

Class III Obesity and its Relationship with the Nutritional Status of Vitamin A in Pre- and Postoperative Gastric Bypass

Silvia Pereira · Carlos Saboya · Gabriela Chaves ·
Andréa Ramalho

Received: 21 December 2007 / Accepted: 25 February 2008 / Published online: 8 April 2008
© Springer Science + Business Media, LLC 2008

Abstract

Background Epidemiological findings have shown the rise of the prevalence of obesity in several segments of the

S. Pereira (✉)
Vitamin A Research Group,
Universidade Federal do Rio de Janeiro,
Rua Dona Mariana, 143/F11-Botafogo,
Rio de Janeiro, Rio de Janeiro CEP: 22280-020, Brazil
e-mail: se.pereira@gmail.com

C. Saboya
General Surgery, American College of Surgeons,
Chicago, IL, USA

C. Saboya
Bariatric Surgery,
Brazilian Society of Bariatric Surgery and Metabolic,
São Paulo, SP, Brazil

C. Saboya
General Surgery,
International College of Surgeons,
Chicago, IL, USA

C. Saboya
General Surgery,
Brazilian College of Surgeons,
Rio de Janeiro, RJ, Brazil
e-mail: cjsaboya@carlossaboya.com.br

G. Chaves
Vitamin A Research Group,
Universidade Federal do Rio de Janeiro,
Rio de Janeiro, Rio de Janeiro, Brazil
e-mail: gvchaves@gmail.com

A. Ramalho
Instituto de Nutrição Josué de Castro, Vitamin A Research Group,
Universidade Federal do Rio de Janeiro,
Rio de Janeiro, RJ, Brazil
e-mail: aramalho@rionet.com.br

world population, and more recent evidences point to a possible association with vitamin A deficiency (VAD). The aim of this study was to investigate vitamin A nutritional status in individuals with class III obesity in the preoperative period and 30 and 180 days after Roux-en-Y gastric bypass, correlating these findings with lipid profile and body mass index (BMI).

Methods The sample was composed of class III obese individuals, males and females, in the preoperative period (T1) and 30 (T2) and 180 days (T3) after bariatric surgery with 5,000 IU of supplementation of retinol acetate. Vitamin A nutritional status was assessed through biochemical indicators (retinol and β -carotene serum levels), which were quantified by high-performance liquid chromatography with an inadequacy cutoff of $<1.05 \mu\text{mol/L}$ and $\geq 40 \mu\text{g/dL}$, respectively.

Results Out of the 114 assessed patients, the mean age was 36.9 ± 11.6 years and BMI was $\geq 40 \text{ kg/m}^2$. The prevalence of VAD was 14%, being 37.5% in T1, 50.8% and 67.8% in T2, and 52.9% and 67% in T3, according to retinol and serum β -carotene, respectively. A decrease of retinol and β -carotene serum levels was observed with BMI increase in T1. An increase of very-low-density lipoprotein cholesterol (VLDLc) and triglycerides was verified with the increase of retinol, being VLDLc in T3 and triglycerides in T1 and T2. In T1, high-dense lipoprotein cholesterol presented a positive and significant correlation with β -carotene and a negative and significant correlation in T3.

Conclusion High prevalence of VAD in the preoperative period and in 30 and 180 postoperative days, even during supplementation, with higher inadequacy of β -carotene in all the three studied time periods, probably occurred because of its bioconversion to retinol due to the increased demand to which those individuals were exposed. It is suggested that the oral intake supplementation does not

present the expected impact, and the need of assessing the nutritional status of vitamin A in the pre- and postoperative Roux-en-Y gastric bypass is emphasized. High prevalence of VAD in T1, with severity in T2 and T3, corroborates the utilization of the cutoff of $<1.05 \mu\text{mol/L}$ as a VAD marker in the studied segment and reinforces its utilization as a tool in clinical practice to identify VAD in morbid obese patients who underwent bariatric surgery.

Keywords Vitamin A · Retinol · β -carotene · Morbid obesity · Gastric bypass

Introduction

Obesity is recognized as a public health issue, and it is characterized by the excessive accumulation of body fat in such a way that it brings damages to health [1] because it favors cardiovascular diseases (CVDs), dyslipidemias, diabetes mellitus, and some types of cancer [2]. According to *Instituto Brasileiro de Geografia e Estatística* [3], 40.6% of the Brazilian population presents overweight, with 11.1% being related to obesity.

