

Volume-6, Issue-1, January-March, 2016

ISSN: 2220-7562

# JBUMDC

Recognized by PMDC

The Journal of Bahria University Medical and Dental College



Bahria University Medical & Dental College,  
Adjacent PNS Shifa, DHA Phase II, Karachi.

**JBUMDC**

**ISSN 2220-7562**

**The Journal of Bahria University Medical and Dental College  
Karachi, Pakistan**

**Peer Reviewed Multidisciplinary Quaterly Published Journal  
Indexed with PakMediNet**

**Patron-in-Chief**

Vice Admiral (Retd) Tanveer Faiz HI (M)  
Rector Bahria University, Pakistan.

**Patron**

Vice Admiral (Retd) Tahseen Ullah Khan HI (M)  
Director General Bahria University Medical & Dental College, Karachi.

**Editor-in-Chief**

Shaheen Moin

**Editor**

Nasim Karim

**Associate Editor**

Iqbal Hussain

**Assistant Editors**

Asad Ullah Khan, Irfan Ali Mirza, Kulsoom Fatima

**Members Advisory Board**

Fatema Jawad  
Huma Qureshi

Kamran Hameed  
Khalid Mehmood

Samad Shera  
Syed Tipu Sultan

**Members Editorial Board - National**

Aafia Zafar (AKUH)

Abid Azhar (KIBJE)

Ahmed Danyal (NM&DC)

Ambreen Usmani (BUMDC)

Anis Jaffery (BUMDC)

Hasan Ali (BUMDC)

Imran Shaikh (BUMDC)

Khalida Nasreen (BUMDC)

Khalid Mustafa (BUMDC)

Masood Qureshi (DUHS)

Mehreen Latif (BUMDC)

Mohiuddin Alamgir (BUMDC)

Munawar Ansari (LUMHS)

Mushtaque Memon (BUMDC)

Naheed Sultan (BUMDC)

Nighat Huda (LNH)

Nighat Rukhsana (BUMDC)

Qamar Jamal (ZMU)

Razia Korego (BUMDC)

Saeeda Baig (ZMU)

Sameer Shahid Ameen (BUMDC)

Sajid Abbas Jafri (BUMDC)

Shazia Shakoor (BUMDC)

Shahid Noor (LNH)

Shakeel Ahmed (BUMDC)

Sher Shah Syed (AH)

Zubair Ahmed Abbasi (BUMDC)

**Members Editorial Board - International**

Aamir Omair (KSA)

Ambreen Ahmed (USA)

Farida Habib (KSA)

Irfanullah Siddiqi (KSA)

Mukhtiar Baig (KSA)

Raheela Hafeez (USA)

Sadiqa Syed (KSA)

Shamaun Razi (KSA)

S. Moazzam Zaidi (Newzealand)

**Editorial Assistants**

Tahira Zamir

Arsalan Ahmed

# CONTENTS Volume-6, Issue-1, January-March, 2016

## EDITORIAL

- Global Threat - Zika Virus 01  
Syed Ijaz Hussain

## REVIEW ARTICLE

- Increasing Burden of Abdominal Obesity in Females and its Aftermaths 03  
Khola Noreen, Nadia Khalid, Imran Shaikh

## ORIGINAL ARTICLES

1. Knowledge, Attitude and Practice Regarding Oral Hygiene among Private School Children 09  
Samreen Mazhar, Ashghar Ali, Mahwish Bano, Muhammad Ali Leghari, Aukif Ali Sheikh
2. Efficacy of 50 g Glucose Challenge Test as a Screening Tool for Gestational Diabetes Mellitus 14  
Ayesha Arif, Sarwat Nazar, Sadia Arif
3. Effects of Combined Regimens of Gabapentin and Verapamil with Diazepam on Kindled Model of Epilepsy in Mice. 18  
Itefaq Hussain Qureshi, Shabana Usman Simjee
4. Evaluation of S-T Resolution by Streptokinase Therapy in Patients of Myocardial Infarction among the Age Group of more than 60 Years 23  
Shiekh Nadeem Ahmad, Musarrat Sultana, Fuad Shiekh, Syed Saud Hasan, Shams -ul -Arfeen Qasmi
5. Prevalence of Vitamin- D Deficiency among Women with GDM 30  
Habiba Sharaf, Quratul Ain Zahid
6. The Safety and Efficacy of Percutaneous Trigger Finger Release under Local Anaesthesia 34  
Abdur Rehman Qureshi, Faaiz Ali Shah, Shahab-ud-Din, Wali Mohammad Khan
7. Relationship of Cardiac Disease with Oral Health: A Single Centre Study 38  
Javed Ashraf, Rana Modassir Shamsher Khan, Khawaja Rashid Hassan, Muhammad Rizwan, Ali Saad Tariq Sarah Ashraf
8. Comparative Study of Lipid Profile in Multibacillary and Paucibacillary Leprosy Patients 43  
Ghulam Sarwar, Viqar Sultana, Ali Gul, Jehan Ara
9. Causes of Male and Female Sub Fertility in the Couples who Underwent 'In Vitro Fertilization' at Life Clinic; a Statistical Study from Lahore, Pakistan 47  
Haroon Latif Khan, Yousaf Latif Khan, Nighat Mahmood, Mariam Mustanser, Saba Sardar, Abdul Rahman Khawaja
10. Is Gender Matters In Paediatric Cardiac Surgery 51  
Iqbal Hussain Pathan , Sohail Khan Bangash , Saad Bader Zaki

## COMMENTARY

- Workplace Based Assessment : A Step Towards Competency Based Training 55  
Shafaq Sultana

## STUDENT CORNER

- Oral Health Education Poster Competition 2015 by Department of Community Dentistry at BUMDC 58  
Raima Bashir, Kulsoom Fatima Rizvi

## CASE REPORT

- Endodontic Therapy of Mandibular Canines with Two Canals in a Single Root 60  
Shama Asghar, Mahwesh Hasan, Asghar Ali

## LETTER TO EDITOR

- Medical Brain Drain - An Increasing Social Stigma 63  
Maria Shoaib

## JBUMDC INSTRUCTION TO AUTHORS

64

# Global Threat – Zika Virus

Syed Ijaz Hussain

Zika virus is an emerging mosquito-borne virus that was first identified in Zika forest of Uganda in 1947<sup>1</sup> in rhesus monkeys through a monitoring network of sylvatic yellow fever. It was subsequently identified in humans in 1952 in Uganda and the United Republic of Tanzania<sup>2</sup>. Outbreaks of Zika virus (Figure 1a) disease have been recorded in Africa, America, Asia and Pacific. It belongs to the family of Flavi virus<sup>3</sup> and its vector is Aedes mosquitoes<sup>4</sup> (which usually bite during the morning and late afternoon/evening hours) (Figure 1b). The reservoir is yet unknown.

Figure: 1a<sup>4</sup>

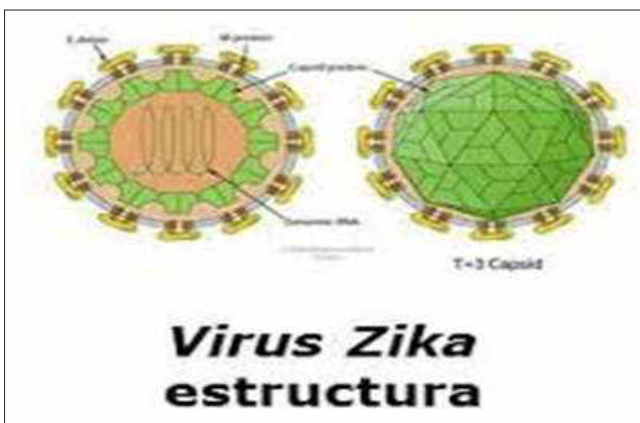


Figure: 1b<sup>4</sup>



The incubation period (the time from exposure to symptoms) is not clear, but is likely to be few days. The symptoms are similar to other arbovirus infections such

as dengue<sup>5</sup>, which include fever, skin rashes, conjunctivitis, muscle and joint pain, malaise, and headache. These symptoms are usually mild and last for 2-7 days. The virus is transmitted to people through the bite of an infected mosquito from the Aedes genus, mainly Aedes aegypti in tropical regions. This is the same mosquito that transmits dengue, chikungunya<sup>6</sup> and yellow fever. Zika virus disease outbreaks were reported for the first time from the Pacific in 2009 and 2013<sup>7</sup> (Yap and French Polynesia, respectively), and in 2015 from America (Brazil and Colombia) and Africa (Cape Verde). In the month of January and February 2016, more than 13 countries including America, Brazil and France reported sporadic spread indicating rapid geographic expansion of this virus. In Colombia it has been reported that 2000 pregnant women are also suffering from this viral disease. The diagnosis is mainly done on the basis of clinical assessment and recent history (e.g. residence or travel to an area where Zika virus is known to be present). Isolation of virus on PCR<sup>8</sup> and other body fluids such as urine and saliva.<sup>9</sup> There is no definite treatment of people suffering from this viral disease and no vaccination is available till to date. The only management is that the patient should be encouraged to take rest, drink plenty of water and to relieve pain and fever by taking analgesic and antipyretic drugs accordingly. During large outbreaks in French Polynesia and Brazil in 2013 and 2014<sup>10</sup> respectively, national health authorities reported potential neurological and auto-immune complications of this virus disease. Recently in Brazil, local health authorities have observed an increase in Guillain-Barré syndrome which coincided with this virus infection in the general public, as well as an increase in babies born with microcephaly in northeast Brazil. Agencies investigating the Zika outbreaks are finding an increasing body of evidence about the link between Zika virus and microcephaly (Figure 2a, 2b). However, more investigation is needed to better understand the relationship between microcephaly in babies and to this virus. Other potential causes are also being investigated.

Figure: 2a<sup>4</sup>



✉ **Dr. Syed Ijaz Hussain**  
Assistant Professor  
Department of Pharmacology  
Bahria University Medical & Dental College  
Karachi.  
Email: col\_zaidi@yahoo.com  
Received: 26-02-2016  
Accepted: 28-02-2016

Figure: 2b<sup>4</sup>



People should be instructed to adhere to the following preventive measures;

1. Rely on reducing mosquitoes through source reduction (removal and modification of breeding sites) and contact between mosquitoes and people.
2. Use insect repellent, wearing clothes (preferably light-coloured) that cover as much of the body as possible; using physical barriers such as screens, closed doors and windows; and sleeping under mosquito nets.
3. Empty, clean or cover containers that can hold water such as buckets, flower pots, so that places where mosquitoes can breed are removed.
4. Give special attention and help to those who may not be able to protect themselves adequately, such as young children, the sick or elderly.
5. During outbreaks, health authorities advise spraying of insecticides to be carried out. Insecticides recommended by the WHO Pesticide Evaluation Scheme may also be used as larvicides to treat relatively large water containers.
6. Travellers should take the basic precautions described above to protect themselves from mosquito bites.

WHO is supporting countries to control this viral disease by:

Defining and prioritizing research regarding this disease by convening experts and partners.

Enhancing surveillance of this virus and potential complications.

Strengthening capacity in risk communication to help countries meet their commitments under the International Health Regulations.

Providing training on clinical management, diagnosis and vector control through a number of WHO Collaborating Centres.

Strengthening the capacity of laboratories to detect the virus.

Supporting health authorities to implement vector control strategies aimed at reducing Aedes mosquito populations such as providing larvicide to treat standing water sites that cannot be treated in other ways, such as cleaning, emptying, and covering them.

Preparing recommendations for clinical care and follow-up of people contracted with this virus, in collaboration with experts and other health agencies.<sup>10</sup>

Measure should be taken through electronic and print media to disseminate information and provide awareness regarding this deleterious virus as prevention is the only armour which we can employ at present to fight against the attack of this virus.

#### REFERENCES:

1. Mitya Underwood, Zika Virus 'new global threat for 2016' *The National* 2016, 1<sup>st</sup> March; sect. World (col- 2)
2. Dick GW, Kitchen SF, Haddock AJ. Zika virus. Isolations and serological specificity. *Trans R Soc Trop Med Hyg.* 1952;46:509-2
3. Duffy MR, Chen TH, Hancock WT, Powers AM. Zika virus outbreak on Yap Island, Federated States of Micronesia. *N Engl J Med.* 2009;360:2536-43 10.1056/NEJMoa0805715
4. Virus Zika - información práctica profesional sanitarios ... [fundacionio.org/264x261](http://fundacionio.org/264x261) Search by image Imágenes de interés sobre Virus Zika: Accessed date 25-02-2016
5. Foy BD, Kobylinski KC, Foy JLC, Blitvich BJ, da Rosa AT, Haddock AD et al. Probable non-vector-borne transmission of Zika virus, Colorado, USA. *Emerg Infect Dis* 2011;14: 880-2
6. Dupont-Rouzeyrol M. *Infect Dis* 2015;21: 381-2
7. Baronti C, Piorkowski G, Charrel RN, Boubis L, Leparc-Goffart I, de Lamballerie X 2013 Complete sequence of Zika virus from a French Polynesia outbreak *Genome Announc.* 2014;2:e00500-14
8. Faye O, Faye O, Dupressoir A, Weidmann M, Ndiaye M, Sall AA. One-step RT-PCR for detection of Zika virus. *J Clin Virol* 2008; 43: 96-101
9. Gourinat AC, O'Connor O, Calvez E, Goarant C, Dupont-Rouzeyrol M. Detection of Zika virus in urine. *Emerg Infect Dis* 2015;21: 84-6
10. Ios S, Mallet HP, Goffart IL, Gauthier V, Cardoso T, Herida M. Current Zika virus epidemiology and recent epidemics. *Med Mal Infect* 2014; 44: 302-7



# Increasing Burden of Abdominal Obesity in Females and its Aftermaths

Khola Noreen<sup>1</sup>, Nadia Khalid<sup>2</sup>, Imran Shaikh<sup>3</sup>

## ABSTRACT:

Obesity is now recognized as one of the major public health issues all over the world. In Pakistan, it is a “silent epidemic” striking significantly because we are still struggling with health and economic burdens of malnutrition, infectious diseases and high infant mortality rates. In epidemiological studies age, sex and ethnic background all have to be taken into consideration, particularly when determining the health risk with obesity. Females are more vulnerable to be affected by obesity related health issues. Body Mass Index (BMI) is a surrogate measure of assessing obesity in terms of height and weight. It does not give any insight into regional body fat distribution. BMI is not a reliable measurement of body composition in individuals particularly in females having high body fat, rather more specifically it is excess abdominal fatness, quantified by waist circumference measurement, which is a better considered measure for assessing abdominal obesity in females.

**Keywords:** Abdominal obesity, Body mass index, Waist circumference, Over weight

## INTRODUCTION:

Obesity is now recognized as one of the major public health issues all over the world. WHO called urgent action to halt global obesity epidemic which is now labeled as “GLOBESITY”. Overweight and obesity is defined as abnormal excessive accumulation of fat within the body that impairs the body functions.<sup>1</sup> South Asian countries are currently affected by obesity epidemic which is a leading cause of various chronic non communicable diseases, their associated mortality and loss of life due to premature deaths.<sup>2</sup> The prevalence of overweight and obesity in Pakistan taking Asian-specific cut off levels reported approximately 25% of adult population as overweight and about 10.3% as obese.<sup>3</sup> Amongst various methods available for assessment of obesity, body mass index (BMI) is the most commonly used method of assessment of overweight and obesity because of its general application and feasibility.<sup>4</sup>

However, it tends to underestimate the prevalence of both conditions. BMI calculates obesity level in terms of height and weight of an individual and does not differentiate between body fat content, muscles and bone mass.<sup>5</sup> It may lead to misclassification of level of obesity as it is not necessary that overweight person has increased body fat as excessive weight. It can be due to increase muscle mass as in athletes or it can be due to increase body fat. It is just a mathematical calculation and not a direct estimation of adiposity.<sup>6</sup> It is an index for weight excess, rather than body fat composition.<sup>7</sup> Racial and ethnic disparities exist in distribution of body fat among different populations and ethnic subgroups.<sup>8</sup> There is different relationship between BMI and body fat distribution among different population and these disparities being more pronounced among women.<sup>9</sup> In Europeans, BMI of 30kg/m<sup>2</sup> corresponds to 25% of body fat in males and 30% of body fat in females<sup>10</sup>, while in South Asians of same gender, age and BMI have increased body fat percent and less muscle mass along with increased risk of cardio metabolic disorders. Evidence has supported the fact that these changes are more pronounced in females as compared to males.<sup>11, 12</sup> The disparities associated gender and ethnicity with regards to obesity assessment should be kept in consideration and instead of uniform BMI cut off population specific assessment of obesity indices based on distribution of body fat should be purposed.<sup>13</sup> Keeping in view BMI related error in measurement of obesity and its associated health risks, researchers are considering for some better tool for measuring the obesity. Since obesity has now become a serious public health issue, accurate level of estimation of obesity has become extremely important because of major health issues associated with excessive body fat. In the past two decades, visceral or abdominal obesity, as reflected anthropometrically by an increased waist circumference, has also emerged as an important predictor of risk of obesity-related diseases. Moreover, as discussed in a review<sup>5</sup> addressing anthropometric indices recommended waist circumference over other indices, as it is simpler to measure and interpret and correlates well with visceral fat measured by computed tomography. Yet, waist

✉ **Dr. Khaula Noreen**

Lecturer

Department of Community Health Sciences  
Bahria University Medical & Dental College  
Karachi

Email: dr\_khula@yahoo.com

✉ **Dr. Nadia Khalid**

Lecturer

Department of Community Health Sciences  
Bahria University Medical & Dental College  
Karachi

✉ **Dr. Imran Shaikh**

Professor & HOD

Department of Community Health Sciences  
Bahria University Medical & Dental College  
Karachi

Received: 19-01-2016

Revised: 18-02-2016

Accepted: 20-02-2016

circumference is also highly correlated with BMI and thus reflects general, and abdominal, obesity.<sup>14</sup>

**MATERIALS AND METHODS:**

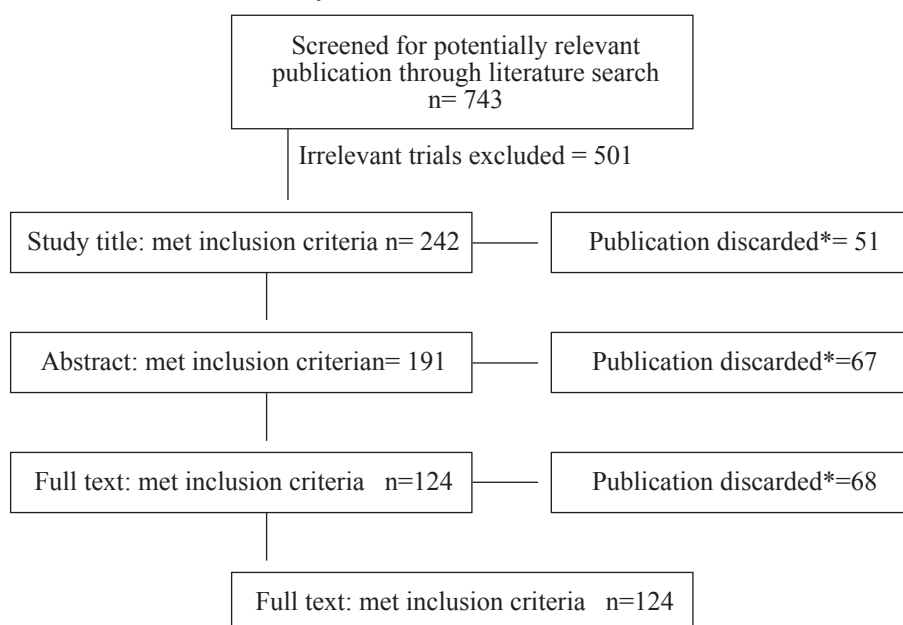
Articles were identified by using multiple electronic databases like Pub Med /MEDLINE /EMBASE Science direct, Google.com and Google scholar from the year 2001 to February 2016. Literature search was done by using key words, terminologies and phrases of obesity, overweight, abdominal obesity, waist circumference, body mass index, anthropometric indices. This review was done according to Preferred Reporting Items for Systematic Reviews and Meta-Analysis (PRISMA).

**Inclusion Criteria:** (1) Studies published in a peer-

reviewed journal(2) Studies using measured or self-reported body mass index (BMI), waist circumference (WC) waist hip ratio (WHR), body fat and Triceps Skin fold thickness (TSF)(3) Studies that must have examined the association between abdominal fat and its associated health hazards particularly in females(4) Study that can be case report, case series, abstract, original article or systematic review or meta-analysis(5) Study that must have been published within the last 15 years.

**Exclusion Criteria:** Studies that used categorizations of measures of obesity indices including Body Mass Index (BMI), waist circumference (WC) or waist hip ratio (WHR) different from the WHO or another internationally comparable classification.

Figure 1  
Flow of study identification, inclusion and exclusion



\* Papers which have not used standard (WHO criteria) categories of BMI classification, obesity indices and other measures of body fat composition. Papers not from peer reviewed journals and not fulfilling the eligibility criteria.

**LITERATURE REVIEW:**

Obesity is an epidemic of the 21<sup>st</sup> century and major causative factor for many other metabolic disorders. According to a global estimate by the World Health Organization (WHO), in 2005 there were about 1.6 billion overweight persons aged 15 years and above and among them at least 400 million adults were obese. World Health Organization uses BMI calculated as kg/m<sup>2</sup>, and it defines obesity as BMI above 30 and overweight, as BMI s above 25. The Indo-Asian specific definition of obesity is set as, BMI above 27 and overweight, BMI above 23. The revision of definition of obesity taking in account the racial differences has resulted in increase prevalence. According to that approximately 1.7 billion people are classified as overweight. The WHO further projects that by 2020, approximately 2-3 billion adults will be overweight and more than 700 million will be obese.<sup>15</sup> World Health

Organization (WHO) estimated that globally obesity prevalence has doubled since 1980; with over all increased prevalence in females as compared to males. Between 1980 and 2013, obesity prevalence in males have increased from 28.8 to 36.9%, while in females have increased from 29.8% to 38%.<sup>16</sup> Obesity prevalence is unacceptably high among Asian women.<sup>17</sup> Obesity in this age group not only predispose them to various reproductive health issues but also increases the risk of chronic health issues in later years.<sup>18</sup> National Nutrition survey of Pakistan 2011 reported that 19.4% of women of reproductive age were overweight and 9.5% of women of reproductive age were obese (15.7% from urban areas compared to 6.5% from rural areas).<sup>19</sup> There is dearth of literature in this regard in our part of world. Moreover, there is limited data available regarding screening modalities for obesity diagnosis and its associated health risks.<sup>20</sup>

### Why abdominal obesity?

Abdominal obesity or central obesity is the accumulation of abdominal fat resulting in an increase in waist size. There is abnormal deposition of fat in abdominal areas with resultant health consequences such as cardiovascular diseases, type 2 diabetes and cancers. It is recognized as a major health issue among the adults both in developed and developing countries.<sup>21</sup> In general, women tend to have higher rates of abdominal obesity than men, which become more prominent with ageing especially after the menopause.<sup>22</sup> Furthermore, South Asian women in particular experience a severe form of abdominal obesity at normal BMI leading to devastating consequences.<sup>23</sup>

Waist circumference is measured by standard WHO STEPS Protocol. The WHO STEPS is a standardized instrument that allows the collection, analysis and dissemination of data regarding risk factor surveillance for non communicable disease in standardized manner. As per this protocol waist circumference is measured by placing measuring tape approximately at mid of upper border of iliac crest and lower margin of last palpable rib.<sup>24</sup>

According to Asian cut offs waist circumference of < 80 cm for females and < 90cm for males is considered as normal.<sup>25</sup> According to data of National Action Plan –Non Communicable diseases (NAP-NCD) First Round of Surveillance, 34.2% males and 60% females in the urban areas and 35.7% males and 55.5% females in the rural areas are reported to have central obesity.<sup>26</sup> This is a grave trend since central obesity is a more important risk factor for chronic non communicable diseases than overall adiposity as measured by BMI in studies on the Pakistani population.<sup>27</sup> In females, gynecoid type of obesity is more common among which fat is deposited at hips, thigh and buttocks.<sup>28</sup> Waist circumference is considered as better tool for abdominal fat assessment specially for the gynecoid type of obesity.<sup>29</sup> Moreover, evidence has supported the fact that it is the the best anthropometric index in predicting a chronic disease risk and related health conditions.<sup>30</sup>

### Abdominal Obesity and reproductive abnormalities:

Reproductive abnormalities are relatively common in women with abdominal obesity, of which the most common abnormality encountered is polycystic ovarian syndrome (PCOS). South Asian women are more vulnerable to this disease that manifests at a younger age when compared with their counterparts in the West.<sup>31</sup> They are also found to have higher fasting insulin concentrations and lower insulin sensitivity than Caucasian women.<sup>32</sup> Polycystic ovarian syndrome (PCOS) is closely associated with abdominal obesity in South Asian women who also have more severe symptoms associated than white Europeans.<sup>33</sup> Obese women are particularly vulnerable to adverse effects of pregnancy including 3 to 10 times the risk of pre-eclampsia, gestational diabetes mellitus, difficulties in labour and delivery, higher rates of caesarean delivery and other peri-natal morbidity and mortality.<sup>34</sup> Obesity has a significant adverse impact on reproductive

outcome. It influences not only the chances of conception but also the response to infertility treatment and increases the risk of miscarriage and pregnancy complications.<sup>35</sup> When fertility is a problem, the primary goal of treatment is to normalize serum androgens and restore reproductive function, which can be achieved by reducing insulin resistance through a decrease in weight and abdominal fat.<sup>36, 37</sup> Studies of weight loss through lifestyle modification have indicated that improvements in fertility rate, hormonal profiles, menstrual regularity, ovulation, and conception occur with modest weight loss (5% of initial body weight) along with parallel improvement in anthropometric indices.<sup>38, 39</sup>

**Cancers:** After the heart disease cancer is ranked as second leading cause of death in developing countries.<sup>40</sup> Current estimates of association of obesity with risk of cancer has documented that 5% of all cancers among postmenopausal women are attributable to being overweight and that 4% to obesity.<sup>41</sup> Evidence has shown the association of obesity with cancer.<sup>42</sup> Increase amount of adipose tissue in obese females lead to release of several hormones like factors adipokines which are pro inflammatory in nature and promotes the cancer development. Common obesity related malignancy include cancer of breast, endometrial, colon and esophageal, kidney.<sup>43,44</sup> Obesity is one of the modifiable risk factor that predispose females to development of cancer Throughout the life span there is particular time period referred as "windows of susceptibility" during which various contributing factors can predispose female body towards the development of cancer among all these factors obesity has significant influence in cancer predisposition.<sup>45</sup> In Pakistan, the data for cancer prevalence in females is in accordance with the trend found worldwide that is breast cancer was reported to be the most prevalent cancer among females specially those having BMI beyond normal limits.<sup>46</sup> Obese females are most vulnerable and have poor prognosis.<sup>47</sup> The most probable underlying mechanism involved in the development of breast cancer due to obesity is hormonal changes such as elevated estrogen levels. Another possible mechanism may be insulin resistance and leading to increase level of circulating insulin that may lead to stimulation of cancers cells.<sup>48,49</sup>

**Cardiovascular disorders:** Evidence has reported that for cardiovascular risk stratification and evaluation of various metabolic disorders it is recommended to measure both BMI and WC together.<sup>50</sup> Previous researches have reported the WC to be modestly stronger predictors of cardiovascular disease risk than BMI.<sup>51, 52</sup> Women are more prone to the health hazards associated with overweight and obesity as compared to their males. Risk associated with obesity is proportional to degree of waist circumference. Women with waist circumference >80 cm have three times more mortality rate as compared to women with < 80 cm. Obesity increases the level of triglycerides and lower the level of HDL promotes the narrowing of small arterioles which lead to coronary artery disease, and associated health risk.<sup>53</sup>

**Obesity and mental health:** Despite of overall increase



in obesity throughout the world, obese people are still facing low acceptance and negative attitude.<sup>54</sup> Females are more vulnerable group to be effected by negative attitudes and disapproval from friends and relatives, they have to face social constrains like criticism, taunting remarks from strangers and discrimination from their normal weight peers.<sup>55</sup> They are also stigmatized and bullied for their physical appearance. Paradoxically such people eat more due to stress and guilt and vicious circle of eating and weight gain continue. Research have proved that on various psychological assessment scales, obese people score very less ranging from sadness, weeping tendencies and severe depression, anxiety, mood swings, insomnia, eating disorders including bulimia and anorexia nervosa.<sup>56</sup>

**Prevention:** Obesity has now become a global epidemic, yet significant reduction in mortality associated with this disease is possible, with millions of lives can be saved and uncountable disabilities can be reduced through primary prevention of risk factors, early diagnosis and prompt treatment. In an era of over consumption of food and obesity, healthy food habits are essential in keeping away a huge variety of chronic diseases. The portion size in pre-packaged, ready-to-eat and restaurant foods is increasing. Many people cannot accurately estimate portion size, and this leads to an underestimation of intake. Life style intervention involving modification in diet and physical activity can be successful in achieving weight loss in severely obese females. There is dire need to create public health awareness by organizing awareness lectures in order to make general public sensitized about deleterious effects of obesity, targeting the important demographical groups, like women, adolescent and children. Young generation should be addressed to make them aware about benefits of healthy life style and importance of primary prevention in development of chronic diseases. Education programs should be arranged in educational institution. The environments in which people live are complex and their individual and combined elements have a marked effect on people's behaviors and dietary intakes. Individuals interact in a variety of micro-environments or settings such as schools, workplaces, homes, restaurants and fast food outlets. These in turn are influenced by the broader macro-environments or sectors such as the food industry, all levels of government, and society's attitudes and beliefs.

There is dire need of integration of all concerned stakeholders including health, public, private, health, education to integrate in order to make effective strategies to halt this problem. Fiscal food policies should be implemented by Government. Food prices have a marked influence on food buying behavior and consequently nutrient intakes. Governments should promote the healthy eating behavior by controlling the prices of natural food products. Nutrition 'signposting' programs should be implemented. Nutrition 'signposts' are signals (such as logos) at point of choice which indicate to the consumer that a food meets certain nutrition standards. Full nutrition information panels should be on food products. Nutrition

information panels appear to facilitate the food choices of those who are trying to reduce their fat intake, greater impact among women, higher educated people and those with established beliefs and knowledge about diet-disease relationships.

