IRON NUTRITIONAL STATUS AMONG FEMALE UNIVERSITI MALAYSIA SABAH (UMS) STUDENTS

PERPUSTAKAAN UNIVERSITI MALAYSIA SABAH

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FACULTY OF FOOD SCIENCE AND NUTRITION UNIVERSITI MALAYSIA SABAH 2014



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DECLARATION

I hereby declare that the material in this thesis is my own except for the quotations, excerpts, summaries and references, which have been duly acknowledged.

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ABSTRACT

IRON NUTRITIONAL STATUS AMONG FEMALE UNIVERSITI MALAYSIA SABAH STUDENTS

The objectives of the present study were to determine iron nutritional status among female Universiti Malaysia Sabah (UMS) students and the differences in dietary iron intake between respondents with and without anaemia. A total of 436 female UMS students of aged 19-26 years were recruited in this study. Iron nutritional status was determined by measuring haemoglobin concentration via gravimetric technique while dietary intake data was obtained through semi-quantitative questionnaire (FFQ) modified from Malaysian Adults Nutrition Survey (MANS) 2003. Anaemia was determined using a single biochemical indicator, which was Hb<12.5g/dL. Prevalence of anaemia was 50.5% in the present study, which was higher than the estimated global prevalence of 30.2%. Most anaemic female students (n=207, 94.1%) were unaware they had Hb<12.5g/dL prior to this study. Malays recorded the highest prevalence of anaemia (53.40%), followed by Kadazans (50.8%) and Chinese (44.5%). The mean dietary iron intake of 14.9±9.3mg corresponded to 74.7% of the recommended nutrient intake. The difference in mean dietary iron intake between anaemic and non-anaemic respondents was not statistically significant (p=0.126). Dietary iron intake was also found to be positively correlated to total energy intake ($r_s=0.889$, p<0.001). Primary iron sources of dietary iron was malted beverages (25.5% of total reported iron intake), followed by grains or cereal products (20.0%) and meat (18.1%). In conclusion, strategies to improve poor iron nutrition status in female undergraduates need to address promoters and inhibitors of non-haem iron absorption given these main iron intake sources.



ABSTRAK

STATUS ZAT BESI DALAM KALANGAN PELAJAR WANITA UNIVERSITI MALAYSIA SABAH

Objektif-objektif kajian ini adalah untuk mengenalpasti status zat besi dalam kalangan pelajar wanita Universiti Malaysia Sabah (UMS) serta perbezaan pengambilan zat besi antara pelajar anemik dan bukan anemik. Seramai 436 wanita UMS pelajar berumur antara 19 hingga 26 tahun telah terlibat dalam kajian ini. Status zat besi subjek ditentukan melalui pengukuran kepekatan hemoglobin melalui teknik gravimetrik manakala data pengambilan makanan diperoleh melalui borang kekerapan pengambilan makanan semi-kuantitatif (FFQ) yang diubahsuai daripada Malaysian Adults Nutrition Survey (MANS) 2003. Anaemia dikenalpasti melalui satu petunjuk biokimia, iaitu Hb<12.5g/dL. Prevalen anemia dalam kalangan responden adalah 50.5%, jauh lebih tinggi berbanding dengan prevalen global, iaitu 30.2%. Kebanyakkan pelajar wanita (n=207, 94.1%) yang mengalami anaemia tidak sedar bahawa mereka mempunyai Hb<12.5g/dL sebelum kajian ini dijalankan. Responden Melayu mencatatkan prevalen anemia tertinggi (53.4%), diikuti oleh etnik Kadazan (50.8%) dan Cina (44.5%). Min pengambilan zat besi sebanyak 14.9±9.4mg merupakan 74.7% daripada saranan RNI 2005 bagi zat besi. Perbezaan dari segi min pengambilan zat besi antara responden anemik dan bukan anemik didapati tidak signifikan (*p*=0.126). Walau bagaimanapun, korelasi positif wujud antara pengambilan zat besi dengan jumlah pengambilan tenaga dalam kajian ini (rs=0,889, p<0.001). Sumber zat besi utama adalah daripada minuman malt (25.5%), diikuti oleh bijirin dan produk bijirin (20.0%) serta daging (18.1%). Kesimpulannya, strategi-strategi untuk memperbaiki status zat besi prasiswazah wanita perlu mengambil kira pengambilan penggalak dan perencat penyerapan zat besi bukan heme memandangkan ia merupakan sumber zat besi utama mereka.



Table of Contents

DECLARATION	i
CERTIFICATION	
ACKNOWLEDGEMENT	
ABSTRACT	iv
ABSTRAK	······································
LIST OF FIGURES	Y
LIST OF TABLES	vi
LIST OF ABBREVIATIONS	vii
LIST OF SYMBOLS	•••••••
LIST OF APPENDICES	viv
	····· XIV

Chapte	er 1: Introduction	1
1.1	Overview	1
1.2	Problem statement	4
1.3	Study objectives	
1.4	Study rationale	······ ۲
1.5	Hypothesis	5

