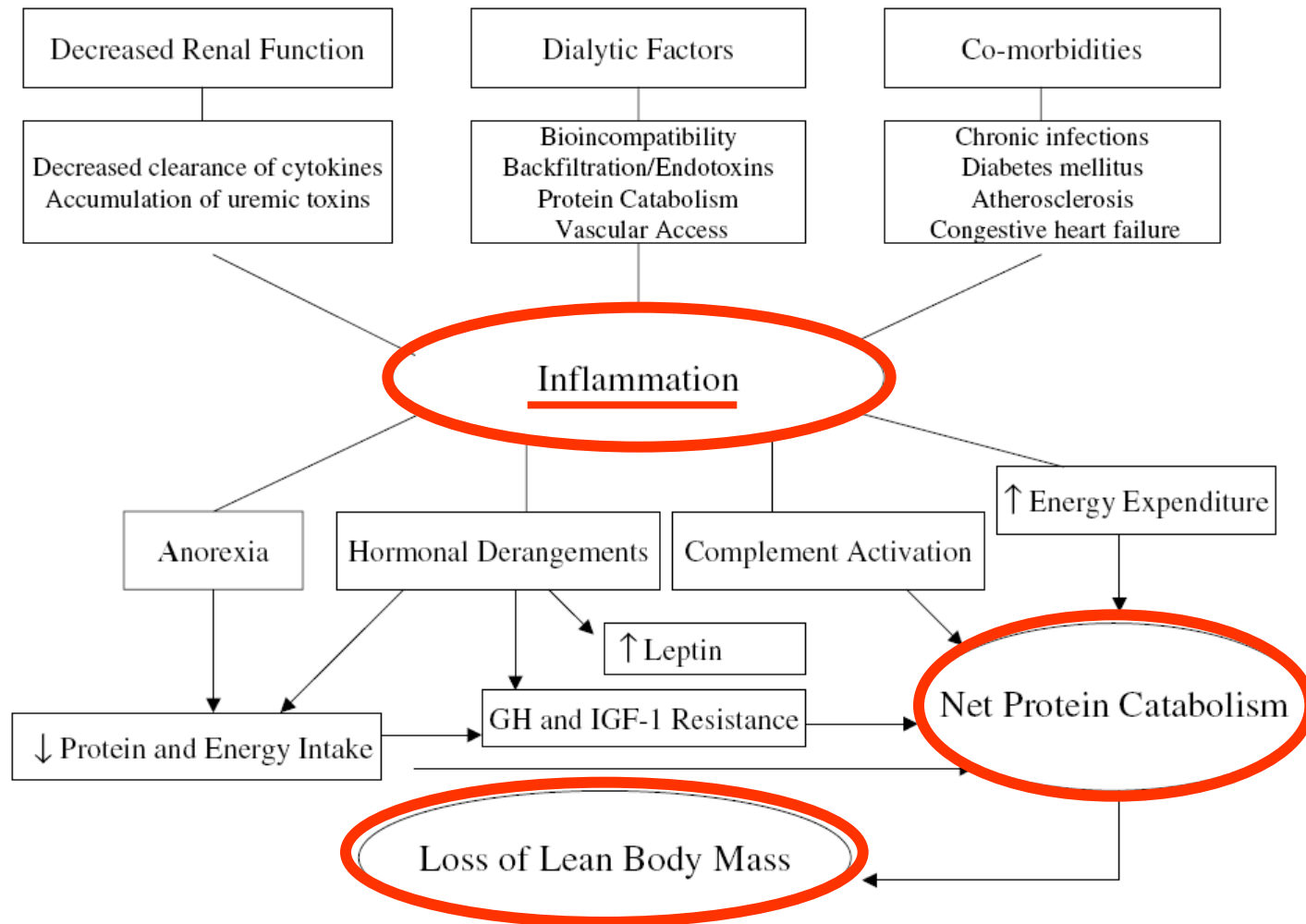
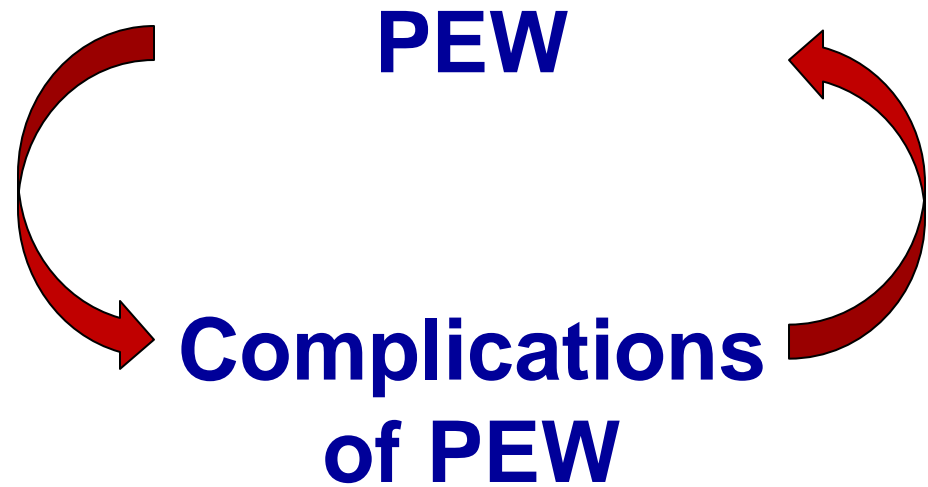
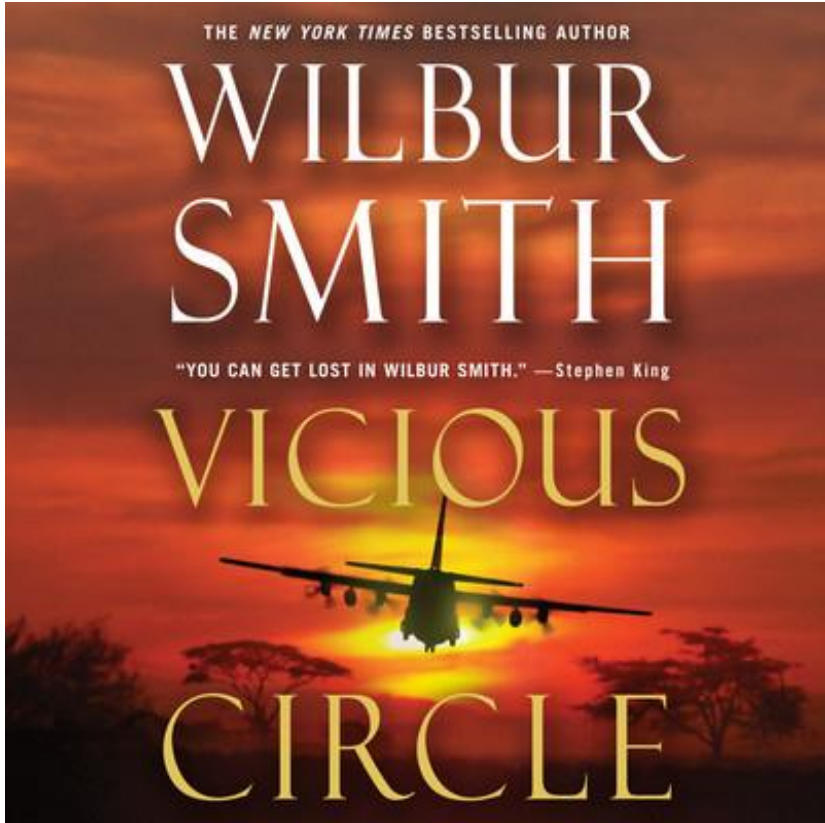
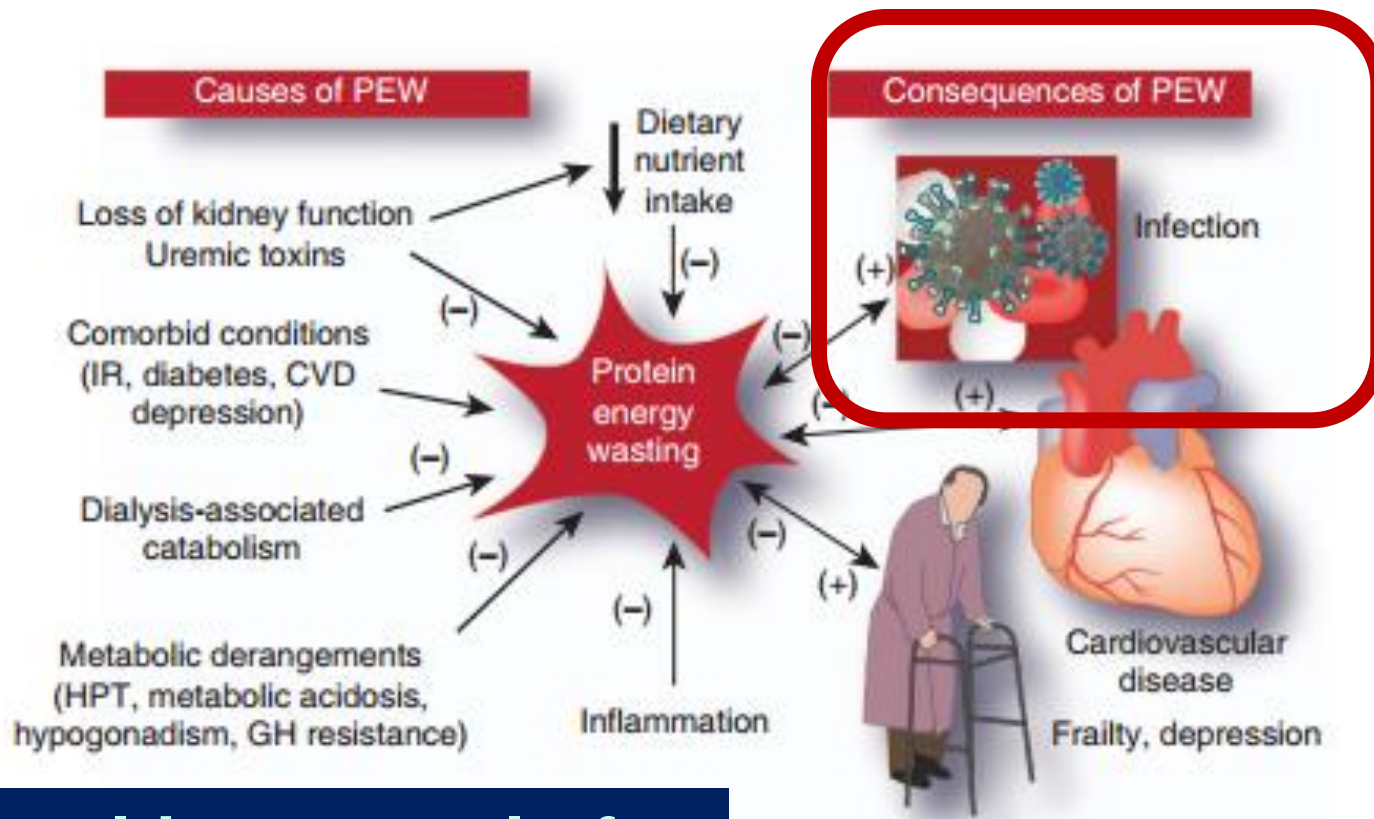


