

Glutamine force-feeding effect on plasma amino-acid concentrations in growing rats fed a cafeteria diet

Mj Salvadó, Li Arola

▶ To cite this version:

Mj Salvadó, Li Arola. Glutamine force-feeding effect on plasma amino-acid concentrations in growing rats fed a cafeteria diet. Reproduction Nutrition Development, 1994, 34 (2), pp.165-173. hal-00899647

HAL Id: hal-00899647

https://hal.science/hal-00899647

Submitted on 11 May 2020

HAL is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers.

L'archive ouverte pluridisciplinaire **HAL**, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d'enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.

Original article

Glutamine force-feeding effect on plasma amino-acid concentrations in growing rats fed a cafeteria diet

MJ Salvadó, LI Arola

Departament de Bioquímica i Biotecnologia, Universitat Rovira i Virgili, Facultat de Química, Pl Imperial Tarraco 1, 43005 Tarragona, Spain

(Received 29 July 1993; accepted 3 February 1994)

Summary — The effect of a glutamine force-feeding on plasma amino-acid levels in rats fed either a reference diet or a cafeteria diet was studied during weaning. The increases in plasma amino-acid levels shown by rats eating the cafeteria diet were related to the force-feeding and/or the age studied. The glutamine solution decreased the levels of proline, ornithine and tyrosine in the plasma of rats eating the cafeteria diet. In rats fed the reference diet, glutamine solution increased the plasma concentrations of threonine and cysteine. A major effect of diet over force-feeding was shown.

glutamine / force-feeding / cafeteria diet / growing rat / amino acid

Résumé — Effets d'un gavage de glutamine sur les concentrations plasmatiques en aminoacides chez des rats en croissance alimentés en cafeteria. L'effet de l'administration orale
d'une solution de glutamine sur les concentrations des acides aminés plasmatiques chez le rat
nourri avec une alimentation de référence ou une alimentation de type cafeteria a été étudié
pendant leur développement. Les variations des acides aminés chez le rat nourri avec une
alimentation en cafeteria dépendent du gavage et/ou de l'âge étudiés. Nous avons observé une
diminution des concentrations de proline, ornithine et tyrosine dans le plasma des rats nourris avec
l'alimentation en cafeteria en utilisant un gavage avec la glutamine. En revanche, chez les rats
nourris avec l'aliment de référence, le gavage avec la glutamine a provoqué une augmentation des
concentrations plasmatiques de thréonine et de cystéine. L'effet de l'alimentation en cafeteria est
plus important que celui du gavage avec la glutamine.

qlutamine / qavage / alimentation en cafeteria / rat en croissance / acides aminés

INTRODUCTION

The cafeteria diets were introduced as a way to generate an animal model of obesity that is closer than genetic obesity to the situation in humans consuming highly palatable choice diets (Sclafani and Springer, 1976). Cafeteria diets may differ somewhat in composition, but it has been shown that despite of a highly variable selection of food, the nutrient composition of the diet ingested is fairly constant (Prats et al, 1989). The diets actually selected by the rats have a common high lipid content (Naim et al, 1985), with relatively unchanged, and proportionally lower, protein and carbohydrate contents (Prats et al, 1989). Constant exposure to these diets from birth results in high growth rates (Salvadó et al, 1986), with high increases in energy intake (Prats et al, 1989) and increased thermogenesis (Rothwell and Stock, 1982). A cafeteria diet increased the retention of dietary nitrogen (Barr and McCracken, 1984) and lowered urinary nitrogen losses (Barber et al, 1985); nitrogen accretion in the body was the highest for the younger animals (Esteve et al, 1992). The amino-acid composition of the diet ingested by reference and cafeteria diet-fed rats has been analyzed: a cafeteria diet results in a higher proportion of amino acids extracted from the diet, although this diet had a very similar amino-acid composition to that of the standard reference diet (Esteve et al, 1993; Rafecas et al, 1993). The individual amino acids best absorbed by young Wistar rats fed a cafeteria diet were Ala, Ser, Hyp, Tyr, Thr and Lys (Esteve et al, 1993).