Chapter 2:	Literature review	6
2.1 Int	roduction	6
2.2 Iro	n metabolism and bioavailability	·····0
2.2.1	Dietary iron absorption metabolism	/
2.2.2	Bioavailability of haem and non-haem iron	······ / o
2.3 Fac	tors leading to decrease iron absorption	o
2.3.1	Phytates	9
2.3.2	Phytates in Malaysian diet	9
2.3.3	Individual iron nutritional status	10
2.3.4	Polyphenols	11
2.3.5	Dietary fibre	11
2.3.6	Calcium	13
2.3.7	Phosphorus	14
238	Endurance physical activities	15
2.3.0	Smoking and altitude	16
2.3.3	Shoking and dillude	17

vi





•

2.4	Nut	rient Deficiencies
2.4	.1	Vitamin B12
2.4	.2	Folate
2.4	.3	Vitamin A
2.4	.4	Copper
2.4	.5	Obesity and inflammation
2.4	.6	Genetic diseases
2.5	Cor	nsequences of decreased iron absorption – Anaemia
2.5	.1	Iron deficiency anaemia
2.5	.2	Pernicious anaemia (Megaloblastic anaemia)
2.5	.3	Thalassemia
2.5	.4	Diagnosis of different forms of anaemia
2.6	Fac	tors leading to increased iron absorption
2.6	.1	Ascorbic acid (Vitamin C)27
2.6	.2	Meat, fish and poultry (MFP)28
2.6	.3	Alcohol
2.6	.4	Iron supplements
2.7	Iror	overload
2.8	Iror	n nutritional concerns in females of reproductive ages
2.8	.1	Menstrual blood loss
2.8	.2	Pregnancy
2.8.	.3	Dietary and energy intake
2.9	Veg	etarianism diet
2.10	Indi	cators of iron nutritional status
2.1(0.1	Haemoglobin
2.1().2	Serum ferritin (SF)
2.1().3	Mean cell volume (MCV)
2.1().4	Zinc protoporphyrin
2.10).5	Transferrin saturation / TIBC / Serum iron
2.11	Asse	essment of iron nutritional status at population level
2.12	Mea	surement of haemoglobin
2.13	Diet	ary Assessment
2.13	3.1	Food frequency questionnaire (FFQ)



Chapte	er 3:	Materials and methods	47
3.1	Mater	ials and consumables	47
3.2 9	Samp	ling method and sample size	47
3.2	2.1	Compensation for data loss	17 49
3.3	Dat	a collection methods	50
3.3	3.1	Haemoglobin concentration measurement	50
3.3	3.2	Food frequency questionnaire (FFO)	50
3.3	3.3	Socio-demographic questionnaire	JZ
3.4	Pilo	t study	
3.5	Stu	dy location	J-T E-A
3.6	Stu	dy methodology	54
3.7	Stat	tistical analyses	55
			30

Chapte	r 4:	Results and discussions	
4.1	Pilo	ot study	58
4.2	Val	idity of HemoCue B microcuvettes	58
4.3	Soc	cio-demographic characteristics of respondente	59
4.4	Hea	alth Status of Respondents	60
4 5	Dro	valence of anaomia	62
4.6	Dro		62
4.0	rie Cum	vious diagnoses and treatments of anaemia	65
л./ ЛО	Sup	ppiement intake	67
4.8	Ind	ependent factors for anaemia	69
4.8.	.1	Menstrual blood loss (MBL)	69
4.8.	2	Thalassemia	70
4.9	Тур	bes of diet consumed	71
4.10	Ene	ergy and nutrient intake	7 <u>4</u>
4.10	0.1	Energy	77 78
4.10).2	Macronutrient energy intake	/4 70
4.10).3	Micronutrients	78
	a.	Iron	/9
	b.	Calcium	79
	с.	Vitamin C	82
	с. д		83
A 1 4	u.	Dietary fibre	85
4.11	Diet	tary nabits among respondents with and without anaemia	86
4.11	l .1	Dark green leafy vegetables	86



4.11.2	Meat, fish, poultry and egg8	7
4.11.3	Citrus fruits	8
4.11.4	Legumes and high fibre food8	9
4.11.5	Milk, tea and coffee8	9

Chapte	r 5: Conclusions	92
5.1	Conclusion	92
5.2	Limitations and improvements of the study	94
5.3	Suggestions for future studies	95

References	
Appendices	106



`

LIST OF FIGURES

		Page
Figure 4.1	Prevalence of anaemia reported by previous studies	63
Figure 4.2	Time-lapse from the last anaemia diagnosis among respondents.	66
Figure 4.3	Reasons for choosing vegetarian diet among respondents.	73
Figure 4.4	Contribution of energy from different food groups.	76
Figure 4.5	Percentage of respondents achieving RNI 2005 requirements.	76
Figure 4.6	Iron intake between anaemic and non-anaemic respondents	80
Figure 4.7	Contribution of dietary iron from different food groups.	82



LIST OF TABLES

		Page
Table 1	Altitude adjustments to measured haemoglobin	18
	concentrations.	
Table 2	Adjustments to measured haemoglobin concentrations for	18
	smokers.	
Table 3	Comparison between biomarkers.	26
Table 4.1	Mean haemoglobin level measured by HemoCue B	60
	photometer and full blood count analyzer in Hospital Likas.	
Table 4.2	Socio-demographic characteristics of respondents.	61
Table 4.3	Prevalence of anaemia by ethnicity of respondents.	63
Table 4.4	Prevalence of anaemia based on quantitative haemoglobin	64
	measurements	
Table 4.5	Types and duration of supplement intake.	67
Table 4.6	Independent factors for anaemia.	71
Table 4.7	Types of diet consumed by respondents.	72
Table 4.8	Percentage of energy contribution from macronutrients.	75
Table 4.9	Mean energy and nutrients intake of respondents and their	77
	respective 95% confidence interval.	
Table 4.10	Prevalence of anaemia among respondents with different	83
	Calcium intake.	
Table 4.11	Dietary intake habits among respondents.	91



LIST OF ABBREVIATIONS

BMI	Body mass index
DHS	Demographic and Health Surveys
FFQ	Food frequency questionnaire
Hb	Haemoglobin
Hct	Haematocrit
IDA	Iron deficiency anaemia
MANS	Malaysian Adults Nutrition Survey
MCV	Mean corpuscular volume
MEASURE	Monitoring and evaluation to assess and use results
NCCFN	National Coordinating Committee on Food and Nutrition
RNI	Recommended Nutrient Intake
SF	Serum ferritin
ТІВС	Total iron binding capacity
WHO	World Health Organization



.