In Pakistan, it is a "silent epidemic," striking significantly because we are still struggling with the health and economic burdens of malnutrition, stunting, infectious disease, and high childhood mortality rates. One paradox of this so-called "nutrition transition" is that even as obesity rates rise, underweight persists, sometimes within the same household. We are currently facing a dual burden-the infectious diseases that accompany malnutrition and, increasingly, the debilitating chronic diseases linked to obesity and Western lifestyles. Given the huge costs of obesity, prevention is key. Slowing the increases in obesity and turning around the epidemic will take large-scale, multifaceted efforts, within individuals and across the nation, to improve people's food choices and increase physical activity. Pakistan needs to develop a national strategy to control obesity in its population by implementing the recommendations of the WHO global strategy on diet, physical activity and health. The implementation program should integrate all stakeholders like health department, print and electronic media, nongovernmental organizations, and private sector. Community based primary prevention programs and clinical trial with more resources and manpower should be introduced with aim of achieving maximum benefit with cost effectiveness.

#### CONCLUSION:

The evidence for the adverse effects of obesity on women's health is overwhelming and indisputable. Obesity, especially abdominal obesity, is central to the reproductive health problems and is strongly related to development of cardio-metabolic risk factors in women. Therefore, more attention should be paid to abdominal obesity both in clinical practice and in epidemiological studies. Moreover, the disparities associated gender and ethnicity with regards to obesity assessment should be kept in consideration and instead of uniform BMI cut off population specific assessment of obesity indices based on distribution of body fat should be purposed.

#### REFERENCES:

1. World Health Organization: Obesity and overweight: fact sheet N0 311. 2012
2. Ramachandran A, Snehalatha C. Rising burden of obesity in Asia. *Journal of obesity*. 2010
3. Jafar TH, Chaturvedi N, Pappas G. Prevalence of overweight and obesity and their association with hypertension and diabetes mellitus in an Indo-Asian population. *Canadian Med Assoc J* 2006; 175:1071-7
4. Chasan-Taber L, Marcus BH, Rosal MC. Proyecto Mamá: a lifestyle intervention in overweight and obese Hispanic women: a randomised controlled trial - study protocol. *BMC Pregnancy and Childbirth*. 2015;15:157. doi:10.1186/s12884-015-0575-3
5. Javed A, Jumean M, Murad MH, Okorodudu D, Kumar S, Somers VK et al. Diagnostic performance of body

- mass index to identify obesity as defined by body adiposity in children and adolescents: a systematic review and meta-analysis. *Pediatric obesity*. 2015;10(3):234-44
6. Bergman, Richard N. A Better Index of Body Adiposity. *Obesity (Silver Spring, Md.)* 19.5 (2011): 1083-9. Accessed on *PMC*. Web. 18 Feb. 2016
  7. Shah NR, Braverman ER. Measuring Adiposity in Patients: The Utility of Body Mass Index (BMI), Percent Body Fat, and Leptin. *PLoS ONE*. 2012; 7(4): e33308
  8. Djibo DA, Araneta MR, Kritz-Silverstein D, Barrett-Connor E, Wooten W. Body adiposity index as a risk factor for the metabolic syndrome in postmenopausal Caucasian, African American, and Filipina women. *Diabetes & Metabolic Syndrome: Clinical Research & Reviews*. 2015; 9(2):108-13
  9. Rush EC, Freitas I, Plank LD. Body size, body composition and fat distribution: comparative analysis of European, Maori, Pacific Island and Asian Indian adults. *Br J Nutr* 2009;102:632-41
  10. Lear SA, Birmingham CL, Chockalingam A, Humphries KH. Study design of the Multicultural Community Health Assessment Trial (M-CHAT): *Ethin Dis* 2006; 16: 96-100
  11. Phillips CM, Tierney AC, Perez-Martinez P, Defoort C, Blaak EE, Gjelstad IM et al. Obesity and body fat classification in the metabolic syndrome: impact on cardio-metabolic riskmetabotype. *Obesity*. 2013;21(1):E154-61
  12. Carpenter CL, Yan E, Chen S, Hong K, Arechiga A, Kim WS et al. Body fat and body mass index among a multi-ethnic sample of college-age men and women. *Journal of obesity*. 2013
  13. Heymsfield, S B, Peterson C M, Thomas D M, Heo M Jr. Schuna, JM Why are there race/ethnic differences in adult body mass index–adiposity relationships? A quantitative critical review. *Obesity Reviews*, 2016; 17: 262-75
  14. Kahn HS, Bullard KM. Beyond Body Mass Index: Advantages of Abdominal Measurements for Recognizing Cardiometabolic Disorders. *The American journal of medicine*. 2016;129(1):74-81
  15. WHO. Non communicable diseases country profiles. Geneva, Switzerland: WHO press; 2013
  16. Akter J, Shahjahan M, Hossain S. Determinants of overweight and obesity among Bangladeshi diabetic women of reproductive age. *BMC Research Notes*. 2014;7:513. doi:10.1186/1756-0500-7-513
  17. Sengupta P, Chaudhuri P, Bhattacharya K. Screening obesity by direct and derived anthropometric indices with evaluation of physical efficiency among female college students of Kolkata. *Ann Med Health Sci Res* 2013;3:517-22
  18. Cardozo ER, Dune TJ, Neff LM, Brocks ME, Ekpo GE, Barnes RB, et al. Knowledge of obesity and its impact on reproductive health outcomes among urban women. *Journal of community health*. 2013; 38(2):261-7
  19. National Health Survey of Pakistan; 2011 Government of Pakistan, Ministry of Population and Development. 2011
  20. Wimalawansa SJ. Visceral adiposity and cardiometabolic risks: epidemic of abdominal obesity in North America. *Research and Reports in Endocrine Disorders* 2013.3:1-14
  21. Johnston LM, Finegood DT. Cross-sector partnerships and public health: challenges and opportunities for addressing obesity and noncommunicable diseases through engagement with the private sector. *Annual review of public health*. 2015;36:255-71
  22. Lovejoy JC, Champagne CM, de Jonge L, Xie H, Smith SR. Increased visceral fat and decreased energy expenditure during the menopausal transition. *Int J Obes* 2008; 32: 949-58
  23. Gray LJ, Yates T, Davies MJ, Brady E, Webb DR, Sattar N, et al. Defining Obesity Cut-Off Points for Migrant South Asians. *PLoS ONE*. 2011; 6(10):e26464
  24. World Health Organization. WHO STEP wise approach to chronic disease risk factor surveillance- Instrument v2.0. Department of Chronic Diseases and Health Promotion. World Health Organization. 20 Avenue Appia, 1211 Geneva 27, Switzerland. Available from: <http://www.who.int/chp/steps>
  25. Gupta S, Kapoor S. Body adiposity index: its relevance and validity in assessing body fatness of adults. *Obesity*, 2014:243294
  26. Nishtar S. Health Indicators of Pakistan – Gateway Paper II. Islamabad, Pakistan: Heartfile; 2007
  27. Alberti K, Zimmet P, Shaw J. Metabolic syndrome—a new world-wide definition. A consensus statement from the international diabetes federation. *Diabetic Medicine*. 2006;23(5):469-80
  28. Noroozi M, Rastegari Z, Paknahad Z. Type of body fat distribution in postmenopausal women and its related factors. *Iranian Journal of Nursing and Midwifery Research*, 2010; 15(1), 27-31
  29. Dagan SS, Segev S, Novikov I, Dankner R. Waist circumference v/s body mass index in association with cardio respiratory fitness in healthy men and women: a cross sectional analysis of 403 subjects. *Nutrition Journal*. 2013;12:12doi: 10.1186/1475-2891-12-12
  30. Hirose K, Tajima K, Hamajima N, Takezaki T, Inoue M, Kuroishi T, et al. Effect of body size on breast-cancer risk among Japanese women. *Int J Cancer*. 1999;80:349-55
  31. Wijeyaratne CN, Seneviratne R A, Dahanayake S, Kumrapeli V, Palipane E, Kuruppu N, et al. Phenotype and metabolic profile of South Asian women with polycystic ovary syndrome (PCOS): results of a large database from a specialist Endocrine Clinic. *Hum Reprod* 2011; 26: 202-13
  32. Misra A, Khurana L. Obesity-related non-communicable diseases: South Asians vs. White Caucasians. *Int J Obes* 2011; 35: 167-87
  33. Kumrapeli V, Seneviratne Rde A, Wijeyaratne C. Health-related quality of life and psychological distress in polycystic ovary syndrome: a hidden facet in South Asian women. *Br J Obstet Gynaecol* 2011; 118: 319-28
  34. Belen A H, Richard A. Impact of obesity on female reproductive health: British fertility society; policy and practice guidelines . *J Hum Fertil*. 2007;10:195-206
  35. Coyne K, Whigham LD, O'Leary K, Yaklic JK, Maxwell RA, Lindheim SR. Gestational carrier BMI and reproductive, fetal and neonatal outcomes: are the risks the same with increasing obesity? *Int J Obes*. 2016;40(1):171-5
  36. Diabetes Prevention Program Research Group. Reduction in the incidence of type 2 diabetes with lifestyle intervention or metformin. *N Engl J Med* 2002; 346: 393-403
  37. Nawaz FH, Rizvi J. Continuation of metformin reduces early pregnancy loss in obese Pakistani women with polycystic ovarian syndrome. *Gynecol Obstet Invest* 2010; 69: 184-9
  38. Irwin ML, Yasui Y, Ulrich CM, Bowen D, Rudolph RE,

- Schwartz RS, et al. Effect of exercise on total and intra-abdominal body fat in postmenopausal women: a randomized controlled trial. *JAMA* 2003; 89: 323-30
39. Crosignani PG, Colombo M, Vegetti W, Somigliana E, Gessati A, Ragni G. Overweight and obese anovulatory patients with polycystic ovaries: parallel improvements in anthropometric indices, ovarian physiology and fertility rate induced by diet. *Hum Reprod* 2003; 18, 1928-32
40. Reeves GK, Pirie K, Beral V, Green J, Spencer E, Bull D. Cancer incidence and mortality in relation to body mass index in the Million Women Study: cohort study. *Bmj*. 2007 Nov 29;335(7630):1134
41. Jemal A, Bray F, Center MM, Ferlay J, Ward E, Forman D. Global cancer statistics. *CA Cancer J Clin* 2011;61:69-90
42. Singh P, Kapil U, Shukla NK, Deo SV, Dwivedi SN. Association of overweight and obesity with breast cancer in India. *Indian Journal of Community Medicine*. 2011 Oct 1;36(4):259-62
43. Murthy N S, Mukherjee S, Ray G, Ray A. Dietary factors and cancer chemoprevention: An overview of obesity-related malignancies. *J Postgrad Med* 2009;55:45-54
44. Neamat-Allah J, Wald D, Sing A, Teucher B, Wendt A, Delorme S, et al. Validation of Anthropometric Indices of Adiposity against Whole-Body Magnetic Resonance Imaging "A Study within the German European Prospective Investigation into Cancer and Nutrition (EPIC) Cohorts. *PloS one* 2014 9(3):e91586
45. Undaram S, Johnson AR, Makowski L. Obesity, metabolism and the microenvironment: Links to cancer. *J Carcinog* 2013;12:19doi: 10.4103/1477-3163-119606
46. Youlten DR, Cramb SM, Yip CH, Baade PD. Incidence and mortality of female breast cancer in the Asia-Pacific region. *Cancer biology & medicine*. 2014;11(2):101-15
47. Stockwell S. Online First: Evidence that Obesity Raises Risk of Death in Young Women with ER-Positive Breast Cancer. *Oncology Times*, 2014
48. Sparano JA, Wang M, Zhao F, Stearns V, Martino S, Ligibel JA, et al. Obesity at diagnosis is associated with inferior outcomes in hormone receptor-positive operable breast cancer. *Cancer* 2012;118:5937-46
49. Roberts DL, Dive C, Renehan AG. Biological mechanisms linking obesity and cancer risk: new perspectives. *Annual review of medicine* 2010.61:301-16
50. IfardMN, Nazem M, Sarrafzadegan N, Nouri F, Sajjadi F, Maghroun M, et al. Body Mass Index, Waist circumference and Cardiovascular Disease Risk Factors in Iranian Adults: Isfahan Healthy Heart Program. *Journal of health, population, and nutrition* 2013.31(3):388-97
51. Klein S, Allison DB, Heymsfield SB, Kelley DE, Leibel RL, Nonas C, et al. Waist circumference and cardiometabolic risk: a consensus statement from shaping America's health: Association for Weight Management and Obesity Prevention; NAASO, the Obesity Society; the American Society for Nutrition; and the American Diabetes Association. *Obesity*. 2007;15(5):1061-7
52. Nazare JA, Smith J, Borel AL, Aschner P, Barter P, Van Gaal L, et al. Usefulness of measuring both body mass index and waist circumference for the estimation of visceral adiposity and related cardiometabolic risk profile (from the INSPIRE ME IAA study). *The American journal of cardiology*. 2015 Feb 1;115(3):307-15
53. Lindquist R, Witt DR, Boucher JL. Preventing cardiovascular disease in women: how can we do better? *Current opinion in cardiology* 2012;27(5):542-9
54. Evaluating a brief anti-weight bias intervention *British journal of health psychology* 2011; 16: 846 -61
55. Claudia S, Melanie L, Marie K, Heide G, Georg S, Hans-Helmut K et al. The stigma of obesity in the general public and its implications for public health -A systematic review *BMC Public Health* 2011; 11:661-6
56. Kasen S P, Cohen H, Chen A. Obesity and psychopathology in women. *Int J Obes* 2008; 32: 558-66



## ORIGINAL ARTICLE

# Knowledge, Attitude and Practice Regarding Oral Hygiene among Private School Children

Samreen Mazhar<sup>1</sup>, Ashghar Ali<sup>2</sup>, Mahwish Bano<sup>3</sup>, Muhammad Ali Leghari<sup>4</sup>, Aukif Ali Sheikh<sup>5</sup>

### ABSTRACT:

**Objective:** To evaluate the knowledge, attitude and behavior of private schools children regarding oral hygiene status of Gadap Town Karachi, Pakistan.

**Materials and Methods:** This descriptive cross sectional study was carried out in two private schools of Gadap town UC-2, Karachi Pakistan. A self-structured questionnaire related to KAP of oral hygiene was used and data was collected through questionnaire. This research was conducted in 290 school children aged 11-19 years who were examined and fulfilled the inclusion criteria. Questionnaire was designed and three house officers were trained for obtaining the data. The knowledge about oral hygiene and gingival index was determined and recorded from each student. Data was analyzed using SPSS version 20

**Results:** There were 67.9% male and 32.1% female students. About 81.4% of students had the knowledge regarding oral health, 12.8% individuals had no knowledge and 5.9% students reported I don't know about oral health. Students with knowledge of brushing of teeth twice daily was 81.7%, and without knowledge were 13.1% and 5.2% individuals said I don't know. 73.8% study participants had the knowledge of sweets / candy as harmful for oral health.

**Conclusion:** Private schools children of Gadap town Karachi Pakistan had knowledge of oral hygiene practices but oral hygiene instructions should be given to further improve the present status.

**Keywords:** Knowledge, Attitude, Practice, Oral hygiene status, Questionnaire, Gingival index

### INTRODUCTION:

Oral diseases qualify as a major community health problem, although these diseases can be prevented by

a positive dental health behavior.<sup>1</sup> Most of the oral diseases in everyday are directly related to lifestyle. They can be considered an important public health issue due to its high prevalence and significant social impact.<sup>2</sup> Oral health knowledge is considered to be an essential pre requisite for health related behavior.<sup>3</sup> Oral health and general health share common risk factors like use of tobacco and the excessive consumption of sweets, chocolates, etc. The solutions to control oral diseases are to be found through shared approaches with integrated chronic disease prevention. Dental caries and periodontal diseases have historically been considered as most important global oral health burdens. At present, the distribution and severity of oral diseases vary in different parts of the world and within the same country.<sup>4</sup> In developed countries, dental caries and periodontal diseases affects 60-90% of school children and adults.<sup>5</sup> It is also a most prevalent oral disease in several Asian and American countries, while it appears to be less common and less severe in most African countries. However, it is expected that the incidence of dental caries will increase in the near future in many developing countries of Africa, particularly as a result of growing consumption of sugars and inadequate exposure to fluorides.<sup>6</sup> While in some industrialized countries there has been a positive reduction in tooth loss among adults in recent years, the proportion of edentulous persons amongst the elderly is still high in some countries. In most developing countries, access to oral health services is limited and teeth are often left untreated or are extracted because of pain and discomfort. Tooth loss and impaired oral function are therefore expected to grow as a public health problem in many developing countries. The tooth loss in adult life may also be due to poor periodontal health. Severe periodontitis which may result in tooth loss is found in 5-15% of most populations. In industrialized countries, studies show that tobacco use is a major risk factor for periodontal disease. With the

#### ✉ Dr. Samreen Mazhar

Lecturer  
Department of Community Dentistry  
Baqai Dental College  
Baqai Medical University  
Near Toll Plaza Karachi  
Email: tayyaba-samreen@hotmail.com

#### ✉ Dr. Asghar Ali

Associate Professor & Head  
Department of Community Dentistry  
Baqai Dental College  
Baqai Medical University  
Near Toll Plaza  
Karachi

#### ✉ Dr. Mahwish Bano

Demonstrator  
Baqai Dental College  
Baqai Medical University  
Near Toll Plaza Karachi

#### ✉ Dr. Muhammad Ali Leghari

Senior Lecturer  
Department of Community Dentistry  
Baqai Dental College  
Baqai Medical University,  
Near Toll Plaza  
Karachi

#### ✉ Dr. Aukif Ali Sheikh

Dental Doctor  
Baqai Dental College  
Baqai Medical University,  
Near Toll Plaza  
Karachi

Received: 18-09-2015

Revised: 01-11-2015

Accepted: 04-01-2016

growing consumption of tobacco in many developing countries the risk of periodontal disease and tooth loss, therefore, may increase. Periodontal disease and tooth loss are also related to general chronic diseases.<sup>7</sup> Dental health educators are frequently invited by different school establishments to deliver lectures on oral health, and to provide preventive services. School teachers traditionally have played a role in educating children about how to prevent oral diseases and promote oral health.<sup>8</sup> Health education programs in schools may be conducted by groups such as public health professionals, health educators, school nurses and teachers. The advantages of using school personnel are the potential for improved continuity of instruction and lowered cost of the service.

According to previous studies in Pakistan, the government spending on public sector education is only 12% of its federal budget. Overall there are 256,088 educational institutions in our country out of which 71% are in public sector. The total student enrollment is 37,462,884 out of which 25,213,894 students are enrolled in public institutes.<sup>9,10</sup> The purpose of this study was to evaluate the knowledge, attitude and practice regarding oral hygiene among two private schools children of Gadap town Karachi, Pakistan.

#### **MATERIALS AND METHODS:**

This descriptive cross sectional study was conducted for a period of six months from September 2014 to February 2015. A self administered close ended questionnaire to assess the oral hygiene related to knowledge, attitude and practices among students of two private schools in Gadap town UC-2 Karachi. There were 197 males and 93 females with age group of 11-19 years. This age group was chosen as the baseline data collected could be utilized for future planning of a school oral health programs. The students of the school represent population of children belonging to low socioeconomic status.

The questionnaire was filled by the team of dentist and trained house officers. The data for this study were collected by carrying out an interview among participants. The stratified simple random sampling technique was used. The questionnaire was divided into four parts. First part included questions on demographic characteristics of children that is age, gender and educational status. Second part of the questionnaire included questions to test the knowledge of children regarding oral hygiene, brushing frequency, sweets/candies, smoking hazards, dental floss, and mouthwash etc. Third part comprised of questions related to their attitude towards increase brushing frequency, withdraw smoking habits, reduce sweets / candies intake etc. Fourth part consists of cleaning practices of teeth and frequency etc. A written consent was taken from school principals as well as the older children before the collection of data. The data was recorded and analyzed by using Statistical Package for Social Sciences (SPSS) version 20.

#### **RESULTS:**

The sample size of this study was 290. The sample comprised of males 67.9% and females 32.1 % (Table 1). The age range among the students was 11-19. The mean age of the study participants was 13.73.

About 81.4% of individuals had the knowledge regarding oral health, 12.8% individuals had no knowledge of oral health and 5.9% individuals don't had the knowledge of oral health. Those participants who had the knowledge of brushing of teeth twice daily were 81.7%, had no knowledge of brushing twice daily was 13.1% and 5.2% individuals don't had the knowledge of brushing teeth twice daily. 73.8% study participants had the knowledge of sweets / candies is harmful for oral health, 21.4% individuals had no knowledge regarding sweets / candies and 4.8% individuals did not know about the harmful effects of sweets / candies of your oral health. 88.6% individuals had the knowledge of chalya or gutka was bad for oral health, 8.3% individuals had no knowledge regarding chalya or gutka was bad for oral health and 3.1% individuals did not know the harmful effects of chalya or gutka of your oral health. 76.2% individuals know about the smoking hazards. The 61.0% individuals had the knowledge of gums bleeding while brushing teeth, while 36.9% individuals had the knowledge of bad breath in mouth (Table 2a).

Regarding the attitude of oral hygiene status in school children 85.2% individuals had to improve oral hygiene through proper brushing. 77.6% know that the gum disease is bad for oral health so they have a plan to increase brushing frequency, 10.7% individuals did not know that gum disease is bad for oral health and 11.7% individuals did not know about the gum disease so they did not increase their brushing frequency. The 78.6% individuals had a plan to reduce sweets and chocolate intake, 17.9% had no plan to reduce sweets and chocolate intake and 3.4% individual didn't had a plan to reduce sweets and chocolate intake. The 80.7% individuals had to reduce gutka and chalya habit, 13.4% individuals had no plan to reduce gutka and chalya habit and 5.9% said that I don't have a plan to reduce gutka and chalya habit that is harmful for oral health. The 72.8% individuals had a plan to withdraw smoking habit (Table 2b)

Frequency of once daily cleaning of teeth was 46.2% and frequency of twice daily cleaning of teeth was 52.4%. Those participants who clean teeth with tooth paste by finger was 4.8%, individuals cleaning teeth with toothpaste by brush was 87.9%, and 5.5% individuals used miswak for cleaning of teeth (Table 2c). 26.6% study participants used dental floss after the meal. The individuals who smoke one cigarette per day were 7.9%, for those who smoke two cigarette per day were 1.7%, who smoke three cigarette per day were 1.7% and the others are included as 88.6%. Frequency distribution of gingival index among school children was also evaluated (Table 3).

Table: 1  
Frequency distribution of participants by age and gender

Age in years	Frequency	Percent
11	11	3.8
12	67	23.1
13	48	16.6
14	74	25.5
15	42	14.5
16	32	11.0
17	7	2.4
18	4	1.4
19	1	.3
Male & Female	Frequency	Percent
Male	197	67.9
Female	93	32.1
Total	290	100.0

Table: 2a  
Frequency distribution of knowledge among school children

Questions	Yes	No	Idon't Know
Do you have knowledge about oral hygiene?	81.4%	12.8%	5.9%
Do you know brushing is necessary twice daily?	81.7%	13.1%	5.2%
Do you know sweets / candies is bad for your oral health?	73.8%	21.4%	4.8%
Do you know chalia or ghutka is bad for your oral health?	88.6%	8.3%	3.1%
Do you have any knowledge regarding smoking hazards?	76.2%	12.1%	11.7%
Do your gums bleed when you brush your teeth?	61%	34.8%	4.1%
Do you feel bad breath in your mouth?	36.9%	58.3%	4.8%
Do you have any knowledge regarding dental floss?	25.9%	52.8%	21.4%
Do you have knowledge about mouthwash?	30%	53.1%	16.9%

Table: 2b  
Frequency distribution of attitude among school children

Frequency	Yes	No	I don't know
I know I have to improve my oral hygiene through proper brushing	85.2%	6.2%	8.6%
I know gum disease is bad for my oral health, that's why I have to plan to increase brushing frequency:	77.6%	10.7%	11.7%
I have a plan to reduce sweets & chocolate intake	78.6%	17.9%	3.4%
I have a plan to reduce gutkha /chalia habit	80.7%	13.4%	5.9%
I have plan to withdraw my smoking habit	72.8%	10%	17.2%

Table: 2c  
Frequency distribution of practice among school children

Variables	Once	Twice	No
Do you clean your teeth	46.2%	52.4%	1.4%
Do you get your teeth clean?			
Tooth Paste with finger	4.8%		
Tooth Paste with brush	87.9%		
Miswak	5.5%		
Not clean	1.7%		

Table: 3  
Frequency distribution of gingival index among school children

Gingival index	Frequency	Percent
No inflammation	121	41.7
Mild inflammation	126	43.4
Moderate inflammation	39	13.4
Severe inflammation	4	1.4
Total	290	100.0

## DISCUSSION:

In Pakistan being a third world country, has its major bulk of population residing in the rural areas. There is lack of health services and personnel in such areas thus leading to the deficiency of knowledge and awareness of dental hygiene being given by the doctors. This study presented an overview of the oral health behavior in terms of knowledge, attitudes and practice of school children ages eleven to nineteen years<sup>10</sup>. Dental plaque initiates reaction in tissues which starts in early ages especially during infancy and results in bacterial challenge in the host. The balance between microbial challenges in the host response is impaired and causes inflammation that results in loss of periodontal attachment. Usually males have a poor oral hygiene. Periodontal disease progression depends on age, sex, socioeconomic status, brushing habits and their frequency.<sup>11,12</sup> Previous studies on gingivitis had been conducted in many parts of the world in different ethical and cultural background. Majority of the students examined in our study used tooth brush and paste to clean their teeth, some used finger or miswak as a method of cleansing. When age wise prevalence was seen it was found 80% in 5-7 years, 79% in 8-10 years and 78% in 11-13 years.<sup>13</sup> When gingival index was considered, children examined had gingivitis out of which 13.4% had moderate gingivitis, 43.4% had mild gingivitis and 1.4% had severe gingivitis while 41% were found to be healthy.

Previous studies showed contrast results in comparison to our results, reason behind may be the difference of socio economic and geographical conditions. There was no periodontitis noted, results were concurrent with previous studies. Previous international studies involving Jordanian school children showed that oral hygiene, gingival conditions, have improved since the early 1990s although gingival disease and dental caries among Jordanians were found to be more prevalent than in developed countries.<sup>10,11</sup> Another previous study conducted among elementary school children 74.9 subjects agreed that fluoride protects the teeth and 84.9 were agreed that clean mouth everyday is the best way to prevent from gum diseases.<sup>14</sup>

In our study, 81.4% had knowledge of oral hygiene but 5.9% in a category of I don't know, 12.8% had no knowledge about oral hygiene. Different authors had explained effects of this type in terms of inequality of access to oral healthcare services. This survey found that 81.7% had high percentage of children in this study.<sup>15</sup>

A high proportion of the subjects reported that they did not attend dental clinics due to fear from dental treatment; this coincided with previous study on Jordanian private and public school children.<sup>16</sup> This might be attributed to the lack of proper oral health education programs for both children and parents. In addition to the above dental treatment undesired, high costs of dental care, and lack of toothache. Lack of parental encouragement and advice to visit the dentist might also contribute to the irregular dental attendance. Lack of parent's regular dental attendance might be reflected in their children.<sup>17,18</sup> The participants demonstrated positive attitudes toward

their dentists and high awareness of the link between oral health and systemic well-being. This might be explained by the fact that schools in Gadap town had been consciously promoting the role of prevention.<sup>19</sup> Unfortunately, these efforts are limited and insufficient nationwide; there is a need for comprehensive national educational programs to improve the oral health practice, knowledge, and attitudes of the general population.<sup>20</sup> Health education, since they indicate that social factors need to be taken into account in public education programs aimed at improving oral health practices<sup>21</sup>. Observed that daily tooth brushing became more frequent after a community education programs about oral hygiene. In other studies based on the KAP model as applied in health education, the educational intervention significantly improved oral health practices.<sup>22</sup> According to WHO periodontal disease is one such chronic diseases for which evidence is available on efficacy of prevention, which has been emphasized by other authors.<sup>23,24</sup> WHO Global Oral Health Program formulated the policies and the necessary actions for the improvement of oral health. The strategy is that oral disease prevention and the promotion of oral health need to be integrated with chronic disease prevention and general health promotion as the risks to health are linked (like tobacco use and the standard of hygiene).<sup>23</sup> It is imperative that dental hygiene awareness is imparted and measures for improvement in oral hygiene are undertaken in all age groups across rural areas of Pakistan as this constitutes the major portion of the population and community oral hygiene promotion must attempt to maximize opportunities for oral health for all and reduce inequalities by removing financial and other barriers.<sup>24,25</sup>

## CONCLUSION:

Students had knowledge of oral hygiene practices like change of brush, frequency of brushing, time period for brushing and brushing techniques but to improve the oral hygiene status of this population, health schemes like free dental checkups; health education and motivation about oral hygiene and free distribution of samples should be made available for this needy population. The sample size of this study is too small to be conclusive. More researches are required in this field with other parameters related to dental fluorosis, water fluoridation, prevalence and risk factors of oral cancer.