Glutamine is now classified as a conditionally essential amino acid (Smith, 1990) indicating that a dietary source is required under metabolic stress (Meijer *et al*, 1993) or certain pathologic conditions (Mobrahan, 1992), especially those concerning the intestinal tract (Souba, 1993). However, in normal physiological conditions glutamine

is not an essential amino acid, and we have supplied it here as a source of non-essential nitrogen. The supplementation of a cafeteria diet with an essential amino-acid mixture showed a synergism between the 2 treatments on nitrogen retention (Salvadó and Arola, 1992a), growth (Salvadó and Arola, 1992a) and amino-acid availability (Salvadó and Arola, 1993) in weaning rats. The present study was designed to investigate the effects of administering oral glutamine solution on the plasma concentrations of amino acids in cafeteria-diet-fed or reference-diet-fed young rats in comparison with a previous force-feeding with essential amino acids (Salvadó and Arola, 1992a; Salvadó and Arola, 1993).

MATERIALS AND METHODS

Sixty weaning rats (14, 20 and 30 d of age) were used. Litters were randomized to 8 pups at delivery and kept with their mothers until sacrifice. Primiparous Wistar rats were kept individually in polypropylene-bottomed cages with wood shavings as absorbent material. The cages were housed in a room with controlled temperature (21–22°C), humidity (75-85%) and lighting (lights on from 8 to 20 h). From mothers fed one of the 2 diets ad libitum (reference diet or cafeteria diet) pups were divided randomly into 4 groups (n = 5) on d 11 after birth: saline-reference; saline-cafeteria; glutamine-reference; glutamine-cafeteria. The pups were given a daily solution at a dose of 10 ml/kg body weight by means of a stomach canula between 9 and 10 am on postnatal d 11. The composition of the force-fed solutions was: a) 9 q/l sodium chloride (saline) in Tween 90 solution; or b) 30 g/l glutamine in Tween 90. The composition of the 2 diets were: a) reference diet: standard pellet (A04 from Panlab, Barcelona) and tap water; and b) cafeteria diet: fresh pastry, cookies, chocolate, bacon, Swiss cheese, candy, roasted hazelnuts, banana, liver pate, chow pellets (as indicate above), tap water and whole milk complemented with 250 g/l sucrose and 15 g/l of a protein and mineral supplement (Gevral Proteina, Cyanamid Iberica, Barcelona). All the materials were presented daily in excess. The total energy intake of reference diet was 1438.5 kJ/d.kg rat

(1036 kJ/d.kg from carbohydrate; 98.5 kJ/d.kg from fat; and 304.5 kJ/d.kg from protein). The total energy intake of the self-selected cafeteria diet was 2463 kJ/d.kg rat (1191 kJ/d.kg from carbohydrate; 932.5 kJ/d.kg from fat; and 339.5 kJ/d.kg from protein). A more detailed description of the diet was given previously (Prats *et al*, 1989).

Pups were killed by decapitation on postnatal d 14, 20 or 30. Blood samples were collected in dry heparinized beakers. Plasma was obtained by centrifugation, and small aliquots were deproteinized with cold acetone. The individual amino acids were determined in clear supernatants, after their derivation with dansyl chloride labelled with 14C, chromatographic separation and evaluation of the radioactivity present (Arola et al, 1976; Arola et al. 1977). Glutamate plus glutamine, aspartate plus asparagine and leucine plus isoleucine, are given as composite values, due to methodological considerations. Statistical comparisons between groups were performed with 3-factor ANOVA test (Dixon et al, 1983). The 3way ANOVA implies the factors: diet, age and force-feeding. When there were no significant interactions between the 3 factors, the results obtained from this analysis were considered; otherwise, 2-way ANOVAs were carried out. All data are expressed as mean ± SEM. In all analyses, differences were considered significant at P < 0.05.