LIST OF SYMBOLS

%	Percentage
g	Gram
mg	Milligram
ml	Milliliter
kcal	kilocalorie
n	Number of respondents
<	Less than
≥	Greater or equal to



LIST OF APPENDICES

		Page
Appendix A	Haemoglobin concentration cutoffs to diagnose anaemia	106
	at sea level (g/dl)	
Appendix B	Classification of public health significance of anaemia in	107
	population on the basis of prevalence estimated from blood	
	levels of haemoglobin	
Appendix C	Medical research ethics approval letter	108
Appendix D	Study information sheet	110
Appendix E	Subject information sheet	112
Appendix F	Informed consent form	114
Appendix G	Socio-demographic and Semi-quantitative food frequency	115
	questionnaire	
Appendix H	Borang maklumat kajian	128
Appendix I	Borang maklumat subjek	130
Appendix J	Borang makluman persetujuan	132
Appendix K	Borang soal selidik sosio-demografik dan kekerapan	134
	pengambilan makanan	
Appendix L	Comparison between haemoglobin concentration taken	148
	using HemoCue B and full blood count analyzer in Hospital	
	Likas	



Chapter 1

Introduction

1.1 Overview

The Malaysian economy has seen a tremendous growth in recent years as it gears towards high income nation by the year 2020. Similar to other developing countries, Malaysia is now facing the double burden of malnutrition where cases of undernutrition coexist with increasing prevalence of over-nutrition in its population (Khor and Zalilah, 2003). Several local studies indicated that iron nutritional problem is still a concern among high risk groups including children, adolescents, pregnant women and elderly (Foo *et al.*, 2004; Rosline *et al.*, 2005; Ngui *et al.*, 2011).

Iron nutritional status refers to the equilibrium of iron absorption, storage and utilization in human body. Individual iron status is classified based on the amount of iron store, starting from negative equilibrium and all the way to positive equilibrium of iron store. When presented in ascending order, they are iron deficiency anaemia (IDA), iron deficiency without anaemia, normal iron store and lastly iron overload (WHO, 2001). However, iron status is normally skewed to the negative end of the scale as a result of inadequate dietary iron intake, low bioavailability of iron and frequent blood loss. In fact, iron deficiency anaemia (IDA) is the most common nutritional problem globally, affecting approximately 40% of the world population (WHO, 2011).

Iron is an essential micronutrient required by human body for a battery of metabolic activities and functions. For instance, it is needed for the synthesis of iron-containing enzymes, cytochrome P450 and plays a vital role in electron transport chains as electron carrier (Gibson, 2005). However, most important function of iron is perhaps in the synthesis of haemoglobin for aerobic respiration (Pettit *et al.*, 2011). Compromised iron supply would result in decrease haemoglobin synthesis, thereby



leads to the development of microcytic anaemia (Zimmermann and Hurrell, 2007). Dietary iron can be derived in its haem form from animal sources such as meat, fish, poultry and egg. Apart from these, the iron can also be acquired in its non-haem form from plants such as dark green leafy vegetables, legumes and fruits.

Anaemia can be defined as a health condition where insufficient mass erythrocytes are synthesized in the body as a result of low haemoglobin level (WHO, 2011). Across the world, approximately 41.8% of pregnant women and 30.2% of non-pregnant females were suffering from varying degree of anaemia. The statistics was even more staggering in Asia where 68% of non-pregnant women were reported to be anaemic (Badham, 2007). In view of this, the World Health Organization (WHO) pledged to strengthened its commitment in cutting down the prevalence of anaemia in females of reproductive age by half by the year 2025 (Stevens *et al.*, 2013). Interestingly, 50% of the reported anaemia cases were caused by iron deficiency alone, making it a major risk factor for the health condition (WHO, 2008).

Females generally face greater risk for iron deficiency or iron deficiency anaemia (IDA). This is especially so for women age 15-49 years who are classified as females of reproductive age by the World Health Organization (WHO and MEASURE-DHS, 2004). The demand for iron during puberty and pregnancy (Hanafi *et al.*, 2013). On top of that, there is also a need to compensate daily obligatory iron loss of 0.6mg/d as well as menstrual blood loss (Patterson *et al.*, 1998). Menstrual blood loss between 30-40ml per cycle can result in an iron loss between 0.4-0.5mg/day (Gibson, 2005). Other risk factors for iron deficiency in females include parasitic infection, inflammation and the usage of intrauterine birth control devices (Zimmermann and Hurrell, 2007). Haemoglobin concentration below 12g/dl in non-pregnant women and 11mg/dl in pregnant women serve as indicator for anaemia (WHO, 2011a). Anaemia can be further classified based on the degree of severity. The specific cutoffs for each category are listed in Appendix I.

Assessment of iron deficiency anaemia (IDA) at population level is often conducted using haemoglobin level as reference cutoff similar to that of anaemia (WHO, 2011a). Besides, WHO (2007) also recognized the estimation of prevalence



of iron deficiency anaemia (IDA) using the prevalence of anaemia as a proxy as it has been known to cause half of the anaemia cases.

