

Review Article

Potential Hematology and Nutritional Complications of Bariatric Surgery

Al-Jafar H^{1*}, Al-Zamil K², Al Ageeli M³, Alhaifi M⁴ and Al-Sabah S⁴

¹Department of Hematology, Amiri Hospital, Kuwait

²Department of General Surgery, Amiri Hospital, Kuwait

³Department of Clinical Biochemistry, Amiri Hospital, Kuwait

⁴Department of General Surgery & Bariatric Surgery, Amiri Hospital, Kuwait

*Corresponding author: Hassan Al-Jafar, Consultant Hematologist, Department of Hematology, Amiri Hospital, Kuwait

Received: June 08, 2018; Accepted: July 02, 2018;

Published: July 11, 2018

Abstract

Bariatrics is a specialized field of medicine that deals with the etiology, prevention and management of patients with morbid obesity. Surgical and nutritional complications are two adverse effects associated with bariatric surgery. For patients in recovery, post-bariatric surgical treatment and management often involves the care of a hematologist. Long term nutritional problems may develop following bariatric surgery and should be considered by physicians, although long-term nutritional complications are often rare or just temporary. Multivitamin and iron supplementation are required as part of a patient's prophylactic post-bariatric surgical management protocol; therefore, the patient's adherence to vitamin supplementation should be checked during follow up appointments. This review aims to emphasize the potential multi systemic hematological and nutritional complications of bariatric surgery. Vitamin deficiencies such as iron and vitamin B12 sometimes affect patients and therefore physicians should be aware of this in the treatment of patients with a history bariatric surgery. Inquiring about a patient's history of bariatric surgery needs to be one of the first questions to ask a patient who is presenting with vitamin deficiencies as this can cause multi system abnormalities. Many patients are lost to follow up after bariatric surgery and come back with complications. Therefore, the importance of maintaining follow up appointments should be emphasized.

Keywords: Bariatric; Hematology; Nutrition; Vitamins

Introduction

The term "bariatric" was used for the first time on 1965 [1]. Bariatrics in medicine deals with the etiology, prevention and management of morbid obesity. Body Mass Index (BMI) is derived from a patient's weight and height to assess the degree of obesity [2]. The BMI is widely applicable in various clinical fields as it is necessary to assess the dose of many drugs [3]. Bariatric surgery is currently the most used and effective measure to treat patients with morbid obesity and to help reduce metabolic disorders such as Diabetes mellitus, dyslipidemia and hypertension [4]. The U.S. National Institutes of Health recommends bariatric surgery for obese people with a BMI of 40 kg/m² [5]. Indication for bariatric surgery could be appropriate for those individuals with a BMI of 35-40 kg/m² without comorbidities or a BMI of 30-35 kg/m² with significant comorbidities [6]. Obesity is associated with severe fat deposition in many organs that may cause serious, adverse effects on health [7] and individuals with BMIs exceeding the healthy range have a much greater risk for medical issues [8]. Usually, exercise, diet and behavior therapy should be the first-line of treatment for individuals with obesity [9], as medical therapy for severe obesity has limited short and long-term success [10]. Bariatric surgery often leads to improvements in a patient's quality of life and obesity-related diseases [11]; moreover, improvements in psychological health have also been observed [12]. Major complications due to bariatric surgery have been reported in 3.3% of patients [13]. Surgical complications may occur as adverse events early in the post-operative period, especially in those excessively overweight [14]. Nutritional complications are considered late side effects and require attention by the treating surgeon.

These complications are often associated with gastrointestinal malabsorption which needs to be addressed to prevent chronic health conditions such as neurological complications associated with long-standing severe vitamin B₁₂ deficiency [15].

I-Potential Surgical Complications

Bleeding

Bleeding is an adverse outcome in many types of surgical procedures. However, the amount of bleeding can be reduced by adequate pre-operative assessment to identify and correct disorders due to other comorbidities. Surgical bleeding can arise from technical causes as well as the presence of a bleeding disorder [16]. The onset of early bleeding is usually ≤ 24 hours after the end of the surgery, the location is either intraluminal or extra luminal, and the severity of the bleeding may be either mild or severe [17]. Bleeding due to a disorder can be categorized into three groups: disorders of platelet function or number, disorders of clotting factors, or a combination of these, and thirdly, the bleeding could be due to vascular or pathological localized lesions [18]. The incidence of bleeding ranges between 0% and 4.4% and varies according to the different procedures performed. However, bleeding remains a known and limited complication in bariatric surgery [19].

Thrombosis

Deep Vein Thrombosis (DVT) is one of the serious complications of bariatric surgery in patients with morbid obesity. Patients with postoperative DVT could be asymptomatic [20] or DVT could simply manifest as chronic venous hypertension, which subsequently develops into varicose veins. Factors that could influence thrombus

Table 1: The effect of trace elements deficiency on hematology aspect.

Trace Element deficiency	Hematological manifestations
Thiamine (B ₁)	Megaloblastic anemia [39]
Riboflavin (B ₂)	normochromic normocytic anemia, megaloblastic anemia [43]
Niacin (B ₃)	Anemia [46]
Pantothenic (B ₅)	Anemia [46,51]
Pyridoxine (B ₆)	sideroblastic microcytic anemia [53].
Biotin (B ₇)	Anemia [56]
Folate (B ₉)	Macrocytic, megaloblastic anemia [46], hemorrhage [59]
Vitamin B ₁₂	Megaloblastic anemia, microcytic and hypochromic anemia. [59,64]
Vitamin A	Normocytic and normochromic anemia [70,71]
Vitamin C	Bleeding disorder and anemia [74]
Vitamin D	Anemia [76]
Vitamin E	Anemia Hemolysis of RBC [48,74,83]
Vitamin K	Bleeding disorder [84,87]
Iron	Microcytic and hypochromic anemia, mild thrombocytosis, chronic bleeding [29,91,93]
Copper	Microcytic, macrocytic and neutropenia [104], Thrombocytopenia is unusual [105]
Magnesium	Iron deficiency anemia [112]
Selenium	Anemia [115]
Chromium	Iron deficiency anemia [117]
Calcium	Iron deficiency anemia [91]
Phosphorus	Anemia [102]
Zinc	Anemia [70]

formation are alterations in blood flow, changes in the vessel wall, alterations in blood constituents and viscosity, and inhibition of the fibrinolysis system. Once DVT occurs, the fate of the thrombus depends on the persistence of factors involved in its formation. Many thrombi will spontaneously lyse or will shrink, but others may extend and embolize, posing a threat to the patient [21]. The reported incidence of DVT after bariatric surgery varies widely, from 0.2% to 1.3% at 30 days [22] and to 0.42% at 90 days [23].

