الآية

بِسْمِ اللَّهِ الرَّحْمَنِ الرَّحِيمِ

:قال تعالى

إِنَّ فِي خَلْقِ السَّمَاوَاتِ وَالْأَرْضِ وَاخْتِلَافِ اللَّيْلِ وَالنَّهَارِ وَالْفُلْكِ الَّتِي تَجْرِي فِي الْبَحْرِ بِمَا يَنْفَعُ النَّاسَ وَمَا أَنْزَلَ اللَّهُ مِنَ السَّمَاءِ مِنْ مَاءٍ فَأُحْيَا بِهِ الْأَرْضَ بَعْدَ مَوْتِهَا وَبَثَّ فِيهَا مِنْ كُلِّ دَابَّةٍ وَتَصْرِيفِ الرِّيَاحِ وَالسَّحَابِ الْمُسَخَّرِ بَيْنَ الشَّمَاءِ وَالْأَرْضِ لَآيَاتٍ لِقَوْمٍ يَعْقِلُونَ

صدق الله العظيم

سورة البقرة الآية 164

Dedication

To my mother

To my father

To my **brothers**

They gave me my life, and everything else in between, but they truly shut me up when the words comes to describe how much I love them and appreciate the efforts they have put into giving me the life I have now.

To my best friends

Who are always in my heart.

Acknowledgements

Firstly we thank **Allah** for His continuous blessing that make this work accomplished and possible.

Then, I would like to express my deep gratitude to **Dr. Amar Mohammed Ismail** who was abundantly helpful and offered invaluable assistance, support and guidance.

I want to extend my deep thanks to **my faculty** for providing the laboratory facilities.

I cannot find word to express my gratitude to **my colleages** for sharing knowledge and assistance

Abstract

A descriptive cross-sectional hospital based study was conducted in the Radiation & Isotopes Centre Khartoum (RICK), during the period from March to August 2013. In order to estimate Vitamin D and phosphate levels in Sudanese patients with breast cancer, sixty histologically confirmed breast cancer patients and thirty apparently healthy females as control group. Vitamin D was estimated using Euroimmun 25-OH vitamin D competitive ELISA kits, phosphate was estimated using phosphomolybdic acid method, fully automated analyzer used for washing and reading processes, spectrophotometer analyzer for reading of phosphate concentration, questionnaire and results data obtained were analyzed using (SPSS version 16). The results showed, that majority of breast cancer patients at age 41-60 which compose 45%. There was decrease in mean of serum vitamin D level among patients compared with control group resulting in (*P*-value=0.000). This study showed that , no significant increase in mean serum phosphate levels between patients in comparison with control group (*P*-value=0.19). The results showed a decrease in vitamin D levels in postmenopause compared with premenaupose breast cancer patients (P-value=0.046). There was no significant decrease in the mean of serum vitamin D and no significant increase in the mean of serum phosphate level in breast cancer patients when classified according to their parity, body mass index and duration of disease, the results showed week negative correlation between vitamin D and phosphate. The study concluded that, there was a decrease in vitamin D level among Sudanese women with breast cancer especially in menopause women. Vitamin D supplement trail to evaluate if breast

cancer patients in Sudan needs vitamin D supplement and

monitoring to avoid vitaminosis is recommended. study the vitamin D and phosphate in correlation with breast cancer stages and metastasis.

اجريت هذه الدراسة بمستشفي الذرة للعلاج بالاشعة النووية بولايه الخرطوم خلال الفترة من مارس الي اغسطس 2013 وذلك بهدف قياس مستوى فيتامين د والفوسفات في مصل مرضي سرطان الثدي .تم جمع ستين عينة دم من مريضات سرطان الثدي بالاضافة الى ثلاثين عينة دم اخري جمعت من نساء اصحاء غير مصابات . قيس معدل فيتامين د باستخدام الطري قة المناعية الانزيمية بينما قيس الفوسفات باستخدام طريقة المولييبدات الفوسفاتية.تم جمع النتائج وحللت البيانات احصائيا حيث اوضحت الدراسة ان خمسة واربعون من النساء المصابات بسرطان الثدي تقع اعمارهن مابين الاربعين والستين عام . كما وجدت الدراسة انخفاضا في متوسط تركيز فيتامين د لدى المرضى خاصة من زادت اعمارهن عن الخمس واربعين. لم تكن هنالك اي فروق معنوية في متوسط تركيز فيتامين د والفوسفات لدى النساء المصابات بسرطان الثدي عندما قسمن الى مجموعات حسب عدد مرات الانجاب ووزنهن وفترة اصابتهن بالمرض . كما وجدت الدراسة علاقة عكسيه غير معنوية بين .فيتامين د الفوسفات

استنادا على هذه النتائج اوصت الدراسة بضرورة اعطاء مرضى سرطان الثدي جرعات فيتامين د مع ضرورة متابعة الجرعة .كما اوصت بمزيد من الدراسات .. فيما يختص بفيتامين د والفوسفات وعلاقتهما بسرطان الثدي