Apart from the physiological conditions, individual iron nutritional status is also associated with dietary patterns and habits. Traditionally, vegetarian diet has been linked to greater risk for iron nutritional problem due to its reliance on nonhaem iron and high exposure to various iron inhibitors. Achieving the recommended dietary iron intake does not guarantee one from having a healthy iron store as demonstrated by the lack of correlation between dietary iron intake and serum ferritin level (Yen *et al.*, 2008).

Iron deficiency has been demonstrated to affect cognitive functions in young children, immunity, individual endurance level, reproductive health and causes anaemia (Patterson *et. al.*, 1998). Fatigue, shortness of breath and impaired work capacity are some of the early symptoms of the micronutrient deficiency (Gibson, 2005). As the condition progresses, other clinical symptoms including angular stomatitis, dysphagia, hypochlorhydria, spoon nails and anemia may developed. Iron deficiency is also associated with higher risk of maternal and perinatal mortality in both moderate and severe cases of IDA (RNI, 2005). Low birth weight of infant is also common among women with IDA.

Looking from the economic perspective, decrease in cognitive functions and productivity may translate into significant economic loss. Horton and Ross (2002) reported that the annual physical productivity losses due to iron deficiency were approximately \$2.32 per capita. This amounted to about 0.57% of a country's gross domestic product (GDP) (Horton and Ross, 2002). A substantial amount of money was spent by authorities worldwide to address the issue of iron deficiency through healthcare system and food supply chain. In United States for instance, \$0.10 to \$1.00 was spent food fortification per person annually (Badham *et al.*, 2007).

At the other end of the scale, excessive iron intake or iron overload is also part of the iron nutritional problem though its prevalence is lower than iron deficiency. Iron overload or haemochromatosis can arise as a result of excessive iron supplement



intake as well as genetic diseases such as mutation at iron receptor HFE-1 (Gibson, 2005). Injection of therapeutic iron and frequent blood transfusion especially in patient with thalassemia are also the risk factors for iron overload. One of the negative impacts of excessive iron intake is the increase risk for cardiovascular disease (Sabate, 2001). Free iron catalyzed Fenton reaction is responsible for the synthesis of oxygen radicals. Free radicals have been known to cause tissue damage and possibly DNA mutation (Kohlstadt, 2013).

1.2 Problem statement

Data on iron nutritional status among females of reproductive age in Malaysia remains limited despite the fact that they are facing higher risk of iron deficiency. Studies conducted in the past indicated that Asian diets generally contain higher amount of iron absorption inhibitors such as phytic acids due to greater consumption of grains and legumes. Thus, Malaysia as an Asian country may also face similar nutritional problem. On top of that, data on the degree of effects of these iron inhibitors in Malaysian food to individual iron nutritional status is also limited.

1.3 Study objectives

Three objectives have been identified and delineated below:

- 1. To measure iron nutritional status and dietary intake among female students of Universiti Malaysia Sabah.
- 2. To determine the differences in mean dietary iron intake between students with and without anaemia.
- 3. To identify major sources of dietary iron among female students of University Malaysia Sabah.



1.4 Study rationale

Iron deficiency anaemia being the most common nutritional problem worldwide has received a wealth of attention from health practitioners across the world (WHO, 2008). However, focus had always been placed on young children, pregnant women and female athletes. At the same time, there is limited information on iron nutritional status of female undergraduate students in Malaysia. Optimum iron nutrition is crucial in students as compromised iron nutritional status causes fatigue and decreased cognitive performance.

The proposed study would assess female university students' iron nutritional status as well determining its relationship with dietary intake pattern. The outcome of the study may provide insights to the prevalence of iron deficiency among young Malaysian females and appropriate measures can be planned to address the issue at the population level.

1.5 Hypothesis

It is hypothesized that dietary iron intake among female students with anaemia is significantly lower than their non-anaemic counterparts and that grains as well as other cereal products are the main sources of dietary iron amongst these students.

The hypothesis was derived from the available literature which indicated higher prevalence of iron deficiency among females of reproductive age due to physiological characteristics and dietary patterns. This is especially in Malaysia where the main sources of dietary iron come from grains and legumes. Absorption of non-haem iron in these food is greatly influenced by the presence of iron inhibitors found abundantly in Asian diet. Subjects with iron deficiency are anticipated to be reflected through lower than normal haemoglobin level among the female subjects (<12g/dl) (WHO, 2011a).



Chapter 2

Literature Review

2.1 Introduction

The indispensable functions of iron in human body had led to extensive studies of the micronutrient in the past. Unlike other nutrients, the body does not possess a specific set of metabolism to excrete excess iron. As a result, iron is carefully regulated via the dietary iron absorption by the epithelial lining of duodenum (Gubler, 1956).

The human body acquires most of the iron (90%) from internal sources such as degradation of worn out erythrocytes every 120 days. Despite so, dietary iron absorption is still being considered of great importance as it compensates obligatory iron loss as well as menstrual blood loss in females of reproductive age. Females of reproductive age are at greater risk for iron nutritional problems due to iron loss induced by obligatory and menstrual blood loss (WHO, 2001).

The absorption of dietary iron is influenced by a several factors related to diet pattern, bioavailability of iron, iron absorption inhibitors and promoters as well as individual health. These factors would be discussed further in this chapter. Particular interests are placed on vegetarian diet and females in reproductive age group in light of the present study objectives. Association of vegetarian diet with higher risk of iron deficiency is often read about and understandable considering the higher dependency on non-haem iron sources as well as greater exposure to iron inhibitors.