Embolism

All patients who undergo surgery are at risk for Pulmonary Embolism (PE). During surgery, PE often initially manifests as hemodynamic instability [24]. PE occurs in approximately 0.3% to 1.6% of the general population that undergoes surgery [25]. The initial hemodynamic insult in PE is the obstruction of blood flow, which causes emboli in the pulmonary vasculature that disrupt pulmonary outflow, cause an acute increase in right ventricular impedance, and initiate neural reflexes and the release of pulmonary vasoconstrictors into the circulation [26]. The reported incidence of PE in patients undergoing open bariatric surgery with prophylaxis measures ranges from 0.36% to 3.0% [27].

II-Potential Nutritional Complications

A-Vitamin Deficiencies

Micronutrients, including trace elements, water, and fat-soluble vitamins can become depleted in patients during the post-bariatric surgical recovery period, and such elements are essential factors that mainly serve as enzymatic cofactors in biochemical pathways and

metabolic processes [28]. Micronutrient deficiencies vary in frequency according to the type of surgery performed [29,30]. Adverse events after bariatric surgery can lead to a wide range of symptoms, most commonly anemia (10%-74%) and neurological dysfunction (5%-9%) [31,32]. Determining the exact risk for developing micronutrient deficiencies is challenging as there is no consensus on the appropriate amount of vitamin and mineral supplementation required across bariatric surgery programs, and therefore, supplementation practices vary widely [33]. It is clear that micronutrient deficiencies are relatively common in patients before and after all types of bariatric surgery; therefore, it is important to screen patients at baseline and at least annually [34].

Thiamine (vitamin B₁) deficiency

Thiamine deficiency is common after bariatric surgery lead to the combination of a reduction in acid production by the gastric pouch, restriction of food intake, and frequent episodes of vomiting [35,36]. Thiamine deficiency accompanied by peripheral neuropathy is characterized by polyneuropathy with paresthesia of the extremities, especially the legs, reduced knee-jerk and other tendon reflexes, severe progressive weakness, muscle wasting, and increased susceptibility to infections [37]. Thiamine deficiency is associated with a condition called Beriberi syndrome, which presents as mental confusion, anorexia, muscular weakness, ataxia, peripheral paralysis, tachycardia; thiamine deficiency associated with edema is known as wet beriberi [38]. Red blood cell megaloblastic changes in thiamine deficiency could cause megaloblastic anemia [39], which is reversible and could be corrected with pharmacologic doses of thiamine

(vitamin B₁) [40] (Table 1).

Riboflavin (vitamin B₂)

Riboflavin deficiency can cause neurodegenerative disorders [41,42]. Owing to its effects on iron, mild to moderate riboflavin deficiency results in anemia despite average cell sizes and normal intra corpuscular hemoglobin content (normochromic normocytic anemia). This type of anemia caused by the combined deficiency of riboflavin (vitamin B₂) and folic acid (B₉) or cyanocobalamin (B₁₂), usually causes megaloblastic anemia [43].

Niacin (vitamin B₃) deficiency

Niacin deficiency can cause pellagra and alterations in protein metabolism, and eventually malnutrition. Symptoms include eczema, intestinal and stomach distress, depression, headache, and thinning of the hair [44,45]. Additional possible symptoms of mild vitamin B₃ deficiency include anemia, skin lesions, diarrhea, insomnia, forgetfulness, irritability, nervousness, and other psychological symptoms [46].

Pantothenic acid (vitamin B₅) deficiency

Pantothenic acid is an essential component of the coenzyme A, which is a cofactor in many enzymatic reactions crucial to the metabolism of carbohydrates, fats, corticosteroids, and sex hormones. Pantothenic acid deficiency is rare in humans and depends on the proper functioning of the adrenal glands and nervous system. Nevertheless, this vitamin is important for healthy growth and development throughout life [47,48]. Symptoms of deficiency are similar to those observed in other vitamin B deficiencies: irritability, fatigue, apathy, and hypoglycemia or increased sensitivity to insulin. In cases of pantothenic acid deficiency, more insulin will bind to receptors, thus causing hypoglycemia [49]. Additional symptoms could include restlessness, malaise, sleep disturbances, nausea, vomiting, abdominal cramps, insomnia, anemia, vomiting, muscle contractions, and abnormal skin features. Deficiency of pantothenic acid is also associated with a condition called burning feet syndrome [50,51].

Pyridoxine (vitamin B₆)

Pyridoxine acts as a coenzyme in many reactions including fatty acid and amino acid metabolism [52]. Therefore, it is important for the blood, central nervous system, and skin metabolism. Vitamin B₆ deficiency could lead to seborrheic dermatitis, glossitis, cheilosis, peripheral neuropathy, and lymphopenia [48]. Severe vitamin B₆ deficiency may cause sideroblastic microcytic anemia due to decreased hemoglobin synthesis, seizures that are refractory to conventional medications, convulsions, and peripheral neuropathy [53].

Biotin (vitamin B₇) deficiency

Biotin deficiency typically occurs due to the dietary absence of the vitamin or malabsorption but can be addressed via nutritional supplementation [54,55]. Lack of vitamin B₇ may cause symptoms of anemia [56]; brittle and thin fingernails; hair loss; conjunctivitis; dermatitis; and red rashes around the eyes, nose, mouth, and genital area. In adults, neurological defects manifest as depression, lethargy, numbness, hallucinations, and tingling of the extremities [55].

Folate (vitamin B₉) deficiency

The prevalence of folate deficiency after bariatric surgery has been

reported to be 9%-39% and is often reported in women who become pregnant after bariatric surgery [57,58]. Folate deficiency may be a consequence of vitamin B₁₂ deficiency since the latter plays a vital role in the conversion of the inactive methyltetrahydrofolic acid to active tetrahydrofolic acid [59]. The deficiency of folic acid can lead to megaloblastic anemia. Folate is essential for the prevention of neural tube defects in infants; therefore, women considering pregnancy following bariatric surgery to treat obesity should receive counseling before conception and prophylactic supplementation of folate and vitamin B₁₂ [60]. Folate deficiency can easily be corrected via oral vitamin supplementation [61].