A cross-sectional study showed that the highest rise of prevalence of obesity in the last decade was in the group of class III obesity or morbid obesity [4]. In these cases, the clinical approach is generally inefficacious, and bariatric surgery presents itself as an important option for treatment because it brings a significant improvement of comorbidities [5]. Even with this improvement and the maintenance of overweight loss in the long run, the countless metabolic alterations due to surgical procedures generate nutritional disorders. This is mainly caused by malabsorption, which is secondary to alterations in the gastrointestinal tract that both associate with the decrease in the intake and the tendency of the individual to avoid consumption of food sources when gastrointestinal disorders occur, such as dumping syndrome [6].

Deficiencies of macronutrients in these individuals frequently associate with those of micronutrients leading, on account of a combination of factors, to the development of anemias, bone demineralization, and several hypovitaminosis. This occurs because of the lack of understanding of the magnitude of metabolic abnormalities, which results in the lack of a specific recommendation of supplements for this group [6–8].

Only a few studies have reported vitamin A deficiency (VAD) after malabsorptive surgeries, and it is even harder to find reports about this deficiency in patients submitted to Roux-en-Y gastric bypass [9]. VAD is one of the most prevalent health public issues in the world, being more severe in developing countries. This deficiency may bring several damages to health, including death [10–12].

Vitamin A participates in several important functions in the human system, such as visual acuity, immunological activity, cellular proliferation, and differentiation [13]. Lately, this vitamin, as well as its precursors, has received special attention for its role against free radicals because it protects the body against oxidative stress and, consequently, it prevents damages and tissue lesions related to several nontransmissible chronic diseases [3, 14, 15].

Although literature does not present many works that assess the nutritional status of vitamin A in individuals with obesity, Viroonudomphol et al. [16], Switzer et al. [17], and Silva et al. [18] showed a negative correlation between the degree of obesity and the nutritional status of vitamin A. Despite the lack of knowledge about the clinical consequences of VAD after bariatric surgery, reports of cases have shown the occurrence of ophthalmic complications, such as night blindness and corneal xerosis [19–22].

As has been shown, many factors may justify VAD in patients submitted to surgical treatment of obesity because these individuals are more susceptible to oxidative stress, which alone much increases the intake of substances with antioxidant function [23]. Besides, surgical procedures reduce the area of the sites of absorption of vitamin A, which, when associated with the decrease in food ingestion followed by a frequent lipid restriction in the postoperative period, including foods that are sources of vitamin A, may also be a contributing risk factor for VAD in this group. The aim of this article was to assess the relationship of class III obesity with the nutritional status of vitamin A in the preoperative period and 30 and 180 days after Roux-en-Y gastric bypass, relating these findings to lipid profile and body mass index (BMI).

Materials and Methods

This study included patients attended at *Clinica Cirúrgica Carlos Saboya* in the city of Rio de Janeiro with class III obesity [24], $\text{BMI} \geq 40 \text{ kg/m}^2$, in the age range of 19–60 years in the pre- and postoperative periods of Roux-en-Y gastric bypass [25], followed up by dietitians of the multidisciplinary team of the clinic. The patient inclusion in the project was done through the signing of an informed consent. For assessment of the nutritional status of vitamin A, samples of 5 ml of blood were obtained by vein puncture from patients after a 12-h overnight fast for determining the serum concentrations of retinol and β -carotene. Analysis was performed by high-performance liquid chromatography as recommended by the World Health Organization (WHO) [26].

The obtained serum values of retinol were compared with the cutoffs for normality proposed by WHO [26], and accordingly, they were presented in interval classes of

Table 1
General characteristics of the sample

Characteristics	<i>n</i>	%
Age (years)		
19–30	36	31.5
31–50	64	56.1
51–70	14	12.4
Total	114	100
Gender		
Male	28	24.5
Female	86	75.5
Total	114	100
BMI (kg/m ²)		
40–44.9	73	64
45–49.9	27	23.7
50–54.9	8	7
55–59.9	6	5.3
Total	114	100

0.35 $\mu\text{mol/L}$. Thus, VAD was classified as severe ($<0.35 \mu\text{mol/L}$), moderate ($\geq 0.35 < 0.70 \mu\text{mol/L}$), and light deficiency ($\geq 0.70 < 1.05 \mu\text{mol/L}$). In the present study, the value of serum retinol that was considered as adequate was $\geq 1.05 \mu\text{mol/L}$, and $< 1.05 \mu\text{mol/L}$ was the cutoff utilized to indicate VAD, as suggested for investigation of marginal deficiency of vitamin A due to the increased risk of morbidity associated with the subclinical or prepathological deficiency of this vitamin [26–28]. The cutoff utilized to indicate inadequacy of β -carotene serum values was under or equal to 40 $\mu\text{g/dL}$, as suggested by Sauberlich et al. [29].