2.2 Iron metabolism and bioavailability

2.2.1 Dietary iron absorption metabolism

Iron absorption occurs throughout the entire length of gastrointestinal tract with higher sensitivity at the duodenum region (Brown, 1963). In order to compensate for the lack of iron excretion metabolism, iron absorption is carefully regulated where only 10% of ingested dietary iron is absorbed into the body. It has been theorized that this is part of the protective mechanism against pathogens as increase virulence was observed when free iron atoms are available (Badham, 2007).

Non-haem iron naturally occurs as ferric salts in plants, animals and dairy products. However, trivalent ferric iron in the salts is not readily absorbed by the intestinal mucosa (Gubler, 1956). As a result, the bioavailability of non-haem iron is much lower than haem iron despite accounting for approximately 80% of dietary iron consumed by human. The reduction of ferric iron (Fe³⁺) to ferrous iron (Fe²⁺) is catalyzed by enzyme ferrireductase (Dcytb) located at the duodenum mucosa (Schmaier and Lazarus, 2011). Acidic condition in the duodenum region greatly favors the reduction process and at the same time inhibiting the dissociation of iron complexes (Gubler, 1956).

Reduced ferrous iron atoms are then transported across the enterocytes via iron transporter known as divalent metal transporter-1 (DMT-1) together with a proton. The bioavailability of non-haem iron can also be interfered by iron absorption inhibitors and individual iron status (Schmaier and Lazarus, 2011). Examples of iron inhibitors include phytate, dietary fibre, polyphenols, calcium and phosphorus.

Unlike non-haem iron, the absorption mechanism of haem iron is yet to be fully established (Anderson *et al.*, 2005). Haem iron derived from animals tissues is absorbed directly into the intestinal mucosa together with protoporphyrin rings attached (Beals, 2013). Degradation of haem moiety within the enterocyte's cytoplasm frees iron atoms from the protoporphyrin rings, allowing them to enter into blood circulation and bind to serum transferrin. Interestingly, haem iron accounts



References

- Aeberli, I., Hurrell, R.F. and Zimmermann, M.B. 2009. Overweight children have higher circulating hepcidin concentrations and lower iron status but have dietary iron intakes and bioavailability comparable with normal weight children. *International Journal of Obesity.* 33: 1111-1117.
- Anchuetz, S., Rodgers, C.D. & Taylor, A.W. 2010. Meal composition and iron status of experienced male and female distance runners. *Journal of Exercise Science and Fitness.* 8(1): 25-33.
- Anderson, G.J., Frazer, D.M., McKie, A.T., Vulpe, C.D. & Smith, A. 2005. Mechanisms of haem and non-haem iron absorption: Lessons from inherited disorders of iron metabolism. *BioMetals.* **18**: 339-348.
- Arabatzis, K., Barbosa, J.C., Johnson, M., Nieman, D.C., Sherman, K.M. & Shultz, T.D. 1989. Dietary status of Seventh-Day Adventist vegetarian and non-vegetarian elderly women. *Journal of the American Dietetic Association*. **89**(12): 1763-1774.
- Badham, J., Zimmermann, M.B. & Kraemer, K. 2007. *The guidebook nutritional anemia. Switzerland*: Sight and Life Press.
- Baels, K.A. 2013. Nutrition and the female athlete from research to practice: Florida. CRC Press.
- Ball, M.J. & Bartlett, M.A. 1999. Dietary intake and iron status of Australian vegetarian women. *The American Journal of Clinical Nutrition*. **70**: 353-358.
- Barr, F., Brabin, L., Agbaje, S., Buseri, F., Ikimalo, J. & Briggs, N. 1998. Reducing iron deficiency anaemia due to heavy menstrual blood loss in Nigerian rural adolescents. *Public Health Nutrition.* 1(4): 249-257.
- Barton, J.C., Lee, P.L. West, C. & Bottomley, S.S. 2006. Iron overload and prolonged ingestion of iron supplements: clinical features and mutation analysis of hemochromatosis-associated genes in four cases. *Journal of HematologyI.* 81: 760-767.
- Beals, K.A. 2013. Nutrition and the female athlete from research to practice: United States: CRC Press.
- Beck, N. 2009. Diagnostic Hematology. London: Springer-Verlag London.
- Bendich, A. 2001. Calcium supplementation and iron status of females. *Nutrition*. 17: 46-51.



- Bjorn-Rassmussen, E. & Hallberg, L. 1979. Effect of animal protein on the absorption of iron in man. Applications of the two-pool extrinsic tag method to measure haem and non-haem iron absorption from the whole diet. *Journal of Clinical Investigation*. 53: 247-255.
- Blattner, R.J. 1948. Comments on current literature: megaloblastic anemia. *Journal of Pediatrics*. **32**(4): 473-474.
- Brown, E.B. 1963. The absorption of iron. *The American Journal of Clinical Nutrition*. **12**: 205-213.
- Burger, S. & Pierre-Louis, J. 2003. *A procedure to estimate the accuracy and reliability of HemoCuetm measurements of survey workers*. Washington: International Life Sciences Institutes.
- Celada, A., Rudoff, H. & Donath, A. 1987. Effect of a single ingestion of alcohol on iron absorption. *American Journal of Hematology*. **5**: 225-237.
- Chapmann, R.W., Morgan, M.Y., Boss, A.M. & Sherlock, S. 1983. Acute and chronic effects of alcohol on iron absorption. *Digestive Diseases and Sciences*. **28**(4): 321-327.
- Charlton, R.W., Jacobs, P., Seftel, H. & Bothwell, T.H. 1964. Effect of alcohol on iron absorption. *British Medical Journal.* 2: 1427-1429.
- Chin, Y.S. & Mohd Nasir, M.T. 2009. Eating behaviours among female adolescents in Kuantan District, Pahang, Malaysia. *Pakistan Journal of Nutrition*. 8(4): 432-432.
- Craig, W.J. 1994. Iron status of vegetarians. *The American Journal of Clinical Nutrition*. **59**: 1233S-1237S.
- Daniel, J. 2011. *Sampling essentials: Practical guidelines for making sampling choices*. United States: SAGE Publications Inc.
- Daniel, W.W. 1999. *Biostatistics: A foundation for analysis in the health sciences* (7th edition). New York: John Wiley & Sons.
- Donovan, U.M. & Gibson, R. 1996. Dietary intakes of adolescent females consuming vegetarian, semi-vegetarian, and omnivorous diets. *Journal of Adolescent Health*. 18: 292-300.
- Drewnowski, A. & Specter, S.E. 2004. Poverty and obesity: the role of energy density and energy costs. *The American Journal of Clinical Nutrition*. **79**(6): 6-16.