Cobalamin (vitamin B₁₂) deficiency

Cobalamin deficiency could occur after bariatric surgery procedures that bypass the lower stomach [62,63]. Potential complications from vitamin B₁₂ deficiency include megaloblastic anemia, generalized weakness, neuropathy, and cognitive difficulties [64]. Bariatric surgery patients are at increased risk of developing vitamin B₁₂ deficiency because their digestive tracts have been altered in such way that it interferes with the natural absorption of this vitamin [65]. In healthy adults, an intrinsic factor released by the parietal cells in the stomach binds with vitamin B₁₂ in the duodenum. The bound vitamin B₁₂ is absorbed in the ileum [66]. The daily recommended oral supplementation dose is 350-600 ug per day, which has shown to correct the deficiency in 81%-95% of patients [67]. Intramuscular vitamin B₁₂ injections are another option for patients who have trouble adhering to daily oral supplements [28].

Vitamin A deficiency

Vitamin A deficiency after bariatric surgery to manage obesity can be observed in 61%-69% of patients [68,69]. Being a fat-soluble vitamin, an approximately one year's supply of vitamin A can be stored in the liver, resulting in deficiencies to manifest much later; this is also true for other fat-soluble vitamins, proteins, and zinc. Vitamin A appears to be involved in the pathogenesis of anemia through diverse biological mechanisms such as enhancement of growth, differentiation of erythrocyte progenitor cells, potentiation of immunity to infection, and reduction of iron reserves in tissues [70]. Symptoms of vitamin A deficiency include pathological manifestations affecting the eye, skin problems, mucous membranes, dry hair, broken nails, and increased risk for infections. Exophthalmia is a condition characterized by abnormal dryness of the conjunctiva and cornea of the eye with inflammation and ridge formation, which is typically associated with vitamin A deficiency. Vitamin A deficiency also may contribute to iron deficiency [71].

Vitamin C deficiency

Vitamin C or ascorbic acid is essential for the hydroxylation of proline and lysine in collagen [72,73]. Vitamin C deficiency results in scurvy, which is characterized by the degeneration of capillaries, bone, and connective tissue, leading to perifollicular petechial rash, poor wound healing, and bleeding of the gums. Early symptoms of vitamin C deficiency, occurring within 1 to 3 months, include fatigue and myalgia, weakness, anemia, gum disease, and skin hemorrhage. This condition usually occurs in malnourished adults [74]. Biochemical evidence of vitamin C deficiency is common, it reported prevalence 10%-50% after bariatric surgery [65]. Vitamin C deficiency can be successfully treated with supplementation of 500 mg once daily and

complete remission can be achieved after one month [28]. However, a significant adverse effect due to vitamin C deficiency after bariatric surgery is rare [74].

Vitamin D deficiency

Vitamin D is mainly responsible for the formation and health of red blood cells. Since vitamin D is needed for iron metabolism a deficiency of vitamin D may cause anemia [75,76]. Severe vitamin D deficiency can lead to fatigue, weakness, general muscle pain, joint pain, muscle cramps, chronic pain, restless sleep, headaches, poor concentration, bladder problems, and gastrointestinal problems [77]. Vitamin D deficiency is common following bariatric surgery and has been reported to occur in 50%-80% of patients [78]. Coates, et al. studied bone metabolism in 25 patients at 9 months following bariatric surgery and found that bone mineral density and content were significantly diminished when the patients were compared to 30 obese control patients [79]. Brolin, et al. found that 51% of post-bariatric surgery patients had significant vitamin D deficiencies at the 2-year follow-up [80].

Vitamin E deficiency

Vitamin E is involved in several processes including antioxidant immunomodulation [81,82]. Vitamin E deficiency has been reported in up to 22% of bariatric surgery patients. Post-bariatric surgery neurological symptoms could manifest in the form of ataxia, muscle weakness, peripheral neuropathy, or unexplained anemia [83]. Vitamin E deficiency also may hemolysis red blood cells, defective embryogenesis, and capillary permeability disorders [48].

Vitamin K deficiency

Vitamin K acts as a critical cofactor in the carboxylation of glutamic acid residues in several proteins as well as proteins involved in bone homeostasis (e.g., osteocalcin) [84]. Vitamin K has a rapid turnover and minimal body reservoirs [85]. Traditionally, vitamin K status is assessed via functional assays such as prothrombin time [86]. Symptoms of vitamin K deficiency include bleeding, arteriosclerosis, and osteoporosis [87]. Bleeding is a primary manifestation of vitamin K deficiency. Easy mucosal bleeding especially epistaxis, gastrointestinal hemorrhage, menorrhagia, and hematuria could occur [48]. Vitamin K deficiency should be treated with 10 mg injection supplements, followed by 1-2 mg/week administered parenterally or orally [88].

B-Mineral Deficiencies

Iron deficiency

Iron deficiency is common and the earliest nutritional deficiency to occur following bariatric surgery, occurring in up to 12%-47% of patients [89,90]. Symptoms of iron deficiency anemia include fatigue, palpitations, anxiety, hair loss, feeling cold [29]. Laboratory tests for determining iron deficiency should include serum ferritin levels, serum iron, transferrin saturation, total iron binding capacity, mean corpuscular hemoglobin, and hemoglobin level. The typical presentation of iron deficiency with anemia is elevated total iron binding capacity, with low values for all other hematological parameters [91]. Iron deficiency has been reported in up to 44% of adults before bariatric surgery, which may contribute to the persistence of iron deficiency post-operatively if not identified and treated [92]. Hypoferritinemia without Anemia (HWA) seen in many

post-bariatric patients is a disorder that requires correction of serum ferritin levels [93]. Post-bariatric surgery iron supplementation is necessary in most cases [30,94]. Refractory responses to oral iron supplementation might necessitate parenteral iron therapy or even blood transfusions [95].

Calcium deficiency

Calcium is a necessary element for human life; 99% of the calcium in the body is stored in teeth and bones [96,97]. Calcium helps the heart, brain, nerves, and muscles, and promotes blood clotting. Calcium deficiency develops in up to 48% of patients that undergo malabsorptive bariatric surgery. Common manifestations include muscle spasms, back and leg cramps, depression, aching joints, eczema, insomnia, cognitive impairment, and convulsions. If calcium deficiency is severe or acute, it can cause muscle spasm or cramping, tingling, burning sensation around the mouth and fingertips, seizures, and tremors [98]. The chances of osteoporosis in older patient's post- bariatric surgery are higher [99]. Iron deficiency anemia could also be related to calcium intake in such patients. Treatment includes ensuring a balanced phosphorous to calcium ratio. Products containing phosphoric acid may worsen the calcium deficiency, as it increases calcium excretion, such as dark colas, bottled teas, and flavored waters [91].