Other laboratory tests were carried out for assessment of the lipid profile. Total cholesterol and triglycerides were assessed by the enzymatic colorimetric method, whereas very-low-density lipoprotein (VLDL), low-density lipoprotein (LDL), and high-density lipoprotein (HDL) were assessed by the selective inhibition method, being classified according to the IV Brazilian Guideline on Dyslipidemia [30]. The anthropometric assessment included weight and height calculation for classification of BMI, and the adopted cutoffs were those recommended by WHO [24] for classification of eutrophy; overweight; and class I, II, and III obesity.

Statistical analyses were performed in the statistical package SPSS for Windows, version 10.0. In the sample description, data were expressed in mean \pm standard deviation for numeric variables and in percentage variables for qualitative variables. Measures of central tendency and of dispersion of the continuous variables were calculated. Comparison of the numeric variables between the groups was conducted through Mann–Whitney test. Correlations were assessed by Spearman coefficient and Pearson chi-square test. The significance level adopted was $p \geq 0.05$.

Results

The sample was composed of 114 patients, with 28 (24.5%) males and 86 (75.5%) females. Mean age of the individuals was 36.9 ± 11.6 years, ranging from 19 to 60, with 56.1% of the sample in the 31–50 age range (Table 1). There was no significant difference between the basal concentrations of serum retinol and β -carotene of the age ranges ($p=0.9$ and 0.3) and between females and males of the sample ($p=0.07$ and 0.09) (Table 1). Out of the individuals studied before the surgical procedure (T1), mean serum concentration of retinol was $1.21 \mu\text{mol/L} \pm 0.58 \mu\text{mol}$, with 14.0% below the cutoff point of $< 1.05 \mu\text{mol/L}$, thus characterizing inadequate levels of vitamin A. Dividing them into interval classes, 62.5% presented marginal VAD, 25% moderate VAD, and 12.5% severe VAD (Table 2). Assessment of serum concentrations of β -carotene showed that 37.5% of the patients presented inadequacy ($\leq 40 \mu\text{g/dL}$) and mean was $62 \pm 43.7 \mu\text{g/dL}$.

When the means of the serum concentrations of retinol and β -carotene were assessed within the first 30 postoperative days (T2), they were $1.17 \pm 0.54 \mu\text{mol/L}$ and $30.95 \pm 19.45 \mu\text{g/dL}$, respectively, and 50.8% of the patients showed VAD and 67.8% showed deficiency of β -carotene. In T3, the means of the serum concentrations of retinol and β -carotene were $1.23 \pm 0.49 \mu\text{mol/L}$ and $36.02 \pm 23.50 \mu\text{g/dL}$, and 52.9% presented inadequate levels of retinol, whereas 67.0% presented inadequate levels of β -carotene (Table 2). In T2, dividing into interval classes, 39.0% showed

Table 2 Means of retinol in the three times analyzed according to interval classification

Times	Severe VAD $< 0.35 \mu\text{mol/L}$		Moderate VAD $0.35\text{--}0.69 \mu\text{mol/L}$		Light VAD $0.70\text{--}1.05 \mu\text{mol/L}$		Adequate serum retinol $\geq 1.05 \mu\text{mol/L}$		Total	
	<i>n</i>	%	<i>n</i>	%	<i>n</i>	%	<i>n</i>	%	<i>n</i>	%
	T1	2	1.7	4	3.5	10	8.8	98	86	114
T2	5	4.4	11	9.6	39	34.3	59	51.7	114	100
T3	0	0	12	10.5	32	28.1	70	61.4	114	100

T1, preoperative; T2, 30 postoperative days; T3, 180 postoperative days

Table 3 Correlation between BMI and serum concentrations of retinol and β -carotene

Variable	Serum retinol ($\mu\text{mol/L}$)			β -carotene ($\mu\text{g/dL}$)		
	<i>n</i>	r_s	<i>p</i> value	<i>n</i>	r_s	<i>p</i> value
BMI T1	114	-0.21	0.02 ^a	114	-0.22	0.01 ^a
BMI T2	114	0.22	0.81	114	0.56	0.55
BMI T3	114	-0.10	0.27	114	-0.06	0.48

r_s = Spearman correlation coefficient

^a Statistical significance

marginal, 11.0% moderate, and 5.0% severe deficiency, whereas in T3 (Table 2), 52.9% showed marginal deficiency and 36.4% moderate deficiency.

