ASSESSMENT OF THE ROLE OF DAIRY FOODS IN DIABETES AND LACTOSE INTOLERANCE INTERVENTION

By

ASMAA AHMED ESMAT MOHAMED

B.Sc.Agric.Sc. (Dariy Science and Technology), Ain shams University, 2006

A thesis submitted in partial fulfillment of the requirements for the degree of

MASTER OF SCIENCE in Agricultural Science (Dairy Science and Technology)

> Department of Food Science Faculty of Agriculture Ain Shams University

Approval Sheet

ASSESSMENT OF THE ROLE OF DAIRY FOODS IN DIABETES AND LACTOSE INTOLERANCE INTERVENTION

By

ASMAA AHMED ESMAT MOHAMED

B.Sc. Agric. Sc. (Dairy Science and Technology), Ain-shams University, 2006

This thesis for M.Sc. degree has been approved by:

Dr. Effat Gouda Fouad Prof. of Dairy Science and Technology, Faculty of Agriculture, Alexandria University.
Dr. Amaal Ahmed Mohamed Hassan Prof. of Food Science and Technology, Faculty of Agriculture, Ain Shams University.
Dr. Azaa Mahmoud Farahat Associate Prof. of Dairy Science and Technology, Faculty of Agriculture, Ain Shams University.
Dr. Mohamed Abd Allah El- Hofi Prof. of Dairy Science and Technology, Faculty of Agriculture, Ain Shams University.

Date of Examination: 23 /12 /2013

ASSESSMENT OF THE ROLE OF DAIRY FOODS IN DIABETES AND LACTOSE INTOLERANCE INTERVENTION

By

ASMAA AHMED ESMAT MOHAMED

B.Sc.Agric.Sc.(Dariy Science and Technology), Ain-shams University, 2006

Under the supervision of:

Dr. Mohamed Abd Allah EL- Hofi

Prof. of Dairy Science and Technology, Department of Food Science, Faculty of Agriculture, Ain Shams University (Principal Supervisor)

Dr. Amr El-Sayed Metwally

Associate Prof. of Dairy Science and Technology, Department of Food Science, Faculty of Agriculture, Ain Shams University

Dr. Azaa Mahmoud Farahat

Associate Prof. of Dairy Science and Technology, Department of Food Science, Faculty of Agriculture, Ain Shams University

ABSTRACT

Asmaa Ahmed Esmat: Assessment of the Role of Dairy Foods in Diabetes and Lactose Intolerance Intervention. Unpublished M.Sc. Thesis, Department of Food Science, Faculty of Agriculture, Ain Shams University, 2014

Diabetes mellitus (DM) is one of the most common metabolic diseases. Milk and milk product play an important role in the human diet. The aim of this study is to throughout some light on Intervention between diabetic and lactose intolerance as carbohydrate disorder and to assess the impact of selected dairy products on blood glucose. The glycemic index was determined. Milk proteins (whey, casein, skim milk, and lactose) were evaluated concerning their influence on postprandial responses of glucose. Pasteurized skimmed milk, whey were tested as a drink, whereas casein was administrated in the form of cheese (kariech type).

Second part was designed to examine the effect of administration of dietary lactose and cow's milk in healthy and alloxan-induced diabetic rats. On the body weight, serum biochemical profiles (serum glucose, triglycerides, cholesterol, HDL, LDL, VLDL, HTR % and AI), intestinal lactase activity and histopathological examination of pancreas tissue was. One hundred and forty-five male albino rats were randomly divided into six groups each of thirty six rats and were fed on experimental diet for 28 days, G1 (control negative), healthy rats fed on basal diet; G2, healthy rats administrated repeated dose of dietary lactose (5 ml/rat/twice daily); G3, healthy rats administrated repeated dose of cow's milk (5 ml/rat/twice daily); G4 (control positive, alloxanized diabetic rats), fed on basal diet; G5, diabetic rats administrated repeated dose of dietary lactose (5 ml/rat/twice daily); G6, diabetic rats administrated repeated dose of cow's milk (5 ml/rat/twice daily).

Obtained results revealed that, the GI has a value of 100 for glucose, 45 for lactose, 15 for whole buffalo's milk, 28 for whole cow's milk, 22 for skim milk, 24 for yoghurt, 43 for flavored yoghurt, 34 for ice cream, 17 for karish cheese and 10 for whey + whey protein. Glycemic

load of milk and milk products ranged from 3 to 11. Whey + whey protein recorded the lowest GL followed with Whole buffalo's milk and yoghurt gives moderate GL. The highest GL was recorded for flavored yoghurt and ice cream due to its high sugar contents.