Phosphorus deficiency

Phosphorus has several functions in the human body. It is a major component of bones, DNA-RNA, ATP (the primary energy producer in the body), and phospholipids. Phosphorus is also essential for maintaining the acid-base balance in the body [100]. Deficiency is rare because phosphorus is widely available in food. However, bariatric surgery, alcoholism, and certain medicines can cause low levels of phosphorus, known as hypophosphatemia [101]. Symptoms of deficiency include anorexia, anemia, muscle weakness, bone pain, confusion, increased susceptibility to infection, difficulty walking and, in severe cases, death [102].

Copper deficiency

Copper deficiency is a well-documented cause of hematologic abnormalities, including anemia with neutropenia [103,104]. Copper is required for iron mobilization in the body; thus, copper deficiency manifests as iron deficiency, which results in low concentrations of red blood cells and white blood cells [105]. Studies have shown that the level of serum copper decreases following bariatric surgery [106]. The prevalence of copper deficiency ranges from 4%-18% [107]. Another concern is that anemia due to copper deficiency is often misdiagnosed as iron and vitamin B₁₂ deficiency. The neurological symptoms such as unsteady gait, extremity numbness, tingling in the hands and feet [108], leukopenia, pain, impaired wound healing, paresthesia, or paralysis may be irreversible [109].

Magnesium deficiency

Magnesium is an important element that activates more than 300 enzymes in the human body [110] and is often found to be deficient in patients after bariatric surgery. Magnesium assists in maintaining the muscle and nerve function in the human body. People with magnesium deficiency are often tired, irritable, nervous, experience muscle stiffness and difficulty in concentrating. Magnesium deficiency is linked to various pathological conditions such as those

affecting bones, heart, muscular system, emotional and dental health, as well as stroke, diabetes, stress, depression, and anxiety. Therefore, magnesium deficiency disrupts biochemical functions in the human body, especially in the nervous system [111]. Magnesium deficiency could lead to anemia, as low magnesium levels cause the red blood cell membranes to become more fragile and easily damaged. Heaton, et al. revealed that anemia could develop during magnesium deficiency due to changes in the red blood cell membrane, which is the primary reason for the underlying disturbances in cellular metabolism [112].

Selenium deficiency

Selenium is a component of the glutathione peroxidase enzyme. It helps prevent cell damage caused by free radicals. The prevalence of selenium deficiency following bariatric surgery has been estimated at up to 20% [113]. Currently, no sufficient evidence supports the need for routine selenium screening or supplementation after bariatric surgery to treat selenium deficiency [114]. Selenium levels need to be checked in patients who have undergone malabsorptive surgical procedures who develop anemia, fatigue, cardiomyopathy, heart palpitations, impaired immunity, infertility, and myalgia [115].

Chromium deficiency

Patients who undergo bariatric surgery are at risk for low chromium intake. Even a mild chromium deficiency could generate problems in blood sugar metabolism and contribute to other symptoms such as anxiety or fatigue [116]. Chromium is considered to work synergistically with iron, and its deficiency can thus lead to iron deficiency [117].

Zinc deficiency

Zinc is a trace mineral co-factor used in various enzymes involved in protein synthesis, digestion, and regulation of gene transcription [118]. Zinc is also essential to maintain immune function. Specifically, zinc is vital for T-cell growth, differentiation, and apoptosis, which kill dangerous bacteria, viruses, and cancer cells. It is also a critical structural component of hormone receptors and proteins that contribute to a healthy, balanced mood [119]. Manifestations of zinc deficiency include respiratory, gastrointestinal symptoms and bacterial infections due to low immunity [120], skin lesions, poor wound healing, dermatitis, hair loss, alopecia, and glossitis - symptoms similar to vitamin B₁₂ deficiency [121]. During inflammation, there is an increased cellular demand for zinc [122]. Zinc deficiency has been reported in 36%-51% of patients who underwent bariatric surgery [123,124].

Other micronutrients such as proteins are vehicles for the functions of metabolism, digestion, and DNA replication [125,126]. They frame the structure of cells such as those of the hair, skin, and nails. Proteins also help the body combat infection by strengthening the immune system [127]. Low intake of protein during post-bariatric surgical recovery may lead to mild to moderate anemia [128]. Anemia-like symptoms of protein deficiency include lethargy, easy fatigue, headaches, and muscle wasting. Following bariatric surgery, women require 60-80 g of protein daily, and men 70-90 g per day. Measurement of albumin levels is a useful approach for assessing the protein status [91,125].

Prophylactic and Supplementary Measures

A prophylactic anticoagulants protocol for post-bariatric

surgery could be a useful tool to prevent thrombosis in such morbid obesity patients; also a supplementary protocol with mega doses of micronutrients should be implemented before bariatric surgery especially for patients with nutritional deficiencies [129]. Multi-vitamin supplementation is recommended after bariatric procedures [130]. The importance of postoperative dietary supplementation for bariatric surgery has been widely recognized by researchers and clinicians [131]. Recent studies continue to report nutrient deficiencies after bariatric surgery, which should always be monitored and treated [132]. Difficulties in swallowing pills were identified as a primary barrier to adherence to multivitamins and other trace elements besides malabsorption [61]. Such patients could benefit from intravenous administration of multivitamins [133].

Conclusion

Bariatric surgery is a new approach for management of diabetes mellitus, various metabolic disorders and psychological problems due to morbid obesity. Over the past three decades, a substantial amount of research has been conducted regarding the complications associated with bariatric surgery; however, the authors think it is a safe procedure when the hematological and nutritional precautions are put into place and the necessary investigations and prophylactic measures are performed. As this type of surgery has become very popular, it is important that physicians perform thorough medical history for the possibility that the patient with obscure complaints may have history of bariatric surgery because some patient forget to mention that in their medical history and being aware of the potential complications will help to ensure safe and effective outcomes.