BMI variable presented a negative correlation coefficient with statistical significance only in T1, as in retinol and β -carotene. In the other two studied periods of time, no significant correlation was observed between these variables (Table 3). VLDL variables and triglycerides presented a positive correlation coefficient with statistical significance with regard to retinol, being VLDL in T1 and T3 and triglycerides in T1 and T2. With regard to β -carotene, the HDL variable presented a significant positive correlation coefficient in T1 and a significant negative correlation in T3. Triglycerides also presented a significant negative correlation in T3.

Discussion

In view of the rise of malnutrition in the last decades, obesity is considered one of the main issues of public health, especially because of the risk of associated diseases, such as arterial hypertension, CVDs, and type 2 diabetes, among others. Nowadays, the approximate rate of mortality due to obesity is around 200,000 people annually, which is one of the highest indices of mortality in the world, with most of the deaths caused by comorbidities [2, 4, 5]. Besides this, it is assumed that in the year 2020, two-thirds of the overall number of diseases will be attributed to nontransmissible chronic diseases [1, 3, 4].

A higher percentage of obesity was observed in females (75.5%) in the present study. In Brazil, the prevalence of obesity has increased much in the last decade [31], especially in adult females, reaching 13.3% and causing a huge impact on public health. When the rate of obesity rise was assessed in Brazil, the results were 0.36% for the female population and 0.20% for the male population annually. It is important to point out that data showed that the prevalence of obesity occurred in nearly all social groups, in urban and in rural populations, and in both genders, affecting 6.9% of males and 12.5% of females [3].

In the present study, high means of BMI were found in the young population. When the weight history of women belonging to class III obesity was compared with that of women from classes I and II, Crerand et al. [32] observed that women in class III obesity had a longer history of obesity, which had generally begun in their adolescence; hence, they maintained a higher body weight along life than those belonging to obesity classes I and II. Studies have shown that a longer period of obesity may bring a higher risk for development of chronic diseases, such as CVDs, due to the longer time of exposure to risk factors [33].

In obesity, besides BMI and the time of exposure, other factors, such as distribution of body fat mainly in the abdominal area (android obesity), keep a relationship with metabolic disorders that may be pointed out as a secondary cause of nutrient deficiency with antioxidant action [34]. The main etiological factor in epidemiological level related to VAD is the inadequate intake of food sources to satisfy the physiological needs of the individual [35]. Although the present study did not assess the dietary intake of the patients, evidences demonstrated that obesity, defined as the positive balance between what is ingested and what is spent [36, 37], is directly related to the quantity and not to the quality of what is consumed; that is, it is associated with a rich intake of calories and a poor intake of micronutrients [38, 39].

Such fact reinforces the thesis that the increase of total intake of foods does not necessarily increase the consumption of foods that are sources of vitamin A, which suggests that, in obesity, there is an insufficient intake of the sources of this vitamin that may lead to a deficiency in the liver store if it remains for a long period, thus increasing the risk for development of metabolic disorders [13].

Studies showed lower serum concentrations of retinol and carotenoids in obese individuals when compared with individuals with normal weight with no significant difference as regards the dietary ingestion of sources of these nutrients, as it was shown by data obtained by an inquiry of food history [18, 31, 40]. Therefore, the higher serum inadequacy of antioxidant nutrients observed in the present study may still be derived from their higher metabolic utilization against oxidative stress because it is known that these individuals are more exposed to it than eutrophics [41]. Studies with obese individuals presented a negative correlation between retinol, β -carotene, and BMI [16, 18], and in the present study, a statistical significance was observed when the means of BMI in T1 were compared with the levels of retinol and β -carotene.

The prevalence of inadequate serum levels of retinol found in the present study was 14.0%, and the inadequacy of β -carotene was 37.5%. In adults, a significantly higher risk for CVD in individuals with decreased serum concentrations of β -carotene was reported [46]. Low concentrations of

circulating antioxidants may contribute to an increased risk and to severity of atherosclerotic disease. In Brazil, information on the prevalence of VAD in individuals with disease profiles similar to the present study is not available; nonetheless, the VAD found in the present study, even being lower than that described for the classic groups of risk in Brazil in the preoperative period (14.0%), was high and rose in the periods of 30 and 180 postoperative days, 50.8% and 52.9%, respectively [42].

The results presented here are alarming and point to the need of conducting more studies addressed to this population group. As has been corroborated by literature, in the present study, a predominance of inadequacy of β -carotene was found, which is considered the most powerful free radical sweeper, hence suggesting that its largest part was converted into retinol. This finding is similar to that reported by Mecocci et al. [43], who demonstrated that the adequate nutritional status of vitamin A decreased the conversion of carotenoids into retinol, therefore evidencing the existence of a relationship between the nutritional status of retinol and carotenoids and that β -carotene is recognized as the most potent precursor of retinol [44].