FIG. 1. Proposed interrelationship between inflammation and "uremic malnutrition."

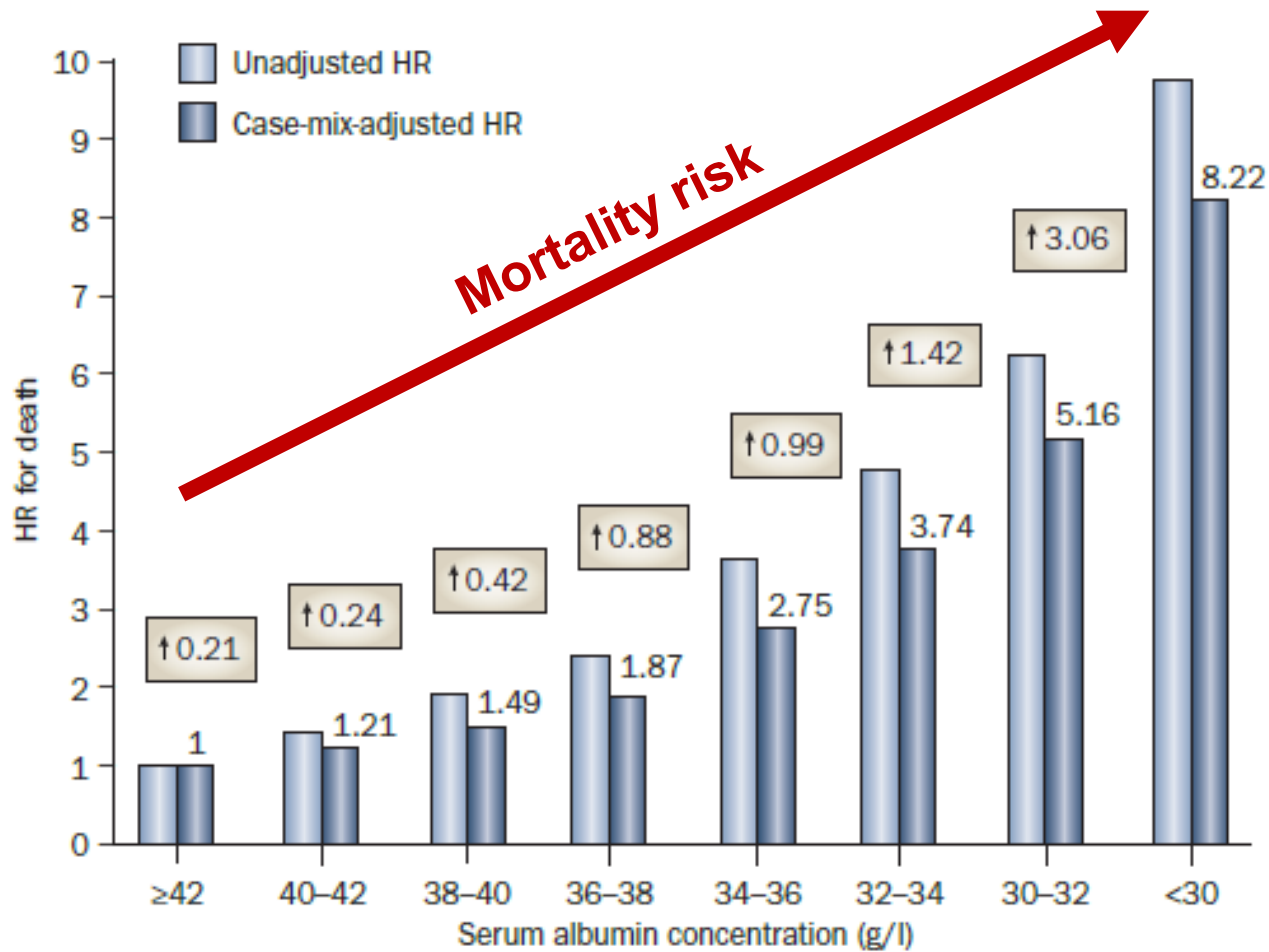


# the concept of «vicious circle» of PEW in CKD/ESRD: PEW leads to complications, complications lead to PEW



**The ultimate goal of nutritional support is to break this vicious circle**

# Protein-Energy Wasting (PEW) is associated with increased mortality risk in HD patients



Kalantar-Zadeh, K. et al. Nephrol Dial Transplant 2005; 20:1880-1888



# What are the targets suggested for ESRD patients on dialysis?

**Table 5.** Recommendations for protein and energy supply in adult patients on haemodialysis [8–10] and peritoneal dialysis (PD) [8–11]