References

1. Low VH, Tan J, Lu J. Complications of laparoscopic adjustable gastric banding: Our local experience. *J Med Imaging Radiat Oncol*. 2012; 56: 432-441.
2. Gray DS, Fujioka K. Use of relative weight and Body Mass Index for the determination of adiposity. *J Clin Epidemiol*. 1991; 44: 545-550.
3. Must A, McKeown NM. The Disease Burden Associated with Overweight and Obesity. South Dartmouth (MA): MDText.com, Inc, 2000.
4. Livingston EH. Bariatric surgery in the new millennium. *Arch Surg*. 2007; 142: 919-922.
5. Robinson MK. Surgical treatment of obesity--weighing the facts. *N Engl J Med*. 2009; 361: 520-521.
6. Fajnwaks P, Ramirez A, Martinez P, Arias E, Szomstein S, Rosenthal R. P46: Outcomes of bariatric surgery in patients with BMI less than 35 kg/m2. *Surgery for Obesity and Related Diseases*. 2008; 4: 329.
7. World Health Organization. Obesity: Preventing and Managing the Global Epidemic. WHO Technical Report Series 894. 2000.
8. Gregg EW, Cheng YJ, Cadwell BL, Imperatore G, Williams DE, Flegal KM. Secular Trends in Cardiovascular Disease Risk Factors According to Body Mass Index in U.S. Adults. *Obstetrical & Gynecological Survey*. 2005; 60: 660-661.
9. De Re AC, Frayne SM, Harris AH. Antiobesity Medication Use across the Veterans Health Administration: Patient-Level Predictors of Receipt. *Obesity (Silver Spring)*. 2014; 22: 1968-1972.
10. Gastrointestinal surgery for severe obesity: National Institutes of Health Consensus Development Conference Statement. *Am J Clin Nutr*. 1992; 55: 615S-619S.
11. Major P, MatAok M, PÄ™dziwiatr M, Migaczewski M, Budzyński P, Stanek M, et al. Quality of Life After Bariatric Surgery. *Obes Surg*. 2015; 25: 1703-1710.

12. Jeremy FK, Richdeep SG, Michael L, Shahzeer K. The Impact of Bariatric Surgery on Psychological Health. *Journal of Obesity*. 2013.
13. Algahtani HA, Khan AS, Khan MA, Aldarmahi AA, Lodhi Y. Neurological complications of bariatric surgery. *Neurosciences (Riyadh)*. 2016; 21: 241-245.
14. Acquafresca PA, Palermo M, Rogula T, Duza GE, Serra E. Early Surgical Complications after Gastric BY-Pass: A Literature Review. *Arq Bras Cir Dig*. 2015; 28: 74-80.
15. Marcia Nelms, Kathryn P. Sucher, Karen Lacey, Sara Long Roth. *Nutrition Therapy and Pathophysiology*. 2011, 2007.
16. *Pharmacology, Toxicology & Therapeutics*. 2016.
17. Wente MN, Veit JA, Bassi C, Dervenis C, Fingerhut A, Gouma DJ, et al. Postpancreatectomy hemorrhage (PPH): an International Study Group of Pancreatic Surgery (ISGPS) definition. *Surg*. 2007; 142: 20-25.
18. Rachid Baz, Tarek Mekhail. *Bleeding Disorders*. The Cleveland Clinic Foundation. 2015.
19. Abraham F, Samuel S, Raul JR. Postoperative Bleeding in the Bariatric Surgery Patient. *The ASMBS Textbook of Bariatric Surgery*. 2014; 9: 241-247.
20. Murugesan A, Srivastava DN, Ballehaninna UK, Chumber S, Dhar A, Misra MC, et al. Detection and Prevention of Post-Operative Deep Vein Thrombosis [DVT] Using Nadroparin Among Patients Undergoing Major Abdominal Operations in India; a Randomised Controlled Trial. *Indian J Surg*. 2010; 72: 312-317.
21. Frederick A. Anderson, Anne-Marie Audet. *Preventing Deep Vein Thrombosis and Pulmonary Embolism: A Practical Guide to Evaluation and Improvement*. 1998. Center for Outcomes Research, University of Massachusetts Medical Center.
22. Lancaster RT, Hutter MM. Bands and bypasses: 30-day morbidity and mortality of bariatric surgical procedures as assessed by prospective, multi-center, risk-adjusted ACS-NSQIP data. *Surg Endosc*. 2008; 22: 2554-2563.
23. Winegar DA, Sherif B, Pate V, DeMaria EJ. Venous thromboembolism after bariatric surgery performed by Bariatric Surgery Center of Excellence Participants: analysis of the Bariatric Outcomes Longitudinal Database. *Surg Obes Relat Dis*. 2011; 7: 181-188.
24. Desciak MC, Martin DE. Perioperative pulmonary embolism: diagnosis and anesthetic management. *J Clin Anesth*. 2011; 23: 153-165.
25. Geerts WH, Heit JA, Clagett GP, Pineo GF, Colwell CW, Anderson FA Jr, et al. Prevention of venous thromboembolism. *Chest*. 2001; 119: 132S-175S.
26. Wood KE. Major pulmonary embolism: review of a pathophysiologic approach to the golden hour of hemodynamically significant pulmonary embolism. *Chest*. 2002; 121: 877-905.
27. Balsiger BM, Kennedy FP, Abu-Lebdeh HS, Collazo-Clavell M, Jensen MD, O'Brien T, et al. Prospective evaluation of Roux-en-Y gastric bypass as primary operation for medically complicated obesity. *Mayo Clin Proc*. 2000; 75: 673-680.