Although there was significant weight loss with regard to T1, suggesting improvement of comorbidities and reduction of oxidative stress, the surgical procedure to which the patient was submitted consisted in the reduction of the gastric capacity and malabsorption on account of exclusion of the duodenum and of 30–50 cm of jejunum of food transport, besides 1 to 2 m of proximal jejunum without the presence of digestive enzymes [45].

It is worth pointing out that diets administered on the 30 initial days after surgery were aimed at gastric rest and hydration to reduce risks such as fistules, usually offering an intake of 300 to 500 kcal daily, which is recognizably deficient in macro- and micronutrients [46]. The literature is consensual with regard to vitamin and mineral supplementation in the postoperative period of bariatric surgery, but the actual vitamin recommended daily intake (RDI) that would indicate the best route of administration and the form of presentation of the nutrients has not been established yet [8, 47].

Usually, the polyvitamins that are available in the market and are utilized by the teams of medical doctors and professionals responsible for bariatric surgery offer a dose of 5,000 IU of retinol acetate, which corresponds to twofold RDI for a normal individual [48]. Nonetheless, in the same way that vitamin A and carotenoids are solubilized and absorbed in the lean intestine, the dietary supplement also has its absorption damaged because it is absorbed by passive diffusion. In the intestinal tract, retinol and carotenoids are incorporated into chylomicrons, which leave the intestine and migrate to the liver, the main organ of synthesis and storage [49]. It is calculated that the

adequate store of vitamin A might promote protection against the deficiency for a 4-week period in adult individuals with a deficient diet [13].

In the sixth month of the postoperative period (T3), even with a mean intake of 1,200 kcal and vitamin and mineral supplement offering 5,000 IU of retinol acetate, the prevalence of retinol and deficiency of β -carotene were 52.89% and 67.0%, respectively, maintaining themselves equivalent to those found on postoperative day 30, which suggests that the oral intake did not seem to have had the expected impact.

Low intake of proteins, lipids, iron, and zinc in the diet may contribute to the development of VAD on account of the interference in the absorption and transport of this micronutrient [50]. Another relevant aspect is that polyvitamins contain vitamin C, vitamin E, selenium, and zinc, which are nutrients with antioxidant action [48, 51], and the prevalence of VAD was kept high and rising even considering the time of exposure to the supplement and the expected increase in the intake of nutrients in the sixth postoperative month. When assessing the impact of surgery on BMI, it was verified that there was a significant decrease in all three studied periods of time.

Many studies showed that bariatric surgery, due to the intense and maintained loss of overweight, effectively controls comorbidities [52, 53], which may be considered secondary causes of VAD because they interfere in the absorption, storage, and transport of vitamin A or in the conversion of pro-vitamin A into its metabolically active form [34]. Dyslipidemia, which relates to cardiopathies, was found among the secondary causes of VAD, being hypertriglyceridemia recognized as a marker of increased risk for CVDs [54].

Among its numberless functions, HDL protects LDL from lipid oxidation [55] and obese individuals move swiftly to dyslipidemias, which results in an increase of oxidative stress. Serban et al. [56] suggested that individuals with dyslipidemia presented higher levels of lipid peroxidation. With the aim of assessing the hypothesis that low concentrations of HDL interfered in the endothelial function and in lipoprotein oxidation, Toikka et al. [57] found significantly lower levels of LDL oxidized in the group with high levels of HDL when compared with low levels of HDL.

Experimental studies with genetically obese rats evidenced that serum levels of triglycerides and total cholesterol were significantly higher in animals with VAD, and a significant decrease was found in the levels of triglycerides with the supplementation of this vitamin but not in total cholesterol. The explanation given to such a finding was that, in obese individuals with VAD, an increase in the synthesis of free fatty acids and triglycerides occurs on account of the lower antioxidant protection [58].

The present study did not corroborate literature because a positive and significant correlation between retinol and triglycerides was found in T1 and T2; however, a study with diabetic children showed similar results, which the authors justified stating that, due to hypertriglyceridemia, there had been an increase in the serum concentrations of retinol as a factor of atherogenic protection [59].

With regard to β -carotene, a significant and positive correlation with HDL in T1 was found, which suggests that it exerted its protective function against LDL oxidation. This antioxidant acts as an important protector against the oxidative attack of LDL and has also been associated with the increase of HDL, as in vitro as in vivo [55–57]. Nonetheless, triglycerides, as well HDL, presented a significant and negative correlation in T3, which suggests that an increase in oxidative stress may occur in view of the possible increase in calorie intake and of the possible return to food habits and reduction in the speed of weight loss.