RESULTS

The final body weight was significantly lower in glutamine than saline force-fed rats; it was also lower in the reference-diet-fed than in cafeteria-diet-fed rats. On d 30, body weights were 76.50 g for saline-reference, 56.60 g for glutamine-reference, 82.67 g for saline-cafeteria and 65.38 g for glutamine-cafeteria.

Table I shows the amino-acid concentrations in the plasma of the 4 groups, measured on d 14, 20 and 30. An overall increase in plasma amino-acid levels is shown by rats fed the cafeteria diet as was previously seen (Calles-Escandon et al, 1984; Rafecas et al, 1991; Salvadó et al, 1991; Salvadó and Arola, 1993). This

increase is especially evident in glucogenic amino acids and in imino acids and is also shown in some basic amino acids: arginine and citrulline, in the branched-chain amino acid valine and in the sulphur amino acid cysteine. However, these increases, with the only exception of glutamine + glutamate concentrations, are affected by the force-feeding and/or the age studied.

Glutamine force-feeding decreased the levels of proline, ornithine and tyrosine in the plasma of rats eating the cafeteria diet; alanine, serine, hydroxyproline, arginine, histidine and leucine + isoleucine were also lower in these rats, but only at 14 d or, in the case of valine, at 20 d. Conversely, glutamine force-feeding increased the plasma levels of threonine at 20 and 30 d, those of citrulline at 20 d and those of leucine + isoleucine and tryptophan at 30 d, when rats were fed the cafeteria diet. On the other hand, in rats fed the reference diet, glutamine force-feeding increased the plasma concentrations of threonine and cysteine, and also those of histidine only at 20 d. Glutamine force-feeding also increased the plasma levels of aspartate + asparagine at 20 d, but after a decrease at 14 d. Methionine and phenylalanine showed no significant difference for the analyzed factors.

Plasma concentrations of alanine, leucine + isoleucine and tryptophan of cafeteria-diet-fed rats changed with age when they were force-fed with glutamine but not when the force-feeding was with the saline solution. Conversely, the variations with age of hydroxyproline and arginine levels seen in the plasma of saline force-fed rats, disappeared when glutamine solution was administered. The developmental patterns showed by serine, threonine, histidine, citrulline and valine also depended on the forcefeeding supplied in rats eating the cafeteria diet. However, the changes shown with age by glycine, proline, ornithine, cysteine, taurine and tyrosine were independent of force-feeding in the 2 diets supplied, as

Table I. Amino-acid concentrations in the plasma of 14, 20 and 30 d glutamine or saline force-fed rats eating a reference or cafeteria diet.