28. Stein J, Stier C, Raab H, Weiner R. Review article: the nutritional and pharmacological consequences of obesity surgery. *Aliment Pharmacol Ther*. 2014; 40: 582-609.
29. Donadelli SP, Junqueira-Franco MV, de Mattos Donadelli CA, Salgado W Jr, Ceneviva R, Marchini JS, et al. Daily vitamin supplementation and hypovitaminosis after obesity surgery. *Nutrition*. 2012; 28: 391-396.
30. Gehrler S, Kern B, Peters T, Christoffel-Courtin C, Peterli R. Fewer nutrient deficiencies after laparoscopic sleeve gastrectomy (LSG) than after laparoscopic Roux-Y-gastric bypass (LRYGB) - a prospective study. *Obes Surg*. 2010; 20: 447-53.
31. Brolin RE, LaMarca LB, Kenler HA, Cody RP. Malabsorptive gastric bypass in patients with superobesity. *J Gastrointest Surg*. 2002; 6: 195-203.
32. Berger JR. The neurological complications of bariatric surgery. *Arch Neurol*. 2004; 61: 1185-1189.
33. Brolin RE, Leung M. Survey of vitamin and mineral supplementation after gastric bypass and biliopancreatic diversion for morbid obesity. *Obes Surg*. 1999; 9: 150-154.
34. Xanthakos SA. Nutritional deficiencies in obesity and after bariatric surgery. *Pediatr Clin North Am*. 2009; 56: 1105-1121.
35. Raziel A. Thiamine deficiency after bariatric surgery may lead to Wernicke encephalopathy. *Isr Med Assoc J*. 2012; 14: 692-694.
36. Manzo G, De Gennaro A, Cozzolino A, Serino A, Fenza G, Manto A. MR Imaging Findings in Alcoholic and Nonalcoholic Acute Wernicke's Encephalopathy: A Review. *Biomed Res Int*. 2014.
37. Michael Zimmermann. Diet, nutrition and the prevention of chronic diseases: by the World Health Organisation, 1991, 203 pages, softcover. WHO, Geneva. *The American Journal of Clinical Nutrition*. 1994; 60: 644-645.
38. Genetic and Rare Diseases Information Center (GARD) -an NCATS Program. 2015. 2017.
39. Boros LG, Steinkamp MP, Fleming JC, Lee WN, Cascante M, Neufeld EJ. Defective RNA ribose synthesis in fibroblasts from patients with thiamine-responsive megaloblastic anemia (TRMA). *Blood*. 2003; 102: 3556-3561.
40. Oishi K, Diaz GA. Thiamine-Responsive Megaloblastic Anemia Syndrome, 2003 Oct 24. In: Adam MP, Ardinger HH, Pagon RA, et al., editors. *GeneReviews®* [Internet]. Seattle (WA): University of Washington, Seattle. 2018.
41. Matrana MR, Davis WE. Vitamin deficiency after gastric bypass surgery: a review. *South Med J*. 2009; 102: 1025-1031.
42. Jaeger B, Bosch AM. Clinical presentation and outcome of riboflavin transporter deficiency: mini review after five years of experience. *J Inherit Metab Dis*. 2016; 4: 559-564.
43. Lane M, Alfrey CP Jr. THE ANEMIA OF HUMAN RIBOFLAVIN DEFICIENCY. *Blood*. 1965; 25: 432-442.
44. Kelly O'Donnell. Severe Micronutrient Deficiencies in RYGB Patients: Rare but Potentially Devastating. *Practical Gastroenterology*. 2011.
45. Fukuwatari T, Shibata K. Nutritional Aspect of Tryptophan Metabolism. *Int J Tryptophan Res*. 2013; 6: 3-8.
46. Cahrtotte R, Leonard G, Tina M, Chris P. 1st Edition. *Best of Five MCQs for the Gastroenterology SCE*, Oxford University Press. 2013.
47. Manzoni AP, Weber MB1. Skin changes after bariatric surgery. *An Bras Dermatol*. 2015; 90: 157-166.
48. O Wenker. *Vitamin Deficiencies: An Overview*. The Internet Journal of Nutrition and Wellness. 2004.
49. Gropper SS, Smith JL, Groff JL. *Advanced nutrition and human metabolism*. Belmont, CA: Wadsworth, Cengage learning. 2009.
50. Otten JJ, Hellwig JP, Meyers LD. *Dietary reference intakes: The essential guide to nutrient requirements*. Washington, DC: The National Academies Press. 2009.
51. Smith CM, Song WO. "Comparative nutrition of pantothenic acid". *Journal of Nutritional Biochemistry*. 1996; 7: 312-321.
52. Xanthakos, Stavra A. Nutritional Deficiencies in Obesity and After Bariatric Surgery. *Pediatric clinics of North America*. 2009; 56: 1105-1121.
53. Stover PJ, Field MS. Vitamin B-6. *Adv Nutr*. 2015; 6: 132-133.
54. Greenway FL, Ingram DK, Ravussin E, Hausmann M, Smith SR, Cox L, et al. Loss of taste responds to high-dose biotin treatment. *J Am Coll Nutr*. 2011; 30: 178-181.
55. Biotin. *Medline Plus Supplements*. Retrieved. 2013.
56. Biotin. *Medline Plus*. U.S. National Library of Medicine, 2010.
57. Weng TC, Chang CH, Dong YH, Chang YC, Chuang LM. Anaemia and related nutrient deficiencies after Roux-en-Y gastric bypass surgery: a systematic review and meta-analysis. *BMJ Open*. 2015; 5: e006964.