## Acknowledgement:

Special thanks are offered to my colleagues, my computer assistant Mr. Rehan Ansari and all those who guided, helped and encouraged me for this study

## REFERENCES:

1. Axellson P. Caries and periodontal disease // in: An introduction to risk prediction and preventive dentistry. 2000. p.39-77
2. Sheiham A. In the chemical prevention of gingivitis necessary to prevent severe periodontitis. *Periodontol* 2000. 1997; 15:15-24

3. Al-Ansari J, Honkata E, Honkata S. Oral health knowledge and behavior among male health sciences college students in Kuwait. *BMC Oral Health* 2003;3:2-8
4. Worthington HV, Hill KB, Mooney J, Hamilton FA, Blinkhorn AS. A cluster randomized controlled trial of a dental health education program for 10-year-old children. *J Public Health Dent.* 2001 Winter;61(1):22-7
5. Melnick SL, Nowjack-Raymer R, Kleinman, DV, Swango PA. 1993, Geneva: WHO.
6. *Community Dentistry and Oral Epidemiology* 31 (Suppl. 1), 3–24. Petersen, P.E. (2003b): Tobacco and oral health - the role of the World Health Organization. *Oral Health and Preventive Dentistry*2003;31(1):309-15
7. Arnlaugsson S, Magnusson TE. Prevalence of gingivitis in 6-year-olds in Reykjavik, Iceland. *Acta Odontol Scand* 1996;54:247-50
8. Petersen PE, Kwan S. Evaluation of communitybased oral health promotion and oral disease prevention - WHO recommendations for improved evidence in public health practice. *Community Dental Health* 2004; 21 (Supplement), 319-21
9. Education statistics 2007-2008. Ministry of Education, Pakistan. <http://www.moe.gov.pk/education/alstatistics.htm> Accessed November 2011
10. Taani DQ. Periodontal awareness and knowledge and pattern of dental attendance among adults in Jordan. *Int Dent J* 2002;52:94-8
11. Taani DQ. Trends in oral hygiene, gingival status and dental caries experience in 13-14-year-old Jordanian school children between 1993 and 1999. *Int Dent J* 2001;51: 277-81
12. Harikiran AG, Pallavi SK, Hariprakash S, Ashutosh, Nagesh KS. Oral health-related KAP among 11 – to 12-year-old school children in a Government-aided Missionary School of Bangalore city. *Indian J Dent Res [serial online]* 2008;19(3):236-42
13. Mahesh Kumar P, Joseph T, Varma RB, Jayanth M. Oral health status of 5 years and 12 years school going children in Chennai city. An epidemiological study. *J Indian Soc-PedoPrev Dent* 2005;23:17-22
14. Bjarnason S. High caries levels: problems still to be tackled. *Acta Odontol Scand* 1998;56:176-8
15. Jalevik B, Sjostrom O, Noren JG. Evaluation of three years of dental care of adolescents in the Public Dental Service in west Sweden. *Swed Dent J* 1999;23:141-8
16. Redmond CA, Blinkhorn FA, Kay EJ, Davies RM, Worthington HV, Blinkhorn AS. A cluster randomized controlled trial testing the effectiveness of a school-based dental health education program for adolescents. *J Public Health Dent.* 1999;59(1):12-7
17. Tewari A, Gauba K, Goyal A. Evaluation of KAP of oral hygiene measures following oral health education through existing health and educational infrastructure. *J Indian Soc Pedod Prev Dent.* 1992 Mar;10(1):7-17
18. AlmerichSilla JM, Montiel JM. Oral health survey of the child population in the Valencia Region of Spain (2004). *Med Oral Patol Oral Cir Bucal.* 2006; 11(4):E369 -81
19. Lueveswanij S, Nittayananta W, Robison VA. Changing knowledge, attitudes, and practices of Thai oral health personnel with regard to Aids: an evaluation of an educational intervention. *Community Dent Health.* 2000; 17(3) :165-71
20. Worthington HV, Hill KB, Mooney J, Hamilton FA, Blinkhorn AS. A cluster randomized controlled trial of a dental health education program for 10-year-old children. *J Public Health Dent.* 2001; 61(1):22-7
21. Choo A, Delac, DM, Messer L B. Oral Hygiene Measures and Promotion: Review and Considerations. *Australian Dental Journal.* 2001; 46: 166-73
22. Kunzel W. Trends in coronal caries prevalence in Eastern Europe: Poland, Hungary, Czechoslovakia, Slovak R Romania, Bulgaria and the former States of the USSR. *Int Dent J* 1996; 46(Suppl):204-10
23. Al-Omiri MK, Al-Wahadni AM, Saeed KN. Oral health attitudes, knowledge, and behavior among school children in North Jordan. *J Dent Educ* 2006;70(2):179-87
24. Tubaishat RS, Darby ML, Bauman DB, Box CE. Use of miswak versus toothbrushes: Oral health beliefs and behaviours among a sample of Jordanian adults. *Int J Dent Hyg.* 2005;3:126-36
25. Ahmed S, Solaiman F, Islam MR, Akhter SM, Nizami MZ, Khatun MA. Attitude on Oral Hygiene among the school going children in selected schools at Dhaka city. *City Dent Col J.* 2013;10:41-6





# Efficacy of 50g Glucose Challenge Test as a Screening Tool for Gestational Diabetes Mellitus

Ayesha Arif<sup>1</sup>, Sarwat Nazar<sup>2</sup>, Sadia Arif<sup>3</sup>

## ABSTRACT:

**Objective:** To evaluate the validity of 50 g oral Glucose Challenge Test as a screening tool for GDM in our population.

**Materials and Methods:** This cross sectional study was carried out in Obstetrical clinic, Combine Military Hospital (CMH) Lahore. 100 women carrying singleton pregnancy between 20-35 years of age, booked in first trimester were included while patients with risk factor of GDM or with established type I or II DM were excluded from study. 50 g GCT was administered to patients between 24-28 weeks of gestation after informed consent. Venous plasma glucose levels after 1 hour of glucose load, were taken, using 140 mg/dl as a cut off value. Regardless of results of screening, all patients were tested with 100 g OGTT as a "gold standard" of diagnosis of GDM.. Validity of 50g GCT was calculated for sensitivity, specificity, positive and negative predictive value. Data was analysed by SPSS version 16.

**Results:** Out of 100 patients, 19% were screen positive and 81% screened negative with 50g GCT. With 100 g OGTT, true positive were 10 out of 19(52.6%) screen positive, and false positive were 9 out of 19(47.4%) screen positive. False negative were 3 out of 81(3.7%) screen negative, whereas true negative were 78 out of 81(96.3%) screen negative. Validity of 50 g GCT has been calculated to be having sensitivity of 76.92%, specificity of 89.6%, positive predictive value of 52.6% and negative predictive value of 96.2%.

**Conclusion:** 50 g GCT is an effective screening tool for GDM between 24-28 weeks of gestation with adequate sensitivity and specificity.

**Key words:** GDM, OGTT, 50 g GCT, Screening tool

## INTRODUCTION:

Gestational Diabetes Mellitus (GDM) is defined as glucose intolerance with onset or first detection during pregnancy.<sup>1,2</sup> It usually disappears immediately after delivery or upto 6 weeks postpartum. Prevalence of GDM varies among different racial, ethnic groups<sup>3</sup> and with prevalence of type II diabetes mellitus (DM). It is more common in African, Latino, Hispanics and Asian (Indian subcontinent) women. Risk factors for GDM are family history,<sup>4</sup> body mass index (BMI) > 25,<sup>5,6</sup> age >35 yrs, grand multi parity, macrosomia in previous pregnancy, intra uterine demise (IUD),<sup>7</sup> foetal anomaly<sup>8</sup> and black race.<sup>9</sup> But 12% of patients with GDM have no risk factors.<sup>10</sup> Overall worldwide its prevalence is 1-14% depending upon the population studied and diagnostic tools applied. Overall GDM affects 2-5% of pregnancies in USA and 4-5% in UK.<sup>11</sup> Reported incidence in Asian population is 2-10%. In Pakistan prevalence of type II DM is around 10-14%<sup>12</sup> and even younger population is getting afflicted with it.<sup>13</sup> A study conducted at Karachi

observed 8% prevalence of GDM.<sup>14</sup> Other studies conducted in different cities of Pakistan showed a range of 15.7% to 24%.<sup>15,16</sup> Early detection and treatment of GDM is of utmost importance to prevent obstetrical and perinatal implications like miscarriages, birth defects, macrosomia, unexplained IUDs, shoulder dystocia, polycythemia, respiratory distress syndrome (RDS), hypoglycaemia, hypocalcaemia, hyperbilirubinemia; childhood risks like dyslipidemias and adiposity.<sup>17</sup> Maternal risk factors include preeclampsia, repeated urinary tract infections (UTIs),<sup>18</sup> vaginal infections, polyhydramnios, instrumental deliveries, perineal tears and increased chances of C sections. One third of women develop type II DM later in life.

For early detection of GDM screening is required as it is an asymptomatic metabolic syndrome. Screening recommendations about tests applied and timing of screening and whether to do universal or selective screening vary among different organizations due to lack of properly conducted randomised controlled trials (RCTs).<sup>19</sup> Screening methods include risk factors based screening, fasting plasma glucose, timed random blood sugar (RBG), HbA1C and 50 g GCT. Yet no screening test is validated.

American Diabetes Association (ADA) and American college of obstetricians and gynaecologists (ACOG) recommend screening by 50 g GCT (threshold 7.2mmol/l or 7.8mmol/l can be used).<sup>20,21</sup> Either of the thresholds can be used. Even for diagnostic, there is lack of universally accepted "gold standard". ADA and ACOG recommend 100 g oral glucose tolerance test (OGTT) while WHO recommends 75 g OGTT as diagnostic test. The question whether selective or universal screening is better is still unanswered. In Canada and USA, universal screening is done as recommended by ACOG and in UK risk factor based screening is practised. ADA recommends universal screening. Australasian Carbohydrate Intolerance Study (ACHOIS) demonstrated

✉ **Dr. Ayesha Arif**  
Associate Professor & Classified gynaecologist  
Department of Gynaecology and Obstetrics  
Combined Military Hospital  
Nowshera  
E-mail: drayshaarif2013@yahoo.com

✉ **Dr. Sarwat Nazar**  
Assitant Professor  
Department of Gynaecology and Obstetrics  
Shifa International Hospital  
Islamabad

✉ **Dr. Sadia Arif**  
Assitant Professor  
Department of Gynaecology and Obstetrics  
PNS Hafeez Hospital  
Islamabad  
Received: 04-11-2015  
Revised: 24-01-2016  
Accepted: 02-02-2016

improved perinatal outcome by formal screening of whole obstetrical population. A commentary on this trial published in BJOG 2006, also supports this recommendation.<sup>22</sup> Canadian task force on preventive health care does not support for or against universal screening for GDM. General recommendation is to conduct risk assessment and then glucose testing for high risk women on 1<sup>st</sup> antenatal visit followed by retesting at 24-28 weeks of gestation. Average risk women should be screened at 24-28 weeks of gestation as recommended by ACOG. In short a single approach to testing of GDM cannot be recommended at present because of lack of evidence based data.

Purpose of this study was to establish efficacy of 50 g GCT as a screening test of GDM in our population as very few local studies are available to guide us in this regard.

**MATERIALS AND METHODS:**

This cross sectional study was conducted from 1<sup>st</sup> Aug 2012 to 30<sup>th</sup> July 2013 at the obstetrical outpatient clinic of Combined Military Hospital, Lahore. Patients were selected through non probability convenient sampling. 100 patients carrying singleton pregnancy either primigravida or multigravida within age group of 20-35 yrs, booked in 1<sup>st</sup> trimester were included in this study. Patients with history of type I or II DM, history of glucose intolerance in the past, with bad obstetrical history, family history of DM, IUDs, still births or early neonatal deaths, congenital anomalies, macrosomic babies and patients with polyhydramnios were excluded. After taking the consent, patients between 24-28 weeks of gestation were tested with 50 g GCT, regardless of previous state of fasting. Venous plasma glucose levels were measured by taking sample of blood one hour after administering the glucose drink using glucose oxidase hexokinase method. A glucose value of 140 mg/dl was taken as cut off. Regardless of the results, all patients were further evaluated with 3 hours 100 g OGTT. Patients with two or more values of blood glucose equal to or exceeding the proposed values were labelled as having GDM and those with one abnormal value were labelled to have impaired glucose tolerance. Values proposed by Carpenter and Coustan and adapted by 4<sup>th</sup> international workshop conference on GDM were used, which are: fasting- 95 mg/dl, 1 hour after glucose load- 180 mg/dl, 2 hours after glucose load - 155 mg/dl and 3 hours after glucose load - 140 mg/dl. Data was collected on a pre designed proforma and was analysed using computer software (SPSS 10). Validity of 50 g GCT was measured in terms of sensitivity, specificity, positive and negative

predictive value.

**RESULTS:**

Total of one hundred patients were evaluated in this study. 19 were screen positive, whereas 81 were screen negative (Table 1). Among screen positive, majority were of greater than 28 years of age i.e. 57% (11 out of 19) and multi or grand multi gravidas i.e. 78.9 % (15 out of 19). Screening was negative mostly in primigravidas i.e. 69.1% (56 out of 81) and in patients with age of less than 28 years i.e. 71.6 % (58 out of 81). All the patients were put to 100 g 3 hrs OGTT as gold standard diagnostic test. Out of these, 13 (13%) were labelled to have GDM due to either one impaired glucose tolerance (IGT) or two abnormal values (frank DM) according to Carpenter and Coustan’s criteria. 76.9% of patients labelled with GDM on 100 g OGTT were screen positive initially, while 23 % of these patients were not picked up on initial screening alone. Out of 13 patients with GDM on OGTT, 69.2% had only impaired glucose tolerance while 30.7% had frank Diabetes (Table 2). Out of 19 screen positive patients, 10(52.6%) had abnormal OGTT as well; so were labelled as true positive. 9 out 19 (47.3%) had normal OGTT; so were labelled as false positive. Out of 81 screen negative patients, 78 (96.3%) came out to be true negative. They had normal OGTT as well. Whereas 3 out of 81(3.7%) had abnormal OGTT, so were labelled as false negative.

According to this study, sensitivity of 50 g GCT was calculated to be 76.92%, specificity of 89.6%, positive predictive value of 52.6% and negative predictive value of 96.2%. (Table 3).

Table: 1  
Results of 50 g GCT

Results of screening	No of patients	%
Screen positive	19	19%
Screen negative	81	81%

Table: 2  
Patients with GDM on OGTT

Total no of patients labelled as Gestational Diabetics	Screen positive	Screen negative	GDM	IGT
13(13%)	10 (76.9%)	3 (23.07%)	4/13 (30.7%)	9/13 (69.2%)

Table: 3  
Validity of 50 g GCT

	Formula	Result	Percentage
Sensitivity	TP/(TP+FN)*100	10/(10+3)*100	76.92%
Specificity	TN/(TN+FP)*100	78/(78+9)*100	89.6%
Positive Predictive Value	TP/(TN+FP)*100	10/(10+9)*100	52.6%
Negative Predictive Value	TN/(FN+TN)*100	78/(3+78)*100	96.2%

**DISCUSSION:**

The high frequency of GDM in Asian (Pakistani population)<sup>14,15,16</sup> and its foetal and maternal implication emphasize the significance of timely diagnosis and management of GDM. As it is an asymptomatic metabolic syndrome so for detection of preclinical disease, screening is required. There is no consensus about time of screening, test to be applied, various thresholds for screening tests and which population should be screened (universal or selective).<sup>23</sup> Systematic screening of pregnant population is still not common in Pakistan despite of the fact that subcontinent is included in high risk population for GDM by most of authorities. There is a need to conduct study about how to screen, which population to screen in Pakistan and to develop a country wide protocol.

In this study, we have evaluated the validity of 50 g GCT as a screening tool for GDM. This was a small study and on low risk patients. Although this study showed very encouraging results to apply GCT as a screening tool between 24-28 weeks of gestation but a larger scale study is still required for on average high risk patients to validate its results. In this study, 19% came out to be screen positive and 81% were negative. This is consistent with many international studies which show that 14 -18% of patients were screen positive if threshold of GCT was taken as = 140mg/dl and 20 - 25% with 130 mg/dl. Most of the screen positive patients were multigravidas (47.3%) in patients with age >28 years, whereas screen was negative in primigravidas. This is consistent with the study of Maresh.<sup>24</sup> All the patients were put to 100 g OGTT regardless of results of screening and 13 patients were labelled as having GDM. It is important to note that all the 3 patients who were screen negative initially but had abnormal 100 g OGTT results had only IGT (Impaired Glucose Tolerance) and none had frank DM. Among the patients who were screen positive, 6 out of 10 had impaired glucose tolerance and 4 had two abnormal values (criteria for GDM). The validity of 50 g GCT was calculated in term of sensitivity, specificity, positive predictive value and negative predictive value which came out to be 76.9%, 89.6% 52.6% and 96.2% respectively. This is in consistence with a study that showed sensitivity of 80% and specificity of 90%.<sup>25</sup> According to these results, we recommend that 50 g GCT, as a screening test for GDM, should be applied to all pregnant ladies between 24-28 weeks of gestation with a threshold of 140 mg/dl, with high sensitivity and specificity and also as a simple method, suitable for all pregnant women. Difficulties encountered were to convince the ladies for these special tests, extra financial burden, and nausea/vomiting associated with glucose intake. Some of large hospitals have instituted this test as an essential part of antenatal clinic services. But still a lot of work is required in this regard to create a nationwide strategy. This study was a small effort to develop a fixed framework of screening of GDM for Pakistani population.

**CONCLUSION:**

50 g GCT is an effective screening tool for GDM between 24-28 weeks of gestation with high sensitivity and specificity. It picked up almost all the cases with GDM or IGT between 24-28 weeks of gestation. This can not only help us to improve perinatal outcome, but also to identify ladies who are at high risk of developing type II DM in future. A large scale population based study is recommended to further validate the findings of this study.

**REFERENCES:**

1. Diagnosis and classification of diabetes mellitus. American Diabetes Association. *Diabetes Care* 2006; 29 Suppl: 1: S43-8
2. WHO. Diabetes Mellitus Fact sheet No. 312. WHO fact sheet No.138 [Online] 2008 (Cited 12 Jan 2009); Available from URL: <http://www.who.int/mediacentre/factsheets/fs312/en/index.html>
3. Shirazian N, Emdadi R, Mahboubi M, Motevallian A, Fazel-Sarjuei Z, Sedighpour N, et al. Screening for gestational diabetes: usefulness of clinical risk factors. *Archives of Gynecology and Obstetrics* 2009; 280: 933-7
4. Hadaegh F, Tohidi M, Harati H, Kheirandish M, Rahimi S. Prevalence of gestational diabetes mellitus in southern Iran (Bandar Abbas City). *Endocrine Practice* 2005; 11: 313-8
5. Coustan DR: Gestational diabetes. In *Diabetes in America*. 2nd ed. Harris MI, Ed. Bethesda, Maryland, National Institutes of Health, 1995, p. 703-16
6. Zargar AH, Sheikh MI, Bashir MI, Masoodi SR, Laway BA, Wani AI, et al. Prevalence of gestational diabetes mellitus in Kashmiri women from the Indian subcontinent. *Diabetes research and clinical practice* 2004; 66: 1 39-45
7. Metzger BE, Gabbe SG, Persson B. International Association of Diabetes and Pregnancy study groups recommendations on the diagnosis and classification of hyperglycemia in pregnancy. *Diabetes Care*. 2010;33:676-82
8. Senanayake H, Seneviratne S, Ariyaratne H, Wijeratne S. Screening for gestational diabetes mellitus in Southern Asian women. *Journ Obstet Gynecol Res* 2006; 32(3): 286-91
9. Friedman S, Khoury-Collado F, Dalloul M, Sherer D M, Abulafia O. Glucose challenge test threshold values in screening for gestational diabetes among black women. *AJOG* 2006; 194: e46-e48
10. Cypryk K, Szymezak W, Czupryniak L, Sobezak M, Lewinski A. Gestational diabetes mellitus – an analysis of risk factors. *Pol Journ Endocrinol* 2008; 59(5): 393-7
11. Nicholson W K, Fleisher L A, Fox H E, Powe N R. Screening for gestational diabetes mellitus. *Diabetes Care* 2005; 28(6): 1482-4
12. Shera AS, Rafique G, Khawaja IA, Baqai S, King H. Pakistan National Diabetes Survey: prevalence of glucose intolerance and associated factors in Baluchistan province. *Diabetes Res Clin Pract* 1999; 44: 49-58
13. Nishtar S, Bile KM, Ahmed A, Faruqui AM, Mirza Z, Shera S, et al. Process, rationale, and interventions of Pakistan's National Action Plan on Chronic Diseases. *Prev Chronic Dis* 2006; 3: A14

14. Iqbal R, Rafique G, Badruddin S, Qureshi R, Cue R, Gray-Donald K. Increased body fat percentage and physical inactivity are independent predictors of gestational diabetes mellitus in South Asian women. *Eur J Clin Nutr* 2007; 61: 736-42
15. Shaukat A, Arain T M, Abid S, Mahmud R. Criteria for detecting gestational diabetes mellitus: National Diabetes Data Group versus World Health Organization. *Journ Coll Physicians Surg Pak* 1999; 9(5): 211-4
16. Naheed F, Kammeruddin K, Hashmi H A, Narijo S. Frequency of impaired oral glucose tolerance test in high risk pregnancies for gestational diabetes mellitus. *JCPSP* 2008; 18(2): 82-5
17. Wright CS, Rifas-Shiman SL, Rich-Edwards JW, Taveras EM, Gillman MW, Oken E. Intrauterine exposure to gestational diabetes, childhood adiposity and blood pressure. *Am J Hypertens* 2009; 22: 215-20
18. Sicaardi DC. Gestational Diabetes. *Obstetrics and Gynecology-Med Students*. Available at: Accessed March 15, 2005
19. Dornhorst A, Williamson C. Diabetes and endocrine disease in pregnancy. In: Edmond D K, ed. *Dewhursts Textbook of Obstetrics and Gynaecology*. 7th edition. London. Blackwell Publishing 2007; 246-59
20. American College of Obstetricians and Gynaecologists. *Gestational Diabetes*. ACOG practice bulletin # 30. American College of Obstetricians and Gynaecologists, Washington, DC 2001.
21. American Diabetes Association. Standard of medical care in diabetes. *Diabetes Care* 2005; 28: 4-36
22. Tuffnell D, West J, Walkinshaw S. Time to screen for and treat gestational diabetes. *Br Journ Obstet Gynaecol* 2006; 113: 3-4
23. Dornhorst A, Williamson C. Diabetes and endocrine disease in pregnancy. In: Edmond D K, ed. *Dewhursts Textbook of Obstetrics and Gynaecology*. 7th edition. London. Blackwell Publishing 2007; 246-59
24. Maresh M. Screening for gestational diabetes mellitus. *Seminars in Fetal and Neonatal Medicine* 2005; 10: 317-23
25. Jarrett RJ. Should we screen for gestational diabetes? *BMJ* 1997;315:736-7



# Effects of Combined Regimens of Gabapentin and Verapamil with Diazepam on Kindled Model of Epilepsy in Mice

Itefaq Hussain Qureshi<sup>1</sup>, Shabana Usman Simjee<sup>2</sup>

## ABSTRACT:

**Objective:** To compare the in vivo effects of anticonvulsant combined regimens of Gabapentin / Verapamil with diazepam on kindled model of Epilepsy in Mice.

**Materials and Methods:** This experimental study was carried out in Hussain Ebrahim Jamal (H.E.J.) Research Institute of Chemistry, International Center for Chemical and Biological Sciences, University of Karachi, Karachi from May 2009 to July 2011. Gabapentin and Verapamil were used as tested drugs while Diazepam was used as a reference drug. Kindling was produced by repeated administration of Pentylentetrazole in a dose of 50 mg/kg by subcutaneous route every 48 hours for 20 days. Six doses of Gabapentin from 50mg/kg to 300mg/kg and six doses of Verapamil from 5mg/kg to 30mg/kg in combination regimen were administered by intraperitoneal route. Diazepam was administered by intraperitoneal route in a dose of 7.5mg/kg. Both tested drugs Gabapentin and Verapamil with reference drug Diazepam were administered once daily, however on the day of Pentylentetrazole treatment the tested and reference drugs were injected 40 minutes before injecting Pentylentetrazole. The anticonvulsive effects of tested drugs were then compared to reference drug Diazepam.

**Results:** Combination regimens of Gabapentin and Verapamil exhibited synergistic dose dependent anti-seizure effects up to 100%. The maximum dose of combined regimen exhibited antiseizure effects which were superior to the reference drug Diazepam.

**Conclusion:** Combination regimens of Gabapentin and Verapamil showed synergistic effect superior to diazepam on kindled model of epilepsy in mice.

**Keywords:** Antiepileptic drugs (AED), Diazepam (DZ), Gabapentin (GBP), Verapamil (VP), Pentylentetrazole (PTZ)

## INTRODUCTION:

Epilepsy is one of the most common neurological disorders which has no age, racial, social, sexual and geographical boundaries. The International League Against Epilepsy (ILAE) defines epilepsy as: "A transient occurrence of signs and symptoms due to abnormal excessive or synchronous neuronal activity in the brain."<sup>1</sup> About 50 million people around the world are suffering from Epilepsy.<sup>2</sup> In Pakistan 1.38 million epileptic patients have been reported.<sup>3</sup> The incidence of Epilepsy in Pakistan is 9.9 per 1000 of the general population.<sup>4</sup> Epileptic foci are a potential site for the generation of epileptic seizures and have high levels of neuronal activity either due to abnormally increased excitatory neurotransmitters or abnormally decreased levels of inhibitory neurotransmitters. The clinical manifestation of epileptic seizures depends upon the affected cortical area from where the seizures originate. There are various pathological causes of Epilepsy resulting in abnormal neuronal discharge or neuronal

and synchronicity such as trauma, infections, neoplasm, abscesses, cysts, metabolic abnormalities, stroke, chronic degenerative disease of the central nervous system and genetics.

It is essential to differentiate between epileptic seizures and general non-epileptic seizures like febrile seizures, hypoglycemic seizures, alcohol and narcotic withdrawal seizures, seizures due to poisoning and diabetic ketoacidosis. In non-epileptic seizures the mechanism of initiation of seizures is almost the same that is hyperexcitability; however, the neurons are hyperexcitable for a limited period of time and not permanent like epileptic foci. The genetics of epilepsy is a very complex phenomenon and new progress has been made in the last 2 decades which shall provide new possibilities for the diagnosis and treatment of inherited disorders of epilepsy. Mutations in the genes which code for the different ionic channels are one of the main causes of inherited epileptic syndromes.<sup>5</sup> These mutational channelopathies result in defective voltage gated ionic channel formation which control flow of sodium, potassium and calcium ions in and out of the neuronal cells.<sup>6,7</sup>

The Pharmacoresistant or refractory epilepsy presents in about 30 percent of the epileptic patients.<sup>8</sup> This pharmacoresistant epilepsy is the most difficult type of epilepsy to be treated.<sup>9,10</sup> The clinical condition can be improved by add-on therapy with newer antiepileptic drugs like GBP and calcium channel blockers like Verapamil (VP). Gabapentin (GBP) is approved as adjunct therapy for partial as well as for generalized tonic clonic seizures. High doses of GBP are needed for improvement in seizure control, however, the high doses are mostly tolerable and its safety and tolerability is rated as good

✉ **Dr. Itefaq Hussain Qureshi**

Assistant Professor  
Liaquat College of Medicine & Dentistry  
Karachi  
Email: drittafaq@gmail.com

✉ **Dr. Shabana Usman Simjee**

Assistant Professor  
H.E.J. Research Institute of Chemistry  
International Center for Chemical and Biological Sciences  
University of Karachi  
Karachi

Received: 14-12-2015

Revised: 12-01-2016

Accepted: 15-01-2016

to excellent.<sup>11,12,13,14</sup> VP is a calcium channel blocker which is widely used as antianginal, antiarrhythmic and antihypertensive drug in patients of coronary heart disease. In heart it acts on rapidly firing L-type voltage gated calcium channels and blocks T-type voltage gated calcium channels in central nervous system. Antiseizure effects of VP have been noted in some clinical randomized trials in pharmaco-resistant epilepsy.<sup>15,16,17</sup> It affects P-glycoprotein expression at various sites including blood brain barrier. Some patients of refractory epilepsy were also found suffering from severe myoclonic epilepsy of infancy. VP when used as an adjunctive therapy successfully controlled seizures. Even on long term usage it has given promising results by its modulating effects on calcium channels.<sup>18,19</sup> Defective calcium channels due to inherited genetic mutation causes various types of epilepsies.<sup>20,21</sup> Until today, there are no satisfactory and approved treatment regimens for pharmaco-resistant refractory epilepsy.<sup>22</sup> GBP and VP are voltage-gated calcium channel blockers; therefore, they can be a potential candidate for the treatment of different kinds of epilepsies.<sup>23,24,25,26,27,28</sup> Present study was designed to compare the anticonvulsant effects of combined regimens of GBP/ VP with DZ on kindled model of epilepsy in mice.

#### MATERIALS AND METHODS:

This experimental study was carried out at Hussain Ebrahim Jamal (H.E.J) Research Institute of Chemistry, International Center for Chemical and Biological Sciences, University of Karachi, Karachi from May 2009 to July 2011. The use of animals in this study was approved by the Scientific Advisory Committee on Animal Care, Use, and Standards, International Center for Chemical & Biological Sciences, University of Karachi, Pakistan, in accordance with the international guidelines for the care and use of laboratory animals. Male NMRI albino mice weighing 20-25 g were used. The group size of 12 were used which had 80% power to detect differences in the means.

The anti- epileptic activity of the GBP and VP were evaluated in vivo by chemically-kindled model of epilepsy. Total duration of study was forty days. The mice were divided into nine groups that is G-I to G-IX. Each group had twelve mice. G-I (normal control) was given only 0.9 percent normal saline. G-II was given PTZ only. G-III to G-VIII was given tested drugs GBP and VP. G-IX was given DZ and PTZ. The kindling was produced in GII group by repeated administration of sub-convulsive dose of PTZ (50 mg/kg, s.c.) every 48 hours for 20 days. The test drugs VP and GBP were administered to the mice intraperitoneally (i.p.) in six different doses. The reference drug DZ was also administered intraperitoneally (i.p.) in a dose of 7.5mg/kg in G-IX. The test drugs GBP and VP and the reference drug were given daily, however, on the day of PTZ-

treatment which was given on every alternate day, the drugs were administered 40 minutes before injecting PTZ. The resultant kindling scores were classified as numerical 1 to 5. The animals showing the score 4-5 on 20<sup>th</sup> day of treatment in G-II PTZ treated were considered to be fully kindled. The mean of the seizure scores were calculated showing results of all groups mean of seizure scores of kindling standard scores from 1 to 5  $\pm$  SEM in 12 mice (n = 12 per group). The mean of the seizure scores were converted into percentage of seizure scores and seizure protection in all nine groups.

**Statistical Analysis:** The statistical analysis was performed using Statistical Package for the Social Sciences (SPSS) version 10 and Graph Pad Prism. Results were reported as mean  $\pm$  SEM. Data of seizure activity was analyzed by nonparametric Student's t-test and ANOVA with post hoc Dennett's multiple comparison tests. The sequential differences among means were calculated at the level of  $p < 0.05$ .

#### RESULTS:

The results of kindling scores were classified as numerical 1 to 5 (Table 1). The data from combined usage of GBP and VP when analyzed showed a dose dependent synergistic anti-epileptic activity exhibiting 33.4%, 41.8%, 45%, 55%, 77% and 100% seizure protections in six different dose regimens respectively (Table 2). Thus, at the maximum dose employed, the combination regimen of GBP and VP exhibited superior anti-epileptic activity in terms of seizure protection capability compared to the reference drug DZ. (Figure 1). GBP and VP at the maximum dose at GVIII group exhibited 100% seizure inhibition and seizure score was 0.00%, while DZ exhibited 91.8% seizure inhibition. Combination regimen of GBP and VP exhibited 8.2% more seizure protection with almost zero seizure score (100% seizure inhibition) compared to DZ. Though the effect is dose dependent, however, the therapeutic index of GBP is much higher than DZ and human maximum dosage of GBP is 4.5 gm per day (Table 2, Figure 1).

Table:1  
Five distinct seizure patterns used for scoring kindling stages

Seizure scoring in kindling by PTZ	Seizure Pattern
0	No Response
1	Ear and Facial Twitching
2	Convulsive Wave through the body
3	Myoclonic Jerks
4	Clonic-Tonic Convulsions, Turnover into Side Position
5	Generalized Clonic-Tonic Seizures, Loss of Postural Control.

