Pancreatectomy Abolishes the Renal Hemodynamic Response to a Meat Meal in Man

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Dear Sir,

According to Brenner et al. [1], the hyperfiltration response which follows intravenous amino acid administration [2–4] of ingestion of a meat meal [5, 6] may be due to a circulating hormone or some other intermediate effector among which glucagon may be a good candidate.

We report on the absence of the hyperfiltration response to a meat meal in man after total pancreatectomy. This finding points to the role of a factor of pancreatic origin in the genesis of the renal hemodynamic response to protein load. We studied a 45-year-old man, weighing 72 kg, who, because of severe acute pancreatitis, had undergone total pancreatectomy 4 years earlier. After surgery he had strictly adhered to an appropriate protocol of insulin administration and to a dietary regimen providing 40 g of protein and 2,000 cal/day. Protein intake was controlled before the study on 3 consecutive days by means of urea generation rates. At the time of the study, plasma creatinine was 0.99 mg/dl, creatinine clearance 84 ml/min × 1.73 m², blood urea 48 mg/dl, and fasting blood glucose 1.55 g/l. Microalbuminuria was absent.

The patient was studied before (3 clearance studies (C_1-C_3) and after a meat meal MM 5 clearance studies (C_4-C_8) at 30, 60, 90, 120 and 180 min). Each clearance lasted 30 min with the exception of C_8 lasting 60 min. The meat meal provided 2 g of protein/kg body weight in the form of cooked red meat. All data measured in C_1-C_3

	Baseline	Time after meat meal				
		30 min	60 min	90 min	120 min	180 mii
GFR, ml/min+1.73+m ²	75	73	81	82	84	
RPF, ml/min · 1.73 · m ²	422	.360	382	390	392	408
FF	0.19	0.20	0.19	0.20	0.20	0.20
$U_{NA} \cdot V \mu M/min$	208	288	258	371	357	228
C _{N0} , ml/min · 1.73 · m ²	1.22	2.09	1.77	2.06	2.8	1.9
CN_/GFR-100	1.49	2.79	2.42	2.54	3.43	2.27
$UK \cdot V, \mu M/\min$	143	108	86	139	126	76
CK, ml/min · 1.73 · m ²	24.7	25.7	20.5	27.5	28	19
CK/GFR-100	30	34.2	28.9	34	34	23
C _{1.1} , ml/min · 1.73 · m ²	20.5	18.7	28	27	28	27
GFR - CLI	63	56	45	54	54	57
C _{Li} /GFR	0.25	0.25	0.38	0.33	0.34	0.32
I - C _{LI} /GFR	0.75	0.75	0.61	0.66	0.66	0.68
$(C_{1,i} - C_{Na}) \cdot P_{Na}, \mu M/min$	2641	2299	3850	3448	3358	3395
$(C_{L_1} - C_{N_R})/C_{L_1}$	0.94	0.89	0.94	0.92	0.90	0.93
Ht, %	44	44	44	44	44	44
RVR, mm Hg/min/ml	15.6	17.4	15.7	16	15.7	14.7
PRA, ng/ml/h	1.21	1.29	1.16	1.30	1.10	1.35
Aldosterone, pg/ml	57	58	62	55	64	63
C peptide, ug/ml	0	0	0	0	0	0
Glucagon, pg/ml	180	210	126	118	109	143
Blood glucose, g/l	1.55	1.60	1.58	1.50	1.50	1.62
Amino acids, $\mu M/1$	3600	3819	4200	4500	4906	6421