List of Contents

Topic title	Page
	No
الآية	
Dedication	
Acknowledgements	Ш
Abstract	IV
الخلاصة	V
List of content	VI
List of figures	Х
Abbreviations	ΧI
Chapter One	
Introduction and Literature Review	
1.Introduction	1
1.1 Breast	1
1.1.1 The normal Breast morphology	1
1.1.2 Inflammatory disorders of the breast	2
1.1.3 Breast cancer	2
1.1.3.1 Breast cancer epidemiology	2
1.1.3.2 breast cancer risk	2
1.1.3.3 Clinical Presentations	2 3 3
1.1.3.4 Pathology and Prognosis	3
1.1.3.5 Breast cancer development and	3
types	
1.1.3.5.1 Non-invasive breast cancer	4
1.1.3.5.2 Invasive breast cancer	4
1.1.3.6 Breast cancer screening	4
1.1.3.7 Breast cancer treatment	4
1.2 Vitamin D	6
1.2.1 Nomenclature of Vitamin D	6
1.2.2 Vitamin D biochemistry	6
1.2.3 Synthesis of vitamin D	7
1.2.3.1 Photolysis of Cholesterol	7
1.2.3.2 Metabolism in Liver	7
1.2.3.3 Metabolism in Kidneys	8
1.2.4 Vitamin D Absorption:	9

1.2.5 Vitamin D Transport	9	
1.2.6 Action of Vitamin D		
1.2.6.1 Genomic action of Vitamin D	10	
1.2.6.2 Non genomic action of Vitamin D		
1.2.7 Role of Vitamin D in human metabolic	11	
processes		
1.2.8 Regulation of Vitamin D metabolism	12	
1.3 Phosphate	12	
1.3.1 Regulation Phosphate	13	
1.3.2 Distribution of Phosphate	13	
1.3.3 Clinical Applications	14	
1.3.3.1 Hypophosphatemia	14	
1.3.3.2Hyperphosphatemia	14	
1.4 Rationale	15	
1.5 Objectives	15	
1.5.1 General objective	15	
1.5.2 Specific objectives	15	
Chapter Two		
Materials and Methods		
2.1 Study approach and design	17	
2.2 Study area	17	
2.3 Study period	17	
2.4 Study population	17	
2.5 Selection criteria	17	
2.5.1 Inclusion criteria	17	
2.5.2 Exclusion criteria	17	
2.5 Study variables	17	
2.7 Data collection	17	
2.8 Data presentation	17	
2.9 Sample Collection	18	
2.10 Ethical considerations	18	
2.11 Quality control	18	
2.12 Method protocol	18	
2.12.1Measurement of Vitamin D	18	
2.12.1.1 Principle of the test	18	
2.12.1.2 Test procedure	19	
2.12.1.3 Calculation of results	19	
2.12.1.4 Reference rang	20	
2.12.1.5 Cross reactivity	20	
2.12.1.6 Interference	20	

2.12.1.7 Stability	20	
2.12.2 Measurement of serum Phosphate	20	
2.12.2.1Principle of the test	20	
2.12.2.2 Reagents	20	
2.12.2.3 Linearity range	20	
2.12.2.4 Procedure	20	
2.12.2.5 Calculation of results	21	
2.12.2.6 Reference range	21	
2.13 Statistical analysis	21	
Chapter Three		
Results		
3. Results	22	
Chapter Four		
Discussion		
4.1 Discussion	34	
4.2 Conclusion	37	
4.3 Recommendation	37	
References		
References	38	
Appendices		
Appendices	40	

List of figures:

N	Title	Pag e
3. 1	Frequency pie chart shows age of study group expressed as percentage.	22
3. 2	A scatter plot of the correlation between serum phosphate (mg/dl) and Vitamin D (ng/ml)	23
3. 3	shows serum vitamin D concentration in patients and control group	24
3. 4	Shows serum vitamin D concentration in patients classified to group 1 (<45 years) and group 2 (> 45 years).	25
3. 5	Shows mean of serum vitamin D concentration in group of patients classified to group1 (<25 kg/m²) and group 2 (> 25 kg/m²).	26
3. 6	Shows serum vitamin D concentration in groups of patients classified to group1 (< 30 months) and group 2 (> 30 months).	27
3. 7	Shows serum vitamin D concentration in study group classified to group 1 (< 3 Childs) and group 2 (> 3 Childs).	28
3. 8	Shows serum phosphate concentration in patient and control group.	29
3. 9	Shows serum phosphate concentration in study group classified to group 1 (< 45 years) and group 2 (> 45 years).	30
3. 10	Shows serum phosphate concentration in study group classified to group 1 (normal < 25 kg/m²) and group 2 (overweight > < 25 kg/m²).	31
3. 11	Shows phosphate concentration in study group classified to group 1(< 30 months) and group 2 (> 30 months).	32
3. 12	Shows serum phosphate concentration in study group classified to group 1 (<3 Childs) and group 2 (> 3 Childs),.	33

Abbreviations

2, 3 BPG 2, 3 Bisphospho Glycerate

ATP Adenosine Triphosphate

BMI Body Mass Index

BRCA1 and 2 Breast Cancer Associated protein 1

and 2

COPD Chronic Obstructive Pulmonary

Disease

DBP Vitamin D Binding Protein

DCIS Ductal Carcinoma Insitu

DNA Deoxyribonucleic Acid

ER Esterogen Receptor

FNA Fine Needle Aspirate

GMP Guanosine Monophosphate

HRT Hormone Replacement Therapy

ICU Intensive Care Unit

IUPAC International Union of Pure and

Applied Chemistry

LCIS Lobular Carcinoma Insitu

MAARS Membrane Associated Rapid

Response

MAP Mitorgen Activated Protein

MRI Magnetic Resonance Imaging

NADPH Nicotinamide Adenine Dinucleotide

Phosphate-oxidase

P. Value Probability Value

PKC Protein Kinase C

PR Progesterone Receptor

PTA Phosphate Acetyletransferase

RICK Radiation and Isotope Center in

Khartoum

RNA Ribonucleic Acid

SD Standard Deviation

SPSS Statistical Package for Social

Science

TNM Tumor, Node, Metastasis

classification

TPN Total Parental Nutrition

VDR Vitamin D Receptor