Thus, we can conclude that patients with class III obesity showed high percentage of VAD, which was progressive when BMI increased, a fact that may be attributed to the increase of oxidative stress on account of the chronic inflammatory condition presented by patients with obesity. The prevalence of VAD after the first 30 postoperative days increased considerably and may be related to the restriction in the intake of food sources of vitamin A and of lipids, associated with malabsorption, with the exclusion of the main site of absorption, even with a daily oral supplementation of 5,000 IU of retinol acetate. After 180 postoperative days, the progressive increase of prevalence of VAD remained even in view of significant weight loss, with a probable improvement of comorbidities and permanence of oral supplementation of 5,000 IU of retinol acetate and a possible dietary increase, which suggests that the oral intake, either of supplements or dietary, did not present the expected impact.

References

- Halpen A. Fisiologia da Obesidade (Physiology of Obesity). In: Garrido Júnior AB, Ferraz EM, Barroso FL, et al., editors. *Cirurgia da obesidade (Obesity Surgery)*. São Paulo: Atheneu; 2003. p. 9–12.
- Bender R, Zeeb H, et al. Causes of death in obesity: relevant increase in cardiovascular but not in all-cancer mortality. *J Clin Epidemiol*. 2006;59:1064–71.
- Fundação Instituto Brasileiro de Geografia e Estatística (IBGE). *Pesquisa de orçamentos familiares 2002–2003: Análise da disponibilidade domiciliar de alimentos e do estado nutricional no Brasil (Research of family budgets: 2002–2003: Analysis of home availability of foods and of the nutritional status in Brazil)*. Rio de Janeiro: Ministério do Planejamento e Orçamento/IBGE; 2004.
- Freedman DS, Khan LK, Serdula MK, et al. Trends and correlates of class 3 obesity in the United States from 1990 through 2000. *JAMA*. 2002;188:1758–61.
- Coutinho WF, Benchimol AK. Obesidade mórbida e afecções associadas (Morbid obesity and associated infections). In: Garrido-Júnior AB, Ferraz EM, Barroso FL, et al., editors. *Cirurgia da obesidade (Obesity Surgery)*. São Paulo: Ed Atheneu; 2003. p. 13–7.
- Bloomberg RD, Fleishman A, et al. Nutritional deficiencies following bariatric surgery: what have we learned? *Obes Surg*. 2005;2:145–54.
- Brolin RE, Leung M. Survey of vitamin and mineral supplementation after gastric bypass and biliopancreatic diversion for morbid obesity. *Obes Surg*. 1999;2:150–4.
- Malinowski SS. Nutrition and metabolic complications of bariatric surgery. *Am J Med Sci*. 2006;4:219–25.
- Slater GH, Ren CJ, Siegel N, et al. Serum fat-soluble vitamin deficiency and abnormal calcium metabolism after malabsorptive bariatric surgery. *J Gastrointest Surg*. 2004;1:48–55.
- Homer MR, Dorea JG, Pereira MG, et al. Inquérito dietético com base no consumo familiar: o caso de Ilhéus, Bahia, Brasil, em 1979 (Dietary inquiry with basis on family consumption: the case of Ilhéus, Bahia, Brazil, in 1979). *Arch Latinoam Nutr*. 1981; 31:726–39.
- Santos LMP, Ascutti LS, Dricot DC. Exophthalmia in the state of Paraíba, northeast of Brazil clinical findings. *Am J Clin Nutr*. 1983;38:139–44.
- United Nations Children's Fund. *Vitamin A and mineral deficiency: a global assessment*. New York; 2004.
- Institute of Medicine. *Vitamin A*. In: *Dietary reference intakes for vitamin A, vitamin K, arsenic, boron, chromium, copper, iodine, iron, manganese, molybdenum, nickel, silicon, vanadium, and zinc*. Washington, DC: National Academy Press; 2001. p. 82–161.
- Singh RB, Niaz MA, Bishnoi I, et al. Diet, antioxidant vitamins, oxidative stress and risk of coronary artery disease: the Peerzada prospective study. *Int J Cardiol*. 1994;5:453–67.
- Mosca L, Rubenfire M, Mandel C, et al. Antioxidant nutrient supplementation reduces the susceptibility of low density lipoprotein to oxidation in patients with coronary artery disease. *J Am Coll Cardiol*. 1997;30:392–9.
- Viroonudomphol D, Pongpaew P, Tungtrongchitr R. The relationships between anthropometric measurements, serum vitamin A and E concentrations and lipid profiles in overweight and obese subjects. *Asia Pac J Clin Nutr*. 2003;1:73–9.
- Switzer BR, Atwood JR, Stark AH, et al. Plasma carotenoid and vitamins A and E concentrations in older African American women after wheat bran supplementation: effects of age, body mass and smoking history. *J Am Coll Nutr*. 2005;3:217–26.
- Silva LSVL, Veiga GVV, Ramalho RA. Association of serum concentrations of retinol and carotenoids with overweight in children and adolescents. *Nutrition*. 2007;23:392–7.
- Quaranta L, Nascimbeni G, Semenario F, et al. Severe corneoconjunctival xerosis after biliopancreatic bypass for obesity (Scopinaro's operation). *Am J Ophthalmol*. 1994;118:817–8.
- Huerta S, Rogers LM, Li Z, et al. Vitamin A deficiency in a newborn resulting from maternal hypovitaminosis A after biliopancreatic diversion for the treatment of morbid obesity. *Am J Clin Nutr*. 2002;76:426–9.
- Hatzifotis M, Dolan K, Newbury L, Fielding G. Symptomatic vitamin A deficiency following biliopancreatic diversion. *Obes Surg*. 2003;4:655–7.
- Chae T, Foroozan R. Vitamin A deficiency in patients with a remote history of intestinal surgery. *Br J Ophthalmol*. 2006; 90:955–6.
- Sarni RO, Ramalho RA. Serum retinol and total carotene concentrations in obese children. *Med Sci Monit*. 2005;11:510–4.