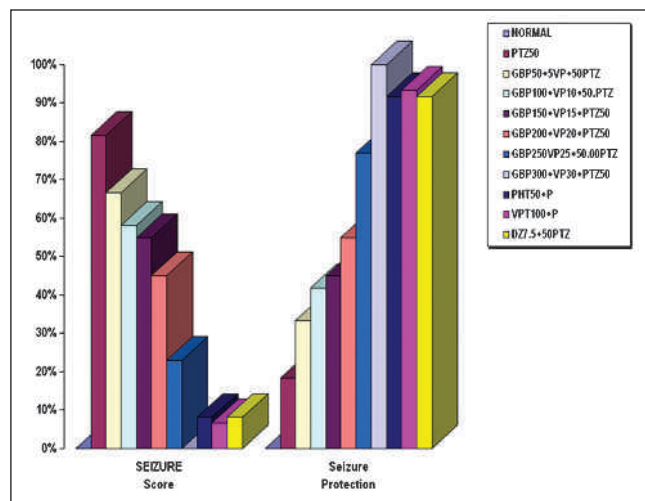
Table: 2  
Effect of Gabapentin and Verapamil treatment on the seizure score and seizure protection against PTZ-induced kindling

Group ID	Treatments	Dosage (mg/kg)	Mean ± SEM of Seizure Score	Seizure Score (%)	Seizure Protection (%)
GI	Normal Control	0.9% Saline	0.00	0.00%	0.00%
GII	PTZ	50	4.08 ± 0.64	81.6%	18.4%
GIII	GBP: VP:PTZ	50: 5:50	3.33 ± 0.74	66.6%	33.4%
GIV	GBP: VP:PTZ	100: 10:50	2.91 ± 0.49	58.2%	41.8%
GV	GBP: VP:PTZ	150: 15:50	2.75 ± 0.43	55%	45%
GVI	GBP: VP:PTZ	200: 20:50	2.25 ± 0.82	45%	55%
GVII	GBP: VP:PTZ	250: 25:50	1.16 ± 0.89	23%	77%
GVIII	GBP: VP:PTZ	300: 30:50	0.00	0%	100%
GIX	DZ:PTZ	7.5:50	0.41 ± 0.14	8.2%	91.8%

Both the seizure score and seizure protection were calculated in % and the data represented as a Mean SEM of n = 12 animals per group

Figure: 1

Graphical representation of synergistic effect of gabapentin and verapamil treatment on the seizure score and seizure protection against PTZ-induced kindling



The mean SEM of seizure score and seizure protection are shown in %

**DISCUSSION:**

Current treatment and management of epilepsy by antiepileptic drugs is not satisfactory and though, availability of newer antiepileptic drugs have widened the choices of the clinicians for the treatment of epilepsy, however; the prognosis and efficacy of these new drugs are still disappointing.<sup>29</sup>The results of our study showed that combined regimens of GBP and VP when analyzed and compared exhibited a dose dependent synergistic anti-epileptic activity starting from 150:15 mg/kg of GBP: withVP exhibiting 45 % seizure protection. This further increased to 55 % seizure protection at the dose of 200: 20 mg/kg of GBP:VP reached maximum that is 100 % seizure protection at 300:30 mg/kg of GBP: VP. When the synergistic effects of GBP and VP were compared to DZ we observed the differences in seizure inhibition of 8.2 % at the maximum dose of combination regimen which was superior to the anti-seizure effects of DZ. Though the effect is dose dependent, however,

the therapeutic index of GBP is much higher than DZ and human recommended maximum dosage of GBP is 4.5 g per day. We are therefore, inclined to hold that the dose dependent superior anticonvulsant synergistic effects of GBP and VP compared to antiseizure effects of standard single dose of DZ had insignificant chances of error. When the synergistic effects of GBP and VP were compared to PTZ control we observed that the seizure inhibition was 17%, 26%, 29%, 39%, 61% and 84% in six dose regimens of combination therapy, demonstrating inhibition of PTZ seizure effects of 67%, 58%, and 55%, 45%, 23% and 0.00 % respectively. At the maximum dose of combination regimen we observed complete inhibition of PTZ induced seizures, which was beyond doubt, superior to the antiseizure effects of reference drug given with PTZ.

In one study, it was observed that animal models using subcutaneous Pentylenetetrazole, is a common model to study the antiseizure effects and mechanism of action of antiepileptic drugs. It was further observed in the same study that animal models can be used to evaluate different combinations of AEDs before their use in humans.<sup>30</sup>In another study it was observed that GBP as monotherapy had not exhibited effective antiseizure effects however, the study revealed that combinations of GBP with other antiepileptic drugs generally exhibited synergistic interactions. The study concluded that GBP had exhibited synergistic effects with other antiepileptic drugs in same dosage.<sup>31</sup>Our results are coinciding with the findings of these studies.

The most important question is how the instant combined regimen of GBP with VP is justified for providing better, safer, synergistic, effective and broader spectrum therapeutic choice for the treatment of various types of epilepsies as compared to other AEDs. Firstly, the mode of action of GBP can be augmented or modified if given in combination with other drugs like calcium channel blockers i.e. verapamil.<sup>32</sup>Secondly, GBP is a very safe drug as compared to other AEDs.<sup>33</sup>Thirdly, the GBP has a very high therapeutic index and safety profile.<sup>34</sup>Fourthly, instant novel regimen of GBP and VP can be employed with adjustment of their doses from minimum to

maximum doses for the treatment of different types of epilepsies without any significant adverse effects. Fifthly, doses of GBP up to 4.5 g/day can be given with low doses of VP to achieve the therapeutic goals which are not possible with other antiepileptic drugs. Sixthly, GBP in various studies has demonstrated its efficacy as monotherapy equivalent to that of carbamazepine (CMZ) for partial and generalized seizures. GBP has also established its efficacy in refractory epilepsy, therefore the combination regimen would have wider spectrum to treat various types of epilepsies.<sup>35</sup> Seventhly, the side effects of GBP are few, tolerable and short term. Eighthly, GBP has been approved by the drug agency FDA as a monotherapy for partial and complex partial seizures with or without generalized tonic-clonic seizures.<sup>36</sup> Lastly most of the antiepileptic drugs have potential for causing hepatitis and bone marrow suppression like Valproate and Carbamazepine.

### CONCLUSION:

The novel combination regimen of GBP with VP has potential for alternate regimen of treatment for different types of seizures including epileptic seizures and would provide better, safer, synergistic and effective therapeutic effect because of its synergistic and channel modulating effects not only in drug resistant epilepsy but also in various other types of epilepsies. This study has provided basic ground-work guidelines for the future clinical use of combination therapies of GBP and VP in different doses combinations in various forms of epilepsies for the long term management of epilepsy.

### REFERENCES:

1. W.H.O, IBE, ILAE, Global Campaign Against Epilepsy, Alas, 2005
2. Khatria A, Iannaccone ST, Ilyas B M S, Abdullah B M , Saleem S. Epidemiology of Epilepsy in Pakistan: Review of literature. *J Pak Med Assoc.* 2003; 53(12): 594-7
3. Sheerani M. Development of a comprehensive epilepsy surgery program in Pakistan. *J. Pak. Med. Assoc.* 2005; 55(1): 32-7
4. Annegers JF, Rocca WA, Hauser WA. Causes of epilepsy: contributions of the Rochester epidemiology project. *Mayo Clin Proc.* 1996;71(6):570-5
5. Khosravani H, Zamponi GW. Voltage-gated calcium channels and idiopathic generalized epilepsies. *Physiol Rev.* 2006;86(3): 941-66
6. Khosravani H, Altier C, Simms B, Hamming KS, Snutch TP, Mezeyova J et al. Gating effects of mutations in the Cav3.2 T-type calcium channel associated with childhood absence epilepsy. *J Biol Chem.* 2004;279(11): 9681-4
7. Rossetti AO. Treatment options in the management of status epilepticus. *Curr Treat Options Neurol.* 2010;12(2):100-12
8. McTague A, Appleton R. Treatment of difficult epilepsy. *Arch Dis Child* 2011;96(2):200-4
9. Summers MA, Moore JL, McAuley JW. Use of verapamil as a potential P-glycoprotein inhibitor in a patient with refractory epilepsy. *Ann Pharmacother.* 2004; 38(10):163 1-4
10. Wahab A, Albus K, Gabriel S, Heinemann U. In search of models of pharmacoresistant epilepsy. *Epilepsia* 2010; 51 (Suppl 3):154-9
11. Dichter MA. Innovative clinical trial designs for future antiepileptic drugs. *Epilepsia* 2007; 48 (Suppl 1): 26-30
12. Goa KL, Sorkin EM. Gabapentin: A review of its pharmacological properties and clinical potential in epilepsy. *Drugs* 1993;46(3):409-27
13. Beydoun A, Fakhoury T, Nasreddine W, Abou-Khalil B. Conversion to high dose gabapentin monotherapy in patients with medically refractory partial epilepsy. *Epilepsia* 1998;39(2):188-93
14. Gee NS, Brown JP, Dissanayake VU, Offord J, Thurlow, Woodruff GN. The novel anticonvulsant drug, Gabapentin (Neurontin), binds to the alpha 2 delta subunit of a calcium channel. *J Biol Chem.* 1996; 271(10): 5768-76
15. Iannetti P, Parisi P, Spalice A, Ruggieri M, Zara F. Addition of Verapamil in the treatment of severe myoclonic epilepsy in infancy. *Epilepsy Res.* 2009;85(1): 89-95
16. Iannetti P, Spalice A, Parisi P. Calcium-channel blocker Verapamil administration in prolonged and refractory status epilepticus. *Epilepsia* 2005;46(6): 967-9
17. Khosravani H, Zamponi GW. Voltage-gated calcium channels and idiopathic generalized epilepsies. *Physiol Rev.* 2006;86(3): 941-66
18. Summers M.A, Moore JL, McAuley JW. Use of Verapamil as a potential P-glycoprotein inhibitor in a patient with refractory epilepsy. *Pharmacotherapy* 2004;38(10):1631-4
19. Giordanetto F, Knerr L, Wällberg A. T-type calcium channels inhibitors: a patent review. *Expert Opin Ther Pat.* 2011;21(1):85-101
20. Fletcher CF. Absence epilepsy in tottering mutant mice is associated with calcium channel defects. *Cell* 87.4 1996: 607-17
21. Miller LC, Drislane FW. Treatment of status epilepticus. *Expert Rev Neurother.* 2008;8(12):1817-27
22. Abend NS, Dennis J. Treatment of refractory status epilepticus: literature review and a proposed protocol. *Pediatric neurology* 38.6 2008 : 377-90
23. Stefani A, Spadoni F, Bernardi G. Gabapentin inhibits calcium currents in isolated rat brain neurons. *Neuropharmacol.* 1998;37(1): 83-9
24. Kwan P, Sills GJ, Brodie MJ. The mechanisms of action of commonly used antiepileptic drugs. *Pharmacol Ther.* 2001;90(1):21-34
25. Macdonald R L, Kelly KM. Mechanisms of action of currently prescribed and newly developed antiepileptic drugs. *Epilepsia.* 1994;35 (Suppl 4): S41-S50
26. Macdonald RL, Kelly KM. Antiepileptic drug mechanisms of action. *Epilepsia.* 1995;36 (Suppl 2): S2-S12
27. Mulley JC, Scheffer IE, Petrou S, Berkovic SF. Channelopathies as a genetic cause of epilepsy. *Curr Opin Neurol.* 2003;16(2):171-6
28. Stephen, J, Martin J. Pharmacotherapy of epilepsy *CNS drugs* 25.2, 2011: 89-107
29. Glauser T. ILAE treatment guidelines: evidence-based analysis of antiepileptic drug efficacy and effectiveness as initial monotherapy for epileptic seizures and syndromes *Epilepsia* 47.7 2006; 1094-120
30. Sills G J. The mechanisms of action of gabapentin and pregabalin *Current opinion in pharmacology* 6.1 2006; 108-13
31. Holmes GL. Animal model studies application to human patients *Neurology* 69.24 suppl 3 2007: S28-S32
32. Borowicz KK. Effect of gabapentin on the anticonvulsant



- activity of antiepileptic drugs against electroconvulsions in mice: an isobolographic analysis. *Epilepsia* 43.9(20-02): 956-63
33. White HS. Comparative anticonvulsant and mechanistic profile of the established and newer antiepileptic drugs *Epilepsia* 40.s5 (1999): s2-s10
34. Beydoun A, Fakhoury T, Nasreddine W, Abou-Khalil B. Conversion to high dose gabapentin monotherapy in patients with medically refractory partial epilepsy. *Epilepsia* 1998;39(2):188-93
35. Gracia-Fleta F. Gabapentin in 50 patients with epilepsy. *Revista de neurologia* 32.1 (2000): 45-9
36. Ramsay RE. Advances in the pharmacotherapy of epilepsy. *Epilepsia* 34.s5 (1993): S9-S16
37. Chadwick D W. A double-blind trial of gabapentin monotherapy for newly diagnosed partial seizures. *Neurology* 51.5 (1998): 1282-8



## ORIGINAL ARTICLE

# Evaluation of S-T Resolution by Streptokinase Therapy in Patients of Myocardial Infarction among the Age Group of more than 60 Years

Shaikh Nadeem Ahmad<sup>1</sup>, Musarrat Sultana<sup>2</sup>, Fuad Shiekh<sup>3</sup>, Syed Saud Hasan<sup>4</sup>, Shams-ul-Arfeen Qasmi<sup>5</sup>

### ABSTRACT:

**Objective:** To evaluate the benefit and efficacy of streptokinase therapy on ST-segment elevation resolution in different types of myocardial infarction in more than 60 years age group.

**Materials and Methods:** This Hospital based cross sectional study was conducted at National Institute of Cardiovascular Diseases (NICVD) of Karachi, Pakistan. The study included patients more than 60 years of age having different types of myocardial infarction. Fifty patients both male & female fulfilling the inclusion criteria for thrombolytic therapy were included. Baseline ECG was recorded before streptokinase infusion and repeated at completion of infusion, at 90 minutes, day 1 and day 2. Effect of streptokinase therapy (SK) on blood pressure, CKMB, and ST-segment resolution was also evaluated at 90 minutes, day 1, and Day 2.

**Results:** The mean systolic blood pressure was  $138.20 \pm 4.57$  and  $125.20 \pm 3.92$  pre and post SK therapy reflecting a percentage decrease of 9.40 and highly significant ( $P < 0.001$ ). The Diastolic blood pressure was decrease to 9.52% with a mean value of  $84.80 \pm 2.46$  and  $76.80 \pm 1.89$  before and after the Streptokinase therapy's, segment resolution at 90 minutes was decreased to 50.69 percent from the baseline and continued to decrease at Day-1 and Day-2 with a percentage reduction of 69.12 and 84.33 % respectively. The P values were highly significant ( $P < 0.001$ ).

**Conclusion:** Thrombolysis when given within 12 hours of the onset of symptoms, improves survival, is beneficial and effective. The magnitude of benefit is greatest when reperfusion is established early. Age itself should not be considered a contraindication for fibrinolysis.

**Keywords:** Streptokinase, ECG, ST-elevation, Myocardial infarction, Efficacy

### INTRODUCTION:

Myocardial infarction is a key component of the burden

of cardiovascular disease. Studying the trends in the incidence and outcome of myocardial infarction and of coronary disease mortality provides crucial insights into the determinants of heart disease which is essential to its treatment and prevention. It is important to recognize that the trends in the incidence and outcome of coronary disease are complex, likely multifactorial and evolve over time.<sup>1</sup>

In developed nations cardiovascular diseases are the leading cause of death and disability and also are increasing rapidly in the developing world. Among 75 years of age and older patients, mortality after acute coronary occlusion approaches 30% at 1 month and exceeds 50% at 1 year. However, despite evidence from several randomized trials that thrombolytic therapy has clear net benefits and is a cost-effective treatment in the elderly.<sup>2</sup>

Intravenous Thrombolysis in Acute Myocardial Infarction "in the Sixth ACCP Consensus Conference on Antithrombotic Therapy" recommended "that patients with ischemic symptoms characteristic of acute myocardial infarction for < 12 h who have ST-segment elevation or left bundle-branch block on the ECG should receive I/V fibrinolytic therapy unless they have contraindications."<sup>3</sup>

Acute myocardial infarction is one of the leading causes of death in the elderly, however clinical data reveals a disproportionately lower use of thrombolytics because of fear of complications especially intracranial hemorrhage. One study has documented that out of one hundred patients 77 (77%) were males and 23 (23%) were females. Mean age was  $73.39 \pm 5.29$  years. No patient developed intracranial hemorrhage. Use of streptokinase for acute myocardial infarction should therefore not be discouraged in the elderly.<sup>4</sup>

Streptokinase is a 1<sup>st</sup> generation fibrin non-specific thr-

✉ **Dr. Shaikh Nadeem Ahmad**  
Professor  
Department of Pharmacology  
Dow Medical College  
Dow University of Health Sciences (DUHS)  
Karachi  
Email: drnadeem\_ahamd@hotmail.com

✉ **Dr. Musarrat Sultana**  
Assistant Professor  
Department of Forensic Medicine  
Dow Medical College  
Dow University of Health Sciences (DUHS)  
Karachi

✉ **Dr. Fuad Shiekh**  
Assistant Professor  
Department of Pharmacology  
Dow Medical College  
Dow University of Health Sciences (DUHS)  
Karachi

✉ **Dr. Syed Saud Hasan**  
Professor  
Department of Pharmacology  
Dow Medical College  
Dow University of Health Sciences (DUHS)  
Karachi

✉ **Dr. Shams-ul-Arfeen Qasmi**  
Professor  
Department of Microbiology  
Baqai Medical University  
Karachi

Received: 07-1-2016  
Revised: 22-1-2016  
Accepted: 10-2-2016

ombolytic and biochemically a serine protease enzyme derived from certain strains of beta hemolytic streptococci.<sup>5</sup> It consists of a single polypeptide chain containing 414 amino acids. It was first used in 1958 in acute myocardial infarction and since then it has revolutionized the management of acute myocardial infarction.<sup>6</sup> Coronary atherosclerosis is by far the most frequent cause of ischemic heart disease and plaque disruption with superimposed thrombosis is the main cause of acute coronary syndrome of unstable angina, myocardial infarction and sudden death.<sup>7,8</sup> The true frequency of atherosclerosis is difficult, if not possible to accurately determine because it is a predominantly asymptomatic condition. More advanced lesions begin to develop when individuals are aged approximately 25 years. Plaque rupture is probably the most important mechanism underlying the unpredictable rapid progression of coronary lesions.<sup>9</sup> The role of platelets in acute coronary syndromes begins with the exposure of the sub-endothelium after plaque rupture. Thrombosis develops on a plaque either because the plaque tear open (rupture) exposing the highly thrombogenic core to blood in arterial lumen.<sup>10</sup>

ST-segment elevation is an excellent marker of acute coronary occlusion in which reperfusion therapy is needed. Patient with non ST elevation of myocardial infarction have a thrombotic stenosis in the affected artery but the artery is usually patent, in contrast ST-elevation myocardial infarction, the artery is occluded and at base line flow cannot be worsen, it can only improve.<sup>11</sup>

The most frequently use electrocardiographic criteria for identifying acute myocardial infarction is ST-segment elevation where ST-segments are (re) emerging as a clinical tool of great importance. Evaluating the response to thrombolytic therapy that early resolution of ST-segment elevation is a useful mean of assessing perfusion.<sup>12</sup>

Thrombolytic therapy is that early and sustained re-canalization prevents cell death, reduces infarct size, preserves myocardial function, and reduces early and late mortality.<sup>13</sup> The current evidences indicate that early thrombolytic therapy can limit extent of myocardial necrosis in evolving myocardial infarction may be early restoration of coronary blood flow, preserve left ventricular function and reduce mortality in patients with acute myocardial infarction (AMI).

Present study was designed with the objective to observe streptokinase therapy, in ST-segment elevation resolution, in age more than 60 years and in different types of myocardial infarction. Moreover also to observe the toxicity of administered streptokinase therapy.

#### **MATERIALS AND METHODS:**

This hospital based cross sectional study was carried out in 2005 for a total duration of 6 months. The study was conducted in the Department of Pharmacology and therapeutics, Basic Medical Sciences Institute Jinnah Post-graduate Medical Centre in collaboration with National Institute of Cardiovascular diseases (NICVD)

of Pakistan, Karachi. The study was approved by the postgraduate committee at NICVD and BASR of Karachi University. Informed consent for administration of thrombolytic drug was obtained from each patient. Inclusion criteria was patients diagnosed with myocardial infarction, more than 60 years of age, with chest pain suggestive of myocardial infarction, ECG findings of ST-Segments elevations. Exclusion Criteria was patients with myocardial infarction having active internal bleeding, cerebro-vascular accident, blood pressure > 200/100 mmHg, pregnancy, allergic reaction to streptokinase, previous coronary artery bypass Graft. Streptokinase (Streptofactor Hakimsons/Eskinase, Medinet), 1500000 units was used

#### **Criteria of ST-segment resolution:**

A positive ST-marker was defined as a reduction in ST-segment elevation of more than 50% within 90 minutes after the start of thrombolytic therapy.

#### **Treatment Plan:**

All patients fulfilling the inclusion criteria for thrombolytic therapy were included and admitted to either coronary care unit or place in the ward with and continuously monitored for arrhythmias. Baseline 12 lead electrocardiogram was taken. Two intravenous lines were maintained, one in each arm. One I/V line used for medication and another for collection of blood samples.

Blood sample for complete blood count, erythrocyte sedimentation rate, urea creatinine, blood glucose, cardiac enzymes and lipid profile, activated partial thromboplastin time.

Tablet aspirin 150 mg was given once for 24 hours. Isosorbide dinitrate I/V infusion 10-20 µg/min followed by oral nitrates

Streptokinase 1.5 million units dissolved in 100 ml 5% dextrose water infused in 60 minutes.

Vital signs 10 minutes during the infusion.

The 12 lead electrocardiograms were recorded. Baseline ECG recorded before streptokinase infusion and repeated at completion of infusion i.e. 90 minutes, day 1 and day 2.

#### **RESULTS:**

During the four months study period 50 patients were included in the study after fulfilling the inclusion criteria for thrombolytic therapy. Demography of patients with acute myocardial infarction exhibited that there were 44 (88%) males and 6 (12%) females, of these 50 patients 30 (60%) had an anterior wall infarction, while 20 (40%) suffered from an inferior wall infarction. No patient had a lateral wall acute myocardial infarction. Two patients died and cause of death was ventricular fibrillation in those patients (Table 1).

The mean systolic blood pressure was 138.20±4.57 and 125.20±3.92 pre and post SK therapy reflecting a percentage decrease of 9.40 and high significant (P<0.001). The Diastolic blood pressure was decrease to 9.52% with a mean value of 84.80± 2.46 and 76.80±1.89 before and after the Streptokinase therapy. ST-segment resolution at 90 minutes was decreased to

50.69 percent from the baseline and continued to decrease at Day-1 and Day-2 with a percentage reduction of 69.12 and 84.33 % respectively. The P values were highly significant (P<0.001) (Table 2)

There were 30 patients out of 50 with anterior wall Myocardial Infarction. The mean value of Systolic Blood Pressure (SBP) before therapy was 146.43± 4.98 and was decreased to 132.86±4.11 after therapy with Streptokinase. The Diastolic Blood Pressure (DBP) was decreased to 10.32 percent post Streptokinase therapy. The ST segment shows a resolution of 62.45% 69.99 and 87.14% at 90 minutes, day-1 and day-2 respectively. Figure-1 The P value for SBP, DBP and ST-segment resolution was highly significant (P<0.001).(Table 3a)

Twenty patients had inferior wall infarction in more than 60-years.(Table-3b)There was highly significant value of SBP, DBP and ST-segment resolution. The mean Systolic Blood Pressure value was 127.73±7.30 before therapy and decrease to 115.45±6.23 post streptokinase therapy which shows a percentage decrease of 9.61. The Diastolic Blood pressure showed a percentage decrease of 8.13. The ST elevation before therapy was 1.40± 0.16, which was resolved to 0.70± 0.13, 0.50 ± 0.13 and 0.33± 0.14 at 90-minutes, day-1 and day-2, showing a percentage decrease of 50.57, 64.28 and 76.92 respectively (Figure-2). The P value was also highly significant (P<0.001).

Table: 1  
Gender, site of myocardial infarction & mortality in age group of > 60 Years

Variables	Age group >60 years n= 50	Total %
Male	44	88%
Female	06	12%
Anterior Wall MI	30	60%
Inferior Wall MI	20	40%
Latral Wall MI	--	--
Death	2	4%

Table: 2  
Percentage changes from Pre to Post Streptokinase (SK) therapy

Variables	No of Observation	(Mean ± SEM)		% change Pre to Post	p-Value
		Pre SK Therapy	Post SK Therapy		
SBP (mmHg)	48	138.20 ± 4.57	125.20 ± 3.92	(-) 9.40	0.001***
DBP (mmHg)	48	84.80 ± 2.46	76.80 ± 1.89	(-) 9.52	0.001***
CKMB (IU)	48	50.92 ± 1.82	154.80 ± 4.25	204.00	0.001***
ST Resolution 90 min	48	2.17± 0.18	1.07± 0.12	(-) 50.69	0.001***
ST Resolution day 1	48	2.17± 0.18	0.67 ± 0.08	(-) 69.12	0.001***
ST Resolution day 2	48	2.17± 0.18	0.34 ± 0.08	(-) 84.33	0.001***

Pharmacological action of Streptokinase therapy on blood pressure, CKMB and ST-Segment elevation resolution  
\*\*\* Highly Significant(-) Shows decrease from pre to post streptokinase therapy

Table: 3a  
Effects of Streptokinase (SK) therapy according to the site of anterior wall myocardial infarction in >60 year of age  
There were 30 patients out of 50 with anterior wall Myocardial Infraction

Variables	No of Observation	(Mean ± SEM)		% change Pre to Post	p-Value
		Pre SK Therapy	Post SK Therapy		
SBP (mmHg)	30	146.43± 4.98	132.86± 4.11	(-) 9.26	0.001 ***
DBP (mmHg)	30	90.00± 4.11	80.71± 2.21	(-) 10.32	0.001 ***
CKMB (IU)	30	54.36 ± 2.51	154.00 ± 7.04	183.29	0.001 ***
ST Resolution 90 min	30	2.69 ± 0.17	1.01 ± 0.12	(-) 62.45	0.001 ***
ST Resolution day 1	28	2.69 ± 0.17	0.80 ± 0.10	(-) 69.99	0.001 ***
ST Resolution day 2	28	2.69 ± 0.17	0.35 ± 0.10	(-) 87.14	0.001 ***

Two patients died because of ventricular fibrillation \*\*\* Highly Significant(-) Shows decrease from pre to post streptokinase therapy

Table: 3b  
 Effects of Streptokinase (SK) therapy according to the site of  
 Inferior wall Myocardial Infarction in < 60 year of age  
 There were 20 patients out of 50 with inferior wall myocardial Infarction

Variables	No of Observation	(Mean ± SEM)		% change Pre to Post	p-Value
		Pre SK Therapy	Post SK Therapy		
SBP (mmHg)	20	127.73 ± 7.30	115.45±6.23	(-) 9.61	0.001 ***
DBP (mmHg)	20	78.18 ± 3.77	71.82 ± 2.63	(-) 8.13	0.001 ***
CKMB (IU)	20	46.55 ± 2.05	155.82±4.01	234.76	0.001 ***
ST Resolution 90 min	20	1.40 ± 0.16	0.70 ± 0.13	(-) 50.57	0.001 ***
ST Resolution day 1	20	1.40 ± 0.16	0.50 ± 0.13	(-) 64.28	0.001 ***
ST Resolution day 2	20	1.40 ± 0.18	0.33 ± 0.14	(-) 76.92	0.001 ***

Pharmacological action of Streptokinase therapy on blood pressure, CKMB and ST-Segment elevation resolution  
 \*\*\* Highly Significant(-) Shows decrease from pre to post streptokinase therapy

Comparative percentage change in the >60 age according to the site of myocardial infarction group, with anterior wall infarction showed a percentage change of 9.26, 10.32, 183.29 and 87.14 of SBP, DBP, CKMB and ST-segment resolution at day2, as compared to a percentage decrease of 9.61, 8.13, 60.4, 234.73, and 76.92 in patients with inferior wall myocardial Infarction at day-2(Table 4a) The mean duration of chest pain was 4.89± 0.28 hours. The minimum chest pain duration was 2 hours and maximum it was 12 hour. The mean stay at hospitals was at hospital was 5.03± 0.24 from a period of 5-days

to 15-days (Table 4b)  
 The complications of Streptokinase therapy in more than 60 years age patients was, that out of 50 patients two died because of ventricular fibrillation, which could have been due to reperfusion arrhythmias or the arrhythmias as a normal cardiac event in Myocardial Infarction.  
 ST-Resolution at Day-1 and Day-2, pre and post SK-therapy according to site of M-1 are mentioned in Figure 2 and 3.