- Free, A.H. & Bing, F.C. 1940. Wheat as a dietary source of iron. *Journal of Nutrition*. **72**(1): 449-460.
- Foo, L.H., Khor, G.L., Tee, E. & Prabakaran, D. 2004. Determinants of iron status in Malaysian adolescents from a rural community. *International Journal of Food Sciences and Nutrition*. **55**(6): 517-525.
- Garson, G.D. 2012. *Sampling*. Carolina: G. David Garson and Statistical Associates Publishing.
- Greger, J.L. 1982. Effect of phosphorus-containing compounds on iron and zinc utilization: a review of the literature. *American chemistry Society Symposium*. **203**: 107-120.
- Gibney, M.J., Margetts, B.M., Kearney, J.M. & Arab, L. 2004. *Public Health Nutrition*. United States: Wiley-Blackwell.
- Gibson, R.S. 2005. *Principles of Nutritional Assessment* (2nd Edition). New York: Oxford University Press Inc.
- Grases, F., Simonet, B.M., Prieto, R.M. & March, J.G. 2001. Phytate levels in diverse rat tissues: influence of dietary phytate. *British Journal of Nutrition*. **86**: 225-231.
- Gropper, S.S., Ken, S. & Barksdale, J.M. 2003. Non-anemic iron deficiency, oral iron supplementation, and oxidative damage in college-aged females. *Biological Trace Elements Research*. **109**: 1-13.
- Gubler, C.J. 1956. Absorption and metabolism of iron. Science. 123(3186): 87-90.
- Hambridge, M. 2003. Biomakers of trace minerals intake and status. *The Journal of Nutrition.* **106**: 948S-955S.
- Hahn, P.F., Bale, W.F., Ross, J.F., Balfour, W.M. & Whipple, G.H. 1943. Radioactive iron absorption by gastro-intestinal tract. *Journal of Experimental Medicine*. **78**(3): 169-188.
- Hallberg, L. & Rossander, L. 1984. Improvement of iron nutrition in developing countries: comparison of adding meat, soy protein, ascorbic acid, citric acid, and ferrous sulphate on iron absorption from a simple Latin American type of meal. *The American Journal of Clinical Nutrition*. **39**: 577-583.
- Hallberg, L. 1998. Daily iron supplementation: why is it necessary?. *American Journal of Clinical Nutrition*. **68**: 213-217.



- Hanafi, M.I., Abdallah, A.R. & Zaky, A. In press. Study of hemoglobin level and body mass index among preparatory year female students at Taibah University, Kingdom of Saudi Arabia. *Journal of Taibah University Medical Sciences*. 13: 122-130
- Harvey, L.J., Armah, C.N., Dainty, J.R., Foxall, R.J., Langford, N.J., Fairweather-Tait, S.J. 2004. Impact of menstrual blood loss on iron deficiency among women in the UK. *British Journal of Nutrition.* **94**: 557-564.
- Hassan, R., Abdullah, W.Z. & Nik Hussain, N.H. 2005. Anemia and iron status of Malays women attending an antenatal clinic in Kubang Kerian, Kelantan, Malaysia. *Southeast Asian Journal of Tropical Medicine and Public Health*. **36**(5): 1304-1307.
- Horton, S. & Ross, J. 2003. The economics of iron deficiency. Food Policy. 28: 51-75.
- Huang, Y., Lin, W., Cheng, C. & Su, K. 1999. Nutrient intakes and iron status of healthy young vegetarian and nonvegetarians. *Nutrition Research*. **19**(5): 663-674.
- Hurrell, R. & Egli, I. 2010. Iron bioavailability and dietary reference values. *The American Journal of Clinical Nutrition*. **91**: 1461S-1467S.
- Hurrell, R.F., Manju, B., Juillerat, M., Cook, J.D. 2006. Meat protein fractions enhances non-haem iron absorption. *Journal of Nutrition.* **136**(11): 2808-2812.
- Hurrell, R.F., Reddy, M. & Cook, J.D. 1999. Inhibition of non-haem iron absorption in man by polyphenolic-containing beverages. *British Journal of Nutrition.* **81**: 289-295.
- Ismail, M., Loh, S.P., Omar, H., Abdullah, A.S. & Dahalan, R. 2001. Bioavailability of iron from Malaysian foods using Caco-2 culture system. *Annals of Nutrition and Metabolism.* 45(S1): 49.
- Johnson-Spear, M.A.. & Yip, R. 1994. Hemoglobin difference between black and white women with comparable iron status: justification for race-specific anemia criteria. *The American Journal of Clinical Nutrition*. **60**: 117-121.
- Kandiah, M., Zalilah, M.S., Chan, Y.M. & Hazizi, A.S. 2007. *Handbook on Nutritional Assessment Methods*. Malaysia: August Publishing.
- Khor, G.L. & Zalilah, M.S. 2003. Dual forms of malnutrition in the same households in Malaysia-a case study among Malays rural households. *Asia Pacific Journal of Clinical Nutrition.* **12**(3): 427-238.