A drastic significant (p<0.05) decrease in body weight (-19.79 %) and increase in serum glucose from 87.69 mg/dl to 381.32 mg/dl was obtained in alloxan-induced diabetic rats fed on basal diet (control positive, G4) during 28 days. However, diabetic rats fed either on dietary lactose (G5) or cow's milk (G6) gained 27.64 % and 35.12 % of their initial body weight, respectively.

Feeding the diabetic rats on dietary lactose (G5) and cow's milk (G6) significantly reduced serum glucose level; (184.63 and 165.97 mg/dl, respectively) than those of the (371.10 mg/dl, control positive group, G4).

The rise in blood sugar is accompanied by increase in serum lipid profile. Diabetes rats administrated cow's milk showed more response towards lowering the serum lipid profile than the dietary lactose. Likewise, significant reduction in serum total cholesterol (TC), triglycerides (TG), low density lipoprotein-cholesterol (LDL-c), very low density lipoprotein-cholesterol (VLDL-c) and increase in high density lipoprotein-cholesterol (HDL-c), was observed in diabetic rats fed with either cow's milk or dietary lactose.

Hyperglycemia is also accompanied by increase in TC/HDL-c, LDL-c/HDL-c ratios and atherogenic index (AI) and decrease in HTR %. Diabetic rats administrated either with cow's milk or dietary lactose (G6 and G5) for 28 days showed significant (p<0.05) reduction in TC/HDL-c, LDL-c/HDL-c ratios and AI and significant (p<0.05) increase in HTR%.

A significant increase in the intestinal lactase activity was observed in diabetic rats alloxanized diabetic rats fed on either cow's milk or dietary lactose (G6 and G6) showed a significant reduction in the intestinal lactase activity, however, the reduction in values of the intestinal lactase activity in diabetic rats administrated cow's milk (G6)

was significantly (p<0.05) higher than that administrated dietary lactose (G5).

Histopathological examination showed oral administrated of cow's milk improved the injury in pancreatic tissue and brought back the normal architecture, however administration of dietary lactose improved the injury in pancreatic tissue.

It can be concluded that milk proteins differ in their capacity to stimulate insulin release. Milk proteins have insulinotropic properties; In addition, administration of cow's milk exhibited significant hypoglycemic activity in alloxanized diabetic rats, and also improvement in lipid profile and body weight as well as regeneration of β -cell of pancreatic tissue and so might be of value in treatment of diabetes. In diabetic rats hyperglycemia directly increases intestinal disaccharidase activities, if patients with diabetes have increased intestinal activities of lactase, those with high lactose consumption would be exposed to greater amounts of the monosaccharides glucose and galactose, and this may result in greater difficulty in achieving adequate blood glucose control, on the other hand lactose intolerance as carbohydrate disorder will be disappeared in diabetic patients.

Key Words: Alloxan, Cow's milk, Diabetic rats, Diabetes, Glycemic index, Lactose, Lactase activity, Lactose intolerance, Proteins.

ACKNOWLEDGEMENT

Ultimate thanks are due to **Allah** who without his aid this work could not be done.

I would like to express my deep gratitude and sincere appreciation to **Prof. Dr. Mohamed Abd Allah El-Hofi,** Professor of Dairy Science and Technology, Department of Food Science, Faculty of Agriculture, Ain Shams University, for his kind help supervision, great helps, valuable guidance and his kind encouragements for me during the present investigation.

Gratefulness and thankfulness are extended to **Dr. Azza Mahmoud Farahat** Associate Professor of Dairy Science and Technology, Department of Food Science, Faculty of Agriculture, Ain Shams University, for her continuous supervision and guidance through the program of the thesis; her kind help and encouragement are greatly appreciated.

Thanks to **Dr. Amr El-Sayed Metwally** Associate Professor of Dairy Science and Technology, Department of Food Science, Faculty of Agriculture, Ain Shams University, for his continuous supervision.

Deep gratitude and sincere appreciation to **Prof. Dr. Amaal Ahmed Mohamed**, Professor of Food Science and Technology, Department of Food Science, Faculty of Agriculture, Ain Shams University, for her guidance and every possible help she kindly offered during this investigation.

I wish also to express my gratitude to **Dr. Safaa Abd El-Aziz Ahmed** Researcher of Food Science, Department of Food Quality Assurance, National Organization for Drug Control and Research, Giza, for her plentiful advice.

Thanks to my friend Marwa Hatem and all staff members of Department of Food Science, Faculty of Agriculture, Ain Shams University, for their continuous support, provides all possible help, both technically and scientifically.