Table: 4a  
 Pharmacological action of Streptokinase therapy on blood pressure, CKMB and ST-Segment elevation resolution

GROUP	Site of MI	SBP	DBP	CKMB	ST Resolution	Day-1	Day-2
		mmHg	mmHg	IU	90-Mints		
More than 60 years	Anterior Wall	(-) 9.26	(-) 10.32	183.29	(-) 62.45	(-) 69.99	(-) 87.14
	Inferior Wall	(-) 9.61	(-) 8.13	234.73	(-) 50.57	(-) 64.28	(-) 76.92

(-) sign shows decrease from pre to post SK-therapy

Table: 4b  
 Duration of chest pain, Cholesterol, Random blood sugar and duration of patients stay at hospital

Group	n	Duration Chest Pain Hours	Cholesterol (mg/dl)	RBS (mg/dl)	Hospital stay days
More than 60 years	50	4.89±0.28	218.84±8.67	132.16±13.82	5.03±0.24
		2-12	132-290	59-392	4-15

Figure: 1  
ST- Resolution according to site of myocardial infarction in more than 60 years age Patients

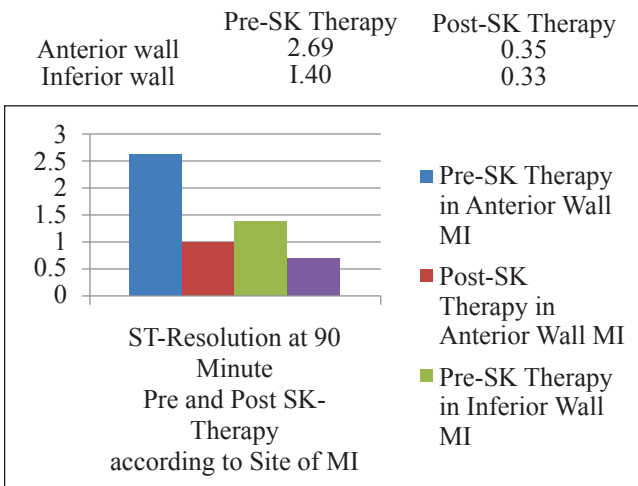


Figure: 2

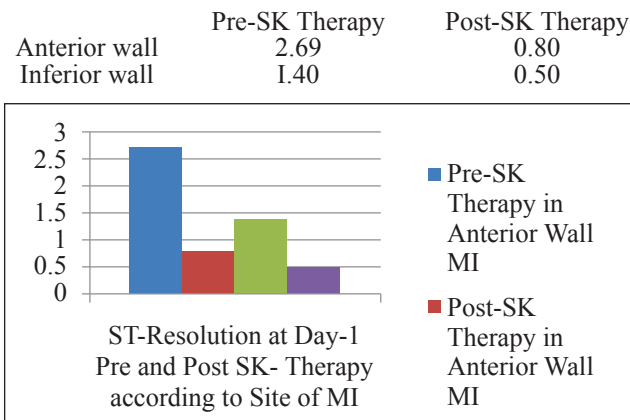
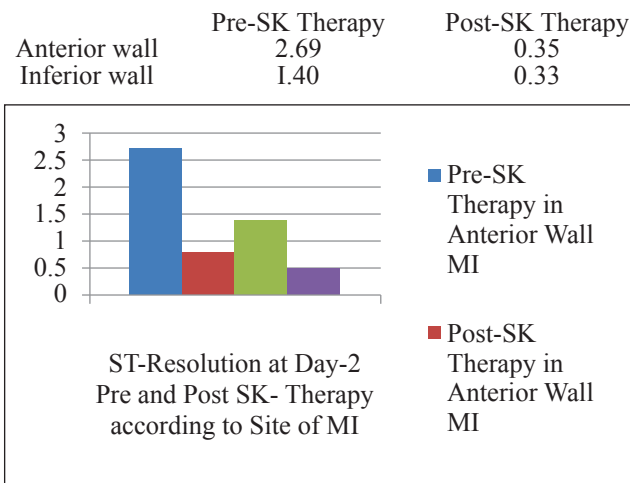


Figure: 3



related artery patency in the largest number of patients, but with the lowest rate of undesirable effects. Emergency management of acute myocardial infarction is evolving at an extremely rapid pace. What nearly all mortality reducing strategies have in common is, prompt restoration of blood flow to ischemic myocardium that has been compromised by intra-coronary thrombosis. Medical therapy alone is the preferred treatment in older patients after myocardial infarction. Indications of revascularization in older patients after myocardial infarction are prolongation of life and relief of unacceptable symptoms despite optimal medical management.<sup>14</sup>

Three clinical criteria have been proposed as markers for myocardial perfusion is reduction of chest discomfort (pain), improvement of electrocardiographic ST-segment elevation, and reperfusion arrhythmias. These clinical signs have been shown to be related to coronary artery recanalization and prognosis. Resolution of chest pain is very subjective and may frequently be related to analgesic medicine, cardiac arrhythmia could be a part of arrhythmias complicating acute myocardial infarction. Resolution of ST-segment elevation has been shown to be a simple and useful predictor of final infarct size, left ventricular function and clinical outcome after thrombolytic therapy.

Though the use of thrombolytic therapy decreases with increased age, but should not be considered a contraindication.<sup>15</sup> This study was conducted to observe the efficacy and complication of streptokinase therapy in more than 60 years age patient. The results of the present study suggest that streptokinase is effective and reduces the percentage resolution of ST-segment elevation. It is also suggested this therapy should be offered to all patients presenting with ST-segment elevation of acute myocardial infarction.

Our study matches with the study of Laurie<sup>16</sup> which provided careful and detailed analysis of trial with specific regard to beneficial-to risk ratio for patients. Our study matches with the GISSI-study<sup>17</sup> in which hospital mortality was 2 to 9 percent for patients 61 to 70 years old as compared to younger patients. In our study the in-hospital mortality was 4 percent in patients more than 60 years.

Present study has demonstrated rapid restoration of coronary blood flow in patients with evolving myocardial infarction. Our study matches with the study of Schroder<sup>18</sup>, who performed short term infusion of streptokinase in 93 patients within six hours after the onset of acute myocardial infarction.

Our study matches with Fibrinolytic Therapy Trials Collaborative (FTT) group study<sup>19</sup>, the data of the study do not provide evidence from withholding fibrinolytic therapy from patients on the basis of age. The excess of death in this study on day 0 to 1 increased with age but so did the reduction in death during days 2 to 35. The absolute mortality reduction seems much the same among younger and older patients. We do have early death in our study, two patients died within twelve hours of the start of therapy, whereas the patients discharged continue to do well.

**DISCUSSION:**

The best reperfusion treatment is one that achieves the highest rate of early, complete, and sustained infarct-

In patients with acute myocardial infarction, quick initiation of thrombolytic therapy is the best strategy for improvement of survival and reduction of morbidity. Since advanced age by itself is certainly not a contraindication to thrombolytic therapy, and because re-infarction occurs frequently, the benefit-risk ratio of re-exposure to streptokinase or its derivative is decreased in the elderly who present with reinfarction.<sup>20</sup> Our study did match with the study of,<sup>21</sup> thrombolytic therapy with streptokinase was found to be a beneficial and cost-effective treatment for suspected acute myocardial infarction in elderly patients in a wide variety of clinical circumstances. Acute myocardial infarction is one of the leading causes of death in the elderly, however clinical data reveals a limited use of thrombolytic because of fear of complications especially intracranial hemorrhage. Use of streptokinase for acute myocardial infarction should not be discouraged in the elderly.<sup>22</sup> This decade has witnessed the establishment of thrombolysis, the most widely available therapeutic intervention that specifically treats the direct cause of myocardial infarction and leads to biologically and clinically compelling patient outcome benefits.<sup>23,24,25</sup>

#### CONCLUSION:

Thrombolytic therapy with streptokinase is found to be a beneficial and cost-effective option for patients having suspected acute myocardial infarction. Thrombolysis when given within 12 hours of the onset of symptoms, improved survival, is beneficial and cost effective. Present study has demonstrated rapid restoration of coronary blood flow in patients with evolving myocardial infarction and the magnitude of benefit is greatest when reperfusion is established early. Age itself should not be considered a contraindication for fibrinolysis. Although intra-coronary application may be somewhat more effective, the advantage of intravenous administration is striking. Considering the experience of others we concluded that I/V short term infusion of streptokinase can be performed safely in patients with evolving myocardial infarction.

One limitation of the administration of an intravenous infusion of streptokinase is that it can cause a significant fall in systemic blood pressure and rapid infusion of high dose intravenous streptokinase frequently causes transient and sometimes severe fall in blood pressure, the magnitude of which is directly related to the rate of infusion of streptokinase.

#### REFERENCES:

1. Véronique L Roger. Epidemiology of myocardial infarction. *Med Clin North Am* 2007; 91(4):537-9
2. Fibrinolytic Therapy Trialists (FTT) Collaborative Group. Indications for fibrinolytic therapy in suspected acute myocardial infarction: collaborative overview of early mortality and major morbidity results from all randomized trials of more than 1000 patients. *Lancet*. 1994; 343:311-22
3. Ohman, EM, Harrington, RA, Cannon, CP Intravenous thrombolysis in acute myocardial infarction. *Chest* 2001;119, 253S-77S

4. Qureshi, AE, Jafri NA, Noeman A, Yasmin S, Khalil H. Streptokinase for acute myocardial infarction in the elderly. *J Ayub Med Coll Abbottabad*. 2014;26(4):535-8
5. Brogden RN, Speight TM, Avery GS. Streptokinase: a review of its clinical pharmacology, mechanism of action and therapeutic uses. *Drugs* 1973;5(5):357-445
6. McNeil JJ, Krum H. Cardiovascular Disorders In Avery's Drug Treatment Eds Speight TM, Halford NHG Adis International Auckland 1997;809-96
7. Fuster V, Badimon L, Badimon J, Chesebro JH. The pathogenesis of coronary artery disease and acute coronary syndromes. *N. Engl. J. Med* 1992; 326:242-50
8. Shah PK, Forrester JS. Pathophysiology of acute coronary syndrome. *Am J Cardiol* 1991; 68(Suppl 6):16C-23C
9. Falk E, Fernandez A. Role of thrombosis in atherosclerosis and its complications. *Am. J. Cardiol* 1995; 75:5B-11B
10. Davies MJ, Thomas AC. Plaque fissuring, the cause of acute myocardial infarction, sudden ischemic death and crescendo angina. *Br. Heart. J* 1985; 53:363-73
11. TIMI III-B investigators. Effects of tissue plasminogen activator and a comparison of early invasive and conservative strategies in unstable angina and non-Q wave's myocardial infarction: Results of TIMI III B trials. *Circulation* 1994; 89:1545-56
12. Cannon CP. Defining acute myocardial infarction by ST-segment elevation. *Euro. Heart. J.* 2000; 21:266-7
13. Collin D, Lijnin HR. Basic and clinical aspects of fibrinolysis. *Blood* 1991; 78(12):3114-24
14. The time investigator. Trial of invasive versus medical therapy in elderly patients with chronic symptomatic coronary artery diseases. (TIME): a randomized trial. *Lancet* 2001; 358: 951-7
15. Jerry H.G, Joel MG, Rebost JG, Michael R, Nisha C, William JR. Recent age related trends in the use of thrombolytic therapy in patients who have had acute myocardial infarction. *Lancet* 1985; 326 (8455):578-81
16. Laurie A O, Aufderheide TP. Evaluation of ST-segment elevation criteria for the pre-hospital electrocardiographic diagnosis of acute myocardial infarction. *Circulation* 1993;70: 606-18
17. GISSI. Effectiveness of intra-venous thrombolytic treatment in acute myocardial infarction. *Lancet* 1986; 1:397-402
18. Schruder R, Biamino G, Enz-Rudiger v, Linderer T. Intravenous short term infusion of streptokinase in acute myocardial infarction. *Circulation* 1983; 67(3):536-48
19. FTT collaborative group: Indication of fibrinolytic therapy in suspected myocardial infarction: collaborative overview of early mortality and major morbidity results from all randomized trials of more than 1000 patients. *Lancet* 1994; 343(8893):311-22
20. Brügemann J, de Graeff PA, van der Meer J, Lie KI. Does the potential for development of streptokinase antibodies change the risk-benefit ratio in older patients? *Drugs Aging*. 1995 Aug;7(2):110-6
21. Krumholz HM, Pasternak RC, Weinstein MC, Friesinger GC, Ridker PM, Tosteson AN et al. Cost effectiveness of thrombolytic therapy with streptokinase in elderly patients with suspected acute myocardial infarction. *N Engl J Med*. 1992; 327(1):7-13
22. Ross AM. Thrombolytic therapy for acute myocardial infarction. *Cardiovasc Clin*. 1989; 20(1): 209-18
23. Franken M, Nussbacher A, Liberman A, Wajngarten M. ST Elevation Myocardial Infarction in the elderly. *J Ger-*

iatrCardiol 2012;9(2):108-14  
25. O'Gara PT, Kushner FG, Ascheim DD. 2013 ACCF/AHA guideline for the management of ST-elevation myocardial infarction: executive summary: a report of the American

College of Cardiology Foundation/American Heart Association Task Force on Practice Guidelines. Circulation 2013; 127:529





# Prevalence of Vitamin- D Deficiency among Women with Gestational Diabetes Mellitus

Habiba Sharaf Ali<sup>1</sup>, Quratul Ain Zahid<sup>2</sup>

## ABSTRACT:

**Objective:** To determine the frequency of vitamin D<sub>3</sub> deficiency among women with Gestational Diabetes mellitus visiting tertiary care hospital.

**Materials and Methods:** This descriptive case series study was carried out in Obstetrics and Gynecology department at Dr. Ziauddin Hospital Karachi for a period of six months. 136 GDM women with age 18 to 30 years having singleton pregnancy and gestational age of 24-40 weeks were enrolled. Patients were offered according to ADA (American Diabetic Association) recommendations, 1 hour 50gm Oral Glucose Challenge Test, without any preparation or fasting. Then 1 hour later plasma glucose measurements were done. If values were >140 mg/dl or 7.8mmol/l then 3 sample 75g Oral Glucose Tolerance Test (WHO criteria) was offered (to diagnose GDM) on next visit. The mothers were advised not to have breakfast on the day of the diagnostic test. Then fasting blood glucose sample was taken. Afterwards hourly samples were taken till 2 hours. If one reading was raised then diagnosis was established as impaired glucose intolerance and if two readings were raised then diagnosis was confirmed as gestational diabetes mellitus. Estimation of vitamin D levels by Electrochemiluminescence technique was done in diagnosed GDM women.

**Results:** Mean age of the patients was 26.46 ± 2.91 years. Mean gestational age of the patients was 33.03 ± 6.14 weeks. There were 57 (41.90%) primiparous and 79 (58.1%) multiparous patients. Frequency of vitamin D deficiency was found in 84 (61.80%) patients with GDM.

**Conclusion:** The frequency of vitamin D<sub>3</sub> deficiency was found higher among women with GDM visiting tertiary care hospital.

**Keywords:** Prevalence, Vitamin D deficiency, GDM, Tertiary care hospital

## INTRODUCTION:

Vitamin D is well known for its primary physiological role of regulation of calcium homeostasis in maintaining bone health. However, mounting evidence indicates that vitamin D is also involved in controlling body composition, energy homeostasis, insulin sensitivity, and immune function. Low levels of vitamin D during pregnancy or breast feeding can have an adverse effect on the baby's growth and development. Studies have shown the prevalence of vitamin D deficiency (defined as <50 nmol/L) or insufficiency (<75-80 nmol/L) during pregnancy.<sup>1,2</sup> One study suggested that 63.3% mothers are affected.<sup>3</sup>

Low levels of vitamin D during pregnancy or breast feeding can have an adverse effect on mother and baby's health and wellbeing. Infants born to mothers with hypovitaminosis D have increased risk of symptomatic hypocalcaemia, small for gestational age and larger fontanelle, suggestive of impaired ossification of skull bones.<sup>4</sup> In addition, vitamin D deficiency has been linked to other adverse effects on pregnancy, such as diabetes

mellitus, preterm deliveries, bacterial vaginosis, pre-eclampsia<sup>5</sup> and small-for-gestational-age babies.<sup>6</sup> Gestational diabetes mellitus (GDM) is hyperglycemia with onset or first recognition during pregnancy. Although the symptoms of GDM are similar to type 2 diabetes mellitus, it is often diagnosed through prenatal screening, rather than reported symptoms.<sup>7</sup> It has also been suggested that vitamin D deficiency may play a role for the occurrence of GDM.<sup>8</sup> The prevalence rates of GDM vary by region. Vitamin D and Lifestyle Intervention for Gestational Diabetes Mellitus Prevention research group indicated that although the prevalence rates differed by regions in Europe (ranges 2.0-6.0%), lower prevalence rates of GDM were found in Northern or Atlantic seaboard parts of Europe (< 4%); while higher prevalence rates (> 6%) predominated in South or Mediterranean seaboard regions.<sup>9</sup> The prevalence of a low vitamin D in pregnancy in USA is reported to be 59%, Ireland 20.8%, Australia 80.5%. United Arab Emirates 40% and in Pakistan 69.9%.<sup>10, 11</sup> There are several studies that suggest a relationship between vitamin D deficiency and GDM risk, however their results appear mixed and inconclusive.<sup>12, 13, 14, 15</sup> The rationale of this study is to determine the current burden of vitamin D deficiency in GDM women during pregnancy at Dr. Ziauddin tertiary care hospital Karachi, so that the preventive strategy can be planned and implemented along with early detection of gestational diabetes mellitus in order to reduce the severity and complications associated with GDM.

## MATERIALS AND METHODS:

This is a descriptive non-probability consecutive case series study conducted in Obstetrics and Gynecology department at Dr. Ziauddin Hospital Karachi for a period of six months. The sample calculation was done using

✉ **Dr. Habiba Sharaf Ali**

Professor

Department of Obstetrics & Gynaecology

Ziauddin University

Karachi

E-mail: rabel5@yahoo.com

✉ **Dr. Quratul Ain Zahid**

Postgraduate trainee

FCPS Part 2

Ziauddin University

Karachi

Received: 11-01-16

Revised: 23-02-16

Accepted: 25-02-16

the raosoft software for "Sample size calculation" by using the proportion (Vitamin D and Lifestyle Intervention for Gestational Diabetes Mellitus Prevention research group indicated that although the prevalence rates differed by regions in Europe (ranges 2.0-6.0%) 8 with 95% confidential interval an 8.5<sup>4</sup> of margin of error, the sample size stands to be n=128). All pregnant women diagnosed as GDM carrying a singleton pregnancy between gestational age 24 - 40 weeks were included in the study. Women with preexisting Diabetes mellitus were excluded from the study. Informed consent was taken from all eligible women for the study. All the patients included were subjected to detailed history taking with special focus on maternal age, parity, gestational age at diagnosis of gestational diabetes, previous history or family history of diabetes, history of gestational diabetes in previous pregnancies. GDM was diagnosed by means of Oral Glucose Tolerance test. Diagnosed women for GDM with OGTT were offered for vitamin D levels using radio immunoassay. Vitamin D deficiency was defined conservatively as <25 nmol/L, insufficiency as 25–50 nmol/L and

sufficiency as >50 nmol/L. Results were documented on proforma. After collection of data, the analysis was conducted by using Statistical Package for Social Sciences (SPSS) software version 17. Mean and standard deviation was calculated for quantitative variable like age, parity, gravida, gestational age was controlled by stratification. Chi square test was applied.

**RESULTS:**

In our results mean age of patient was 26.46 ±2.91 years. Mean gestational age of the patients was 33.03 ±6.14 weeks. Majority of the patients 76 (55.90%) presented at >30 weeks of gestational weeks. There were 57 (41.90%) primiparous and 79 (58.1%) multiparous patients. We found 84(61.80%) patients of GDM suffering from Vitamin D<sub>3</sub> deficiency (Figure 1). Stratification was done to see the effect of age, gestational age, parity and gravida on the outcome. Chi-square test was applied. No significant association was detected between women's age, parity and gestational age and vitamin D deficiency (Table 1).

Figure: 1  
Percentage of women with GDM and Vitamin D<sub>3</sub> deficiency

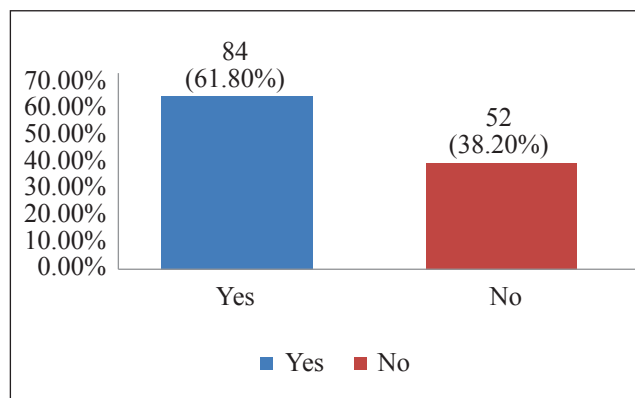


Table: 1  
Comparison of vitamin D deficiency with respect to age, gestational age, parity and gravida

Variables		Vitamin D Deficiency		Total	P-value
		Yes	No		
Age	≤25	38 (61.3)	24 (38.7)	62 (100)	0.917
	>25	46 (62.2)	28 (37.8)	74 (100)	
	Total	84 (61.8)	52 (38.2)	136 (100)	
Gestational Age	≤30	38 (63.3)	22 (36.7)	60 (100)	0.738
	>30	46 (60.5)	30 (39.5)	76 (100)	
	Total	84 (61.8)	52 (38.2)	136 (100)	
Parity	Primiparous	35 (61.4)	22 (38.6)	57 (100)	0.941
	Multiparous	49 (62)	30 (38)	79 (100)	
	Total	84 (61.8)	52 (38.2)	136 (100)	
Gravida	Primigravida	39 (63.9)	22 (36.1)	61 (100)	0.639
	Multigravida	45 (60)	30 (40)	75 (100)	
	Total	84 (61.8)	52 (38.2)	136 (100)	

**DISCUSSION:**

Vitamin D deficiency is associated with a number of adverse pregnancy outcomes. One of the effects of vitamin D<sub>3</sub> deficiency is its association with gestational diabetes mellitus. The probable reason of this relationship could be that vitamin D plays a role in glucose homeostasis and it also improves insulin sensitivity of the target cells (liver, skeletal muscle, and adipos-etissue).<sup>16</sup> However the prevalence of vitamin D<sub>3</sub> deficiency and GDM is different in different parts of the world. The prevalence of GDM in UK, USA and European countries was estimated to be 5%, 3-7% and 6% respectively. Higher prevalence of GDM was noted in African, Asian, Indian and Hispanic women.<sup>17, 18</sup> In London antenatal population, vitamin D level of less than 25 nmol/l was found in 47% of Indian Asian women, 64% of Middle Eastern women, 58% of black women and 13% of Caucasian women.<sup>19</sup> In our study, frequency of vitamin D deficiency among women with GDM was found to be 84(61.80%) patients, which is high. International studies results with regard to the relationship between vitamin D deficiency and GDM risk appears mixed and is not very clear with some of them in favor<sup>12, 13</sup> and others not in favor of relationship between vitamin D deficiency and GDM.<sup>14, 15</sup>

A meta-analysis by Poel<sup>13</sup> has found a significant association between vitamin D<sub>3</sub> deficiency and gestational diabetes mellitus with odds ratio of 1.61 (95% CI 1.19-2.17; p=0.002). Similarly another meta-analysis of 20 independent observational studies done by Zhang provided strong evidence that vitamin D deficiency was associated with an increased risk of gestational diabetes.<sup>20</sup> The levels of serum 25(OH) D have been observed to be inversely associated with levels of HbA1c among women with GDM and this relationship seemed not to be affected by other known risk factors in a study done by Lau.<sup>21</sup> in a tertiary referral Centre. Another recent large study published has found no association between circulating vitamin D<sub>3</sub> levels and GDM. They also looked for other complications such as preterm birth, small for gestational age and fetal growth retardation and found no association with circulating vitamin D<sub>3</sub> levels.<sup>22</sup> Anna has documented in pregnant women with vitamin D<sub>3</sub> deficiency that most of their subjects had no significant association with GDM.<sup>23</sup> A case controlled study on 952 subjects however have found significant association with vitamin D<sub>3</sub> deficiency and GDM after adjusting for risk factors for GDM.<sup>24</sup> A recent meta-analysis on association of vitamin D<sub>3</sub> and GDM showed increased risk of gestational diabetes, pre-eclampsia, and small for gestational age infants but no association with increased risk of caesarean section.<sup>25</sup> Besides finding high levels of vitamin D<sub>3</sub> deficiency in women with gestational Diabetes Mellitus we could not find any association with women's age, parity and gestational age.

**CONCLUSION:**

The frequency of vitamin D deficiency was found to be higher among women with GDM visiting tertiary care

hospital, however we are not sure whether it is a casual or true relationship because of the observational nature of the study. A larger randomized controlled trial is needed to prove this relationship

**REFERENCES:**

1. Karim SA, Nusrat U, Aziz S. Vitamin D deficiency in pregnant women and their newborns as seen at a tertiary-care center in Karachi, Pakistan. *Int J Gynaecol Obstet.* 2011;112:59-62
2. Teale GR, Cunningham CE. Vitamin D deficiency is common among pregnant women in Rural Victoria. *Aust NZ J Obstet Gynaecol.* 2010; 50:259-61
3. Dijkstra S H, Van Beek A, Janssen JW. High prevalence of vitamin D deficiency in newborn infants of bin, risk mothers. *Arch Dis Child.* 2007; 92(9):750-3
4. Javaid MK, Crozier SR, Harvey NC. Maternal vitamin D level during pregnancy and childhood bone mass at age 9 Years: a longitudinal study. *Lancet.* 2006; 367:36-43
5. Robinson CJ, Alanis MC, Wagner CL, Hollis BW, Johnson DD. Plasma 25-hydroxyvitamin D levels in early-onset severe preeclampsia. *Am J Obstet Gynecol.* 2010; 203:366.e1-6
6. Leffelaar ER, Vrijkotte TG, van Eijsden M. Maternal early pregnancy vitamin D status in relation to fetal and neonatal growth: results of the multi-ethnic Amsterdam Born Children and their Development cohort. *Br J Nutr.* 2010; 104:108-17
7. Bellamy L, Casas JP, Hingorani AD, Williams D. Type 2 diabetes mellitus after gestational diabetes: a systematic review and meta-analysis. *Lancet.* 2009; 373:1773-9
8. Dror DK. Vitamin D status during pregnancy: maternal, fetal, and postnatal outcomes. *Curr Opin Obst Gynecol.* 2011;23:422-36
9. Buckley BS, Harreiter J, Damm P, Corcoy R, Chico A. Gestational diabetes mellitus in Europe: prevalence, current screening practice and barriers to screening. A review. *Diabet Med.* 2012; 29:844-54
10. Ponsonby A, Lucas RM, Lewis S, Halliday J. Vitamin D status during Pregnancy and Aspects of Offspring Health. *Nutrients.* 2010; 2:389-407
11. Naqvi KZ, Ali ST, Thontia S, Madiha. Prevalence of Vitamin D deficiency in pregnant population at term attending a tertiary care hospital Karachi, Pakistan. *Pak J Surg.* 2012; 28(2):122-5
12. Soheilykhah S, Mojibian M, Rashidi M, Rahimi-Saghand S, Jafari F. Maternal vitamin D status in gestational diabetes mellitus. *Nutr Clin Pract.* 2010; 25: 524-7
13. Poel YH, Hummel P, Lips P, Stam F, van der Ploeg T. Vitamin D and gestational diabetes: a systematic review and meta-analysis. *Eur J Intern Med.* 2012; 23:465-9
14. Baker AM, Haeri S, Camargo CA JR, Stuebe AM, Boggess KA. First trimester maternal vitamin D status and risk for gestational diabetes (GDM) a nested case-control study. *Diabetes Metab Res Rev.* 2012; 28:164-8
15. Farrant HJ, Krishnaveni GV, Hill JC, Boucher BJ, Fisher DJ. Vitamin D insufficiency is common in Indian mothers but is not associated with gestational diabetes or variation in newborn size. *Eur J Clin Nutr.* 2009;63:646-52
16. Ahmed El Lithy, Abdella RM, El-Faissal YM, The relationship between low maternal serum vitamin D levels and glycemic control in gestational diabetes assessed by HbA1c levels: an observational cross-sectional study. *BMC Pregnancy Childbirth.* 2014; 14: 362doi:

- 10.1186/1471-2393-14-362
17. Carolan M, Davey MA, Biro MA, Kealy M. Maternal age, ethnicity and gestational diabetes mellitus. *Midwifery* 2012; 28(6):778-83
  18. Makgoba M, Savvidou MD, Steer PJ. An analysis of the interrelationship between maternal age, body mass index and racial origin in the development of gestational diabetes mellitus. *BJOG*.2012; 119: 276-82
  19. Yu CK, Sykes L, Sethi M, Teoh TG, Robinson S. Vitamin D deficiency and supplementation during pregnancy. *Clin Endocrinol (Oxf)* 2009; 70:685-90
  20. Zhang MX, Pan GT, Guo JF, Li BY. Vitamin D Deficiency Increases the Risk of Gestational Diabetes Mellitus: A Meta-Analysis of Observational Studies *Nutrients* 2015, 7, 8366-75
  21. Parlea L, Bromberg IL, Feig DS, Vieth R, Merman E, Lipscombe LL. Association between serum 25-hydroxy-vitamin D in early pregnancy and risk of gestational diabetes mellitus. *Diabet Med.* 2012 ; 29 29(7):e25-32. doi:10.1111/j.1464-5491.2011.03550.x
  22. A Rodriguez, R Garcia Esteban. Association of maternal circulating 25 Hydroxyvitamin D<sub>3</sub> Concentration with pregnancy and birth outcome *BJOG* 2015; 122:1695-704
  23. Anna P, Venndula B. Vitamin D<sub>3</sub> status in women Gestational Diabetes Mellitus during pregnancy and postpartum. *Biomed Research International* 2015; 1-7. DOI: 10.1155/2015/260624
  24. Colin Zhang, Chunfang Qiu. Maternal Plasma 25 Hydroxy vitamin D<sub>3</sub> concentrations and risk of gestational Diabetes mellitus. *Plos one* 2008;3:e377753.
  25. Jafari FA, Nagulesapillai T, Ronksley PE. Association between maternal serum 25-hydroxyvitamin D level and pregnancy and neonatal outcomes: systematic review and meta-analysis of observational studies. *BMJ* 2013; 346:f1169



# The Safety and Efficacy of Percutaneous Trigger Finger Release Under Local Anaesthesia

Abdur Rehman Qureshi<sup>1</sup>, Faaiz Ali Shah<sup>2</sup>, Shahab-ud-Din<sup>3</sup>, Wali Muhammad Khan<sup>4</sup>

## ABSTRACT:

**Objective:** To determine the safety and efficacy of trigger finger and thumb released percutaneously with an 18 gauge needle under local anaesthesia.

**Materials and Methods:** This descriptive case series study was conducted at Orthopaedic and Traumatology Unit "A" Medical Teaching Institution (MTI) Lady Reading Hospital (LRH) Peshawar Pakistan from April 2014 to December 2015. All patients of trigger finger or thumb of either gender fulfilling the inclusion criteria were percutaneously released under local anaesthesia with the tip of an 18-gauge hypodermic needle. Post operative assessment of these patients was done weekly for a month and then monthly for 6 months. Clinical results were evaluated in terms of pain, activity level and patient satisfaction after 6 months at follow up and rated as excellent, good and poor.

**Results:** Thirty two fingers in twenty five patients with mean age 38.28 years  $\pm$  11SD (range 18 to 62 years) were included in the study. Post operatively excellent results were achieved in 90.9% (20/22) patients and good in 9% (2/22) patients at six months follow up. There were only 3 (9.3%) failed releases requiring conversion to open release. There was no recurrence of trigger finger and no digital nerve nor tendon injuries reported.

**Conclusion:** Percutaneous trigger finger release under local anaesthesia is a safe and highly effective method for releasing trigger fingers. We recommend it as a treatment of choice for established trigger finger or thumb.