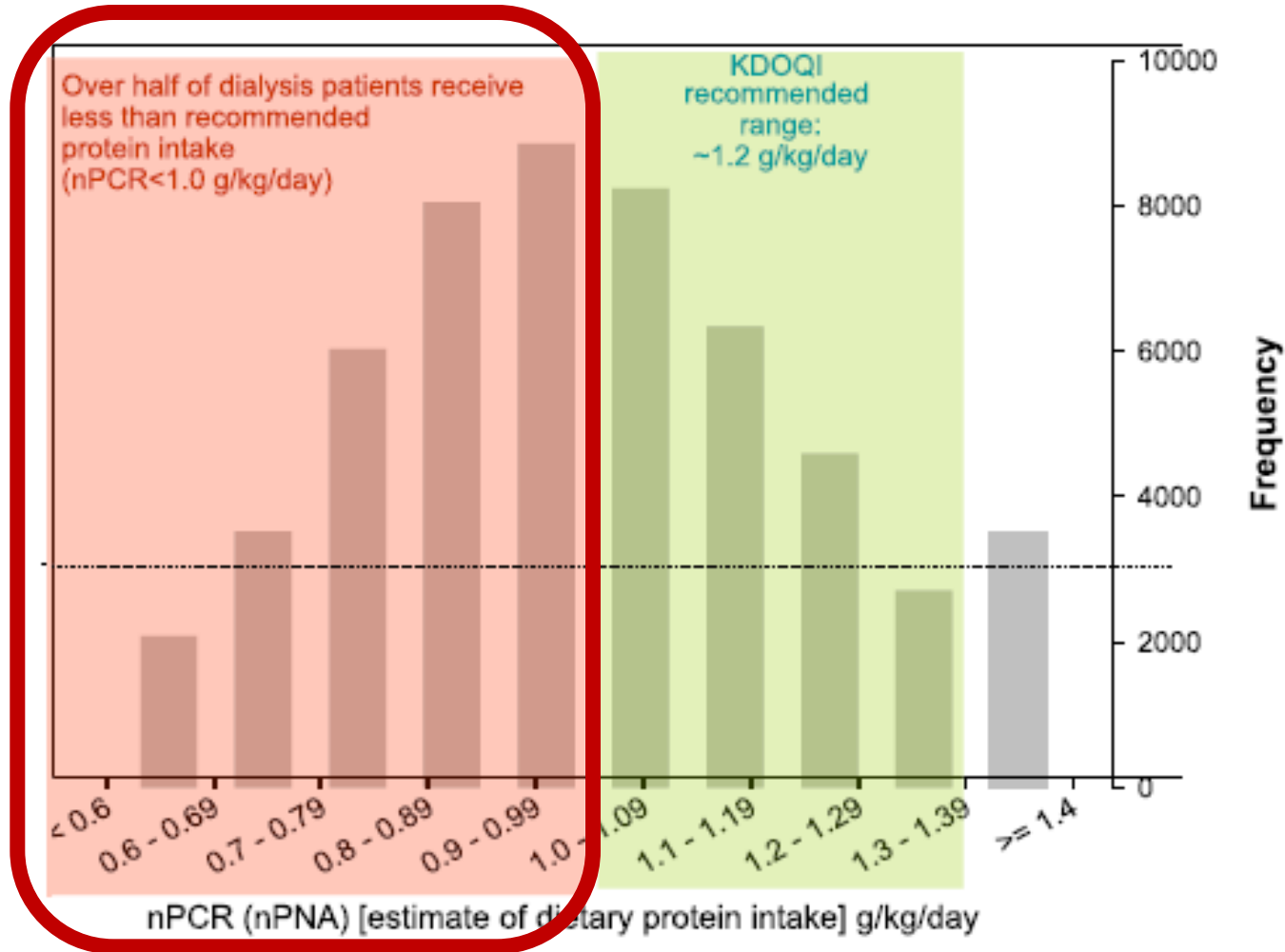
	ESPEN	NKF	EBPG
Protein intake (g/kg/day) (HD)	1.2–1.4 (>50% HBV)	1.2 (>50% HBV)	≥1.1
Protein intake (g/kg/day) (PD)	1.2–1.5 (>50% HBV)	1.2–1.3 (>50% HBV)	1.3
Energy intake (kcal/kg/day) (HD)	35	<60 years 35 <60 years 30	30–40, adjusted to age, gender and activity
Energy intake (kcal/kg/day) (PD)	35	<60 years 35 <60 years 30	<60 years 35 <60 years 30

ESPEN, European Society of Parenteral and Enteral Nutrition; NKF, National Kidney Foundation; EBPG, European best practice guidelines; HD, haemodialysis; DP, peritoneal dialysis.

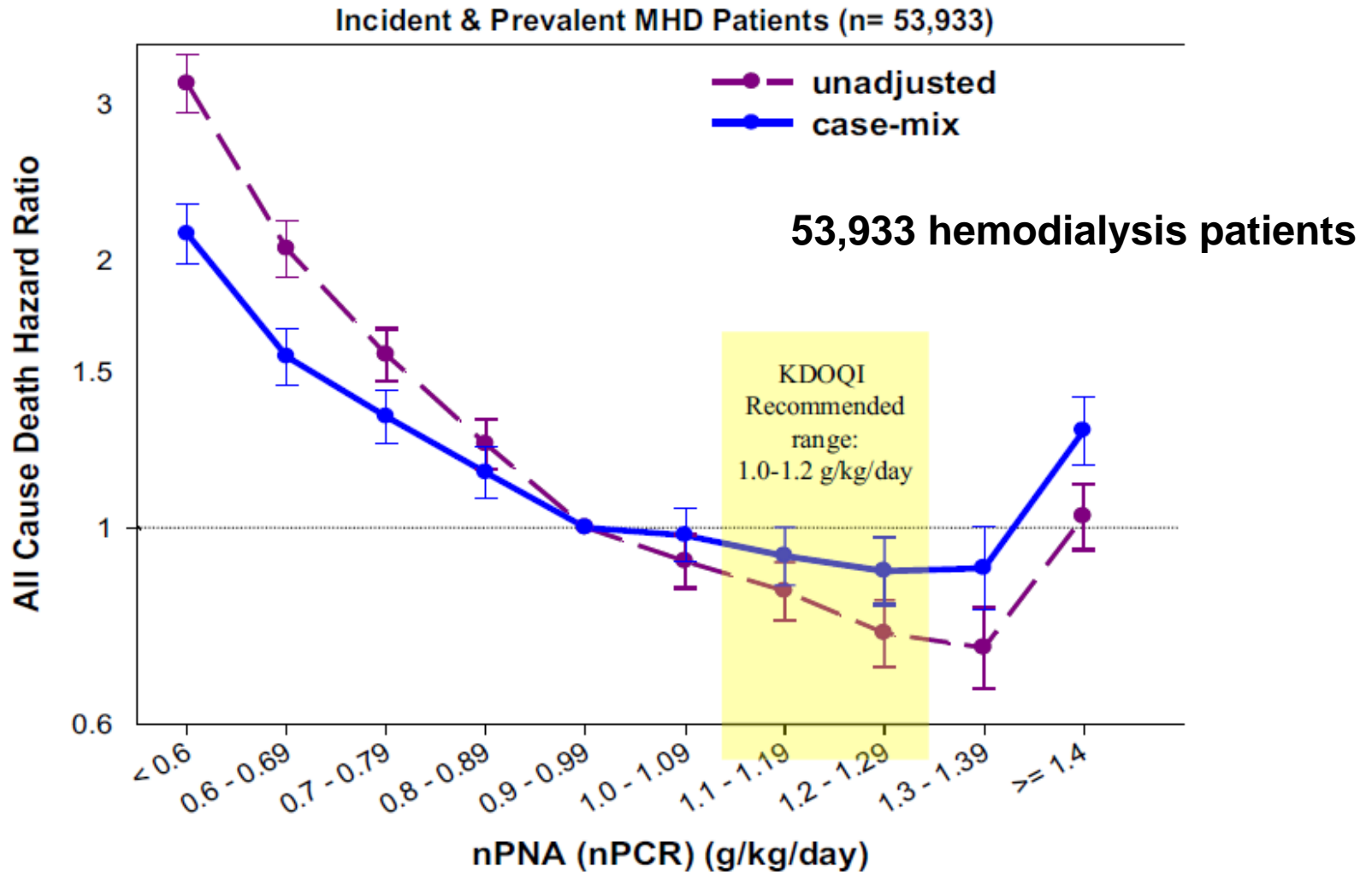


**In many patients it is difficult to reach and maintain nutritional targets**

# Frequency distribution of protein intake in hemodialysis patients



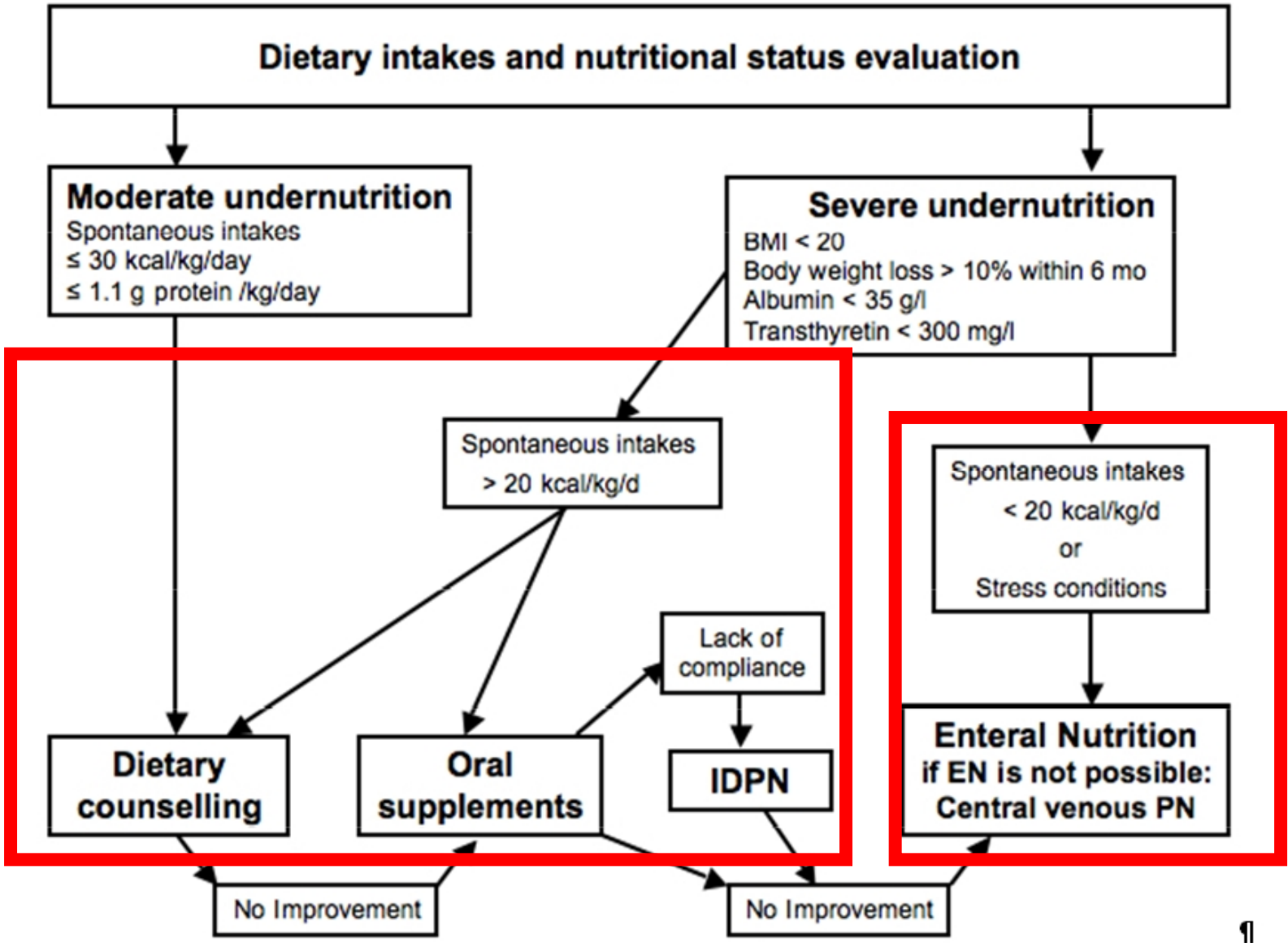
# Association between dietary protein intake, estimated by nPCR (nPNA) and survival