	Age (d)	Saline reference	Saline cafeteria	Glutamine reference	Glutamine cafeteria	ANOVA
Gluconeogenic amino ad	cids	~				
alanine	14 20 30	0.83 ± 0.11 0.80 ± 0.11 1.02 ± 0.10	1.47 ± 0.14 1.46 ± 0.12 1.57 ± 0.14	0.83 ± 0.05 1.22 ± 0.11 1.01 ± 0.09	0.81 ± 0.04↓ 1.69 ± 0.13 1.79 ± 0.14	D,A,I
glutamate + glutamine	14 20 30	0.89 ± 0.07 0.83 ± 0.03 1.20 ± 0.12	1.30 ± 0.15 1.23 ± 0.06 1.70 ± 0.16	0.81 ± 0.03 0.89 ± 0.04 1.17 ± 0.23	0.87 ± 0.05 1.11 ± 0.13 1.54 ± 0.12	D,A
aspartate + asparagine	14 20 30	0.04 ± 0.007 0.02 ± 0.004 0.03 ± 0.002	0.06 ± 0.006 0.04 ± 0.005 0.04 ± 0.007	0.03 ± 0.000 0.05 ± 0.005 0.03 ± 0.007	0.04 ± 0.006 0.05 ± 0.006 0.04 ± 0.006	D,I
glycine	20	0.51 ± 0.04 0.69 ± 0.04 0.87 ± 0.15	0.75 ± 0.07 1.11 ± 0.04 1.76 ± 0.25	0.36 ± 0.02 0.99 ± 0.08 1.00 ± 0.11	0.43 ± 0.02 1.16 ± 0.12 1.42 ± 0.19	D,I
serine	20	0.33 ± 0.04 0.52 ± 0.06 0.41 ± 0.04	0.61 ± 0.07 0.87 ± 0.08 0.69 ± 0.09	0.31 ± 0.03 0.77 ± 0.04 0.42 ± 0.02	$0.34 \pm 0.08 \downarrow$ 0.75 ± 0.07 0.76 ± 0.09	D,A,I
threonine	20	0.15 ± 0.01 0.24 ± 0.03 0.24 ± 0.02	0.25 ± 0.02 0.27 ± 0.02 0.29 ± 0.04	0.16 ± 0.02↑ 0.38 ± 0.02↑ 0.29 ± 0.02↑	0.22 ± 0.01 0.41 ± 0.07↑ 0.42 ± 0.03↑	D,G,A,I
Imino acids						
proline	20	0.43 ± 0.02 0.54 ± 0.02 0.46 ± 0.06	0.66 ± 0.03 0.74 ± 0.09 0.95 ± 0.01	0.41 ± 0.02 0.70 ± 0.05 0.44 ± 0.06	$0.40 \pm 0.04 \downarrow$ $0.68 \pm 0.05 \downarrow$ $0.77 \pm 0.08 \downarrow$	D,A,I
hydroxyproline	20	0.14 ± 0.01 0.07 ± 0.01 0.06 ± 0.00	0.23 ± 0.02 0.11 ± 0.01 0.12 ± 0.02	0.13 ± 0.01 0.13 ± 0.02 0.08 ± 0.02	$0.11 \pm 0.01 \downarrow$ 0.10 ± 0.01 0.11 ± 0.03	D,A,I
Basic amino acids						
lysine	20	0.48 ± 0.01 0.46 ± 0.04 0.47 ± 0.03	0.59 ± 0.02 0.33 ± 0.04 0.40 ± 0.03	0.58 ± 0.13 0.47 ± 0.06 0.49 ± 0.04	0.56 ± 0.08 0.34 ± 0.05 0.38 ± 0.08	Α
arginine	20	0.26 ± 0.02 0.44 ± 0.07 0.27 ± 0.03	0.68 ± 0.15 0.64 ± 0.06 0.32 ± 0.07	0.24 ± 0.01 0.66 ± 0.08 0.26 ± 0.06	$0.32 \pm 0.03 \downarrow$ 0.54 ± 0.03 0.36 ± 0.08	D,A,I
histidine	20	0.21 ± 0.03 0.11 ± 0.01 0.07 ± 0.01	0.28 ± 0.03 0.09 ± 0.01 0.12 ± 0.01	0.16 ± 0.01 0.17 ± 0.02↑ 0.08 ± 0.02	$0.28 \pm 0.02 \downarrow$ 0.12 ± 0.01 0.09 ± 0.01	A,I