- Khor, G.L., Cobiac, L. & Skrzypiec, G. 2002. Gender differences in eating behaviour and social self concept among Malaysian university students. *Malaysian Journal of Nutrition.* **8**(1): 75-98.
- Kim, H, House, W.A. & Miller, D.D. 2004. Habitual tea consumption protects against the inhibitory effects of tea on iron absorption in rats. *Nutrition Research.* **24**: 383-393.
- Kohlstadt, I. 2013. Advancing Medicine with Food and Nutrients (2nd Edition). United States: CRC Press.
- Labbe, R.F., Dewanji, A. & Mclaughlin, K. 1999. Observations on the zinc protoporphyrin/heme ratio in whole blood. *Clinical Chemistry*. **45**: 146-148.
- Loh, S.P. & Khor, G.L. 2010. Iron intake and iron deficiency anaemia among young women in Kuala Lumpur. *Malaysian Journal of Medicine and Health Sciences*. 6(1): 63-70.
- Looker, A.C., Dallman, P.R., Carroll, M.D., Gunter, E.W. & Johnson, C.L. 1997. Prevalence of iron deficiency in the United States. *Journal of the American Medical Association*. 227: 973-976.
- Mahan, L.K., Escott-Stump, S. & Raymond, J.L. 2012. *Krause's Food and the Nutrition Care Process* (13th Edition). United States: Elsevier.
- McClung, J.P. 2012. Iron status and the female athlete. *Journal of Trace Elements in Medicine and Biology*. 26: 124-126.
- Mehdad, A., Siqueira, E.M.A. & Arruda, S.F. 2010. Effect of vitamin A deficiency on iron bioavailability. *Journal of Nutrition & Metabolism*, **57**(35): 35-39.
- Miles, M.P., Keller, J.M., Kordick, L.K. & Kidd, J.R. 2012. Basal, circadian, and acute inflammation in normal versus overweight men. *Medicine & Science in Sports & Exercise*. **44**(12): 2290-2298.
- Mirnalini, K., Zalilah M.S., Safiah, M.Y., Tahir, A., Siti Haslinda, M.D., Siti Rohana, D., Khairul Zarina, M.Y. Mohd Hasyami, S. & Normah, H. 2008. Energy and nutrient intakes: findings from the Malaysian Adult Nutrition Survey (MANS). *Malaysian Journal of Nutrition*. 14(1): 1-24.
- Monsen, E.R. & Cook, J.D. 1976. Food iron absorption in human subjects. IV. The effects of calcium and phosphate slats on the absorption of nonheme iron. *American Journal of Clinical Nutrition*. **29**(10): 1142-1148.



- Morris, E.R. & Ellis, R. 1976. Isolation of monoferric phytate from wheat bran and its biological value as an iron source to the rat. *Journal of Nutrition*. **106**: 753.
- Mulvihill, B., Fidelma, Kirwan, F.M., Morrissey, P.A. & Flynn, A. 1998. Effect of myofibrillar muscle protein on the *in vitro* bioavailability of non-haem iron. *International Journal of Food Sciences and Nutrition*. **49**: 187-192.
- Nair, K.M. & Iyengar, V. 2009. Iron content, bioavailability & factors affecting iron status of Indians. *Indian Journal of Medical Research*. **130**: 634-645.
- National Coordinating Committee for Food and Nutrition. 2005. Recommended Nutrient Intakes for Malaysia. Kuala Lumpur: Ministry of Health Malaysia.
- NCCLS (National Committee on Clinical Laboratory Standards). 1996. Erythrocytes protoporphyrin testing: Approved guideline. C42-A. NCCLS. Vilanova, PA.
- Ngui, R., Lim, Y.A.L., Liam, C.K., Chow, S.C. & Jaffar, S. 2012. Association between anaemia, iron deficiency, neglected parasitic infections and socioeconomic factors in rural children of West Malaysia. *Neglected Tropical Diseases*. **6**(3): e1550-e1558.
- Norhaizan, M.E. & Nor Faizadatul Ain, A.W. 2009. Determination of phytate, iron, zinc, calcium contents and their molar ratios in commonly consumed and prepared food in Malaysia. *Malaysian Journal of Nutrition.* **15**(2): 213-222.
- Norimah, A.K. & Margetts, B.M. 1997. Calibration of a food frequency questionnaire developed for the South Asian community in the United Kingdom. *Malaysian Journal of Nutrition.* **3**: 49-60.
- Norimah, A.K., Jamal, K., Siti Haslinda., Zuhaida, H., Rohida, S., Fatimah, S., Siti Norazlini., Poh, B.K., Kandiah, M., Zalilah, M.S., Wan Manan, W.M., Fatimah, S. & Azmi, M.Y. Food consumption patterns: findings from the Malaysian Adult Nutrition Survey (MANS). *Malaysian Journal of Nutrition*. **14**(1): 25-39.
- Oberleas, D. & Harland, B.F. 1986. Anion-exchange method for determination of phytate in foods: Collaborative study. *Journal of Association of Analytical Chemistry*. **64**(4): 667-669.
- Olsen, C. & St. George, D.M.M. 2004. *Cross-sectional studies and data analysis*. Iowa: College Entrance Examination Board.
- Patterson, A.J., Brown, W.J. & Roberts, D.C.K. 1998. Development, prevention and treatement of iron deficiency in women. *Nutrition Research*. **18**(3): 489-502.