**When  
spontaneous  
nutritional  
intake is not  
enough**

# Nutritional support in ESRD patients

- Oral nutritional supplementation (ONS), as intradialytic or daily ONS
- Enteral nutrition (n.g. tube, PEG)
- Parenteral nutrition (in-hospital PN; home PN; intradialytic parenteral nutrition, IDPN)



## 5.2. Oral supplements and enteral feeding

- Nutritional supplements should be prescribed if nutritional counselling does not achieve an increase in nutrient intake to a level that covers minimum recommendation (see Guideline 3) (Evidence level III).
- Products specifically formulated for dialysis patients should be prescribed in preference to standard supplements for non-renal patients (Evidence level III).
- Enteral tube [naso-gastric or percutaneous entero-gastrostomy (PEG)] feeding using disease specific formulas for dialysis patients should be prescribed if attempts to increase dietary intake with oral supplements fail and nutritional status does not improve (Evidence level IV).



# Intradialytic oral supplementation

- Snacks during dialysis
- Intradialytic (commercial) liquid oral supplements

# Liquid oral supplements for ESRD

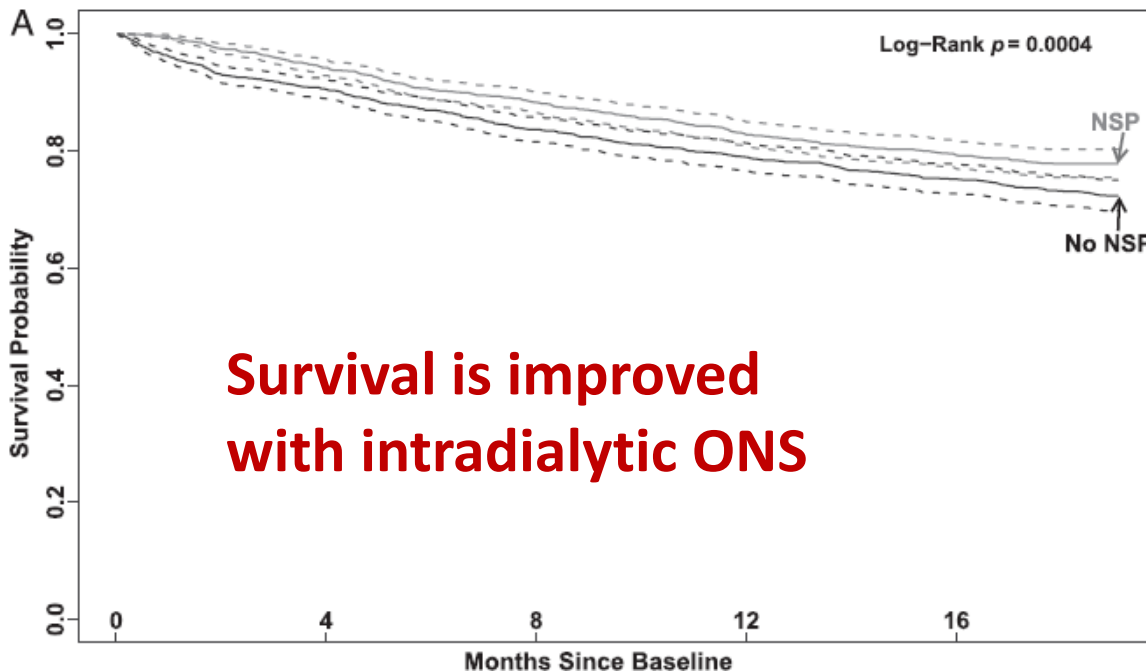
	Standard enteral diet	Renilon 7.5	Dialycare	Nepro HP
		Nutricia	Abbott	Abbott
Kcal/ml	1	2	2	1.8
Prot, g/L	40	75	70	81
Energy ratio %	16 prot 35 fat 49 CHO	15 prot 45 fat 40 CHO	15 prot 43 fat 41 CHO 2g FOS	18.1 prot 48.4 fat 33.5 CHO 8.4 g FOS
Na/K, mmol/L	43.5 Na/38.5 K	25.6 Na/5.6 K	36.5 Na/27.2 K	30 Na/27 K
Fibers	no	no	yes	yes
Omega -3	Not always	no	yes	yes

# What can be expected from ONS using commercial liquid diets

- Single 200 ml can → 360-400 kcal, 14-16 g protein
- Intradialytic administration (one can) → total weekly intake 1100-1200 Kcal, 42-48 g protein
- Daily administration (one can/day, intra- and interdialytic): total weekly intake 2500-2800 Kcal, 98-102 g prot

## Oral Intradialytic Nutritional Supplement Use and Mortality in Hemodialysis Patients

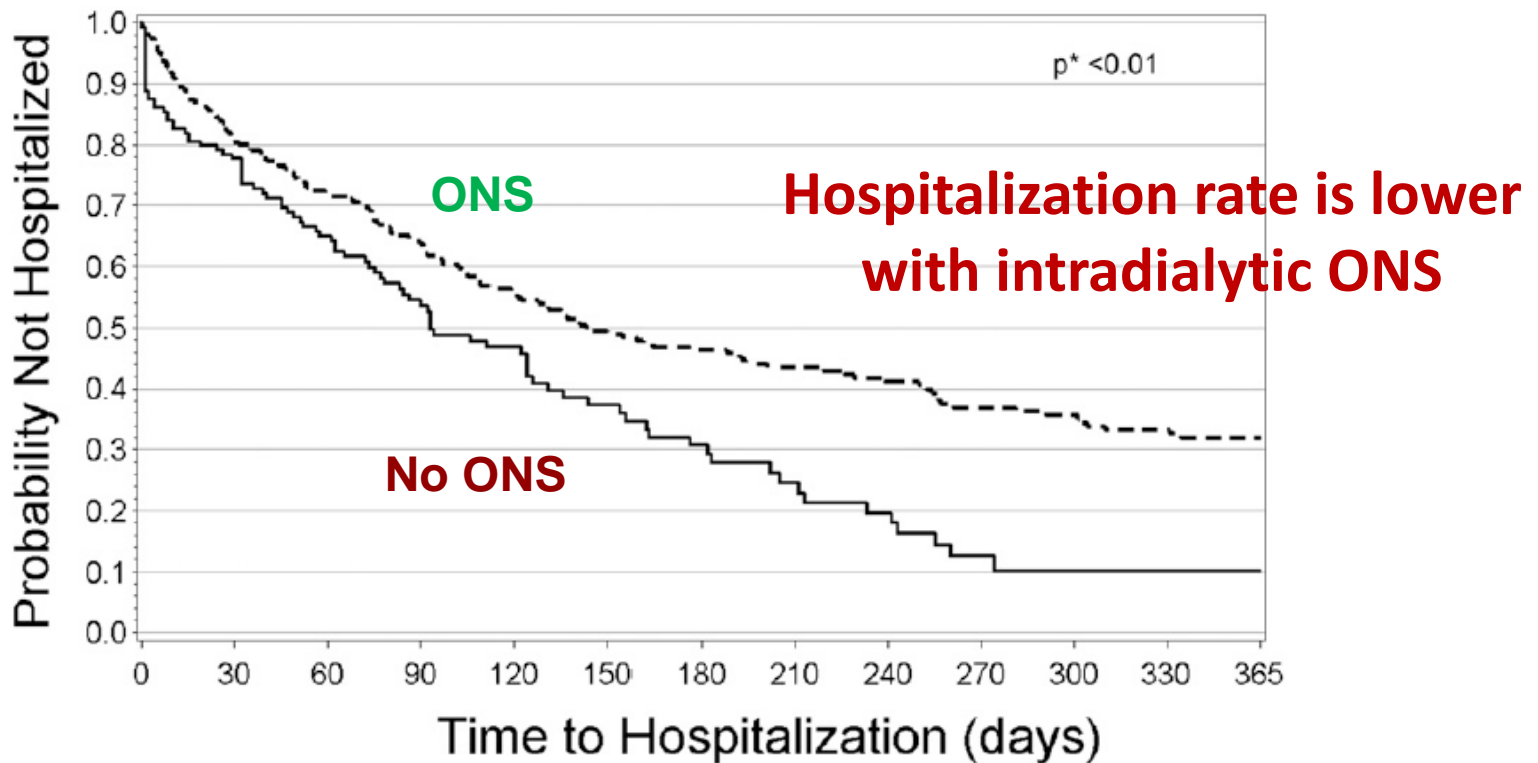
Daniel E. Weiner, MD, MS,<sup>1</sup> Hocine Tighiouart, MS,<sup>2</sup> Vladimir Ladik, MS,<sup>3</sup>  
Klemens B. Meyer, MD,<sup>1</sup> Philip G. Zager, MD,<sup>4</sup> and Douglas S. Johnson, MD<sup>5</sup>