Table I. Cont

	Age (d)	Saline reference	Saline cafeteria	Glutamine reference	Glutamine cafeteria	ANOVA
citrulline	20	0.02 ± 0.002 0.09 ± 0.010 0.06 ± 0.010	0.05 ± 0.003 0.10 ± 0.003 0.08 ± 0.005	0.02 ± 0.002 0.10 ± 0.002 0.07 ± 0.006	0.02 ± 0.004 0.13 ± 0.0131 0.10 ± 0.015	D,A,I
ornithine	20	0.20 ± 0.01 0.17 ± 0.01 0.11 ± 0.02	0.26 ± 0.02 0.16 ± 0.02 0.13 ± 0.01	0.14 ± 0.02 0.17 ± 0.01 0.13 ± 0.01	$\begin{array}{c} 0.17 \pm 0.02 \downarrow \\ 0.16 \pm 0.02 \downarrow \\ 0.07 \pm 0.01 \downarrow \end{array}$	G,A,I
Branched-chain amino	acids					
valine	20	0.20 ± 0.03 0.40 ± 0.04 0.32 ± 0.01	0.31 ± 0.01 0.54 ± 0.08 0.34 ± 0.01	0.21 ± 0.02 0.45 ± 0.05 0.30 ± 0.03	0.24 ± 0.01 $0.37 \pm 0.05 \downarrow$ 0.43 ± 0.05	D,A,I
leucine + isoleucine	20	0.27 ± 0.04 0.19 ± 0.02 0.17 ± 0.02	0.29 ± 0.04 0.22 ± 0.01 0.18 ± 0.02	$\begin{array}{c} 0.22 \pm 0.02 \\ 0.23 \pm 0.01 \\ 0.17 \pm 0.03 \end{array}$	$0.19 \pm 0.01 \downarrow \\ 0.15 \pm 0.01 \\ 0.28 \pm 0.05 \uparrow$	1
Sulphur-containing am	ino aci	ds				
cysteine	20	0.007 ± 0.001 0.009 ± 0.001 0.003 ± 0.005	0.010 ± 0.001 0.020 ± 0.004 0.008 ± 0.001	0.009 ± 0.001↑ 0.017 ± 0.004↑ 0.008 ± 0.001↑	0.015 ± 0.002	
methionine	20	0.04 ± 0.005 0.05 ± 0.007 0.03 ± 0.005	0.06 ± 0.005 0.05 ± 0.010 0.04 ± 0.010	0.05 ± 0.009 0.06 ± 0.010 0.03 ± 0.004	0.03 ± 0.007 0.06 ± 0.012 0.06 ± 0.017	
taurine	20	0.13 ± 0.01 0.77 ± 0.04 0.24 ± 0.03	0.26 ± 0.03 0.77 ± 0.04 0.35 ± 0.06	$\begin{array}{c} 0.14 \pm 0.01 \\ 0.95 \pm 0.08 \\ 0.34 \pm 0.05 \end{array}$	0.17 ± 0.02 0.64 ± 0.11 0.48 ± 0.09	A,I
Aromatic amino acids						
tyrosine	20	0.12 ± 0.01 0.36 ± 0.03 0.06 ± 0.01	0.15 ± 0.01 0.61 ± 0.07 0.12 ± 0.01	0.13 ± 0.01 0.54 ± 0.04 0.10 ± 0.01	$0.13 \pm 0.01 \downarrow \\ 0.43 \pm 0.03 \downarrow \\ 0.10 \pm 0.01 \downarrow$	A,I
tryptophan	20	0.12 ± 0.01 0.15 ± 0.02 0.18 ± 0.02	0.15 ± 0.01 0.19 ± 0.01 0.20 ± 0.01	0.10 ± 0.01 0.21 ± 0.02 0.22 ± 0.04	0.10 ± 0.01 0.16 ± 0.01 0.30 ± 0.02↑	A,I
phenylalanine	20	0.08 ± 0.01 0.06 ± 0.01 0.08 ± 0.01	0.08 ± 0.01 0.17 ± 0.01 0.10 ± 0.02	0.10 ± 0.02 0.20 ± 0.05 0.13 ± 0.03	0.12 ± 0.02 0.12 ± 0.05 0.15 ± 0.05	

Amino-acid concentrations are expressed in mM. Statistical significance (ANOVA) of the comparisons: for diet: D; for age: A; for force-feeding: G, for interaction among the comparisons done: I. The symbols \downarrow and \uparrow represent that the values were lower or higher, respectively, in glutamine than in saline force-fed pups.

were the serine, threonine, hydroxyproline, arginine, citrulline, valine, leucine + isoleucine and tryptophan concentrations in the plasma of animals fed the reference diet. Asparagine + aspartate and histidine changed with age depending on force-feeding in rats fed the reference diet. Glutamine + glutamate and lysine showed a developmental patterns similar for the several diets and solutions supplied.