24. WHO. Obesity: preventing and managing the global epidemic. Report of WHO consultation on obesity. Geneva: World Health Organization; 1998.
25. NIH Consensus Development Conference Draft Statement. Gastrointestinal surgery for severe obesity. *Obes Surg.* 1991;1:257–65.
26. WHO. Global prevalence of vitamin A. Indicators for assessing vitamin A deficiency and their application in monitoring and evaluating intervention programmers. Micronutrient Series, WHO/NUT. 10. Geneva: WHO; 1996.
27. Sommer A. La carencia de vitamina A y sus consecuencias. Guía práctica para la detección y el tratamiento Vitamin A deficiency and its consequences. Practical guide for detection and treatment). Geneva: OMS; 1995.
28. International Vitamin A Consultative Group (IVACG). IVACG Statement. Maternal night blindness: a new indicator of vitamin A deficiency. Washington, DC: IVACG; 2002.
29. Sauberlich HE, Hodges RE, Wallace DL, et al. Vitamin A metabolism and requirements in the human studied with the use of labeled retinol. *Vitam Horm.* 1974;32:251–75.
30. IV Diretriz Brasileira Sobre Dislipidemias e Prevenção da Aterosclerose (IV Brazilian Guideline on Dyslipidemias and Prevention of Atherosclerosis). *Arq Bras Cardiol.* 2007;88:s.1.
31. Ford ES, Giles WH, Mokdad AH. Increasing prevalence of the metabolic syndrome among US adults. *Diabetes Care.* 2004;10:2444–9.
32. Crerand CE, Wadden TA, et al. A comparison of weight histories in women with class III vs. class I–II obesity. *Surg Obes Relat Dis.* 2006;2:165–70.
33. Must A, Jacques PF, Dallal GE, et al. Long-term morbidity and mortality of overweight adolescents: a follow-up of the Harvard Growth Study 1922 to 1935. *N Engl J Med.* 1992; 327:1350–5.
34. Queiróz E, Ramalho RA, et al. Vitamin A status in diabetic children. *Diabetes Nutr Metab.* 2000;4:125–7.
35. West CE. Meeting requirements for vitamin A. *Nutr Rev.* 2000;58:341–5.
36. Baltasar A. Definición: ¿Qué es la obesidad grave? (What is severe obesity?). In: *Obesidad y Cirugía - como dejar de ser obeso (Obesity and Surgery—how not to be obese any longer)*. 1st ed. Madrid: Editora Arán; 2000. p. 15–21.
37. Polacow VO, Lancha AH Jr. Dietas hiperglicídicas: efeitos da substituição isoenergética de gordura por carboidratos sobre o metabolismo de lipídios, adiposidade corporal e sua associação com atividade física e com o risco de doença cardiovascular (Hyperglycemic diets: effects of isoenergetic substitutions of fat by carbohydrates on lipid metabolism, body adiposity and their association with physical activity and with the risk for cardiovascular disease). *Arq Bras Endocrinol Metabol.* 2007;51: 389–400.
38. Sorensen TI. The changing lifestyle in the world: body weight and what else? *Diabetes Care.* 2000;2:1–4.
39. Butte NF. The role of breastfeeding in obesity. *Pediatr Clin North Am.* 2001;1:189–98.
40. Sarni RS, Kochi C, et al. Impact of vitamin A megadose supplementation on the anthropometry of children and adolescents with non-hormonal statural deficit: a double-blind and randomized clinical study. *Int J Vitam Nutr Res.* 2003;73:303–11.
41. Matsuoka H. Endothelial dysfunction associated with oxidative stress in human. *Diabetes Res Clin Pract.* 2001;54:S65–72.
42. Ramalho RA, Flores H, Accioly E, et al. Deficiência de vitamina A no Brasil (Deficiency of vitamin A in Brazil). *Soc Iber Nutr.* 2005.