**Keywords:** Percutaneous release, Trigger finger, Tendon, Stenosing tenosynovitis, Local anaesthesia

## INTRODUCTION:

The term trigger finger for stenosing tenovaginitis or tenosynovitis of the tendon sheaths of flexor muscles of the finger was first proposed by French physician Alphonse Henri Notta in 1850 in his study of four cases of adult patients.<sup>1,2</sup> In recognition of Notta's discovery, a tendon nodule located on the volar aspect of the base of paediatric trigger thumb, mentioned in most studies is now commonly referred to as Notta's node.<sup>3</sup> Histo-

pathologically, proliferative thickening of flexor tendon sheath layer at the first annular (A1) pulley occurs resulting in narrowing of fibrous tunnel leading to tendon impingement when it moves through the narrow tunnel as the patient flexes and extends the affected finger.<sup>3,4</sup> Female patients in their fifth or sixth decades are more commonly affected than men and children rarely affected. It has a prevalence of about 2 percent in general population with diabetes, rheumatoid arthritis and amyloidosis are more prone to develop triggering.<sup>3,5</sup> The exact etiology of trigger finger in majority of cases is unknown<sup>6</sup> but certain occupations that involve constant gripping or repeated activities have been reported to be associated with more frequent triggering.<sup>3</sup> Thumb and ring finger is most commonly affected by triggering but it can involve any finger.<sup>5</sup> Various treatment options for trigger finger are injections of corticosteroid in the sheath of affected tendon<sup>7</sup> and percutaneous<sup>8</sup> or open<sup>9</sup> surgical release of the A1 pulley. In 1958 Lorthioir<sup>8</sup> first demonstrated A1 pulley release percutaneously in 52 patients with 100 percent success rate and no reported complications. A1 pulley release percutaneously with a hypodermic needle is a safe, quick, simple, economically feasible and highly effective technique with a short post-operative rehabilitation than open surgical release and can be performed as a day case or out-patient department (OPD) procedure.<sup>10,11</sup> The possible complications associated with open surgical release like infection, post-operative painful scar, A1 pulley tear resulting in bowstringing of flexor tendons, stiffness of interphalangeal joints and digital neurovascular injuries are minimal with percutaneous release.<sup>12</sup> The objective of this study was to evaluate the clinical results and safety of percutaneous trigger finger release under local anaesthesia in our set up.

### ✉ Dr. Abdur Rehman Qureshi

Assistant Professor  
Department of Orthopaedics & Traumatology Unit "A"  
Medical Teaching Institution (MTI)  
Lady Reading Hospital (LRH)  
Khyber Pakhtunkhwa (KPK)  
Peshawar  
E-mail: hwabannu@yahoo.com

### ✉ Dr. Faaiz Ali Shah

Senior Registrar  
Department of Orthopaedics & Traumatology Unit "A"  
Medical Teaching Institution (MTI)  
Lady Reading Hospital (LRH)  
Khyber Pakhtunkhwa (KPK)  
Peshawar

### ✉ Dr. Shahab-ud-Din

Professor and Head  
Department of Orthopaedics & Traumatology Unit "A"  
Medical Teaching Institution (MTI)  
Lady Reading Hospital  
Peshawar

### ✉ Dr. Wali Muhammad Khan

Senior Registrar  
Orthopaedics & Traumatology Unit "A"  
Medical Teaching Institution (MTI)  
Lady Reading Hospital  
Peshawar

Received: 11-01-2016

Revised: 25-02-2016

Accepted: 26-02-2016

## MATERIALS AND METHODS:

This study was conducted in Orthopaedic and Traumatology Unit "A" Medical Teaching Institution (MTI) Lady Reading Hospital (LRH) Peshawar Pakistan

from April 2014 to December 2015. All Patients of trigger finger or thumb of both gender and Quinnell's<sup>13</sup> grades IV and V (Intermittent locking but actively correctable and complete locking but only passively correctable respectively) were recruited from the outpatient department for this study. All patients had painful finger or thumb flexion and extension and triggering was clearly observed in all patients. Children with trigger fingers, patients of rheumatoid arthritis, previous flexor tendon repairs and patients having bleeding disorders were excluded from the study. The study protocol was approved by the hospital ethics committee and informed written consent was taken from all the participants of the study. Complete history and physical examination and X- rays of affected finger or thumb were done in all the included subjects. Under local anaesthesia in the outpatient department, the tip of an 18-gauge hypodermic needle was used to divide the A1 pulley percutaneously. Post operative assessment of these patients was done weekly for a month and then monthly for 6 months. The clinical results were evaluated in terms of pain, activity level and patient satisfaction at 6 months follow up and rated as excellent, good and poor according to Grundberg's<sup>14</sup> rating system (Table 1). Statistical analysis of the data was done with SPSS (version 16). Frequency and percentages were used for categorical or qualitative variables such as gender. Mean  $\pm$  Standard Deviation (SD) was used for numerical or quantitative variables such as age (in years).

**Operative Technique:**

The procedure was performed in the outpatient department (OPD) as day case surgery. The position of the patient for the procedure was supine with forearm supinated and trigger finger in extension. Affected finger and hand was scrubbed with povidone iodine solution. About 1 ml of plain lignocaine was injected into the skin overlying the A1 pulley. To prevent injury to the digital artery and nerve during the procedure, we utilized the safe anatomical landmarks as the tubercle of the scaphoid bone and the midpoint of proximal palmar crease for the A1 pulley release in the little finger while the radial side of pisiform bone and midpoint of proximal palmar crease was used for index finger A1 pulley release.<sup>15</sup> The middle and ring fingers were released through midpoint of distal palmar crease<sup>16</sup> while the metacarpophalangeal crease was used as a starting landmark for trigger thumb release in all cases.<sup>17</sup> To avoid damaging digital arteries and nerves during the procedure the metacarpophalangeal joint is hyper extended so that the tendon gets closer to the skin and with neurovascular bundle falls to the side of the tendon. An 18 gauge hypodermic needle tip was inserted over A1 pulley and divided in one clean stroke. The disappearance of grating sensations confirmed that the pulley was completely cut. After the procedure, the patient was instructed to flex and extend the finger a few times and a dressing was applied over the area. Post procedure nonsteroidal anti- inflammatory drugs were prescribed for three days in all cases. All the patients were advised weekly follow up visits for a month and then monthly for six months for assessment of any

recurrence, pain, wound infection, digital stiffness etc.

**RESULTS:**

Thirty two fingers in twenty five patients with mean age 38.28 years  $\pm$  11 (range 18 to 62 years) were included in the study. Nine (36%) patients were male while 16 (64%) were female. The frequency of fingers or thumb involvement among our patients is shown in Table 2. Nine (36%) patients (12 fingers) were diabetics. A total of six (18.7%) fingers had failed a trial of treatment by steroid injection at least once before percutaneous release. Eighteen (56.2 %) fingers were Quinnell's Grade III while 14 (43.7%) were grade IV on initial admission. Post operatively excellent results were achieved in 90.9% (20/22) patients and good in 9% (2/22) patients while no poor result was recorded at six months follow up. There were only 3 (9.3%) failed releases requiring conversion to open release at first follow up visit and all the three (thumb, index, little finger) were Quinnell's Grade IV diabetics and had previously failed steroid injection. In all three patients, intra-operative observation revealed incomplete release of the A1 pulley. There were no signs of digital nerve or artery injury nor was there any significant tendon injury in any of these patients. The mean operative time was 12 min (9-15), including the local anaesthesia of the patient. There was no recurrence of triggering. Range of motion was preserved in all cases. There was no wound infection, hematoma formation, digital nerve or tendon injuries reported in our study.

Table: 1  
Grundberg's rating system to evaluate clinical outcome

Rating	Pain	Activity and patient satisfaction
Excellent	No pain Returned to work or activity	Patient satisfied
Good	Pain only with heavy use Returned to work or activity	Patient satisfied
Poor	Pain unchanged	Patient dissatisfied

Table: 2  
Frequency of fingers or thumb involvement among our patients

Finger	Side of trigger finger/thumb		Total
	Right	Left	
Thumb	4	2	6 (18.7%)
Index	4	1	5 (15.6)
Middle	8	3	11 (34.3)
Ring	6	1	7 (21.8)
Little	3	0	3 (9.3%)

## DISCUSSION:

Percutaneous release of trigger finger is easy to perform, economically feasible with excellent results and minimal complications and is therefore preferred than open surgical release.<sup>18</sup> Our study yielded excellent results in majority (90.9%) of patients and good in other (9%) patients while no poor result at six months follow up. Other studies also reported that percutaneous release alone gave excellent functional results.<sup>10,19</sup> Our results of release are therefore comparable with those reported previously by other authors. Mishra<sup>20</sup> used the tip of 20 gauge hypodermic needle for percutaneous release of 27 trigger fingers and reported 95.4% excellent results with no recurrence or complications. They concluded that percutaneous release has a very high success rate and is a safer technique with a very few documented complications rather than open release. Similarly Dahabra<sup>21</sup> used 18 gauge needle tip for A1 pulley release and reported a success rate of 92.8% while failure in only 3(7.2%) fingers. Forty six trigger fingers were percutaneously released by Sahu<sup>11</sup> and excellent, good and poor results were noted in 82.6%(38/46), 13.0%(6/46) and 4.3%(2/46) patients respectively at final follow up visit, by taking into account post op pain, patient activity and satisfaction. In our study no post procedure complications like digital nerve or flexor tendon injury, recurrence, wound sepsis and hematoma formation were reported and this was due to the fact that we carefully utilized the established guidelines for the precise anatomical recognition of the pulleys for needle placement and aseptic technique in each and every case. Ha<sup>17</sup> percutaneously released 185 trigger fingers with no complications noted while Gilbert<sup>22</sup> reported sensory loss on the radial side of the thumb in 3 (1%) patients. Fu<sup>23</sup> reported persistent or recurrent triggering symptom in 4% of his patients. Guler<sup>24</sup> reported an incidence of about 5.7% of digital nerve injury in his series of trigger thumb release and he therefore advised precise anatomical surface markings or use of ultrasound for trigger thumb release. For trigger thumb he suggested open surgery rather than percutaneously. There were only 3(9.3%) failed releases in our study requiring conversion to open release at first follow up visit. All the three (thumb, index, little finger) were diabetics and had previously failed steroid injection as well. These cases with incomplete release were among the first cases in our series. Our last 29 fingers were completely released. Our inability to release trigger finger in a few cases might be due to our anxiety about the proximity of digital neurovascular bundle with hypodermic needle placement site or learning curve for the procedure as the surgeon's skill<sup>25</sup> is of utmost importance for a successful and complication free trigger finger release. Furthermore all the failed cases were diabetics and as Ryxewicz<sup>9</sup> noted that hyperglycemia leads to fibrosis and possibly tenosynovitis which is not only resistant to cure but also has a very high rate of recurrence and therefore requires early open surgical release rather than percutaneous release. Small sample size was the one of the limitations of our study. Although

our study had fewer number of cases however but we achieved excellent results in majority of trigger finger and thumb release percutaneously. Further an analysis was not made based on a comparison with other methods of anesthesia and surgical techniques or steroid injection, because the study would then be more difficult and costly than the present. We believe that percutaneous trigger finger release is a very useful technique and we recommend continued study over its long-term effects.

## CONCLUSION:

Percutaneous trigger finger release is a safe and highly effective technique. Utilizing precisely the safe anatomical landmarks we found it safe for all the fingers including the thumb, index finger and little finger. It is a quick and less invasive technique and can be done as a day care procedure in the outpatient department(OPD). It is easy to perform, economically feasible to all patients, has no major complications and allows the patient to return to his daily activities and work quickly. This technique produced excellent results in majority of our patients. We recommend it as a treatment of choice for established trigger finger or thumb.

## REFERENCES:

1. Clapham PJ, Chung KC. A historical perspective of the Notta's node in trigger fingers. *J Hand Surg Am.* 2009; 34:1518-22
2. Dierks U, Hoffmann R, Meek MF. Open versus percutaneous release of the A1-pulley for stenosing tenosynovitis: A perspective randomized trial. *Tech Hand Up Extrem Surg.* 2008;12:183-7
3. Moore JS. Flexor tendon entrapment of the digits (trigger finger and trigger thumb). *J Occup Environ Med.* 2000;42: 526-45
4. Sbernardori MC, Bandiera P. Histopathology of the A1 pulley in adult trigger fingers. *J Hand Surg Eur* 2007;32: 554-6
5. Saldana MJ. Trigger digits: Diagnosis and treatment. *J Am Acad Orthop Surg.* 2000;9:246-52
6. McAuliffe JA. Tendon disorders of the hand and wrist. *J Hand Surg Am.* 2010;35:846-53
7. Dhal J, Hammert WC. Overview of injectable corticosteroids. *J Hand Surg Am.* 2012;37:1715-7
8. Lorthioir J. Surgical treatment of trigger-finger by a subcutaneous method. *J Bone Joint Surg Am.* 1958;40:793-5
9. Ryzewicz M, Wolf JM. Trigger digits: Principles, management and complications. *J Hand Surg Am.* 2006;31:135-46
10. Gilberts EC, Beekman WH, Stevens HJ, Wereldsma JC. Prospective randomized trial of open versus percutaneous surgery for trigger digits. *J Hand Surg Am.* 2001;26:497-500
11. Blumberg N, Arbel R, Dekel S. Percutaneous release of trigger digits. *J Hand Surg Br.* 2001;26:256-7
12. Sahu RL, Gupta P. Experience of percutaneous trigger finger release under local anaesthesia in the Medical College of Mullana, Haryana Ambala. *Ann Med Health Sci Res.* 2014;4(5):806-9
13. Quinnell RC. Conservative management of trigger finger. *Practitioner.* 1980;224:187-90
14. Grundberg AB, Dobson JF. Percutaneous release of the

- common extensor origin for tennis elbow. *Clin Orthop Relat Res.* 2000;376:137-40
15. Wilhelmi BJ, Mowlavi A, Neumeister MW, Bueno R, Lee WP. Safe treatment of trigger finger with longitudinal and transverse landmarks: An anatomic study of the border fingers for percutaneous release. *Plast Reconstr Surg.* 2003;112:993-9
  16. Eastwood DM, Gupta KJ, Johnson DP. Percutaneous release of the trigger finger: An office procedure. *J Hand Surg.* 1992;17:114-7
  17. Ha KI, Park MJ, Ha CW. Percutaneous release of trigger digits. *J Bone Joint Surg.* 2001;83:75-7
  18. Ucar BY. Percutaneous surgery: A safe procedure for trigger finger? *N Am J Med Sci.* 2012;4(9):401-3
  19. Park MJ, OH I, Ha KI. A1 pulley release of locked trigger digit by percutaneous technique. *J Hand Surg Br.* 2004;29(5):502-5
  20. Mishra SR, Gaur AK, Choudhary MM, Ramesh J. Percutaneous A1 pulley release by the tip of a 20 G hypodermic needle before open surgical procedure in trigger finger management. *Tech Hand Up Extrem Surg.* 2013;17(2):112-5
  21. Dahabra IA, Sawaqed IS. Percutaneous trigger finger release with 18 gauge needle. *Saudi Med J.* 2007;28(7):1065-7
  22. Gilberts EC, Wereldsma JC. Long term results of percutaneous and open surgery for trigger fingers and thumb. *Int Surg.* 2002;87(1):48-52
  23. Fu YC, Huang PJ, Tien YC, Lu YM, Fu HH, Lin GT. Revision of incompletely released trigger fingers by percutaneous release: Results and complications. *J Hand Surg Am.* 2006;31:1288-91
  24. Guler F, Kose O, Ercan EC, Turan A, Canbora K. Open versus percutaneous release for the treatment of trigger thumb. *Orthopaedics.* 2013;36(10):1290-4
  25. Maneerit J, Sriworakun C, Budhraj N, Nagavajara P. Trigger thumb: Results of a prospective randomized study of percutaneous release with steroid injection versus steroid injection alone. *J Hand Surg Br.* 2003;28:586-9





## ORIGINAL ARTICLE

# Relationship of Cardiac Disease with Oral Health: A Single Centre Study

Javed Ashraf<sup>1</sup>, Rana Modassir Shamsher Khan<sup>2</sup>, Khawaja Rashid Hassan<sup>3</sup>, Muhammad Rizwan<sup>4</sup>, Ali Saad Tariq<sup>5</sup>, Sarah Ashraf<sup>6</sup>

### ABSTRACT:

**Objective:** To observe relationship of chronic dental and oral morbidity with cardiovascular disease in Pakistani population. **Materials and Methods:** All indoor cardiac patients aged 40 and above, clinically and angiographically diagnosed with CHD at Islam Central Hospital, Sialkot, were included in the study. Demographic and clinical data (Age, Gender, Smoking, and Diabetes) were noted from patients' hospital record files. Missing teeth were examined and number of teeth missing was estimated from the number of teeth remaining in the mouth upon clinical examination. Attendants without a history of cardiac disease, of the cardiac patients who agreed to be included in the study, were examined for comparison of tooth loss. **Results:** Nine hundred and thirty six cardiac patients and 595 healthy attendants with mean age of  $51.9 \pm 8.4$  years were examined. Chronic periodontal disease and mean ( $\pm$ SD) tooth loss was significantly ( $P < 0.001$ ) higher in cardiac patients. Odds ratio (OR) = 1.543 was found in cardiac patients when compared with healthy controls (95%CI = 1.985–2.851). Tooth loss was significantly ( $P < 0.001$ ) associated with both males and female cardiac patients especially along with diabetes and smoking. **Conclusion:** Chronic periodontal disease and tooth loss were found to be significantly higher in cardiac disease patients in comparison to healthy controls. Other risk factors found were age, gender, smoking and diabetes.

**Keywords:** Cardiac patients, Chronic dental morbidity, Oral morbidity

### INTRODUCTION:

Non-communicable chronic diseases (NCDs) are presently, causing 80% of deaths in low and middle-income countries including Pakistan<sup>1</sup>. Association of poor oral health, periodontal disease and tooth loss with increased risk of cardiovascular diseases (CVD), pulmo-

nary diseases, diabetes and adverse pregnancy outcomes such as low birth weight babies has been observed and established in scientific literature.<sup>2</sup> Chronic systemic and oral diseases share many common risk factors such as age, gender, education; smoking, diet, and obesity. These are also referred to as "shared risk factors".<sup>3,4,5</sup> A healthy mouth is a pre-requisite for overall good general health. When oral health is compromised, overall health is also affected.<sup>6</sup> Tooth loss is common among human beings, and having less than 20 natural teeth is categorized as poor Oral health.<sup>7</sup> Including other systemic diseases, higher incidence of chronic periodontal disease and tooth loss has also been reported to be significantly associated with cardiovascular disorders in various case-control and cross-sectional studies.<sup>8,9,10</sup> Tooth loss is found to be associated with CVD on the basis of chronic oral infections, such as chronic periodontal diseases.<sup>11</sup> Tooth loss may lead to changes in diet and other behaviors which in turn lead to increased risk for CVD.<sup>12</sup> CVDs are expected to rise as an epidemic in developing countries and projected to be a major cause of death by 2020.<sup>13</sup> Despite an expected rise in NCDs in Asia and a high prevalence of oral disease, few studies on their association have been conducted in Asiatic region including Pakistan. The purpose of this study was to observe prevalence of tooth loss in cardiac patients of the Pakistani population and to explore its possible association with coronary heart disease.

### MATERIALS AND METHODS:

This cross-sectional study was done from 1<sup>st</sup> Jan, 2014 to 30<sup>th</sup> June, 2015. Non-probability convenient sampling technique was employed for sample selection. All indoor cardiac patients aged 40 and above clinically and angiographically diagnosed with CHD at Islam Central Hospital, Sialkot, were included in the study. Healthy individuals were taken as control. Demographic and clinical data (Age, Gender, Smoking, and Diabetes) were noted from patients' hospital record files. Missing teeth were examined at bedside with the help of a mouth

#### ✉ Dr. Javed Ashraf

Associate Professor and Head  
Department of Community Dentistry  
Islam Dental College  
Sialkot  
E-mail: dr\_javedansari110@yahoo.com

#### ✉ Dr. Rana Modassir Shamsher Khan

Associate Professor and Head  
Department of Orthodontics  
Islam Dental College  
Sialkot

#### ✉ Dr. Khawaja Rashid Hassan

Assistant Professor and Head  
Department of Dental Materials  
Islam Dental College  
Sialkot

#### ✉ Dr. Muhammad Rizwan

Associate Professor  
Department of Oral Pathology  
Islam Dental College  
Sialkot

#### ✉ Dr. Ali Saad Tariq

Assistant Professor and Head  
Department of Oral Biology  
Islam Dental College  
Sialkot

#### ✉ Dr. Sarah Ashraf

House Officer  
Montmorency College of Dentistry  
Lahore

Received: 15-01-2016

Revised: 17-02-2016

Accepted: 20-02-2016

mirror and tweezers. The number of teeth missing was estimated from the number of teeth remaining in the mouth upon clinical examination. Attendants without a history of cardiac disease, of the cardiac patients who agreed to be included in the study, were examined for comparison of tooth loss. Study subjects with other chronic systemic diseases, such as chronic obstructive pulmonary disease (COPD), chronic arthritis, chronic liver disease, and kidney diseases were excluded from the study. Data was analyzed using SPSS version 19. Summary statistics were calculated through descriptive analysis; independent t-test was applied for comparison of tooth loss between cardiac and healthy subjects. For comparisons of categorical variables, chi-square test was applied. Subjects were grouped into smoker–diabetic and nonsmoker–nondiabetic for a comparison of tooth loss. Multivariate regression models were fit to observe the association of tooth loss and CHD and confounding factors.

## RESULTS:

During a nine-month study period, 1531 subjects were examined. Subjects' age ranged from 40 to 70 years and the mean age was  $52.0 \pm 8.4$  years; 936 were CHD patients with a mean age of  $53.7 \pm 8.4$  years, and 595 were healthy individuals with a mean age of  $49.1 \pm 7.7$  years. Seventy four percent were cardiac patients and 58% healthy subjects were males. Thirty seven percent of cardiac patients and 20.5% healthy individuals were smokers. Thirty six percent were diabetics among cardiac patients as compared with 16.5% among healthy individuals. There was a statistically significant ( $p < 0.001$ ) difference among cardiac patients and their healthy attendants with Odds ratio 2.82 (CI = 2.287-3.512), 2.036 (CI = 1.612-2.572) and 2.840 (CI = 2.202-3.663) healthy genders, smokers and diabetic subjects, respectively (Table 1).

The main variable of this study, tooth loss, was found in 1242 (81%) subjects of the study sample. Subjects with at least one missing tooth showed a mean ( $\pm$ SD) tooth loss  $8.8 \pm 8.5$ . CHD patients showed a mean tooth loss of  $9.8 \pm 9.2$  and healthy subjects had a mean tooth loss of  $7.0 \pm 6.9$ ; the difference was statistically significant ( $p < 0.001$ ). Seventeen percent of CHD patients had all natural teeth as compared with 33% healthy individuals. CHD patients were at OR of 1.54 (CI = 1.192-1.197) for having more tooth loss as compared with healthy individuals. Seventy eight percent of subjects showed tooth loss in the range of 1-15 teeth and 88% of subjects showed 2 teeth losses up to 20 teeth. Thirteen percent CHD patients as compared with 4% of healthy individuals had 21-32 teeth lost (Table 2).

Tooth loss analysis among genders showed that a mean tooth loss in CHD (56%) and healthy (21%) males was  $7.1 \pm 8.4$  and  $4.4 \pm 6.5$ , respectively, with a significant difference ( $p < 0.001$ ). Mean tooth loss among CHD (42%) and healthy (46%) females was  $11.5 \pm 10.2$  and  $6.4 \pm 6.8$  respectively, with a statistically significant difference ( $p < 0.001$ ). CHD males with tooth loss had an OR of 1.78 (CI = 1.307-2.427) and CHD females had an OR of 2.79 (CI = 1.521-5.148) (Table 3).

Among smoker-diabetic subjects, 67% of cardiac patients were presented with a mean tooth loss of  $8.8 \pm 9.3$  as compared with 13% of healthy individuals with a mean tooth loss of  $4.7 \pm 6.00$  ( $p = 0.014$ ). Nonsmoker-nondiabetic CHD (40%) and healthy (40%) subjects were found with a mean tooth loss of  $8.0 \pm 8.6$  and  $7.3 \pm 8.5$ , respectively. Smoker-diabetic patients with tooth loss (OR = 2.246; CI = 1.789-6.394) had higher risk for CHD (Table 3).

Table 4a presents an age-related pattern of tooth loss prevalence in cardiac and healthy subjects. Forty to fifty five year old subjects showed a significantly higher tooth loss ( $p = 0.015$ ) in cardiac patients with an OR of 1.396 (CI = 1.046-1.863). Mean tooth loss showed a steady increase in age groups 40-49 years ( $3.9 \pm 5.2$ ), 50-59 years ( $7.7 \pm 8.2$ ), and P60 years ( $12.9-10.8$ ), however, statistical differences were insignificant in all age cohorts.

In multivariate regression analysis, coefficient was positive and a higher code for smoking was 1, the OR was 1.33; higher code for diabetes was 1 and the OR was 3.50; higher code for genders was 1 and the OR was 3.24; higher code for tooth loss was again 1 and the OR was 1.45. It can be significantly concluded that cardiac patients with smoking, diabetes, male gender and tooth loss were at higher risk as compared with healthy individuals.

Logistic regression model adjusted for all risk factors of CHD noted in this study showed that tooth loss P 1 teeth ( $p = 0.010$ ), 620 teeth ( $p = 0.024$ ) and >20 teeth ( $p < 0.001$ ) are statistically significant predictors of CHD. Adjusted OR for tooth loss 620 teeth and >20 teeth were 1.39 (95%CI = 1.04–1.78) and 3.52 (95%CI = 2.01–6.18) (Table 4b).

## DISCUSSION:

This cross-sectional study on the topic from Pakistan has found a statistically significant difference in tooth loss between cardiac and healthy subjects. An association of tooth loss with prevalent coronary heart disease is observed in this study that supports previous studies on the relationship of tooth loss and cardiac conditions.<sup>8,10,11,14,15</sup>

Demographic data of the study sample shows that males, diabetics and smokers were more than twice (OR P 2.036; CI = 1.612–2.572) at risk of CHD as compared with the healthy individuals. Males were significantly higher than females in the CHD group; whereas there was no difference in male–female ratio in the healthy group. Age is the constant and most commonly reported factor associated with missing teeth.<sup>16,17</sup> Tooth loss difference is found in genders; in particular, males have less number of teeth.<sup>18</sup> This study has noted a monotonous relationship between increasing tooth loss and advancing age in CHD/non-CHD individuals who were closely related with respect to their socioeconomic status (SES) background, and this finding corresponds with another contemporary study.<sup>19</sup> The current study showed that CHD males with tooth loss were twice the number of CHD females; however a mean tooth loss was much higher in females. These findings also correspond with other studies.<sup>20,21</sup> In the current study, incidence of tooth

Table: 1  
Summary statistics of cardiac and healthy subjects

Variables	Cardiac n (%)	Healthy n(%)	Total	P-value/OR (CI=95%)
Study Sample	936(61)	595(39)	1531(100)	
Age				
Mean	53.7 + 8.4	49.1 + 7.7	52 + 8.4	< 0.001a
Range	40-70			
Male	692(74)	298(50)	990(65)	< 0.001/2.827b
Female	244(26)	297(50)	541(35)	
Smokers	348(37)	134(22.5)	482 (31.5)	
Non-Smokers	588(63)	461(77.5)	1049(68.5)	< 0.001/2.036b
Diabetics	336(36)	98(16.5)	434(28)	
Non-Diabetics	600(64)	497(83.5)	1097(72)	< 0.001/2.840b

a = Students T-test.  
b =Chi-Square Test.

Table :2  
Cardiac and Healthy subjects compared for tooth loss

Variables	Cardiac	Healthy	Total	P-value
Tooth loss				
Mean± SD	9.8 ± 9.2	7.0 ± 6.9	8.7 ± 8.5	< 0.001 <sup>a</sup>
Subjects with tooth loss				
n(%)	n(%)	n(%)	n(%)	n(%)
> 1 tooth	784(84)	458(77)	1242(81)	< 0.01 <sup>b</sup>
2-15 teeth	606(65)	373(63)	979(79)	< 0.01 <sup>b</sup>
16-32 teeth	178(19)	85(14)	263(21)	NS

a=T-test.  
b=Chi-square.  
NS = Not-Significant

Table: 3  
Cardiac and healthy subjects with tooth loss compared among genders, smokers, non-smokers, diabetics, non-diabetics

Variables	Cardiac	Healthy	Total	P-value
Males				
N= 990				
N (%)	555(56)	207(23)	762(79)	< 0.01 <sup>b</sup>
Mean ± SD	7.1 ± 8.4	4.4 ± 6.5	6.2 ± 7.9	< 0.01 <sup>a</sup>
Females				
N=541				
N (%)	229(42)	251(46)	480(88.7)	< 0.01 <sup>b</sup>
Mean ± SD	11.5 ± 10.2	6.4 ± 6.8	8.8 ± 8.9	< 0.01 <sup>a</sup>
Smokers and Diabetics				
N (%)	77(67)	15(13)	92(80)	< 0.02 <sup>b</sup>
Mean ± SD	8.8 ± 9.3	4.7 ± 6.0	8.0 ± 8.8	< 0.01 <sup>a</sup>
Non-smokers and Non-Diabetics				
N (%)	290(40)	291(40)	581(79.6)	NS <sup>b</sup>
Mean ± SD	8 ± 8.6	7.3 ± 8.2	7.8 ± 8.5	NS <sup>a</sup>

a = Independent T-test.  
b = Chi-Square test.  
NS= Non-significant.

Table: 4a  
Cardiac and Healthy subjects compared for tooth loss in different age group

Variable	Cardiac	Healthy	Total	P-Value/OR (CI 95%)
40-55 years (n=1046)				
n%	451	354 (34)	805 (77)	<0.050/1.396 (1.046-1.863) <sup>b</sup>
Mean ± SD	5.7±7.0	4.1 ± 5.2	5.0 ± 6.3	<0.001 <sup>a</sup>
56+years (n=485)				
n%	333 (69)	104 (21)	437 (90)	NS <sup>b</sup>
Mean ± SD	12 ± 10.6	10.42 ± 9.6	11.6 ± 10.3	NS <sup>a</sup>
40-49 years (n=615)				
n%	207 (34)	241 (39)	448 (73)	NS <sup>b</sup>
Mean ± SD	3.9 ± 5.2	3.3 ± 4.0	3.6 ± 4.6	NS <sup>a</sup>
50-59years (n=538)				
n%	307 (57)	141 (26)	448 (83)	<0.050/1.665 (1.049-2.643)
Mean ± SD	7.7 ± 8.2	6.3 ± 7.0	7.2 ± 7.8	NS <sup>a</sup>
60+years (n=378)				
n%	270 (71)	76 (20)	346 (91.5)	NS <sup>b</sup>
Mean ± SD	12.9 ± 10.8	11.6 ± 9.9	12.7 ± 10.6	NS <sup>a</sup>

NS: nonsignificant  
a Stands for Independent t-test  
b Stands for Chi square test

Table: 4b  
Multivariate Logistic regression models for CHD/non-CHD subjects.