DISCUSSION

In a previous report (Salvadó and Arola, 1992b) it could be demonstrated that glutamine force-feeding enhanced urea production and liver adenylate deaminase activity and inhibited liver glutamine synthetase and serine dehydratase activities in chowfed pups. Minor effects of glutamine force-feeding were shown in the already modified activities of nitrogen metabolism enzymes in the livers of cafeteria-fed rats (Salvadó and Arola, 1992b).

The amino-acid percentage composition of the diet selected by cafeteria rats was practically identical to that of the reference diet (Rafecas et al, 1993). However, cafeteria-diet-fed rats had a higher dietary protein digestion/absorption efficiency than reference-diet-fed animals (Rafecas et al, 1993). This is probably due to changes induced by diet in the intestinal amino-acid transport systems, and is in agreement with the modulation of intestinal amino-acid transport by energy availability (Karasov et al, 1987) and physiological conditions (Israel et al, 1968; Schedl, 1974). Further, the lower proportion of protein (not amount) in the cafeteria diet could explain the higher intestinal absorption of protein amino acids (Esteve et al, 1993).

In general, the changes in plasma aminoacid concentrations caused by glutamine force-feeding are different when rats are eating cafeteria diet or reference diet. In rats eating the cafeteria diet, the decreases are shown in some plasma amino acids, and in rats eating the reference diet increases are shown in other amino acids. This could be explained because nutrient delivery has been shown to influence glutamine transport activity in the gut mucosa (Souba, 1993). On the other hand, the increased quantities of circulating ketone bodies in cafeteria-fed pups (Salvadó and Arola, 1992b) may have a suppressive effect on the oxidation of glutamine in the intestine (Nagy and Kretchmer, 1988).

A dose-related increase in human serum concentrations was reported for glutamine as well as for alanine, citrulline, and arginine (which are considered as the end products of glutamine metabolism) after oral glutamine administration (Ziegler et al, 1990). The results obtained in the present study disagree with this, the only agreement being the higher levels of citrulline at 20 d in cafeteria-diet-fed rats. Conversely, proline another end product of intestinal glutamine metabolism - is decreased by glutamine force-feeding in cafeteria-diet-fed pups. Further, alanine and arginine are lower at d 14 in the same rats. The decreased concentrations shown by some amino acids only at 14 d could be related to the fact that rat pups begin to nibble solid food at d 14 (Babicky et al, 1973). However a posterior adaptation to glutamine force-feeding is shown on the following studied days. It is also conceivable that the provision of glutamine to the intestine by the oral route may decrease the uptake of glutamine from the circulation (Moundras et al, 1993,). A lowered sulphur intake has been described in rats fed the cafeteria diet (Fernandezlopez et al, 1993). However, we have found higher plasma cysteine levels in cafeteria-fed rats and a similar effect is caused by glutamine supply. In the latter, the results could be related with the described high cysteine levels related to decreased body weight (Hoffer, 1990), but we have no explanation

for the results of cafeteria-fed rats when saline solution is added. Threonine is one of the amino acids that is more absorbed in the cafeteria-diet-fed than in reference-dietfed rats (Esteve et al, 1993). Addition of excess threonine to the diet is reflected in elevated plasma levels (Patten, 1988). Thus, the increased availability of threonine observed in rats fed the cafeteria diet is the result of an increased dietary intake. The higher threonine levels in rats force-fed glutamine is the result of the sustained depression of liver serine (threonine) dehydratase activity seen in pups force-fed this glutamine solution (Salvadó and Arola, 1992b) and is in agreement with the non-concentrative transport of threonine to liver cells (Fafournoux et al, 1990). Many changes with age are due to a peak on d 20, this day being considered the peak of lactation (Arola et al, 1982) and the values found here, for example, in tyrosine, are related to the values found in the literature (Remesar et al. 1980).