Kaplan-Meier curve shows survival probabilities by nutritional supplement protocol (NSP) status, derived from propensity score-matched analyses, for the primary cohort

# Association between Oral Nutritional Supplementation and Clinical Outcomes among Patients with ESRD

Christine Cheu,\* Jeffrey Pearson,\* Claudia Dahlerus,\* Brett Lantz,\* Tania Chowdhury,\* Peter F. Sauer,<sup>†</sup> Robert E. Farrell,<sup>†</sup> Friedrich K. Port,\* and Sylvia P.B. Ramirez\*



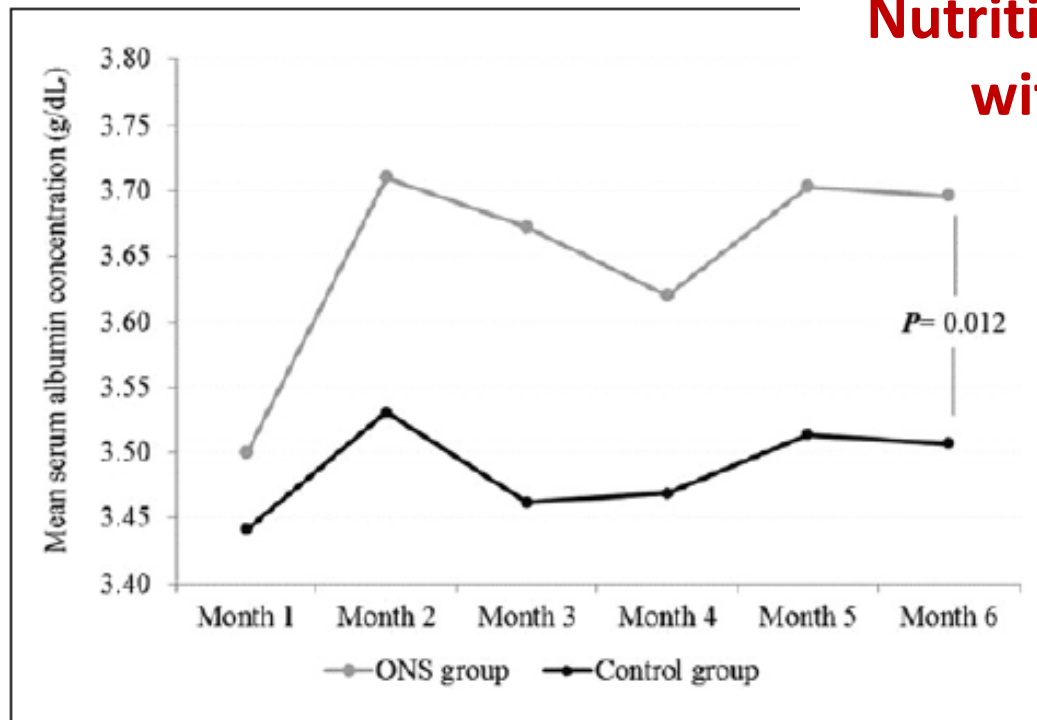
*Clin J Am Soc Nephrol* 8: 100–107, 2013.

# Long-Term Oral Nutrition Supplementation Improves Outcomes in Malnourished Patients With Chronic Kidney Disease on Hemodialysis

Siren Sezer, MD<sup>1</sup>; Zeynep Bal, MD<sup>1</sup>; Emre Tatal, MD<sup>1</sup>; Mehtap Erkmen Uyar, MD<sup>1</sup>; and Nurhan Ozdemir Acar, MD<sup>1</sup>

**Nutritional status is improved with intradialytic ONS**

Two daily servings containing 40 Kcal and 14 g of proteins



**Figure 2.** Mean serum albumin concentration (g/dL) in oral nutrition supplementation (ONS) and control groups during the study period.

# ONS and nutritional outcomes

**Table 2 | Effects of oral nutritional supplements (ONS) on nutritional outcomes in MHD patients in randomized clinical trials**

Reference	n	Design	Days	Nutritional significant effects
Acchiardo <i>et al.</i> <sup>149</sup>	15	RCT: ONS versus control groups	105	↑ Albumin, transferrin, bone density
Allman <i>et al.</i> <sup>150</sup>	21	RCT: ONS versus control groups	180	↑ BW, LBM
Tietze <i>et al.</i> <sup>151</sup>	19	RCT, crossover, ONS versus control periods	120	↑ BW, arm muscle circumference
Eustace <i>et al.</i> <sup>152</sup>	47	RCT: ONS versus control groups	90	↑ Albumin, grip strength, SF12 mental health
Hiroshige <i>et al.</i> <sup>153</sup>	44	RCT, crossover, ONS versus control periods	180	↑ DEI, DPI, fat mass, fat-free mass, albumin
Sharma <i>et al.</i> <sup>154</sup>	40	RCT: ONS versus control groups	30	↑ Albumin
Leon <i>et al.</i> <sup>155</sup>	180	RCT: ONS versus control groups	365	↑ DEI, DPI, albumin
Cano <i>et al.</i> <sup>87</sup>	186	RCT: ONS versus ONS + IDPN groups	365	↑ nPNA, BMI, albumin, prealbumin in both groups
Fouque <i>et al.</i> <sup>156</sup>	86	RCT: ONS versus control groups	90	↑ DEI, DPI, SGA, QOL
Moretti <i>et al.</i> <sup>157</sup>	49	RCT: ONS versus control groups	365	↑ nPNA, albumin

Abbreviations: BMI, body mass index; BW, body weight; DEI, dietary energy intake; DPI, dietary protein intake; IDPN, intradialytic parenteral nutrition; LBM, lean body mass; MHD, maintenance hemodialysis; nPNA, normalized protein nitrogen appearance; QOL, quality of life; RCT, randomized clinical trial; SGA, subjective global assessment.

*Kidney International* (2013) **84**, 1096–1107

**Positive effects of ONS on nutritional status**

**IDPN**



# What is IDPN

Intradialytic parenteral nutrition (IDPN) is a specific form of nutritional supplementation for ESRD patients on hemodialysis, based on the administration of nutrients (usually a mixture of amino acids, dextrose and lipids as lipid emulsions) during each dialysis session

# Modalities of IDPN

## a) Self-made IDPN

- Sequential administration of nutrients in the circuit
- All-in-one bags with nutrients compounded in the dialysis ward

## b) Pharmacy-compounded IDPN

- All-in-one bags with nutrients compounded by hospital pharmacy

## c) Commercial all-in-one admixtures

## Types of IDPN

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(a) IDPN based on compounded all-in-one admixtures

All-in-one admixtures with bags compounded in the dialysis ward

All-in-one admixtures with nutrients compounded by the hospital pharmacy

(b) IDPN based on commercially available all-in-one admixtures

Volume 625–1,250 ml

Osmolarity 1,200–1,600 mOsm/l

Essential and non essential aminoacids + glucose + lipid emulsions

Caloric density about 1 kcal/ml

Aminoacids 40–60 g/l

Non protein kcal ratio (gluc/lip) about 2:1

P 10–16 mmol/l

K 24–40 mmol/l

Available with or without electrolytes

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*IDPN* intradialytic parenteral nutrition