Variable	OR(CI=95)	P-value
Tooth Loss		
No	1	< 0.01
1 tooth	1.45	< 0.01
2-20 teeth	1.39	< 0.03
>20 teeth	3.52	
Smoking		
No	1	< 0.01
Yes	1.33	< 0.01
Diabetes		
No	1	< 0.01
Yes	3.5	< 0.01
Gender		
Female	1	< 0.01
Male	3.24	< 0.01

loss was noted significantly higher in subjects with diabetes and smoking, which are the most important confounding factors associated with cardiac diseases; and the Odds Ratio associated with cardiac patients was more than two times than the non-cardiac subjects. CHD subjects with diabetes and smoking having missing teeth, were five times higher in number and two times higher with a mean tooth loss than the healthy subjects. These results support the previous studies<sup>22,23</sup> showing that smoking and diabetes significantly contribute to tooth loss. However, cardiac patients of this study remained 1.232 times at higher risk for tooth loss, and this association was observed independent of confounding factors. Tooth loss (partial/total) is the dental equivalent of death, and tooth loss diminishes quality of life, often substantially.<sup>11</sup> The findings of studies on tooth loss and systemic diseases provide a clue that tooth loss may be considered as one of the

important components of oral diseases that affects the general health of the people.

Desvarieux<sup>24</sup> reported that greater the number of teeth lost, the greater the extent of severe periodontal disease; in turn the severity of periodontal disease is associated with increased risk of CHD.<sup>25</sup> Correspondingly, other previous studies<sup>8,10,11</sup> have reported on the risk of myocardial infarction, stroke and prevalent coronary heart disease in relation to tooth loss. The findings emerging from this study analysis explained a relationship between tooth loss and cardiac diseases and partially/fully confirmed from other studies on the same topic.<sup>26,27,28,29</sup>

The association of tooth loss, as observed in this study, with CHD and previous periodontal disease may be a significant public health problem because of the higher prevalence of the periodontal disease in the general public of developing countries.<sup>30</sup> Tooth loss distribution and risk association in individuals with and without cardiac diseases in this study provides a good reason for conducting such studies in developing countries like Pakistan where oral health is not a priority for the country stakeholders and the public at large where non-communicable diseases also show a steep rise.<sup>31</sup> This study illustrates that total tooth loss is a risk indicator for established CHD and confirms that some classical risk factors associated with an increase in CHD risk are also associated with the increased likelihood of tooth loss. Other risk factors for tooth loss, such as education and income, could not be included in this study; these may be considered as limitations.

#### CONCLUSION:

Chronic periodontal disease and tooth loss were found to be significantly higher in cardiac disease patients in comparison to healthy controls. Other risk factors found were age, gender, smoking and diabetes.

**Conflict of interest:** No conflict of interest by the authors.

**REFERENCES:**

1. Strong K, Mathers C, Leeder S, Beaglehole R. Preventing chronic diseases: how many lives can we save? *Lancet* 2005;366(9496):1578-82
2. Joshipura KJ, Ritchie C, Douglass C. Strength of evidence linking oral conditions and systemic disease. *Compendium* 2000;21(Suppl 30):13-23
3. Page RC. The pathology of periodontal diseases may affect systemic diseases: inversion of a paradigm. *Annals Periodontol* 1998;3(1):13-23
4. DeBowes LJ. The effects of dental disease on systemic disease. *Vet Clin North Am Small Anim Pract* 1998;28(5):1057-62
5. Seymour RA, Preshaw PM, Steele JG. Oral health and heart disease. *Prim Dent Care* 2002;9(4):125-31
6. Gift H. Issues of aging and oral health promotion. *Gerodontics* 1988;4:194-206
7. Beck JD, Slade G, Offenbacher S. Oral disease, cardiovascular disease and systemic inflammation. *Periodontol* 2000;23:110-20
8. Destefano F, Anda RF, Kahn HS, Williamson DF, Russell CM. Dental Diseases and risk of coronary heart disease and mortality. *Br Med J* 1993;306:668-91
9. Okoro CA, Balluz LS, Eke PL, Ajani UA, Strine TW, Town M, et al. Tooth loss and heart diseases: findings from the Behavioral Risk Factor Surveillance System. *Am J Prev Med* 2005;29(5 suppl 1):50-6
10. Paunio K, Impivaara O, Tiesko J. Missing teeth and ischemic heart disease in men aged 45–64 years. *Eur Heart J* 1993;14(Suppl. K):54-6
11. Loesche WJ. Periodontal disease as a risk factor for heart disease. *Compendium Continuing Edu Dent* 1994;15:976-91
12. Joshipura KJ, Hu FB, Manson JE, Stampfer MJ, Rimm EB, Speizer FE, et al. The effect of fruit and vegetable intake on risk for coronary heart disease. *Ann Intern Med* 2001;134:1106-14
13. Bokhari SAH, Khan AA. Growing burden of non-communicable diseases: the contributory role of oral diseases, Eastern Mediterranean region perspective. *Eastern Mediterranean Health J* 2009;15:1011-20
14. Joshipura KJ, Douglass CW, Willet WC. Possible explanations for the tooth loss and cardiovascular disease relationship. *Ann Periodontol* 1998;3:175-83
15. Ylostalo PV, Jarvelin MR, Laitinen J, Knuutila ML. Gingivitis, dental caries and tooth loss: risk factors for cardiovascular diseases or indicators of elevated health risks. *J Clin Periodontol* 2006;3(2):92-101
16. Katz R, Gustavsen F. Tooth mortality in dental subjects in a US urban area. *Gerodontics* 1986;2:104-7
17. Douglass CW, Jette AM, Fox CH, Tennsted SL, Joshi A, Feldman HA, et al. Oral health status of elderly in New England. *J Gerontol* 1993;48:M 39-46
18. Hamasha AH, Sasa I, Al Qudah M. Risk indicators associated with tooth loss in Jordanian adults. *Community Dent Oral Epidemiol* 2000;28:67-72
19. Andriankaja OM, Genco RJ, Dorn J, Dmochowski J, Hovey K, Falkner KL, et al. The use of different measurements and definitions of periodontal disease in the study of the association between periodontal disease and risk of myocardial infarction. *J Periodontol* 2006;77(6):1067-73
20. Slade GD, Gansky SA, Spencer AJ. Two-year incidence of tooth loss among South Australian aged 60+ years. *Community Dent Oral Epidemiol* 1997;25:429-37
21. Lukacs JR. Gender differences in oral health in South Asia: metadata imply multifactorial biological and cultural causes. *Am J Hum Biol* 2011 May;23(3):398-411
22. Holm G. Smoking as an additional risk factor for tooth loss. *J Periodontol* 1994;65:996-1001
23. Ueno M, Takeuchi S, Oshiro A, Shinada K, Ohara S, Kawaguchi Y. Association between diabetes mellitus and oral health status in Japanese adults. *Int J Oral Sci* 2010;2:82-9
24. Desvarieux M, Demmer RT, Rundek T, Boden-Albala B, Jacobs Jr DR, Papapanou PN, et al. Relationship between periodontal disease, tooth loss, and carotid artery plaque: the Oral Infections and Vascular Disease Epidemiology Study (INVEST). *Stroke* 2003;34(9):2120-5
25. Holmlund A, Holm G, Lind L. Severity of periodontal disease and number of remaining teeth are related to the prevalence of myocardial infarction and hypertension in a study based on 4, 254 subjects. *J Periodontol* 2006;77(7):1173-8
26. Steele JG, Sanders AE, Slade GD, Allen PF, Laltis, Nuttal N, et al. How do age and tooth loss affect oral health impacts and quality of life? A study comparing two national samples. *Community Dent Oral Epidemiol* 2004;32(2):107-14
27. Shah SA, Khitab U, Chughtai MA, Khan AS. Causes of dental extractions among 2000 patients—a study at oral and maxillofacial surgical unit, Khyber College of Dentistry. Peshawar-Pakistan *Pak Oral Dent J* 2004;24(2):209-12
28. Desvarieux M, Schwahn C, Volzke H, Demmer RT, Ludemann J, Kessler C, et al. Gender differences in the relationship between periodontal disease, tooth loss, and atherosclerosis. *Stroke* 2004;35:2029-35
29. Buhlin K, Gustafsson A, Ahnve S, Janszky I, Tabrizi F, Klinge B. Oral Health in women with coronary heart disease. *J Periodontol* 2005;76(4):544-50
30. Desvarieux M. Periodontal disease, race, and vascular disease. *Compend Contin Educ Dent* 2001;22(3 Spec):34-41
31. Boutayeb A, Boutayeb S. The burden of non-communicable diseases in developing countries. *Inter J Equity Health* 2005;4:2 doi: 10.1186/1475-9276-4-2



# Comparative Study of Lipid Profile in Multibacillary and Paucibacillary Leprosy Patients

Ghulam Sarwar<sup>1</sup>, Viqar Sultana<sup>2</sup>, Ali Gul<sup>3</sup>, Jehan Ara<sup>4</sup>

## ABSTRACT:

**Objective:** To evaluate the lipid profile in Multibacillary and Paucibacillary leprosy subjects and compare them with age and sex matched healthy control subjects.

**Materials and Methods:** This observational study was performed after approval from BASR, University of Karachi in the Department of Biochemistry, University of Karachi, from December 2014 to November, 2015. Present study was conducted in 42 newly diagnosed leprosy patients of both sexes and all ages were included in this study. The diagnosis were on clinical ground and bacterial examination by slit skin smear test, and are classified in two groups, Paucibacillary (PB) and Multibacillary (MB), based on the WHO guide lines. 1-5 skin lesions were regarded as PB with no acid fast rods on the smear and skin lesions more than 5 were regarded as MB. A positive bacterial index classifies the patient as MB, regardless of the number of skin lesions with bacteria visible on a smear.

**Results:** A total of 30 control subjects and 42 leprosy patients among 24 Multibacillary and 18 Paucibacillary leprosy were recruited for this study. Biophysical parameters in Multibacillary and Paucibacillary subjects were completely non significant when compared with control group. In biochemical parameters among Multibacillary and Paucibacillary leprosy cases, all the lipid fractions total cholesterol, triglycerides and LDL -cholesterol were significantly decreased ( $p < 0.05$ ) but HDL -cholesterol significantly increased ( $p < 0.05$ ) in both Multibacillary and Paucibacillary leprosy groups when compared with control group.

**Conclusion:** This study showed that, all the lipid fractions except HDL cholesterol were decreased significantly ( $p < 0.05$ ), where as HDL Cholesterol was increased significantly ( $p < 0.05$ ) in both Multibacillary and Paucibacillary leprosy groups when compared with control group.

**Keywords:** Leprosy, Lipid profile, Multibacillary(MB), Paucibacillary (PB)

## INTRODUCTION

Leprosy is a granulomatous, chronic infectious disease caused by *Mycobacterium leprae*.<sup>1</sup> *Mycobacterium leprae* was discovered in 1873, by G. H. Armauer Hansen in Norway, therefore leprosy is referred as Hansen's disease. It is a mutilating, debilitating, devastating and deforming disease. It mainly affects the skin and peripheral nerves, leading to sensory loss in the skin, muscle weakness and often permanent disabilities of hands and feet.<sup>2</sup>

Over the last 25 years with the efforts of leprosy control

programs and multi drug therapy (MDT) leprosy have decreased worldwide dramatically prevalence from approximately 5.4 million registered cases during the start of 2008.<sup>3,4,5</sup> Leprosy remains a significant public health problem in several parts of the world. According to World Health Organization (WHO) by 105 countries, the number of new cases detected during the year 2011 were 219, 075. indeed in year 2012 were 33,955 new cases were detected in Brazil alone (WHO 2012 ).<sup>6</sup> Leprosy is now known to be neither sexually transmitted nor highly infectious after treatment. Approximately 95% of people are naturally immune and sufferers are no longer infectious after as little as 2 weeks of treatment. It is completely curable by using multi drug therapy.<sup>7</sup> Leprosy is not a killing disease, it is a crippling disease and if not treated early and properly, may form permanent deformities.<sup>8</sup> The signs and symptoms may be ignored in the early stages until visible disabilities have not occurred.<sup>9</sup> Leprosy affects both sexes but males are affected more than females and ratio is 2:1. Until coming of AIDS, leprosy was the most feared infectious disease globally. It is still considered to be dreadful infectious disease, so normal healthy people try to avoid and breakup all kind of links to these patients.<sup>10</sup>

Leprosy has struck fear into human beings for thousands of years. In the time of Christ it was considered to be a holy curse conferred upon the people due to their wrong doings and the affected unfortunate was totally isolated and discarded. According to some ancient transcript the patients were confined to huge dungeons or well and even tortured and stone to death if they even tried to enter the cities. Leprosy cases are found world wide, Leprosy remains a public health problem with over 210,000 registered cases in world at the beginning of 2008.<sup>11</sup> The intracellular germ *Mycobacterium leprae* mediate strong inflammatory response in affected

### ✉ Dr. Ghulam Sarwar

Assistant Professor,  
Department of Biochemistry  
Shaheed Mohtarma Benazir Bhutto Medical University @ CMC  
Larkana  
Sindh  
Email: dr\_sarwar@hotmail.com

### ✉ Dr. Viqar Sultana

Professor & Chairperson  
Department of Biochemistry  
University of Karachi  
Pakistan

### ✉ Dr. Ali Gul

Assistant Professor  
Department of Pharmacology & therapeutics  
Shaheed Mohtarma Benazir Bhutto Medical University @ CMC  
Larkana  
Sindh

### ✉ Dr. Jehan Ara

Professor  
Department of Food Science and Technology  
University of Karachi  
Pakistan

Received: 28-01-2016

Revised: 25-02-2016

Accepted: 28-02-2016

individuals and causes gross destruction of tissues during the chronic course of infection.<sup>12</sup> Among all mycobacteria it is likely the most dependent on the host for basic metabolic functions, in part because of its extensive genomic decay.<sup>13</sup> Lipid metabolism in leprosy have been examined in various studies, but there has been limited work using whole metabolite profiles.<sup>14</sup> With this background present study was designed to evaluate the lipid profile in Multibacillary and Paucibacillary leprosy subjects and to compare them with age and sex matched healthy control subjects.

#### MATERIALS AND METHODS:

This observational study was performed after approval from BASR, University of Karachi in the Department of Biochemistry, University of Karachi, from December, 2014 to November 2015. A total of 42 newly diagnosed leprosy patients of both sexes and all ages were included in this study, among them 33 males and 09 females, aged 13 to 70 years (mean  $36.7 \pm 1.71$  years). The diagnosis was made on clinical ground and bacterial examination by slit skin smear test, and are classified in two groups, paucibacillary (PB) and multibacillary (MB), based on the WHO guide lines. 1- 5 skin lesions were regarded as PB with no acid fast rods on the smear and skin lesions more than 5 are regarded as MB. A positive bacterial index classifies the patient as MB, regardless of the number of skin lesions with bacteria

visible on a smear.<sup>15</sup> A total of 30 age, sex matched healthy control subjects were taken from general population for comparison. Informed consent was taken from each patient and control subject for this study. After overnight fasting, 6 ml of blood was drawn from anticubital vein after all aseptic measures, blood was allowed to clot at 37°C, serum was separated after centrifuged at 3000 rpm for 10 minutes then analyzed. Serum cholesterol was estimated by the Enzymatic kit method, serum triglycerides were determined by enzymatic colorimetric (GPO-PAP) kit method, serum HDL-cholesterol was determined by CHOD-PAP kit method<sup>16</sup> and LDL-cholesterol was calculated according to Friedewald's formula.<sup>17</sup>

#### RESULTS:

A total of 30 control subjects and 42 leprosy patients among 24 were Multibacillary and 18 were Paucibacillary leprosy recruited for this study. Biophysical parameters in Multibacillary and Paucibacillary subjects were completely non significant when compared with control group (Table 1). In biochemical parameters among Multibacillary, Paucibacillary leprosy cases, all the lipid fractions Total Cholesterol, Triglycerides and LDL - Cholesterol were significantly decreased ( $p < 0.05$ ) but HDL - Cholesterol significantly increased ( $p < 0.05$ ) in both Multibacillary and Paucibacillary leprosy groups when compared with control group (Table 2, Figure 1).

Table: 1  
Comparison of biophysical parameters of multibacillary, paucibacillary leprosy cases and controls

Biophysical Parameter	Cases		Controls
	MB (n=24)	PB (n=18)	(n=30)
Weight (kg)	51.5 ± 1.44 *	52.3 ± 2.16	56.8 ± 1.36
Height (m)	1.60 ± 0.01	1.58 ± 0.01	1.61 ± 0.01
BMI	20.2 ± 0.55	20.9 ± 0.77	21.2 ± 0.53
BP Systolic(mmHg)	118.9 ± 1.23	116.8 ± 1.78	119.0 ± 1.30
BP Diastolic(mmHg)	77.1 ± 0.75	77.7 ± 1.07	77.8 ± 0.92

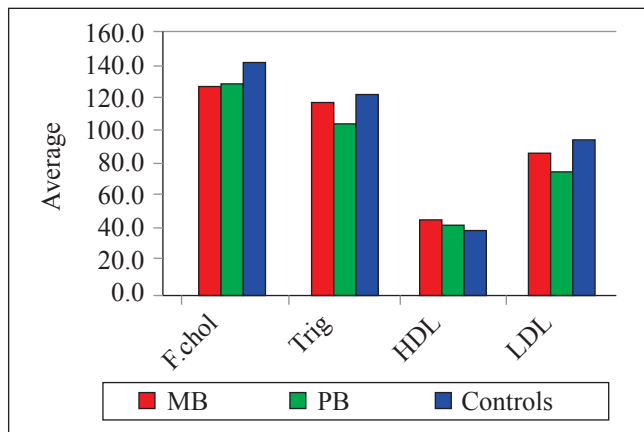
Values are expressed as mean ± s.e.m, No significant difference was observed

Table: 2  
Comparison of biochemical parameters of multibacillary, paucibacillary leprosy cases and controls

Biochemical Parameter	Cases		Controls
	MB (n=24)	PB (n=18)	(n=30)
Total Cholesterol (mg %)	* 146.1 ± 18.90	* 145.2 ± 16.90	148.2 ± 19.50
Triglyceride (mg %)	* 126.2 ± 13.08	* 125.4 ± 12.11	128.2 ± 17.08
HDL Cholesterol (mg %)	* 43.4 ± 3.24	* 44.3 ± 3.87	42.8 ± 4.40
LDL Cholesterol (mg %)	* 85.2 ± 10.28	* 84.4 ± 13.83	87.2 ± 12.6

Values are expressed as mean ± s.e.m, \*  $p < 0.05$  statistically significant

Figure 1  
Comparison of biochemical parameters of multibacillary (mb), paucibacillary (pb) leprosy cases and controls



### DISCUSSION:

Lipids play an important role in all aspects of life. Although every living organism has been found to contain sterols, cholesterol is found almost exclusively in animals, it is also the main sterol. Studies have showed that lipid profile is altered in leprosy. The lipids inside the lepra cells may be of host origin and probably may result in alteration in serum lipids and therefore some research workers used alteration in the lipid profile as diagnostic tool for leprosy. Lipids are found everywhere in the body tissue and have an important role in virtually all aspects of biological life. Serving as hormones or hormone precursors, aiding in digestion, provide energy storage and metabolic fuels, acting as functional and structural components in bio-membranes and forming insulation to allow nerve conduction or to prevent heat loss.<sup>18</sup>

Metabolism of host-derived fatty acids is required for the synthesis of mycobacterial lipids including virulence factors such as phthiocerol dimycocerosate, sulfolipid-1, and polyketide synthase-derived phenolic glycolipid (PGL) and therefore, host lipids are used both for virulence and growth.<sup>19,20</sup> The lipids inside the lepra cells may be of host origin and may result in alteration in serum lipids.<sup>21</sup> In this study we have found significant reduction in total cholesterol in both MB and PB groups ( $p < 0.05$ ), when compared with control, this observation was in accordance with Gupta.<sup>22</sup> Similarly when triglycerides levels in the two test groups were compared with control we found statistically significant reduction in MB and PB Leprosy ( $p < 0.05$ ), whereas Misra<sup>23</sup> have documented an increased in serum triglyceride levels in their studies. These observations were not in agreement with our study.

In contrary when HDL cholesterol levels in both the test groups were compared with control we observed statistically significant increased levels in both groups of leprosy ( $p < 0.05$ ). These observations were in agreement with the findings of Bansal.<sup>24</sup> Whereas LDL cholesterol decrease was statistically significant in both groups when compared with control ( $p < 0.05$ ). These

observations were in accordance with the Kher<sup>25</sup> and Ahaley.<sup>26</sup>

### CONCLUSION:

All the lipid fractions except HDL cholesterol were decreased significantly ( $p < 0.05$ ), whereas HDL cholesterol was increased significantly ( $p < 0.05$ ) in both Multibacillary and Paucibacillary leprosy groups when compared with control group. Increased level of HDL cholesterol as compared to controls are in favour of ailing lepers.

### REFERENCES:

- Swathi M, Tagore R. Study of oxidative stress in different forms of leprosy. *Ind J Derm* 2015; 60(3):321-4
- Prasad PVS and Kaviarasan PK. Leprosy therapy, past and present: Can we hope to eliminate it. *Ind J Der* 2010; 55:316-24
- World Health Organization. 2002. Leprosy. Global situation. *Wkly. Epidemiol. Rec.* 77:1-8
- World Health Organization. Global leprosy situation, beginning of 2008. *Wkly. Epidemiol. Rec.* 2008;83:293-300
- World Health Organization. Global leprosy situation, 2009. *Wkly. Epidemiol. Rec.* 2009;84:333-40
- Henrique J P, Gomes R L R, Flávia S, Prevedello C, Mira MT, Eleidi A. Investigation of association between Susceptibility to Leprosy and SNPs inside and near the BCHE Gene of Butyrylcholinesterase. *J Trop Med Brazil* 2012; doi:10.1155: 1-4
- American Leprosy Missions, Inc. About leprosy frequently asked Questions. Retrieved October 2, 2012
- Ebenso J, Velema JP. Test-Retest Reliability of the Screening Activity Limitation and Safety Awareness (SALSA) Scale in North-West Nigeria. *Lepr Rev* 2009; 80:197-204
- John AS, Rao PSS, Das S. Assessment of needs and quality care issues of women with leprosy. *Lepr Rev* 2010; 81:34-40
- Soomro FR, Shaikh GS, Bhatti NS, Baloch J, Abbasi P, Kumari M et al. Deformity and Disability Index in Patients with Leprosy in Larkana District, Sindh, Pakistan. In: *Studies on New and Old World Leishmaniases and their Transmission, with Particular References to Ecuador, Argentina and Pakistan Kyowa, Japan.* Res Rep Ser No: 7, 2004; 177-81
- Watson CL, Popescu E, Boldsen J, Slaus M, Lockwood DNJ. Single Nucleotide Polymorphism Analysis of European Archaeological *M. leprae* DNA, 2011
- Vijayaraghavan R, Suribabu CS, Oommen PK, Panneerselvam C. Vitamin E reduces reactive oxygen species mediated damage to bio-molecules in leprosy during multi-drug therapy. *Curr Trends Biotechnol Pharm* 2009; 3:428-39
- Cole ST, Eglmeier K, Parkhill J, James KD, Thomson NR, Wheeler PR et al. Massive gene decay in the leprosy bacillus. *Nature* 2001; 409:1007-11
- Al-Mubarak R, Heiden J V, Broeckling C D, Balagon M, Patrick J, Brennan PJ et al. Serum Metabolomics Reveals Higher Levels of Polyunsaturated Fatty Acids in Lepromatous Leprosy: Potential Markers for Susceptibility and Pathogenesis. *PLoS Negl Trop Dis.* Sept 2011; 5(9): 1303-5
- Grossi MAF, Leboeuf MAA, Andrade ARC, Lyon S,



- Antunes CMF, Sekula SB. The influence of ML. Flow test in leprosy classification. *Rev Soc Br Med Trop* 2008 ;41:34-8
16. Rifai N, Bachorik PS, Albers JJ. Lipids, lipoproteins and apolipoproteins In: Tietz Fundamental Clinical Chemistry, 5th ed. Edited by Burtis CA and Ashwood ER, WB Saunders, Philadelphia, 2001;pp.462-93
  17. Friedewald WT, Levy RI, Fredrickson DS. Estimation of the concentration of low density lipoprotein cholesterol in plasma, without use of the preparative ultracentrifuge. *Clin Chem* 1972; 18:499-502
  18. Rifai N, Warnick GR. Lipids, Lipoproteins, Apolipoproteins and other cardiovascular risk factors. In: Tietz Textbook of clinical chemistry. 4th ed. WB Saunders, Philadelphia 2006; pp.903-81
  19. Jain M, Petzold CJ, Schelle MW, Leavell MD, Mougous JD, Bertozzi CR et al. Lipidomics reveals control of Mycobacterium tuberculosis virulence lipids via metabolic coupling. *Proc. Natl. Acad. Sci* 2007; 104:5133-8
  20. Reed MB, Domenech P, Manca C, Su H, Barczak AK, Kreiswirth BN et al. A glycolipid of hypervirulent tuberculosis strains that inhibits the innate immune response. *Nature* 2004; 431:84-7
  21. Imaeda T. Electron microscopic analysis of the components of the laprae cells. *Int J lepr* 1960; 28:22-37
  22. Gupta A, Koranne RV, Kaul N. Study of serum lipids in leprosy. *Ind J Der Ven Lep* 2002; 68:262-6
  23. Misra UK, Venkatasubramanian TA. Serum lipids in leprosy by silicic acid column chromatography. *Ind J Lepr* 1964;32:248-59
  24. Bansal SN, VK Jain, Dayal Sand, RK Nagpal Serum lipid profile in leprosy. *Ind J Der Ven Lep* 1997; 63:78-81
  25. Kher JR, Baji PS, Ganeriwal SK, Reddy BV, Bulakh PM. Serum lipoproteins in lepromatous leprosy. *Lepr Ind* 1983; 55:80-5
  26. Ahaley SK, Sardeshmukh AS, Suryakar AN, Samson PD. Correlation of serum lipids and lipoproteins in leprosy. *Ind J Lep* 1992; 64:91-8



# Causes of Male and Female Sub Fertility in the Couples who Underwent 'In Vitro Fertilization' at Life Clinic; a Statistical Study from Lahore, Pakistan

Haroon Latif Khan<sup>1</sup>, Yousaf Latif Khan<sup>2</sup>, Nighat Mahmood<sup>3</sup>, Mariam Mustanser<sup>4</sup>, Saba Sardar<sup>5</sup>, Abdul Rahman Khawaja<sup>6</sup>

## ABSTRACT:

**Objective:** To explore the causes of male and female sub fertility in the couples undergoing 'In Vitro Fertilization' at LIFE clinic.

**Materials and Methods:** In this retrospective, cross-sectional, observational study from Lahore, Pakistan all couples coming for evaluation and treatment for sub-fertility from 1<sup>st</sup> January to 30<sup>th</sup> April 2015 at Lahore institute of fertility and endocrinology LIFE with n=344 patients were included Fertility and Endocrinology (LIFE). Sampling method was non probability consecutive. The data collection instrument was an especially designed Performa. Causes of Sub fertility male and female were studied and data was extracted from the files of LIFE.

**Results:** Out of 344 patients 138(40.1%) had female factors, 122(35.5%) had male factor, 38(11.0%) had combined factors whereas 46(13.4%) had unexplained infertility. Out of 138, 55(39.85%) females had tubal factor, 2(1.45%) had endometriosis, 21(15.22%) had PCO and 60(43.47%) had unexplained causes of sub-fertility. Out of 122 males, 90 (74.4%) had oligospermia/asthenospermia and 32(25.6%) had azoospermia.

**Conclusion:** Tubal factor was a major cause of sub-fertility in females whereas in 2/3 of the females, cause of sub-fertility was not explained. Among the males, oligospermia was the most common cause and was found in 74.4%. Health education about menstrual hygiene should be imparted early in life to prevent Sub-fertility due to infection. Premarital counseling and testing should be made easily available. Andrology should be made part of the Gynecology courses and curricula. Community based Sub-fertility research should be encouraged to assess the disease burden and frequency of preventable causes.

**Keywords:** Sub fertility, Females, Males, Causes, Lahore, Pakistan

## INTRODUCTION:

Mothering and motherhood have been regarded as issues of prime importance regardless of religion, geography, culture, art, mythology and literature. To become a parent and have children is a desire that leads goals and life plans of a common man and inability to achieve that desire may jeopardize the whole life of a person or a family.<sup>1</sup> Inability to have a kid even after 12 months of unprotected intercourse without use of contraceptives is known as subfertility.<sup>2</sup> Subfertility evaluation is usually started by the gynecologist or the couple is referred to a fertility center where the couple is evaluated for subfertility. If age of the female is more than 35 years

or there is history of menstrual irregularities or there is a known male factor for subfertility then evaluation may be commenced earlier.<sup>3,4</sup> Subfertility has two types, primary and secondary subfertility. Primary subfertility is the inability to have a child despite unprotected vaginal intercourse for twelve months among women of reproductive age (15 to 49 years).