Finally, I would like to thank also my family and my daughter, Gody. Special thanks for my husband, Medhat who has supported me to fulfill my work with no worry.

CONTENTS

	Page
LIST OF TABLES	V
LIST OF FIGURES	IX
LIST OF ABBREVIATIONS	XII
1. INTRODUCTION	1
2. REVIEW OF LITERATURE	4
2.1.Disorders of carbohydrate metabolism	4
2.1.1.Lactose intolerance	4
2.1.2.Sucrose intolerance	5
2.1.3.Galactosemia	5
2.1.4.Diabetes mellitus	6
2.2. Diabetes	6
2.2.1. Definition of diabetes	6
2.2.2. Classification of diabetes	7
2.2.3. Diabetes prevalence	9
2.2.3.1. World wide	9
2.2.3.2. Arab Region	10
2.2.3.3. Egypt	11
2.2.4. Diagnosis and complication of diabetes	11
2.2.5. Milk and diabetes	12
2.2.5.1. Milk protein and diabetes	13
2.2.5.2. Whey	14
2.2.5.3. Dairy peptides in effective blood glucose	
management Human intervention studies with	
protein hydrolysates and combinations thereof	
Protein hydrolysates	16
2.2.6. Dairy consumption and diabetes	21
2.2.7. Vitamin D and diabetes	21
2.2.7. Trans-palmitoleic acid and diabetes	22
2.2.8. Fiber and diabetes	23

2.2.9. Glycemic index	23
2.3. Lactose Intolerance	24
2.3.1. Lactose intolerance prevalence	26
2.3.2. Lactose Intolerance and Diabetes	26
2.3.3. Effect of dietary lactose on intestinal lactase and	
others disaccharides enzymes activity	27
2.3.4. Effect diabetes on others disaccharides enzymes	29
activity	
3. MATERIALS AND METHODS	31
3.1. Materials	31
3.1.1. Milk	31
3.1.2 .Glucose and Lactose	31
3.1.3.Whey	31
3.1.4. Whey protein	31
3.1.5.Karish cheese	31
3.1.6. Ice cream	31
3.1.7. wheat Bran	31
3.1.8. Yoghurt ,Flavored Yoghurt	32
3.1.9.Blood glucose monitoring system	32
3.1.10.Chemicals	32
3.1.11. Experimental animal	32
3.2. Methods	32
3.2.1.Lactose content	32
3.2.2.Protein content	32
3.2.3. Subjects and study design	33
3.2.4. Glycemic Load (GL)	36
3.2.5.Blood analysis	36
3.2.6.The diet experiment	36
3.2.7. Biological experimental design	36
3.2.8.Blood sampling	37
3.2.9. Serum biochemical analysis	38
3.2.9.1. Lipids parameters	38

3.2.10.Gut mucosa	39
3.2.11.Determination of β -D galactosidase (lactase) activity	39
3.2.12. Histological examination	39
3.2.13. Statistical analysis	40
4. RESULTS AND DISCUSSION	41
PART, I Glycemia in Healthy Subjects after Lactose-Equivalent	
Meals of Milk and Milk Products	41
4.1. Effect of milk components on blood glucose level	43
4.1.1. Effect of sugar type	43
4.1.2. Effect of milk	45
4.1.3. Effect of protein on postprandial blood glucose	47
4.1.4. Effects of fat on postprandial blood glucose	52
4.1.5. Effects of bran on postprandial blood glucose	54
4.2. Glycemic index and glycemic load of milk and milk	
products	55
PART, II Biological evaluation of administration of dietary	
lactose or cow's milk in healthy and alloxan-induced	
diabetic rats	68
4.3. Biological evaluation of administration of dietary lactose	
or cow's milk in healthy and alloxan-induced diabetic rats	68
4.3.1. Effect of oral administration of dietary lactose or	
cow's milk on body weight and body weight gain of	
rats	69
4.3.2. Effect of oral administration of dietary lactose or	
cow's milk on serum glucose in healthy and diabetic	
rats	74
4.3.3. Effect of oral administration of dietary lactose or	
cow's milk on serum lipid profile in healthy and	
diabetic rats	78
4.3.3.1. Serum total cholesterol (TC)	78
4.3.3.2. Serum triglycerides level (TG)	91
4.3.3.3. Serum lipoprotein fraction	95