	BBraun		Baxter	Fresenius
	Nutriflex lipid		Olimel	SmofKabiven
	Special	Plus	N9/N9E	
Total volume (ml)	1,250/1,875/ 2,502	1,250/1,875/ 2,500	1,000/1,500/ 2,000	986/1,477/1,970
Total protein (g/l)	57.6	38.4	44.3	50
Total kcal (1 l)	1,180	1,012	1,140	1,100
Non protein kcal (1 l)	936	840	960	900
g of nitrogen/l	8	5.6	7	8
Glucose (g/l)	144	120	140	125
Lipids (g/l)	40	40	40	38
Type of lipid	PUFA + MCT	PUFA + MCT	MUFA + PUFA	PUFA (omega-3) + MCT
Osmolarity (mOsm/l)	1,545	1,215	1,170	1,500
Na (mmol/l)	53.6	40	35	40
K (mmol/l)	37.6	28	30	30
P (mmol/l)	16	12	15	12
Available without electrolytes?	Yes	Yes	Yes (N9E)	Yes (only the 1,500 and 2,000 ml bags)

**Table 3** All-in-one admixtures for IDPN in ESRD

	BBraun	
	Special	Plus
Total volume (ml)	1,250/1,875/ 2,502	1,250/1,875/ 2,500
Total protein (g/l)	57.6	38.4
Total kcal (1 l)	1,180	1,012
Non protein kcal (1 l)	936	840
g of nitrogen/l	8	5.6
Glucose (g/l)	144	120
Lipids (g/l)	40	40
Type of lipid	PUFA + MCT	PUFA + MCT
Osmolarity (mOsm/l)	1,545	1,215
Na (mmol/l)	53.6	40
K (mmol/l)	37.6	28
P (mmol/l)	16	12
Available without electrolytes?	Yes	Yes

*IDPN* intradialytic parenteral nutrition, *ESRD* end stage renal disease, *MCT* medium chain triglycerides, *MUFA* monounsaturated fatty acids, *PUFA* polyunsaturated fatty acids

# Safe IDPN

- **Volume**: not more than 1000 ml/dialysis
- **Infusion rate**: not more than 250 ml/hour
- **Aminoacids**: not more than 50 g/dialysis
- **Glucose**: not more than 500 Kcal/dialysis
- **Lipids**:  $\leq 1$  g/Kg/dialysis, i.e. not more than 500 Kcal/dialysis

**Table 5** Practical aspects of IDPN

	Suggestions	Actions
Formula	Use the most concentrated commercial all-in-one admixtures (energy density about 1.0 kcal/ml)	
Biochemical monitoring	Check serum triglyceride levels before planning an IDPN program Check serum glucose levels at dialysis start, mid-dialysis, end-dialysis and 1 h after the end of the dialysis at each dialysis of the first 3 IDPN weeks	Don't start IDPN if levels >300 mg/dl  Serum glucose levels should be maintained in the 110–180 mg/dl range. If serum glucose >180 mg/dl add subcutaneous insulin administered as rapid action analogues (start with 0.1 UI/kg); Do not give insulin after the 3rd h of dialysis
IDPN administration	Infuse IDPN in the venous drip chamber Start nutrient administration after 15 min of dialysis, when dialysis machine pressures and patient parameters are stable Start slowly, with 1/3 of the targeted amount in the first week, 2/3 in the second week and full amount from the 3rd week	Always use a parenteral infusion pump  If a nutrient admixture with 1 kcal/ml is used, this means not more than 1 ml/kg/h the first week, 2 ml/kg/h the second week, 3 ml/kg/h at full regimen
Dialysis procedures	Remove fluid added with IDPN by adjusting the ultrafiltration rate as per patient's needs Check pre-dialysis electrolytes	  In the case of severe hyperkalemia ( $\geq 6$ mmol/l) and/or hyperphosphatemia (>5.5 mg/dl), use electrolyte-free admixtures
Nutrient intake	Calculate the maximum macronutrient amount given by IDPN per dialysis (4 h) as energy 15 kcal/kg/dialysis and aminoacids: 0.8 g/kg/dialysis;	Calculate the amount of nutrient admixture as ml/dialysis session and divide by the hours of dialysis to have the hourly administration rate of fluids

# Time of onset and duration of different insulin preparations

Type of Insulin	Onset	Peak	Duration
Fast-acting			
Regular	½-1 hr.	2-4 hr.	6-8 hr.
Lyspro/ Aspart/ Glulisine	<15 min.	1-2 hr.	4-6 hr.
Intermediate-acting			
NPH	1-2 hr.	6-10 hr.	12+ hr.
Long-acting			
Detemir	1 hr.	Flat, Max effect in 5 hrs.	12-24 hr.
Glargine	1.5 hr.	Flat, Max effect in 5 hrs.	24 hr.



# What we can expect from IDPN

- 1 L of IDPN /dialysis (1000 Kcal, 50 g of AA) for three HD/week
- Weekly amount of nutrients by IDPN: 3000 Kcal + 150 g
- 10-15% of AA lost through the filter
- Daily supplementation (including non dialysis days): 5-6 kcal/Kg/day, 0.25 g/Kg/day of AA → i.e, not more than 20-25% of ideal daily nutrient intake targets

**Is IDPN safe in daily  
clinical practice?**

## Very low rate of metabolic complications during IDPN in ESRD patients on HD

Table 2. Adverse events observed during 2-yr follow-up<sup>a</sup>

Adverse Event	No. of Events	
	Control Group	IDPN Group
Event		
Deaths	36	40
heart failure	10	8
stroke	7	8
infection	8	7
cancer	1	7
Other causes	10	10
Hospitalizations for arteriovenous care	64	54
vascular access thrombosis	10	10
Hospitalization for other reasons	180	180
Events inducing discontinuation of IDPN	—	11
Nausea and vomiting	34	46
Diarrhea	14	8
Abdominal pain	9	8
Increase in plasma triglycerides >2 mmol/L	2	8
Increase in serum ALAT >1 N	1	0
Increase in serum GGT >1 N	1	9

<sup>a</sup>Some patients had more than one event.

**Table 3** Serum lipids of patients and controls in pre- and postdialysis states at the beginning and after 6 months of the study period

	<i>IDPN group</i>		<i>P value</i>	<i>Controls</i>		<i>P value</i>
	<i>Pre-HD</i>	<i>Post-HD</i>		<i>Pre-HD</i>	<i>Post-HD</i>	
<i>Total cholesterol (mg/dl)</i>						
0 months	183.3 ± 49.9	199.2 ± 55.6	NS	178.0 ± 28.9	192.5 ± 32.5	0.002
6 months	190.8 ± 39.7	190.7 ± 32.6	NS	156.7 ± 25.5	167.7 ± 26.3	0.001
<i>HDL (mg/dl)</i>						
0 months	58.7 ± 17.2	60.3 ± 16.1	NS	51.3 ± 20.3	58.2 ± 23.8	0.011
6 months	52.7 ± 20.8	53.5 ± 20.0	NS	49.7 ± 16.6	56.0 ± 20.4	0.028
<i>LDL (mg/dl)</i>						
0 months	106.3 ± 45.1	113.7 ± 41.1	NS	110.2 ± 37.4	122.7 ± 40.6	0.003
6 months	108.8 ± 39.1	113.3 ± 39.9	NS	92.0 ± 31.4	101.0 ± 32.6	0.001
<i>Triglycerides (mg/dl)</i>						
0 months	106.8 ± 39.7	199.3 ± 125.5	NS	151.8 ± 46.3	198.8 ± 166.2	NS
6 months	112.7 ± 47.3	147.3 ± 69.7	NS	160.7 ± 58.5	167.7 ± 159.9	NS

Abbreviations: HDL, high-density lipoprotein; IDPN, intradialytic parenteral nutrition; LDL, low-density lipoprotein; post-HD, post-hemodialysis; pre-HD, pre-hemodialysis.