Table 1. Effects of a meat meal in a pancreatectomized man

were averaged and the data considered as baseline values prior to protein load. During the study, the patient received 1.5 U/h of regular insulin. We measured glomerular filtration rate (GFR) by inulin, renal plasma flow (RPF) by p-aminohippurate, the delivery of tubular fluid from the proximal straight tubules by lithium clearance (C_{Li}), sodium and potassium clearance (C_{Na} , C_K), absolute and fractional Na and K excretion (U_{Na}·V, U_K·V, C_{Na}/GFR·100, C_K/ GFR-100), hematocrit (Ht), plasma concentrations of C peptide, glucagon, renin, aldosterone, glucose, and total amino acids. We calculated the absolute rate of proximal reabsorption of isotonic fluid (GFR - CLI) by CLI-derived formulae, the proximal fraction reabsorption (1- C_{Li}/GFR), the fractional clearance of lithium ($C_{Li}/$ GFR), the absolute $(C_{L1}-C_{Na}) \cdot P_{Na}$ and fractional $(C_{L1}-C_{Na})/C_{Li}$ distal sodium reabsorption and the renal vascular resistance (RVR) according to the formula MAP. (1-Ht)/RPF, where MAP is the mean arterial blood pressure.

The results of this study are summarized in table 1 and show that: (1) GFR and RPF following a meat meal did not exceed baseline values; (2) inhibition of proximal isotonic reabsorption took place; (3) stimulation of distal sodium reabsorption occurred; (4) plasma amino acid concentration increased over time, and (5) glucagone of gastric/pancreatic origin (7) was not affected by the meat meal.

The data also indicate that after pancreatectomy no hyperfiltration response follows a meat meal in subjects on low-protein alimentation, which points to the lack of a circulating hormone/effector of pancreatic origin [8]; the initial depression of GFR and RPF which followed the meat meal is supported by data obtained in adults and children with diabetes mellitus [9, 10].

References

- Brenner BM, Meyer TW, Hostetter TH: Dietary protein intake and the progressive nature of kidney disease: The role of hemodynamically mediated glomerular sclerosis in ageing, renal ablation and intrinsic renal disease. N Engl J Med 1982; 300:652-659.
- 2 Graf H, Stummvoll HK, Luger A, Prager R: Effect of amino acid infusion on glomerular filtration rate. N Engl J Med 1983; 308:159-160.

- 3 Teer We PM, Geerlings W, Rosman JB, Sluiter WJ, Van der Gest WJ, Donker AJM: Testing the renal reserve filtration capacity with an amino acid solution. Nephron 1985;41:193-199.
- 4 Castellino P, Coda B, DeFronzo RA: Effect of an amino acid infusion on renal hemodynamics in humans. Am J Physiol 1986;251:F132-F140.
- 5 Bosch JP, Saccaggi A, Lauer A, Ronco C, Belledonne M, Glabman S: Renal functional reserve in humans. Am J Med 1983; 75:943-949.
- 6 Alvestrand A, Bergstrom J: Glomerular hyperfiltration after protein ingestion, during glucagon infusion and insulin-dependent diabetes is induced by a liver hormone: Deficient production of this hormone in hepatic failure causes hepato-renal syndrome. Lancet 1984;i:195-197.
- 7 Valverde I, Alarcon C, Ruiz-Grande C, Rovira A: Plasma glucagon and glucagon-like immunoreactivity in totally pancreatectomized humans; in Tiego A, Alberti KGMM, DelPrato S, Vranic M (eds): Diabetes Secondary to Pancreatopathy, Amsterdam, Excerpta Medica/Elsevier Sciences, 1988; pp 51-60.
- 8 Premen AJ, Hall JE, Smith JRMJ: Postprandial regulation of renal hemodynamics: Role of pancreatic glucagon. M J Physiol 1985;248: F656-F662.
- 9 Bosch JP, Lew S, Glabman S, Lauer A: Renal hemodynamic changes in humans. Response to protein loading in normal and diseased kidney. Am J Med 1986;81:809-815.
- 10 Castellino P, DeSanto NG, Capasso G, Anastasio P, Coppola S, Capodicasa G, Perna A, Torella R, Salvatore T, Giordano C: Low protein alimentation normalizes renal haemodynamic response to acute protein ingestion in type one diabetic children. Eur J Clin Invest 1989;19:78–82.

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