When an essential amino-acid mixture was added to the cafeteria diet, an additive effect was shown, pups grew even more (Salvadó and Arola, 1992a), less nitrogen was excreted (Salvadó and Arola, 1992a), and amino acids were more available (Salvadó and Arola, 1993) than when the cafeteria diet is administered alone. Conversely, higher nitrogen excretion was shown in chow-fed pups when they were force-fed glutamine (Salvadó and Arola, 1992b), but these changes were not observed in the deeply modified nitrogen metabolism by the cafeteria diet (Salvadó and Arola, 1992b). The present changes observed in plasma amino-acid concentrations by glutamine supplying are dependent of the diet eaten by the pups; there are differences in handling of glutamine supplied by cafeteria-diet-fed or reference-diet-fed rats and a major effect of diet over force-feeding is postulated. However, the body-weight of pups are decreased in both diets when glutamine solution is supplied, which reinforces the

statement that the mechanisms underlying the development of obesity did not run in parallel to those affecting the control of amino-acid utilization (Rafecas et al, 1993). On the other hand, the differences in nitrogen management caused by both diet and force-feeding may be explained by a differential control of intestinal absorption of nutrients caused by their different proportions. These changes in enteral metabolism will be translated to hepatic and overall metabolism.

ACKNOWLEDGMENT

Thanks are given to R Rycroft for his assistance in the edition of manuscript.

REFERENCES

- Arola L, Herrera E, Alemany M (1977) A new method for deproteinization of small samples of blood plasma for amino-acid determination. *Analyt Biochem* 82, 236-239
- Arola L, Palou A, Herrera E (1976) Determination of plasma amino acids in small samples with the use of dansylchloride. *Biochimie* 58, 1221-1226
- Arola L, Palou A, Remesar X (1982) Amino-acid enzyme activities in liver and kidney of developing rats. Arch Int Physiol Biochim 90, 163-171
- Babicky A, Parizek J, Ostadalova I (1973) Initial solid food intake and growth of young rats in nests of different sizes. *Physiol Bohemoslov* 22, 557-565
- Barber T, Viña JR, Viña J (1985) Decreased urea synthesis in cafeteria-diet-induced obesity in the rat. *Biochem J* 230, 675-681
- Barr HG, McCracken KJ (1984) High efficiency of utilization in 'cafetaria' and force-fed rats kept at 29°C. Br J Nutr 51, 279-387
- Calles-Escandon J, Cunningham J, Felig P (1984)
 The plasma amino acid response to cafeteria feeding in the rat: influence of hyperphagia,
 sucrose intake, and exercise. *Metabolism* 33,
 364-368

- Dixon WJ, Brown MB, Engelman L (1983) BMDP Statistical Software. University of California Press. Berkelev
- Esteve M, Rafecas I, Fernandezlopez JA (1993) Dietary amino-acid balances in young wistar rats fed a cafeteria diet. *Biochem Molec Biol Int* 29, 1069-1081
- Esteve M, Rafecas I, Remesar X (1992) Nitrogen balances of lean and obese Zucker rats subjected to a cafeteria diet. *Int J Obes* 16, 237-244
- Fatournoux P, Remesy C, Dermigne C (1990) Fluxes and membrane transport of amino acids in rat liver under different protein diets. Am J Physiol 259, E614-E625
- Fernandezlopez JA, Rafecas I, Esteve M (1993) Effect of genetic and dietary obesity on sulphur management by the rat. *Nutr Res* 13, 825-830
- Hoffer J (1990) Cysteine and Cystine. In: Report of the Expert Advisory Committee on Amino Acids, Minister of Supply and Services, Ontario. canada
- Israel Y, Salazar I, Rosenmann E (1968) Inhibitory effects of alcohol on intestinal amino-acid transport *in vivo* and *in vitro*. *J Nutr* 96, 499-504
- Karasov W, Solberg D, Diamond J (1987) Dependence of intestinal amino-acid uptake on dietary protein on amino-acid levels. Am J Physiol 252. G-614-G625
- Meijer GAL, Vandermeulen J, Vanvuuren AM (1993) Glutamine is a potentially limiting amino acid for milk production in dairy cows A Hypothesis. *Metabolism Clin Exp* 42, 358-364
- Mobrahan S (1992) Glutamine A conditionally essential nutrient or another nutritional puzzle. Nutr Rew 50, 331-333
- Moundras C, Remesy C, Bercovici D (1993) Effect of dietary supplementation with glutamic acid or glutamine on the splanchnic and muscle metabolism of glucogenic amino acids in the rat. J Nutr Biochem 4, 222-228
- Nagy LE, Kretchmer N (1988) Utilization of glutamine in the developing rat jejunum. J Nutr 118, 189-193
- Naim M, Brand J, Kare MR (1985) Energy intake, weight gain and fat deposition in rats fed flavored, nutritionally controlled diets in multi-