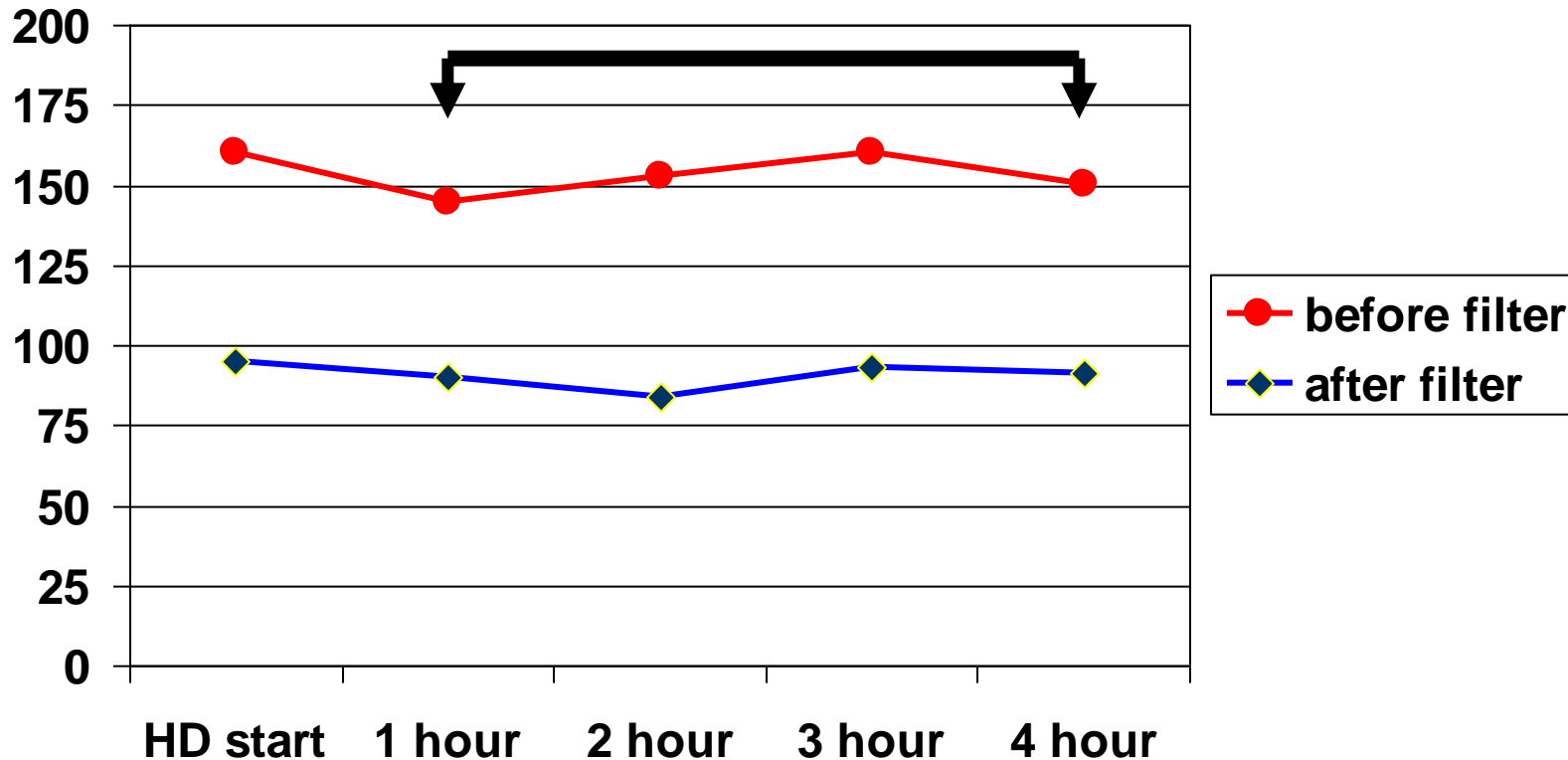
**No difference in serum triglyceride levels  
between IDPN group and controls**

**Any technical problem with filters and circuits when lipid emulsions are infused during the dialysis session?**

# No changes of pressures in the extracorporeal circuit of dialysis during IDPN with 20% lipid emulsion in ESRD

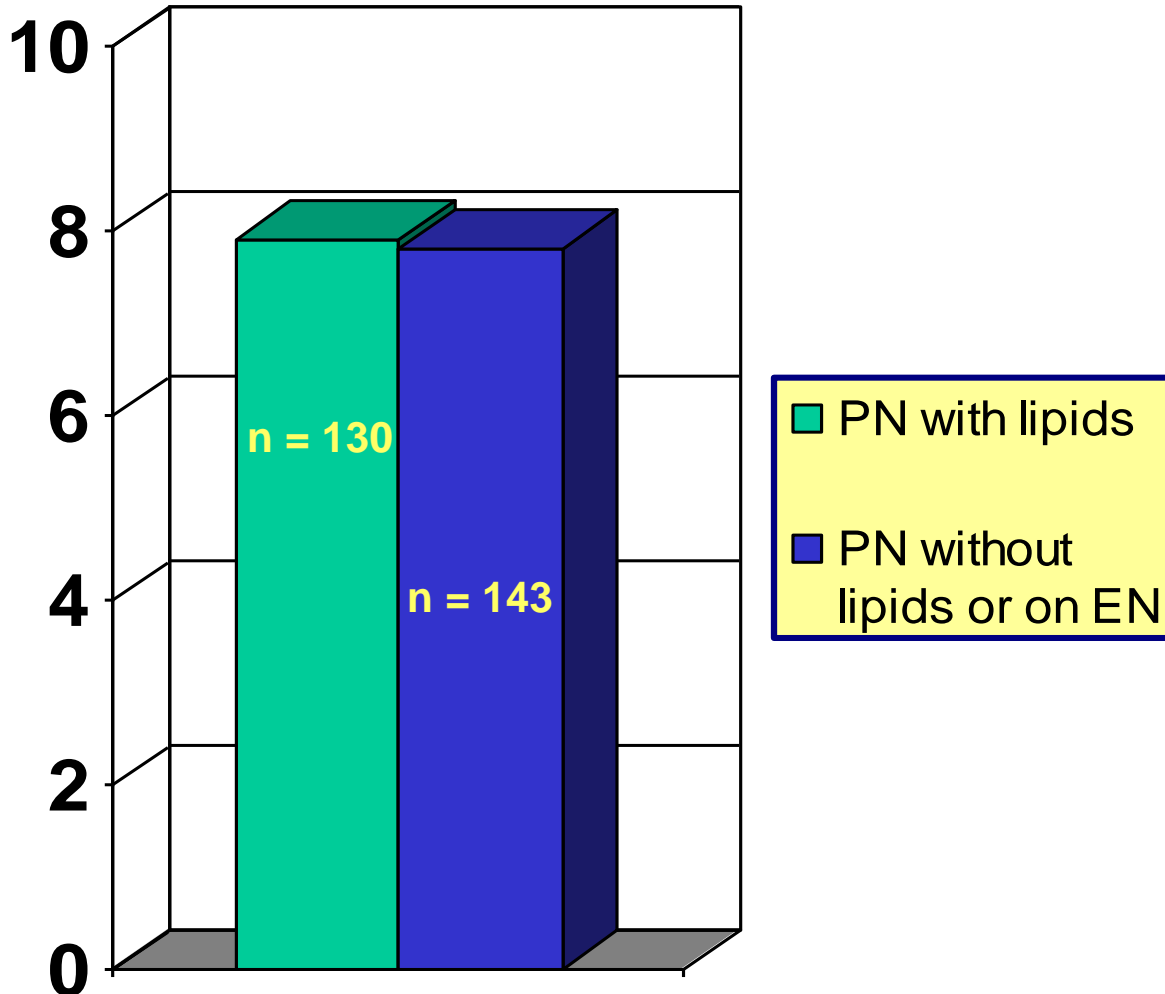
Circuit Pressures  
mmHg

20% MCT/LCT Lipid emulsion 1 ml/Kg/h



# No effects of parenteral nutrition with lipid emulsions on filter duration in sustained low-efficiency dialysis (SLED)

Hours of treatment



- 273 SLED in 37 ICU patients with AKI
- Prescribed duration 8 hours/treatment
- PN with all-in-one system
- (20% lipid emulsion 250-500 ml/24 hours)

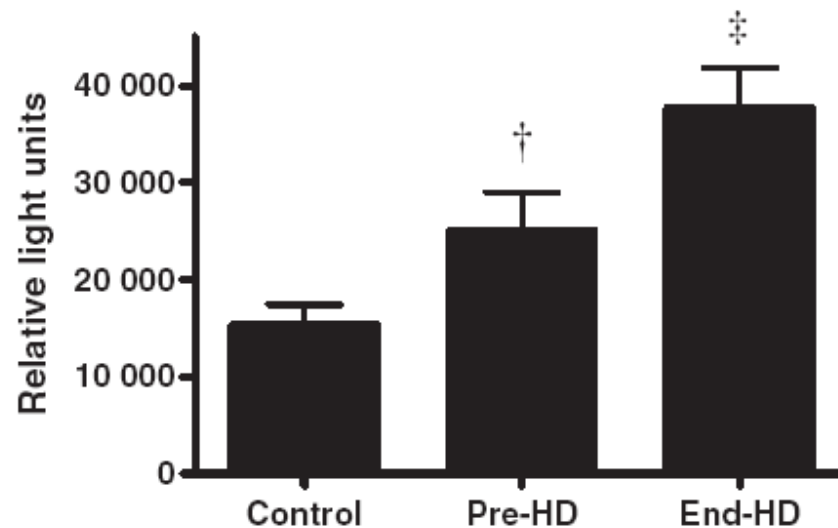


**What are the specific nutritional effects of intradialytic nutritional support in ESRD patients?**



# Activation of caspase-3 in the skeletal muscle during haemodialysis

Michel A. Boivin<sup>\*</sup>, Shadi I. Battah<sup>\*</sup>, Elizabeth A. Dominic<sup>†</sup>, Kamyar Kalantar-Zadeh<sup>‡</sup>, Arny Ferrando<sup>§</sup>, Antonios H. Tzamaloukas<sup>¶</sup>, Rama Dwivedi<sup>\*\*</sup>, Thomas A. Ma<sup>††</sup>, Pope Moseley<sup>‡‡</sup> and Dominic S. C. Raj<sup>\*\*</sup>, <sup>§§</sup>, <sup>¶¶</sup>



**Muscle catabolism is activated during dialysis**

†  $P < 0.01$  Pre-HD vs. Control; ‡  $P < 0.001$  End-HD vs. Pre-HD and control.