1

- Peng, L.S., Omar, H., Abdullah, A.S., Dahalan, R. & Ismail, M. 2005. The effect of calcium, ascorbic acid and tannic acid on iron availability from *Arthrospira platensisi* by Caco-2 Cell Model. *Malaysian Journal of Nutrition*. **11**(2): 177-188.
- Pilch, S.M. & Senti, F.R. 1984. Assessment of the iron nutritional status of the U.S. population based on data collected in the second National Health and Nutrition Examination Survey 1976-1980. Life Sciences Research Office United States: Federation of the American Societies for Experimental Biology.
- Pollard, J., Kirk, S.F.L. & Cade, J.E. 2002. Factors affecting food choice in relation to fruit and vegetable intake: a review, *Nutrition Research Reviews*. **15**: 373-387.
- Rae, C., Furlong, W., Horsman, J., Pullenayegum, R., Demers, C., St-Louis, J., Lilicrap, D. & Barr, R. 2013. Bleeding disorders, menorrhagia and iron deficiency: impacts on health-related quality of life. *Haemophilia*, **19**: 385-391.
- Reddy, S. & Sanders, T.A.B. 1990. Haematological studies on pre-menopausal Indian and Caucasian vegetarians compared with Caucasian omnivores. *British Journal of nutrition.* **64**: 331-338.
- Reinhold, J., Garcia, L.P.M., Arias-Amando, L. & Garyon, P. 1982. *Dietary fibre0iron interactions: fibre-modified uptakes of iron by segments of rat intestine. Dietary Fiber in Health and Disease.* New York: Plenum Press.
- Roberts, P.D., StJohn, J.B., Stewart, S., Baird, I.M., Coghill, N.F. & Morgan, J.O. 1993. Apparent folate deficiency in iron-deficiency anaemia. *British Journal of Haematology*. **30**: 165-176.
- Sabate, J. 2001. Vegetarian Nutrition. Florida: CRC Press.
- Schlesier, K., Kuhn, B., Kiehntopf, M., Winnefeld, K., Roskos, M., Bitsch, R. & Bohm, V. 2012. Comparative evaluation of green and black tea consumption on the iron status of omnivorous and vegetarian people. *Food Research International.* 46: 522-527.
- Schmaier, A.H. & Lazarus, H.M. 2011. Concise guide to hematology. United States: Wiley-Blackwell.
- Senewirate, B., Hettiarachchi, J. & Senewiratne, K. 1974. Vitamin B12 absorption in megaloblastic anaemia. *British Journal of Nutrition.* **32**: 491-501.
- Shamsuddin, A.M. 19999. Metabolism and cellular functions of IP6: a review. *Anticancer Research*. **19**(5A): 3733-3736.



- Shimi, G. & Hasnah, H. 2013. Does cooking affect the phytate content in local soy based dishes. *International Food Research Journal*. **20**(5): 2873-2880.
- Snyder, A.C., Dvorak, L.L. & Roepke, J.B. 1989. Influence of dietary iron source on measures of iron status among female runners. *Medicine and Science in Sports and Exercise*, 21(1): 7-10.
- Subar, A.F. 2004. Developing dietary assessment tool. *Journal of Dietetic Association*. **104**: 769-770.
- Tee, E., Kandiah, M., Norimah, A., Chong, S., Satgunasingam, N., Kamarudin, L & Milani, S. 1999. School-administered weekly iron-folate supplements improve hemoglobin and ferritin concentration in Malaysian adolescent girls. *The American Journal of Clinical Nutrition*. 69: 1249-1256.
- Urrechaga, E., Borque, L. & Escanero, J.F. 2011. Erythrocyte and reticulocyte parameters in iron deficiency and thalassemia. *Journal of Clinical Laboratory Analysis*. **25**: 223-228.
- Von Schenck, H., Falkensson, M. & Lundberg, B. 1986. Evaluation of 'HemoCue', a new device for determining hemoglobin. *Clinical Chemistry*. **331**: 526-529.
- Wauben, I.P.M. & Atkinson, S.A. 1999. Calcium does not inhibit iron absorption or alter iron status in infant piglets adapted to a high calcium diet. *Nutrient Interactions* and Toxicity. **129**(3): 707-711.
- Weinsier, R. 2000. Use of the term vegetarian. *American Journal of Clinical Nutrition*. **71**(5): 1211-1212.
- WHO (World Health Organization) & Centre of Diseases Control (CDC). 2007. Assessing the iron status of population. Geneva: WHO Press.
- WHO (World Health Organization) & MEASURE *DHS+*. 2004. *Infecundity, infertility, and childlessness in developing countries.* Geneva: WHO Press.
- WHO (World Health Organization). 2001. *Iron deficiency anaemia: Assessment, prevention and control: A guide for programme manager*. Geneva: World Health Organization.
- WHO (World Health Organization). 2004. *Iron deficiency anaemia: assessment, prevention and control-A guide for programme managers*. Geneva: WHO Press.
- WHO (World Health Organization). 2008. *Worldwide prevalence of anaemia 1993-2005 WHO global database on Anaemia.* Geneva: WHO Press.