- choice (cafeteria) design. J Nutr 115, 1447-1458
- Patten BM (1988) L-Threonine and the modification of ALS. *Neurology* 38, 354
- Prats E, Monfar M, Castellà J (1989) Energy intake of rats fed a cafeteria diet. *Physiol Behav* 45, 263-272
- Rafecas I, Esteve M, Fernandezlopez JA (1993) Individual amino-acid balances in young lean and obese Zucker rats fed a cafeteria diet. *Mol Cell Biochem* 121, 45-58
- Rafecas I, Esteve M, Remesar X (1991) Plasma amino acids of lean and obese Zucker rats subjected to a cafeteria diet after weaning. *Biochem Int* 25, 797-806
- Remesar X, Arola L, Palou A (1980) Plasma amino-acid concentrations during development in the rat. *Arch Int Physiol Biochim* 88, 443-452
- Rothwell NJ, Stock MJ (1982) Effect of feeding a palatable 'cafeteria' diet on energy balance in young and adult lean (+/–) Zucker rats. *Br J Nutr* 47, 461-471
- Salvadó MJ, Segués T, Alemany M, Arola L (1986) Effects of lactation on circulating plasma metabolites in cafeteria-fed rats. Br J Nutr 55, 139-147
- Salvadó MJ, Arola L (1992a) Effects of diet and essential amino-acid gavage on young rat amino-acid metabolism enzymes. *Comp Biochem Physiol* 103A, 817-822
- Salvadó MJ, Arola L (1992b) Influence of diet and non-essential nitrogen on amino-acid metabolism enzymes of developing rats. Nutr Res 12, 955-963
- Salvadó MJ, Arola L (1993) The plasma aminoacid response to cafeteria feeding and essential amino-acid gavage in weaning rats. *Biochem Molec Biol Int* 29, 613-620
- Salvadó MJ, Segués T, Arola L (1991) Postnatal development of plasma amino acids in hyperphagic rats. Ann Nutr Metabolism 35, 242-248
- Sclafani A, Springer D (1976) Dietary obesity in adult rats: similarities to hypothalamic and human obesity syndrome. Physiology & Behavior 17, 461-471
- Schedl H (1974) Intestinal adaptation in diabetes: amino acid and carbohydrate absorption, 205-216. *In: Intestinal Adaptation* (RH Dowling, OR Piecken, eds). Schattanes Verlag, Stuttgart

- Smith R (1990) Glutamine metabolism and its physiologic importance. *J Paren Ent Nutr* 14, 40S-4S
- Souba WW (1993) Intestinal glutamine metabolism and nutrition. *J Nutr Biochem* 4, 2-9
- Ziegler T, Benfell K, Smith R (1990) Safety and metabolic effects of L-glutamine administration in humans. *J Paren Ent Nutr* 14, 137S-46S