4.3.3.3.1. Low density lipoprotein-cholesterol (LDL-c)	
level	96
4.3.3.3.2. Very low density lipoprotein-cholesterol	
(VLDL-c)	100
4.3.3.3. High density lipoprotein cholesterol (HDL-c)	102
4.3.3.3.4. Lipoprotein fraction ratios TC/HDL-c,	
LDL/HDL-c and HTR (%)	105
4.3.3.5. Atherogenic index (AI)	108
4.3.4. Effect of oral administration of dietary lactose or	
cow's milk on intestinal lactase activity	110
4.3.5. The potential mechanisms of dairy consumption in	
preventing type 2 diabetes	116
4.3.6. Histopathological examination of pancreas tissue of	
healthy, diabetic rats and diabetic rats treated with	
dietary lactose and cow's milk	134
5. SUMMARY AND CONCLUSION	146
6. REFERENCE	152
ARABIC SUMMARY	

LIST OF TABLES

No.	Title	Page
3.1	Nutrient composition and serving size of the test meals and glucose reference meal	34
3.2	Nutrient composition and serving size of the test meals and glucose reference meal	35
3.3	Nutrient composition and serving size of the test meals and glucose reference meal	35
4.1	Blood glucose in response to equal amounts of carbohydrate from a glucose reference meal and test	
4.2	meals of lactose Blood glucose in response to equal amounts of carbohydrate from a glucose reference meal and test meals of whole buffalo milk, skim buffalo milk and cow	44
4.3	milk Blood glucose in response to equal amounts of carbohydrate from a glucose reference meal and test meals of skim milk, whey + whey protein and cheese	46
4.4	lactose Blood glucose in response to equal amounts of carbohydrate from a glucose reference meal and test	47
4.5	meals of whole milk and skim milk Blood glucose in response to equal amounts of carbohydrate from a glucose reference meal and test	53
4.6	meals of whole milk, skim milk, whole milk +bran and skim milk +bran Glycemic index (GI), Glycemic load (GL) and the incremental area under the curve (IAUC) of blood	54
	glucose responses for Buffalo whole milk	59

4.7	Glycemic index (GI), Glycemic load (GL) and the	
	incremental area under the curve (IAUC) of blood	
	glucose responses for Buffalo skim milk	60
4.8	Glycemic index (GI), Glycemic load (GL) and the	
	incremental area under the curve (IAUC) of blood	
	glucose responses for cow milk	61
4.9	Glycemic index (GI), Glycemic load (GL) and the	
	incremental area under the curve (IAUC) of blood	
	glucose responses for yoghurt	62
4.10	Glycemic index (GI), Glycemic load (GL) and the	
	incremental area under the curve (IAUC) of blood	
	glucose responses for flavored yoghurt	63
111	Chromic index (CI) Chromic load (CI) and the	03
4.11	Glycemic index (GI), Glycemic load (GL) and the	
	incremental area under the curve (IAUC) of blood	
	glucose responses for ice cream	64
4.12	Glycemic index (GI), Glycemic load (GL) and the	
	incremental area under the curve (IAUC) of blood	
	glucose responses for lactose	65
4.13	Glycemic index (GI), Glycemic load (GL) and the	
	incremental area under the curve (IAUC) of blood	
	glucose responses for karish cheese +lactose	66
4.14	Glycemic index (GI), Glycemic load (GL) and the	
	incremental area under the curve (IAUC) of blood	
	glucose responses for whey +whey protein	67
4.15	Body weight and body weight gain of healthy and	
	diabetic rats received dietary lactose or cow's milk for 28	
	day	70
4.16	Serum glucose (mg/dl) level of healthy and diabetic rats	
	treated with dietary lactose or cow's milk for 28 days	
	· ·	75

4.17	Serum total cholesterol (mg/dl) level of healthy and diabetic rats treated with dietary lactose or cow's milk for	
	28 days.	79
4.18	Serum triglycerides (mg/dl) level of healthy and diabetic	
	rats treated with dietary lactose or cow's milk for 28 days	92
4.19	Serum LDL-cholesterol (mg/dl) level of healthy and	
	diabetic rats treated with dietary lactose or cow's milk for	
	28 days.	98
4.20	Serum VLDL-cholesterol (mg/dl) level of healthy and	
	diabetic rats treated with dietary lactose or cow's milk for	
	28 days.	101
4.21	Serum HDL-cholesterol (mg/dl) level of healthy and	
	diabetic rats treated with dietary lactose or cow's milk for	
	28 days	103
4.22	Serum total cholesterol/HDL-c ratio, HTR%, LDL/HDL-	
	c and AI in healthy and diabetic rats treated with dietary	
	lactose or cow's milk for 28 days.	106
4.23	Intestinal lactase activity of healthy and diabetic rats	
	treated with dietary lactose or cow's milk for 28 days	112
4.24	Histopathological alterations for the pancreatic tissue of	
	healthy and diabetic rats treated with dietary lactose and	
	cow's milk for 28 days.	143