**Figure 1** Caspase-3 activity in muscle was increased in patients with ESRD at baseline (pre-HD) compared with controls and was further augmented by haemodialysis (End-HD). †  $P < 0.01$  Pre-HD vs. Control; ‡  $P < 0.001$  End-HD vs. Pre-HD and control.

# Negative protein balance in skeletal muscle during hemodialysis

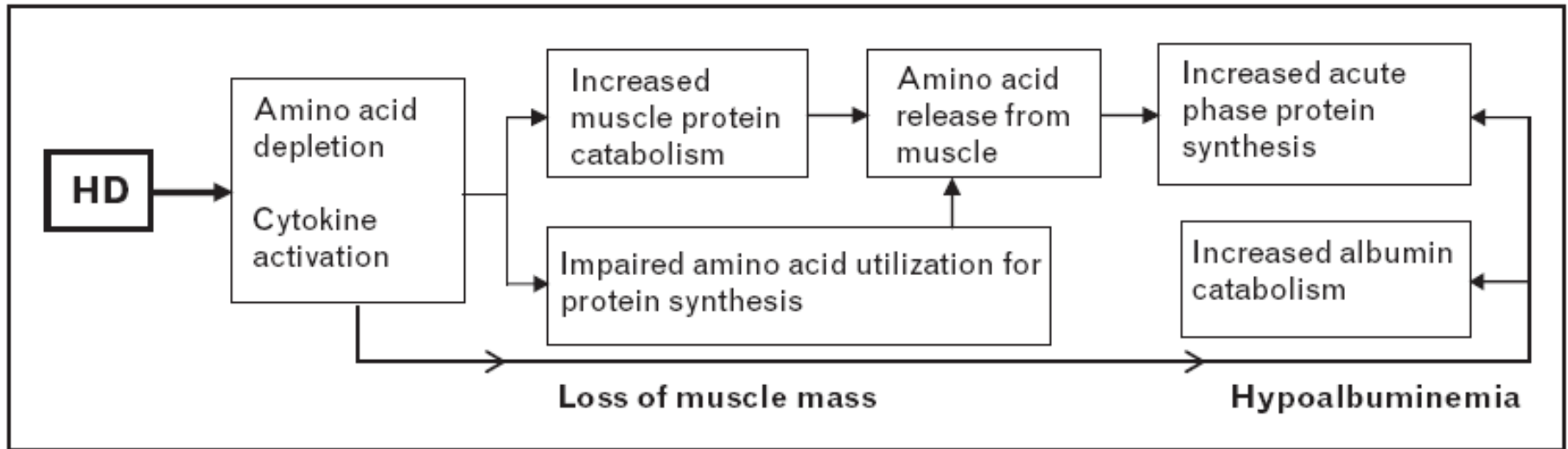
**Table 2** Muscle protein kinetic studies during haemodialysis

	Control	Pre-HD	End-HD
Phenylalanine concentration in artery (Ca; $\mu\text{mol L}^{-1}$ )	89.8 $\pm$ 4.5	84.5 $\pm$ 7.8*	65.9 $\pm$ 5.8
Phenylalanine concentration in vein (Cv; $\mu\text{mol L}^{-1}$ )	92.0 $\pm$ 4.6	85.2 $\pm$ 7.9*	74.6 $\pm$ 5.4 <sup>†</sup>
Leg muscle protein synthesis (Rd; $\text{nmol } 100 \text{ mL}^{-1} \text{ min}^{-1}$ )	42.62 $\pm$ 5.78	41.19 $\pm$ 3.03	55.15 $\pm$ 4.48 <sup>‡</sup>
Leg muscle proteolysis (Ra; $\text{nmol } 100 \text{ mL}^{-1} \text{ min}^{-1}$ )	50.47 $\pm$ 7.69	41.63 $\pm$ 2.47	84.61 $\pm$ 3.65 <sup>††, ‡‡</sup>
Net balance ( $\text{nmol } 100 \text{ mL}^{-1} \text{ min}^{-1}$ )	-7.85 $\pm$ 5.47	-2.28 $\pm$ 1.93	-29.47 $\pm$ 6.03 <sup>††</sup>

\* $P < 0.05$  Pre-HD vs. End-HD, <sup>†</sup>Ca vs. Cv  $P < 0.01$ ; <sup>‡</sup> $P < 0.02$  End-HD vs. Pre-HD, <sup>††</sup>Ra vs. Rd  $P < 0.05$ ; <sup>‡‡</sup> $P < 0.001$  End-HD vs. Control and Pre-HD.

Eur J Clin Invest 2010; 40 (10): 903–910

**AA are released from muscle protein catabolism during hemodialysis: they are in part utilized in the liver for acute phase protein synthesis, and in part they are removed by hemodialysis itself**



**Current Opinion in Nephrology and Hypertension** 2008, 17:589–594

**Free AA losses into dialysate 8.2 gr/dialysis  
Plasma AA levels decreased by 33%**

**Wolfson M et al.,  
Kidney Int 1982; 21:500**

# **Positive effects of intradialytic nutritional support (oral or parenteral)**

- Improved energy balance
- Improved protein (AA) balance
- Improved albumin synthesis rate
- Improved nutritional parameters

Pupim LB et al., Sem Nephrol 2006; 26:134-157

Table 3. Randomized Studies of IDPN

Study	Design	Treatment Duration	No. With PEW	Parameters Measured	Outcome
Toigo et al, <sup>30</sup> 1989	11 pts: 26.5 g of modified EAA 10 pts: 24 g of EAA + NEAA	6 mo	None	Nerve conduction velocity, Alb	Decrease in Alb in EAA + NEAA group
Cano et al, <sup>31</sup> 1990	12 pts: 0.08 g of N/kg (/HD session) from EAA + NEAA, 1.6 g/kg (/HD session) lipids 14 pts: no intervention	3 mo	All	BW, appetite, MAMC	Increase in calorie (9 kcal/kg/d) and protein intake (0.25 g/kg/d) in IDPN-treated pts
McCann et al, <sup>14</sup> 1999	19 pts; 70% glucose, 15% AA, 20% lipids	11 wk	NA	Delivered Kt/V, URR	Decrease in delivered Kt/V in pts who received AA-containing IDPN
Navarro et al, <sup>32</sup> 2000	17 pts	3 mo			Positive net AA balance Increase in PCR, Alb, transferrin
Cano et al, <sup>33</sup> 2006	17 pts: olive oil-based IV lipid emulsion 18 pts: soybean oil-based IV lipid emulsion	5 wk			Both groups showed similar improvement in nutritional status, plasma lipid, oxidative and inflammatory parameters
Cano et al, <sup>34</sup> 2007	89 pts: IDPN 93 pts: control	12 mo	All	Primary end point, all-cause mortality; secondary end points, hosp rate, BW, Karnofsky score, BMI	No difference in hosp rate or mortality between 2 groups

**Positive nutritional and metabolic effects of IDPN on nutritional status in ESRD**

## **Intradialytic Parenteral Nutrition Does Not Improve Survival in Malnourished Hemodialysis Patients: A 2-Year Multicenter, Prospective, Randomized Study**

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**The FINE study from France:**

**One-yr IDPN on top of optimal daily oral supplementation**

## ABSTRACT

Although intradialytic parenteral nutrition (IDPN) is a method used widely to combat protein-calorie malnutrition in hemodialysis patients, its effect on survival has not been thoroughly studied. We conducted a prospective, randomized trial in which 186 malnourished hemodialysis patients received oral nutritional supplements with or without 1 year of IDPN. IDPN did not improve 2-year mortality (primary end point), hospitalization rate, Karnofsky score, body mass index, or laboratory markers of nutritional status. Instead, both groups demonstrated improvement in body mass index and the nutritional parameters serum albumin and prealbumin ( $P < 0.05$ ). Multivariate analysis showed that an increase in prealbumin of  $>30$  mg/L within 3 months, a marker of nutritional improvement, independently predicted a 54% decrease in 2-year mortality, as well as reduced hospitalizations and improved general well-being as measured by the Karnofsky score. Therefore, although we found no definite advantage of adding IDPN to oral nutritional supplementation, this is the first prospective study demonstrating that an improvement in prealbumin during nutritional therapy is associated with a decrease in morbidity and mortality in malnourished hemodialysis patients.

**No advantage per se of adding IDPN to adequate oral supplementation**

**Nutritional supplementation, no matter what was the modality (oral supplementation alone or IDPN+oral supplementation etc.) improved mortality in ESRD patients on HD if nutritional targets are met**

# Take home messages

- Protein-energy wasting (PEW) is frequent among ESRD patients on hemodialysis and represents a negative prognostic factor
- Intradialytic nutritional support is able to improve nutritional status in ESRD and, likely, the vicious circle of PEW
- Along with ONS, IDPN is a safe and effective modality for nutritional supplementation in selected patients