58. Madan AK, Orth WS, Tichansky DS, Ternovits CA. Vitamin and trace mineral levels after laparoscopic gastric bypass. *Obes Surg.* 2006; 16: 603-606.
59. Mahmood L. The metabolic processes of folic acid and Vitamin B12 deficiency. *J Health Res Rev* [serial online]. 2014; 1: 5-9.
60. American College of Obstetricians and Gynecologists. ACOG Committee opinion no. 549: obesity in pregnancy. *Obstet Gynecol.* 2013; 121: 213-217.
61. Modi AC, Zeller MH, Xanthakos SA, Jenkins TM, Inge TH. Adherence to vitamin supplementation following adolescent bariatric surgery. *Obesity* (Silver Spring). 2013; 21: E190-E195.
62. Majumder S, Soriano J, Louie Cruz A, Dasanu CA. Vitamin B12 deficiency in patients undergoing bariatric surgery: Preventive strategies and key recommendations. *Surg Obes Relat Dis.* 2013; 9: 1013-1019.
63. George A Bray, Claude Bouchard. *Handbook of Obesity - Volume 2: Clinical Applications*, Fourth Edition. 2014.
64. Sakly G, Hellara O, Trabelsi A, Doqui M. [Reversible peripheral neuropathy induced by vitamin B12 deficiency]. *Neurophysiol Clin.* 2005; 35: 149-153.
65. Malinowski SS. Nutritional and metabolic complications of bariatric surgery. *Am J Med Sci.* 2006; 331: 219-225.
66. Oh R, Brown DL. Vitamin B12 deficiency. *Am Fam Physician.* 2003; 67: 979-986.
67. Schilling RF, Gohdes PN, Hardie GH. Vitamin B12 deficiency after gastric bypass surgery for obesity. *Ann Intern Med.* 1984; 101: 501-502.
68. Zalesin KC, Miller WM, Franklin B, Mudugala D, Rao Buragadda A, Boura J, et al. Vitamin a deficiency after gastric bypass surgery: an underreported postoperative complication. *J Obes.* 2011; 2011.
69. Clements RH, Yellumhanthi K, Wesley M, Ballem N, Bland KI. Incidence of vitamin deficiency after laparoscopic Roux-en-Y gastric bypass in a university hospital setting. *Am Surg.* 2006; 72: 1196-1202.
70. Semba RD, Bloem MW. The anemia of vitamin A deficiency: epidemiology and pathogenesis. *Eur J Clin Nutr.* 2002; 56: 271-281.
71. Dana Swiley. *Micronutrient and Macronutrient Needs in Roux-en-Y Gastric Bypass Patients* BT Online Editor. 2008.
72. Hansen EPK, Metzsch C, Henningsen E, Toft P. Severe Scurvy After Gastric Bypass Surgery and a Poor Postoperative Diet. *J Clin Med Res.* 2012; 4: 135-137.
73. Padayatty SJ, Levine M. New insights into the physiology and pharmacology of vitamin C. *CMAJ.* 2001; 164: 353-355.
74. Rizzo MR, Abbatecola AM, Barbieri M, Vietri MT, Cioffi M, Grella R, et al. Evidence for anti-inflammatory effects of combined administration of vitamin E and C in older persons with impaired fasting glucose: impact on insulin action. *J Am Coll Nutr.* 2008; 27: 505-511.
75. Chakhtoura, M, Nakhoul N, Akl EA, Mantzoros CS, El Hajj Fuleihan GA. Guidelines on Vitamin D Replacement in Bariatric Surgery: Identification and Systematic Appraisal. *Metabolism: clinical and experimental.* 2016; 65: 586-597.
76. Azizi-Soleiman F, Vafa M, Abiri B, Safavi M. Effects of Iron on Vitamin D Metabolism: A Systematic Review. *Int J Prev Med.* 2016; 7: 126.
77. Grant WB, Holick MF. Benefits and requirements of vitamin D for optimal health: a review. *Altern Med Rev.* 2005; 10: 94-111.
78. Bacci V, Silecchia G. Vitamin D status and supplementation in morbid obesity before and after bariatric surgery. *Expert Rev Gastroenterol Hepatol.* 2010; 4: 781-794.
79. Coates PS, Fernstrom JD, Fernstrom MH, Schauer PR, Greenspan SL. Gastric bypass surgery for morbid obesity leads to an increase in bone turnover and a decrease in bone mass. *J Clin Endocrinol Metab.* 2004; 89: 1061-1065.
80. Matrana MR, Davis WE. Vitamin Deficiency After Gastric Bypass Surgery: A Review. *Southern Medical Journal.* 2009.
81. Cuesta M, Pelaz L, Pérez C, Torrejón MJ, Cabrerizo L, Matía P, et al. Fat-soluble vitamin deficiencies after bariatric surgery could be misleading if they are not appropriately adjusted. *Nutr Hosp.* 2014; 30: 118-123.
82. Kurutas EB. The importance of antioxidants which play the role in cellular response against oxidative/nitrosative stress: current state. *Nutr J.* 2016; 15: 71.
83. Ueda N, Suzuki Y, Rino Y, Takahashi T, Imada T, Takanashi Y, et al. Correlation between neurological dysfunction with vitamin E deficiency and gastrectomy. *J Neurol Sci.* 2009; 287: 216-220.
84. Shearer MJ. Vitamin K. *Lancet.* 1995; 345: 229-234.
85. Olson RE. The function and metabolism of vitamin K. *Annu Rev Nutr.* 1984; 4: 281-337.
86. Strole J, Lovell G, Heubi J. Prevalence of subclinical vitamin K deficiency in cholestatic liver disease. *J Pediatr Gastroenterol Nutr.* 2009; 49: 78-84.
87. Booth SL. Roles for vitamin K beyond coagulation. *Annu Rev Nutr.* 2009; 29: 89-110.
88. Allied Health Sciences Section Ad Hoc Nutrition Committee, Aills L, Blankenship J, Buffington C, Furtado M, Parrott J. *ASMBS Allied Health Nutritional Guidelines for the Surgical Weight Loss Patient.* *Surg Obes Relat Dis.* 2008; 4: S73-108.
89. Jáuregui-Lobera I. Iron deficiency and bariatric surgery. *Nutrients.* 2013; 5: 1595-1608.
90. Bavarese M, Paganini S, Lima TP, Salgado W Jr, Ceneviva R, Dos Santos JE, et al. Nutritional course of patients submitted to bariatric surgery. *Obes Surg.* 2010; 20: 716-721.
91. Mallory GN, Macgregor AM. Folate Status Following Gastric Bypass Surgery (The Great Folate Mystery). *Obes Surg.* 1991; 1: 69-72.
92. Flancbaum L, Belsley S, Drake V, Colarusso T, Tayler E. Preoperative nutritional status of patients undergoing Roux-en-Y gastric bypass for morbid obesity. *J Gastrointest Surg.* 2006; 10: 1033-1037.
93. Al-Jafar HA1. HWA: Hypoferritinemia without anemia a hidden hematology disorder. *J Family Med Prim Care.* 2017; 6: 69-72.
94. Al-Mutawa A, Al-Sabah S, Anderson AK, Al-Mutawa M. Evaluation of Nutritional Status Post Laparoscopic Sleeve Gastrectomy-5-Year Outcomes. *Obes Surg.* 2018; 28: 1473-1483.
95. Gurewitsch ED, Smith-Levitin M, Mack J. Pregnancy following gastric bypass surgery for morbid obesity. *Obstet Gynecol.* 1996; 88: 658-661.
96. Helen O Wilson, Dev BN Datta. Complications from micronutrient deficiency following bariatric surgery. *Annals of Clinical Biochemistry.* 2014; 51: 705-709.
97. Ross AC, Taylor CL, Yaktine AL, et al. Institute of Medicine (US) Committee to Review. *Dietary Reference Intakes for Vitamin D.* Washington (DC): National Academies Press (US). 2011.
98. North London Obesity Surgery Service (NLOSS). About Calcium and Calcium Deficiency.
99. Scott Belsley. Calcium Supplementation Post Bariatric Surgery. 2015.
100. Beleidya AE, Sherbinia S, HebatAllah, Elgebalya F, Ahmed A. Calcium, magnesium and phosphorus deficiency in critically ill children. 2017; 65: 60-64.
101. GB HealthWatch. 2017.
102. Elisa Brantley. Phosphorous. 2015.
103. Griffith DP, Liff DA, Ziegler TR, Esper GJ, Winton EF. Acquired Copper Deficiency: A Potentially Serious and Preventable Complication Following Gastric Bypass Surgery. *Obesity* (Silver Spring). 2009; 17: 827-831.
104. Prodan CI, Holland NR, Wisdom PJ, Burstein SA, Bottomley SS. CNS demyelination associated with copper deficiency and hyperzincemia. *Neurology.* 2002; 59: 1453-1456.