43. Mecocci P, Polidori MC, Troiano L, et al. Plasma antioxidants and longevity: a study on healthy centenarians. *Free Radic Biol Med.* 2000;28:1243–8.
44. Rodriguez-Amaya DB. Food carotenoids: analysis, composition and alterations during storage and processing of foods. *Forum Nutr.* 2003;56:35–7.
45. Higa KD, Boone KD, Ho T. Laparoscopic Roux en Y gastric bypass for morbid obesity: technique and preliminary results of our first 400 patients. *Arch Surg.* 2000;135:1029.
46. Cruz M, Marimoto IMI. Intervenção nutricional no tratamento cirúrgico da obesidade mórbida: resultados de um protocolo diferenciado (Nutrition intervention in the surgical treatment of morbid obesity: results of a differentiated protocol). *Rev Nutr.* 2004;17:263–72.
47. Rhode B, McLean LD. Vitamin and mineral supplementation after gastric bypass. In: Deitel M, editor. (Chapter 19 Update) *Surgery for the morbidly obese patient*. Toronto: F-D Communications; 2000.
48. Cambi MPC. Avaliação nutricional e de qualidade de vida em pacientes submetidos à cirurgia bariátrica (Nutritional and life quality assessment of patients submitted to bariatric surgery). Master thesis of the Post-Graduation. Program - UFSC: Florianópolis; 2001.
49. Blomhoff R. Transport and metabolism of vitamin A. *Nutr Rev.* 1994;52:13–23.
50. Christian P, West JR, et al. Maternal night blindness increases risk of mortality in the first 6 months of life among infants in Nepal. *J Nutr.* 2001;131:1510–2.
51. Zemora S, Diego J. Antioxidantes. Micronutrientes en lucha por la salud (Antioxidants. Micronutrients in fight for health). *Rev Chil Nutr.* 2007;34:17–26.
52. Karlsson J, Sjöström L, Sullivan M. Two-year follow up of health-related quality of life (HRQL) and eating behavior after gastric surgery for severe obesity: Swedish Obese Subjects (SOS)—an intervention study of obesity. *Int J Obes Relat Metab Disord.* 2005;22:113–26.
53. Carvalho OS, Moreira CLCB, et al. Cirurgia bariátrica cura síndrome metabólica? (Does bariatric surgery cure metabolic syndrome?). *Arq Bras Endocrinol Metabol.* 2007;51:78–85.
54. National Cholesterol Education Program (NCEP) Expert Panel on Detection, Evaluation, and Treatment of High Blood Cholesterol in Adults (Adult Treatment Panel III). *National Cholesterol Education Program National Heart, Lung, and Blood Institute*. Bethesda: National Institutes of Health NIH Publication; 2002.
55. Leborgne L, Maziere JC, Maziere C. Oxidative stress, atherogenesis and cardiovascular risk factors. *Arch Mal Coeur Vaiss.* 2002;9:805–14.
56. Serban MG, Negru T. Lipoproteins, lipidic peroxidation and total antioxidant capacity in serum of aged subjects suffering from hyperglycemia. *Rom J Intern Med.* 1998;36:65–70.
57. Toikka JO, Ahotupa M, Viikari JS, et al. Constantly low HDL-cholesterol concentration relates to endothelial dysfunction and increased in vivo LDL-oxidation in healthy young men. *Atherosclerosis.* 1999;147:133–8.
58. Baena RM, Campoy C, et al. Vitamin A, retinol binding protein and lipids in type 1 diabetes mellitus. *Eur J Clin Nutr.* 2002;56:44–50.
59. Santos HS, Cruz WM. A terapia nutricional com vitaminas oxidantes e o tratamento quimioterápico oncológico (Nutritional therapy with oxidant vitamins and the oncological chemotherapy treatment). *Rev Bras Cancerol.* 2001;3:303–8.