105. Gletsu-Miller N, Breanne WN. Mineral Malnutrition Following Bariatric Surgery. *Adv Nutr*. 2013; 4: 506-517.
106. Rojas P, Carrasco F, Codoceo J, Inostroza J, Basfi-fer K, Papapietro K, et al. Trace element status and inflammation parameters after 6 months of roux-en-y gastric bypass. *Obes Surg*. 2011; 21: 561-568.
107. Gletsu-Miller N, Broderius M, Frediani JK, Zhao VM, Griffith DP, Davis SS Jr, et al. Incidence and prevalence of copper deficiency following roux-en-y gastric bypass surgery. *Int J Obes (Lond)*. 2012; 36: 328-335.
108. Griffith DP, Liff DA, Ziegler TR, Esper GJ, Winton EF. Acquired copper deficiency: a potential serious and preventable complication following gastric bypass surgery. *Obesity (Silver Spring)*. 2009; 17: 827-831.
109. Moize V, Deulofeu R, Torres F, de Osaba JM, Vidal J. Nutritional intake and prevalence of nutritional deficiencies prior to surgery in a Spanish morbidly obese population. *Obes Surg*. 2011; 21: 1382-1388.
110. Kupetsky-Rincon EA, Li Q, Uitto J. Magnesium reduces carotid intima-media thickness in a mouse model of pseudoxanthoma elasticum: a novel treatment biomarker. *Clin Transl Sci*. 2012; 5: 259-264.
111. Qais Faryadi. The Magnificent Effect of Magnesium to Human Health: A Critical Review. *International Journal of Applied Science and Technology*. 2012.
112. Heaton FW, Tongyai S, Motta C, Rayssiguier Y, Guex E. Changes in the erythrocyte membrane during magnesium deficiency Original Research Article. *Nutrition Research*. 1987; 7: 655-663.
113. Shankar P, Boylan M, Sriram K. Micronutrient deficiencies after bariatric surgery. *Nutrition*. 2010; 26: 1031-1037.
114. Morin B, Bonnamy C, Maurel J, Samama G, Gignoux M. [Late intestinal fistula following implantation of parietal abdominal prostheses]. *Ann Chir*. 2001; 126: 876-880.
115. Mechanick JI, Youdim A, Jones DB, Garvey ET, Hurley DL, McMahon M, et al. Clinical practice guidelines for the perioperative nutritional, metabolic, and nonsurgical support of the bariatric surgery patient-2013 update: cosponsored by American Association of Clinical Endocrinologists, The Obesity Society, and American Society for Metabolic & Bariatric Surgery. *Obesity (Silver Spring)*. 2013; 21: S1-27.
116. Andrew Weil and Brian Becker. *Chromium*. 2012.
117. D. L. Watts. The nutritional relationships of Iron. *Journal of Orthomolecular Medicine*. 1988; 3: 110-116.
118. Cousins RJ, Blanchard RK, Moore JB, Cui L, Green CL, Liuzzi JP, et al. Regulation of zinc metabolism and genomic outcomes. *J Nutr*. 2003; 133: 1521S-6S.
119. Baum MK, Shor-Posner G, Campa A. Zinc status in human immunodeficiency virus infection. *J Nutr*. 2000; 130: 1421S-3S.
120. Kumar P, Clark ML. *Kumar & Clark's clinical medicine (8th ed.)*. Edinburgh: Elsevier/Saunders. 2012.
121. Yamada T, Alpers DH, Kalloo AN, Kaplowitz N, Owyang C, Powell DW. *Textbook of gastroenterology (5th ed.)*. Chichester, West Sussex: Blackwell Pub. 2009.
122. Foster M, Samman S. Zinc and regulation of inflammatory cytokines: implications for cardiometabolic disease. *Nutrients*. 2012; 4: 676-694.
123. Slater GH1, Ren CJ, Siegel N, Williams T, Barr D, Wolfe B, et al. Serum fat-soluble vitamin deficiency and abnormal calcium metabolism after malabsorptive bariatric surgery. *J Gastrointest Surg*. 2004; 8: 48-55.
124. Madan AK, Orth WS, Tichansky DS, Ternovits CA. Vitamin and trace mineral levels after laparoscopic gastric bypass. *Obes Surg*. 2006; 16: 603-606.
125. Aron-Wisniewsky J, Verger EO, Bounaix C, Dao MC, Oppert JM, Bouillot JL, et al. Nutritional and Protein Deficiencies in the Short Term Following Both Gastric Bypass and Gastric Banding. *PLoS ONE*. 2016.
126. Geetha A, Thiru PNV, Sheela Devi R, Subramanian S. *Biochemistry Higher Secondary-Second Year-Chapter: Protein Metabolism*. Government of Tamilnadu - 1st Edition - 2005.
127. Protein Deficiency after Gastric Bypass Surgery - My Bariatric Life. 2017.
128. Abazar Habibinia. Protein Deficiency Anemia. *Canadian Academy of Sports Nutrition*. 2014.
129. Bordalo LA, Teixeira TF, Bressan J, Mourão DM. Bariatric surgery: how and why to supplement. *Rev Assoc Med Bras*. 2011; 57: 111-118.
130. Brolin RE, Gorman JH, Gorman RC, Petschenik AJ, Bradley LJ, Kenler HA, et al. Are vitamin B12 and folate deficiency clinically important after roux-en-Y gastric bypass?. *J Gastrointest Surg*. 1998; 2: 436-442.
131. Qianxiong Zheng. Adherence to Micronutrient Supplementation in Bariatric Patients.
132. Saif T, Strain GW, Dakin G, Gagner M, Costa R, Pomp A. Evaluation of nutrient status after laparoscopic sleeve gastrectomy 1, 3, and 5 years after surgery. *Surg Obes Relat Dis*. 2012; 8: 542-547.
133. Ferguson TI, Emry S, Price-Davies R, Gosslett AG. A review of stability issues associated with vitamins in parenteral nutrition. *Clinical Nutrition ESPEN*. 2014; 9: e49-e53.