Secondary subfertility means the inability to have a child after at least one pregnancy. Globally, primary subfertility is commoner than secondary subfertility.<sup>5</sup>

The process of conception in human beings is quite complicated and a firm understanding of anatomy and physiology of male and female reproductive system is

### ✉ Dr. Haroon Latif Khan

Embryologist  
Lahore Institute of Fertility and Endocrinology (LIFE)  
Hameed Latif Hospital  
Lahore  
Email: haroon@lifepakistan.com

### ✉ Dr. Yousaf Latif Khan

Professor  
Department of Obstetrics & Gynaecology  
Lahore Institute of Fertility and Endocrinology (LIFE)  
Hameed Latif Hospital  
Lahore

### ✉ Dr. Nighat Mahmood

Embryologist  
Lahore Institute of Fertility and Endocrinology (LIFE)  
Hameed Latif Hospital  
Lahore

### ✉ Dr. Mariam Mustanser

Embryologist  
Lahore Institute of Fertility and Endocrinology (LIFE)  
Hameed Latif Hospital  
Lahore

### ✉ Ms. Saba Sardar

Research Manager & Asst. Biostatistician  
Research Cell  
Lahore Institute of Fertility and Endocrinology (LIFE)  
Hameed Latif Hospital  
Lahore

### ✉ Dr. Abdul Rahman Khawaja

Research specialist  
Research cell  
Lahore Institute of Fertility and Endocrinology (LIFE)  
Hameed Latif Hospital  
Lahore  
Received: 29-01-2016  
Revised: 23-02-2016  
Accepted: 25-02-2016

essential for evaluating clinician to identify the investigations needed for a correct diagnosis of cause of infertility. <sup>6,7,8</sup> Modern laboratory and sonology/radiology can help an intelligent clinician to think about the treatment strategies after reaching a meaningful diagnosis. <sup>9,10</sup> The whole record of the couple is to be scrutinized to look for co-morbid conditions and associated diseases. Sometimes it is very difficult to make the couple understand the mechanics of the male and female genital tract. Conception is the result of interplay between many physical and biochemical processes gauged by the biological clock. <sup>11,12</sup> The journey of the egg starts after well-timed ovulation and successful picking up by the fallopian tubes. <sup>13,14</sup> If semen has been deposited in the vaginal tract the spermatozoa are needed to travel all the way through the cervix and uterus to reach the fallopian tube having the egg. <sup>15,16</sup> After fertilization the embryo travels to the uterus to be implanted there and the process of pregnancy starts. So subfertility evaluation may be a long, tedious and expensive process. A well prepared couple should have the awareness and patience to bear the process. When a couple is labelled to be subfertile stress, anger, jealousy and frustration may be the usual result that may overwhelm the couple due to social response or resentment of close relatives. <sup>17</sup> Subfertility may be an important cause of marital conflict and may lead to an unwanted divorce and polygamy. <sup>18,19,20</sup> In eastern culture motherhood qualifies womanhood and so women are blamed and victimized on being responsible for childlessness of the couple. <sup>21</sup> World Health Organization affirmed in 2001 that if subfertility rate exceeds 15% it may be declared to be a public health problem. <sup>22</sup> Subfertility is a social stigma and a couple is afraid to accept it and here comes into play the role of the family physician. A general practitioner or a family physician has a unique opportunity to help a childless couple. Initial workup and support can help the scared subfertile couple to think about visiting a fertility physician. An early visit to a fertility center may be of great value in terms of right and timely investigation. <sup>23</sup> The sub fertile couple is needed to be evaluated before detailed investigations. A detailed history and physical examination may point towards an area to be investigated if analyzed thoughtfully with an inquisitive mind. A well concerted effort may help the clinician to tailor the treatment strategy to have a fruitful outcome. <sup>24</sup> Couples across the globe are affected by the sub-ability to have children which is a source of personal discontent and social resentment. <sup>2</sup> A well prepared couple has the awareness and patience to bear the process. <sup>25</sup> The couple may need psychosocial counseling which may include family counseling. <sup>26</sup> Anatomy and physiology of the reproductive system may be compromised as a cause of subfertility. Sometimes genes may be implicated. <sup>27</sup> Prevalence of subfertility in different regions is different mainly due to environmental factors as well as reproductive behaviors, smoking and pollution etc. <sup>28</sup> Present study was designed to explore the causes of male and female sub fertility in the couples undergoing

‘In Vitro Fertilization’ at Lahore institute of fertility and endocrinology LIFE clinic.

**MATERIALS AND METHODS:**

In this retrospective, cross-sectional, observational study from Lahore Pakistan all couples coming for evaluation and treatment for subfertility from 1<sup>st</sup> January to 30<sup>th</sup> April at Lahore institute of fertility and endocrinology (n=334) were included. Sampling method was non probability consecutive. The data collection instrument was a specially designed performa validated by biostatistician and epidemiologist of LIFE research center. Causes of male and female subfertility were studied. Data was extracted from the files of LIFE. Data was entered into SPSS version 15.0 and descriptive analysis was done for frequencies and percentages for categorical variables. Mean, S.D and variance were calculated for numerical variables.

**RESULTS:**

In this study unexplained subfertility was seen in 13.4% (46) of the couples, both males and females were implicated in 11.0% (38) of the couples, only females were said to be responsible in 40.1% (138) of the cases where as in 35.5% (122) couple males were implicated (n=344). In the females 43.47% (60) had unexplained subfertility, in 39.85% (55) tubal factors were said to be responsible, 15.22% (21) had PCO and 1.45% (2) had endometriosis (n=138). Amongst the males oligospermia was found in 74.4% (90) and azoospermia in 25.6% (32) (n=122) (Table 1, Figure 1,2 & 3)

Table: 1

Sr.	Variable	Frequency	Percentage
1	Female factor	138	40.1
	Tubal factor	55	39.85
	Endometriosis	2	1.45
	PCOS	21	15.22
	Un-explained	60	43.47
2	Male factor	122	35.5
	Oligospermia	90	74.4
	Azoospermia	32	25.6
3	Both responsible	38	11.0
4	No cause found (female/male/unexplained)	46	13.4
Total		344	100 %

Figure: 1  
Breakup of causes of subfertility

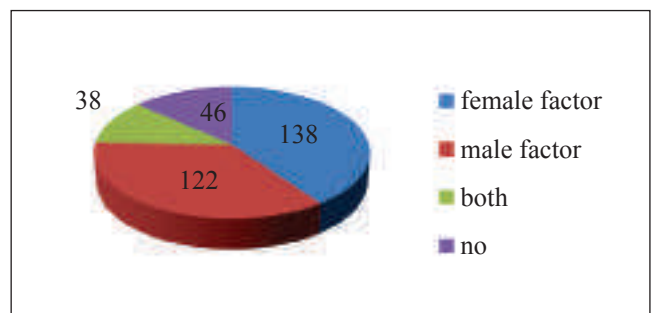


Figure: 2  
Break up of female factors

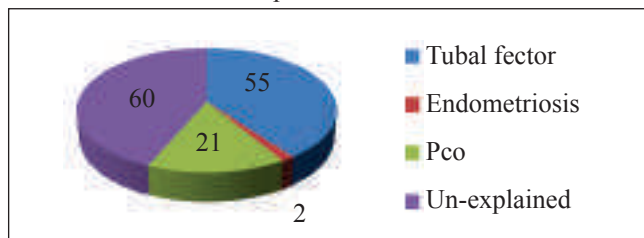
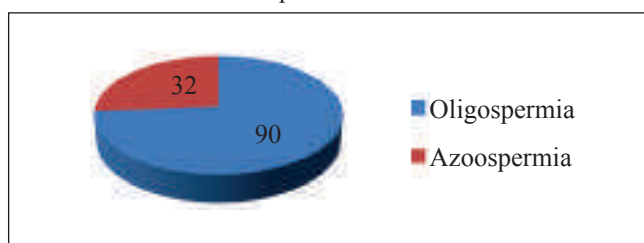


Figure : 3  
Break up of male factors



**DISCUSSION:**

In our study only females were found to be responsible for subfertility in 40.1% of the couples, only males in 35.5% (122), both males and females in 11.0% (38) and unexplained subfertility was seen in 13.4% (46) of the couples. In males oligospermia was found in 74.4% (90) and azoospermia in 25.6% (32). In females unexplained subfertility was found in 43.47% (60), tubal factors in 39.85% (55), PCOS in 15.22% (21) and endometriosis in 25.6% (32).

Subfertility may be primary or secondary depending on the history of pregnancy in the past.<sup>29</sup> Primary subfertility is more common in developed countries as compared to developing countries where secondary subfertility is also prevalent.<sup>30,31</sup>

Zargar has shown that primary subfertility is quite common in India as compared to secondary subfertility but most of the subfertile couples delay to consult the fertility physician. Male factor was found to be responsible in 22.4 % whereas ovulatory dysfunction (17.2%), tubal factor (7.2%), failure of ovaries (8.8%) and hyperprolactinemia (8.4%) were the causes found in sub fertile females.<sup>32</sup>

In a study done in Mongolia, used WHO protocol 'Standardized Investigation of the Infertile Couple' to explore causes of sub fertility, female factor was responsible for subfertility in 45.8% of couples, male factor was responsible in 25.6% of couples and unexplained subfertility was seen in 9.8% of couples whereas in 18.8% of couples both the partners contributed to subfertility. History of STI (sexually transmitted infection) and PID (pelvic inflammatory disease) were found to be 33.5% and 25.1%, respectively. History of STI (sexually transmitted infection) in males was present in 42% whereas previous testicular damage was seen in 27.7% of the males.<sup>33</sup>

Another study done in Mongolia in 2004 also used

WHO protocol. Primary subfertility was seen in 62.4% and secondary subfertility in 37.6%. Female factor was responsible for subfertility in 52.7% of couples, male factor in 6.4% of couples and unexplained subfertility was seen in 2.2% of couples whereas in 38.7% of couples both the partners contributed to subfertility. In the female factor, tubal block was found in 36.5%, pelvic adhesions in 23.6% and endocrine disorders in 32.8. Females showed to have four times more inflammatory complications as compared to males.<sup>34</sup>

Elussein analyzed 710 Sudanese couples at Khartoum Fertility Center in Sudan to explore causes of infertility. Primary subfertility was seen in 62.4% and secondary subfertility in 37.6%. Female factor was responsible for subfertility in 49.3% of couples, male factor in 36.2% of couples and unexplained subfertility was seen in 13.0% of couples whereas in 1.5% of couples both the partners contributed to subfertility. Female subfertility was found to be mostly due to ovulation failure i.e. 60.3%. Male subfertility was found to be due to Oligozoospermia (16.8%) and asthenozoospermia (17.5%).<sup>29</sup>

Whitman-Elia examined the diagnosis of the couples after evaluation and found that male factors caused subfertility in 40% of the subfertile couples, female factors in 40-55% and unexplained factors in around 10%. In a very small percentage i.e. 5% mixed factors were found to be responsible. Female factors responsible for subfertility were tubal disease, endometriosis and pelvic adhesions, ovulatory dysfunction and cervical factors. Hypothyroidism, luteal phase defect and immunologic factors were also implicated in a small percentage i.e. 5%. In 10% of the couples cause of subfertility remained unexplained even after all investigations.<sup>23</sup>

The diagnosis of idiopathic infertility in males reflects a poor understanding of the factors involved in regulation of spermatogenesis. Single gene mutations and chromosomal aberrations are some of the genetic causes of impairment of spermatozoa, which constitute ten to fifteen percent of severe subfertility in the males.<sup>35</sup>

**CONCLUSION:**

Tubal factor is the major cause of subfertility in the females whereas in 2/3rd of the females, cause of subfertility was not explained. Among the males oligospermia was the most common cause and was found in 74.4%. Health education about menstrual hygiene should be imparted early in life to prevent Sub fertility due to infection. Premarital counseling and testing should be made essential. Andrology should be made part of the Gynecology obstetrical courses and curricula. Community based Sub fertility research should be encouraged to assess the disease burden and frequency of the preventable causes.

**Acknowledgment:**

Mrs. Rameen Ahmad, assistant biostatistician in Lahore Institute of Fertility and Endocrinology (LIFE), is acknowledged for her help in data extraction and analysis.

## REFERENCES:

1. Begum BN, Hasan S. Psychological problems among women with infertility problem: a comparative study. *J Pak Med Assoc.* 2014;64(11): 1287-91
2. Mascarenhas MN, Flaxman SR, Boerma T, Vanderpoel S, Stevens GA. National, regional, and global trends in infertility prevalence since 1990: a systematic analysis of 277 health surveys. *PLoS Med.* 2012;9(12):e1001356
3. Maheshwari A, Hamilton M, Bhattacharya S. Effect of female age on the diagnostic categories of infertility. *Hum Reprod* 2008;23(3):424-538
4. Bhattacharya S, Maheshwari A, Mollison J. Factors Associated with Failed Treatment: an Analysis of 121,744 Women Embarking on Their First IVF Cycles. *PLoS ONE* 2013;8(12): e82249
5. Adamson PC, Krupp K, Freeman AH, Klausner JD, Reingold AL, Madhivanan P. Prevalence & correlates of primary infertility among young women in Mysore, India. *Indian J Med Res.* 2011;134:440-6
6. Son WY, Das M, Shalom-Paz E, Holzer H. Mechanisms of follicle selection and development. *Minerva Ginecol.* 2011;63(2):89-102
7. Stouffer RL, Bishop CV, Bogan RL, Xu F, Hennebold JD. Endocrine and local control of the primate corpus luteum. *Reprod Biol.* 2013;13(4):259-71
8. Zeleznik AJ. The physiology of follicle selection. *Reprod Biol Endocrinol.* 2004; 16:2:31
9. Abou-Setta AM, Mansour RT, Al-Inany HG, Aboulghar MM, Aboulghar MA, Serour GI. Among women undergoing embryo transfer, is the probability of pregnancy and live birth improved with ultrasound guidance over clinical touch alone? A systemic review and meta-analysis of prospective randomized trials. *Fertil Steril.* 2007; 88(2):333-41
10. Ammar T, Sidhu PS, Wilkins CJ. Male infertility: the role of imaging in diagnosis and management. *Br J Radiol.* 2012;85 Spec No 1:S59-68
11. Gratton RJ, Nisker JA, Daniel S, Toth S, Gunter J, Kaplan BR, et. al. An aggressive philosophy in controlled ovarian stimulation cycles increases pregnancy rates. *Hum Reprod.* 1993;8(4):528-31
12. Campbell S Ash Monga; *Gynaecology by Ten Teachers* (18th ed.). Hodder Education. 2006 ISBN 0-340-81662-7
13. Psychoyos A. Endocrine control of egg implantation. In: *Handbook of Physiology* 1973 Volume II, Part 2 pp 187-215 Eds RO Greep, EG Astwood and SR Geiger. American Physiological Society, Washington DC
14. Hickey M, Balen A. Menstrual disorders in adolescence: investigation and management. *Hum Reprod Update.* 2003;9(5):493-504
15. Williams M, Hill CJ, Scudamore I, Dunphy B, Cooke ID, Barratt CL. Sperm numbers and distribution within the human fallopian tube around ovulation. *Hum Reprod.* 1993; 8(12):2019-26
16. Suarez SS, Pacey AA. Sperm transport in the female reproductive tract. *Hum Reprod Update.* 2006;12(1):23-37
17. Ali S, Sophie R, Imam AM, Khan FI, Ali SF, Shaikh A, et al. Knowledge, perceptions and myths regarding infertility among selected adult population in Pakistan: a cross-sectional study. *BMC Public Health.* 2011;(4):11:760
18. Chester R. Is there a relationship between childlessness and marriage breakdown? *J Biosoc Sci.* 1972;4(4):443-54
19. Ibeh U O, Obidoa M A. Marital Disharmony: Causes and resolution strategies in Enugu State of Nigeria. *Research on Humanities and Social Sciences;* 2013; 3(22), 40-8
20. Khan HL, Khan YL, Suhail S, Awais A, Khawaja AR. Characteristics of Female Patients Visiting for Assisted Reproductive Technology in a Private Clinic in Lahore Pakistan. *J SZMC.* 2014; 5(4): 715-20
21. Hasanpoor-Azghdy SB, Vedadhir A. The emotional-psychological consequences of infertility among infertile women seeking treatment: Results of a qualitative study. *Iran J Reprod Med* 2014; 12(2): 131-8
22. Bergstrom S. Childlessness. In: Lawson J B, Harrison K A, Bergstrom S, editors. *Maternity care in developing countries.* RCOG PRESS; 2001. pp. 360-8
23. Whitman-Elia GF, Baxley EG. A primary care approach to the infertile couple. *J Am Board Fam Pract.* 2001;14(1):33-45
24. Swerdloff RS, Wang C, Kandeel FR. Evaluation of the infertile couple. *Endocrinol Metab Clin North Am.* 1988; 17(2):301-37
25. Cousineau Tara M, Domar AD. Psychological impact of infertility. *Best Pract Res Clin Obstet Gynaecol.* 2007; 21: 293-308
26. Greil A L, Slauson-Blevins, K, Mc Quillan J. The experience of infertility: A review of recent literature. *Sociol Health Illn.* 2010;32(1):140-62
27. Masoumi SZ, Parsa P, Darvish N, Mokhtari S, Yavangi M, Roshanaei G. An epidemiologic survey on the causes of infertility in patients referred to infertility center in Fatemeh Hospital in Hamadan. *Iran J Reprod Med.* 2015;13(8):513-6
28. Macaluso M, Wright-Schnapp T, Chandra A, Johnson R, Satterwhite C, Pulver A, et al. A public health focus on infertility prevention, detection, and management. *Fertil Steril* 2010; 93: 16
29. Elusse EA, Magid YM, Omer MM, Adam I. Clinical patterns and major causes of infertility among Sudanese couples. *Trop Doct.* 2008;38(4):243-4
30. Templeton A, Fraser C, Thompson B. Infertility-epidemiology and referral practice. *Hum Reprod.* 1991;6(10):1391-4
31. Menuba IE, Ugwu EO, Obi SN, Lawani LO, Onwuka CI. Clinical management and therapeutic outcome of infertile couples in southeast Nigeria. *Ther Clin Risk Manag.* 2014;10:763-8
32. Zargar AH, Wani AI, Masoodi SR, Laway BA, Salahuddin M. Epidemiologic and etiologic aspects of primary infertility in the Kashmir region of India. *Fertil Steril.* 1997;68(4):637-43
33. Bayasgalan G, Naranbat D, Tsedmaa B, Tsogmaa B, Sukhee D, Amarjargal O, et al. Clinical patterns and major causes of infertility in Mongolia. *J Obstet Gynaecol Res.* 2004;30(5):386-93
34. Philippov OS, Radionchenko AA, Bolotova VP, WHO. Estimation of the prevalence and causes of infertility in western Siberia. *Bull* 1998; 76: 183-7
35. Ferlin A, Arredi B, Speltra E. Molecular and clinical characterization of Y chromosome microdeletions in infertile men: a 10-year experience in Italy. *J Clin Endocrinol Metab.* 2007;92:762-70



# Is Gender Matters in Paediatric Cardiac Surgery

Iqbal Hussain Pathan<sup>1</sup>, Sohail Khan Bangash<sup>2</sup>, Saad Bader Zaki<sup>3</sup>

## ABSTRACT:

**Objective:** To observe the potential effect of gender difference on the survival after pediatric cardiac surgery.

**Materials and Methods:** This retrospective cross sectional study was carried out in PCICU of Pediatric Cardiac surgery department at National institute of cardiovascular diseases Karachi, Pakistan from October 2013 till September 2015. Data was evaluated to find out the effect of gender on survival of patient during their post-cardiac surgery PCICU stay for structural heart defects.

**Results:** A total of 518 patients were operated for structural heart defects on pump and including glenn shunt and fonton procedure irrespective of use of pump. 68% of these were boys and 32% were girls. Unadjusted mortality was similar for both boys and girls (9% versus 8%, P=0.87). After adjustment of complex surgeries with Aristotle basic score > 8 like intervention for TGA, univentricular hearts and DVR, more prevalent in male population, the outcome was not significantly different with 5% v/s 4% for boys and girls respectively.

**Conclusions:** Patient's gender has a no significant effect on mortality after pediatric cardiac surgeries.

**Keywords:** Congenital heart surgeries (CHD), Aristotal score, Paediatric cardiac surgery, Gender, Mortality

## INTRODUCTION:

Risk prediction of an intervention always have central role in decision making and counseling as well as quality assessment of care. In adult cardiac surgery different models of risk prediction are used for risk assessment. Literature provide considerable evidence that female gender carries a higher operative morbidity and mortality.<sup>1,2,3,4</sup> Female gender is considered as an independent risk factor in various models like euroscore,<sup>5</sup> parsonnet score<sup>6</sup> and northern new England<sup>7</sup> score. Various reasons were mentioned in literature for adverse outcome in female gender. However there is no consensus on fact behind these observations; some suggest it is inherent other suggest socioeconomic or cultural reason responsible for delayed referral. Nevertheless it is generally accepted that compared to male, female gender present with different risk profile when presented for cardiac surgery.<sup>8,9</sup> Despite of same risk profile any

intervention to medically address the same condition may result in a very different result between the genders and a given postoperative complication have more adverse effect on women.<sup>10</sup> Despite suggestion that female gender may be important risk factor for outcomes of cardiac operations, literature into gender-related influences on outcome of congenital heart disease (CHD) surgery or paediatric cardiac surgeries are sparse. Nevertheless when a review of literature was conducted for risk scoring system in paediatric cardiac surgery, we observed that scoring systems often lack complex integration of factors as in adult cardiac surgery risk scores. Scoring systems like RACHS<sup>11</sup> and the Aristotle score<sup>12</sup> used in paediatric cardiac surgery based on diagnosis and complexity of surgical procedure respectively. Thus predict more on the basis of diagnosis or intervention.

A recent report of risk-adjusted clinical data from the STS-CHSD data center failed to demonstrate effect of gender on the early postoperative mortality.<sup>13</sup> While there are many reports showing; female gender was associated with as high as 50% higher odds of death.<sup>14,15,16</sup> There may be an opposite trend than adults may be observed as it is well established fact that by nature normal female fetuses have a higher survival rate than male fetuses.<sup>17</sup>

In our study, we retrospectively analyzed data from the paediatric cardiac surgery department at NICVD to determine effect of gender on surgical outcome.

## MATERIALS AND METHODS:

Post surgical admission record of PCICU of NICVD was reviewed. All the paediatric patients underwent open heart surgeries and Glenn and Total Cavopulmonary connections for univentricular physiology irrespective of the use of cardiopulmonary bypass pump and were included in the study. Patient under went off pump surgeries were excluded from study. Similarly patients requiring emergency intervention as well as with missing data regarding sex and age were excluded from study. Data was collected from October 2013 to September 2015.

Data was analyzed and presented with percentage for

✉ **Dr. Iqbal Hussain Pathan**  
Fellow Paediatric Cardiac Surgery  
Department of Cardiac Surgery  
National Institute of Cardiovascular Disease (NICVD)  
Rafiquee (HJ) Shaheed Road  
Karachi

Email: driqbalnicvd@gmail.com

✉ **Dr. Sohail Khan Bangash**  
Associate Professor  
Paediatric Cardiac Surgery  
Department of Cardiac Surgery  
National Institute of Cardiovascular Disease (NICVD)  
Rafiquee (HJ) Shaheed Road,  
Karachi

✉ **Dr. Saad Bader Zaki**  
Assistant Professor  
Paediatric Cardiac Surgery  
Department of Cardiac Surgery  
National Institute of Cardiovascular Disease (NICVD)  
Rafiquee (HJ) Shaheed Road  
Karachi

Received: 15-02-16

Revised: 27-02-16

Accepted: 28-02-16

categorical data and numerical data with mean standard deviation. Two by two table was used to assess statistical significance of any association of unadjusted risk factor that is female gender with survival. After adjusting the risk factors like procedures with Aristotal score > 8 and double valve surgery, association of female gender with postoperative survival was also determined.

**RESULTS:**

After reviewing the record a total of 518 patients were included in this study. Their demographic data is shown in Table 1. 518 patients were operated in our department from October 2013 till September 2015. Patients shifted from PCICU to ward were 475 (91.6%) without shifting back to step up again. Among 518 patients total male patients were 350 (67.6%) compare to female patients 168 (32.4%). Incidence of unadjusted postoperative mortality was [30(8.5%) v/s 13(7.7%) p =.87] for males and females respectively. Though incidence of more complex procedures with Aristotal score >8 and double valve replacement were more prevalent in male population but it did not significantly influenced the outcome 12(5%) v/s 7(4%) for males and females respectively.

Table: 1  
Demographic data

Variable	Numbers
Total	518
Male	350 (67.57%)
Female	168 (32.4%)
Age	
Neonates	11 (2.12%)
Children	507 (98%)

Table: 2  
Diagnosis of patients or intervention where diagnosis is undefined and outcome of individual procedure is mentioned

Type of procedure	Numbers	Aristotal basic score	Mortality observed
TOF	276(52.5%)	8 (5 to 10%)	14(5.07%)
VSD	129 (25%)	6(1 to 5%)	6(4.6%)
ASD	58 (11%)	3(<1%)	1(1.7%)
TGA	22 (4.2%)		
TGA (ARTERIAL SWITH)	12	11(10 to 20%)	11(91%)
TGA (ATRIAL S WITCH)	10	11(10 to 20%)	6(75%)
MVR	4 (7%)	7.5(1 to 5%)	0
DVR	2		0
VSD + Aortic Regurgitation	1		0
TRICUSPID ATRESIA	17 (3%)		
TRICUSPID ATRESIA (GLENN)	11	7.5(1 to 5%)	2(18.2%)
TRICUSPID ATRESIA (TCPC)	6	9(5 to 10%)	2(33.3%)
TAPVR	2	9(5 to 10%)	1
PAPVR	5(.9%)	7 to 9(5 to 10%)	0
CAVCD	2	9(5 to 10%)	0
Total procedures	518		43(8.3%)

TOF (Tetralogy of Fallot) VSD (Ventricular Septal Dfect), ASD (Atrial Septal Defect), TGA (Transposition Of Great arteries), MVR (Mitral valve replacement), DVR (double valve replacement) PDA(Persistent Ductous Arteriousis), TCPC(total cavopulmonary connection),CAVSD(Complete Aterioventricular Canal Defect), TAPVR(Total Anamolus Pulmonary Venous Return), PAPVR(Partial Anamolus Pulmonary Venous Return)

**DISCUSSION:**

Gender is very important demographic variable mentioned in almost every study conducted on humans. Differences in outcomes for diseases and interventions between males and females are increasingly being observed . It is still undetermined whether gender affects outcome after pediatric cardiac surgery because before puberty, hormonal differences are less prominent. Nevertheless there is evidence that gender differences might play role in mortality and morbidity early in life. From fetal life to postnatal status there are many biological differences between both genders. Similarly it had been long known observation that male babies are more likely born preterm.<sup>18</sup> Male children are more prone to deaths from respiratory<sup>19</sup> and neurologic complications<sup>20</sup> than female children. Premature girls have higher serum level of catecholamines possibly responsible for their better survival.<sup>21</sup> Effect of gender have been well observed for coronary artery disease , heart failure, valve disease, and pulmonary hypertension.<sup>22,23,24,25</sup> Even females who survive myocardial infarction are more prone for re-infarction and higher mortality than males.<sup>26</sup> Gender related heart defect pattern difference is also common observation like more boys are presented with transposition of great vessels and left sided obstructive lesion than girls who present more with atrial septal defects and Ebstein’s anamoly.<sup>27,28</sup> In children with congenital heart disease , little data exist on differences in operative / health outcomes between males and females. Evidence of gender related difference in surgical outcomes after pediatric heart surgery has been conflicting. Possibly results of different studies were confounded by the differential pattern and severity of CHD. Potential of selection biases and small sample sizes were major limitation of these studies. Chang and colleagues, reviewed large number of inpatient hospital records including almost all congenital heart surgeries with the aim to determine the effect of gender on outcome; females were found to have higher odds of death than males. Nevertheless it involved a single region while excluding low flow hospitals.<sup>29</sup> Same reports from California showed higher in hospital mortality rates for female children following cardiac surgery.<sup>30</sup> Harry has also reported increase association of adverse outcome with female gender with odds ratio 1.31.<sup>31</sup> There are many reports suggestive of opposite trend in paediatric patients with adverse out come for male patients . New England Regional Infant Cardiac Registry presented with data showed that female infants had a 5% lower mortality.<sup>32</sup> A cohort mortality study in patients observed higher death rates in males gender compared to females

with CHD . Pattern was persistent from 10 years of age and onward till adulthood.<sup>33</sup>

Our study failed to demonstrate any difference in survival in either of gender. The predominance of males having CHD surgery compared to females was most significant finding of our study, 68% v/s 32% for males and females respectively. Though males had more severe CHD at birth, female sex failed to demonstrate a protective effect on surgical mortality. Limitations of our study are first it has retrospective study design. Secondly like analysis of any databases it is important to understand effect of documentation of record within the presented diagnostic and procedural report. Thirdly it must be paired with this consideration that is the potential presence of comorbidity profiles not related to surgical disease or intervention. Fourthly, this study does not directly examine the effect of payer status on risk-adjusted outcomes. Finally, inter institutional transfers cannot be tracked. Whereas the strengths of this study are, it is a single centre study and all patients were operated by two surgeons and got same level of care thus helping us in controlling many biases.

#### CONCLUSION:

Patient's gender has a no significant effect on mortality after pediatric cardiac surgeries. Although we have found that a higher proportion of males had high-risk procedures and underwent more CHD surgeries but we have observed no difference on the survival.

#### REFERENCES:

1. Edwards FH, Clark RE, Schwartz M. Coronary artery bypass grafting: The Society of Thoracic Surgeons National Database experience. *Ann Thorac Surg* 1994;57:12-9
2. Hammar N, Sandberg E, Larsen FF, Ivert T. Comparison of early and late mortality in men and women after isolated coronary artery bypass graft surgery in Stockholm, Sweden 1980 to 1989. *J Am Coll Cardiol* 1997;29:659-64
3. Woods SE, Noble G, Smith JM, Hasselfeld K. The influence of gender in patients undergoing coronary artery by-pass graft surgery: an eight year prospective hospitalized cohort study. *J Am Coll Surg* 2003;196:428-34
4. Zitser-Gurevich Y, Simchen E, Galai N, Mandel M. Effect of perioperative complications on excess mortality among women after coronary bypass: The Israeli Coronary Bypass Graft study (ISCAB). *J Thorac Cardiovasc Surg* 2002;123:517-24
5. Nashef SAM, Roques F, Michel P, Gauducheau E, Lemeshow S, Salamon R, Euro SCORE study group. European system for cardiac operative risk evaluation (Euro SCORE). *Eur J Cardiothorac Surg* 1999;16:9-13
6. Bernstein AD, Parsonnet V. Bedside estimation of risk as an aid for decision-making in cardiac surgery. *Ann Thorac Surg* 2000;69:823-8
7. O'Connor GT, Plume SK, Olmstead EM. Multivariate prediction of in-hospital mortality associated with coronary artery bypass graft surgery. Northern New England Cardiovascular Disease Study Group. *Circulation* 1992; 85:2110-8
8. Koch CG, Khandwala F, Nussmeier N, Blackstone EH. Gender profiling in coronary artery bypass grafting. *J Thorac Cardiovasc Surg* 2003;126:2044-51
9. Zindrou D, Taylor KM, Bagger JP. Excess coronary artery bypass mortality among women with hypothyroidism. *Ann Thorac Surg* 2002;74:2121-5
10. O'Connor GT, Morton JR, Diehl MJ. Differences between men and women in hospital mortality associated with coronary artery bypass graft surgery. *Circulation* 1993;88(1): 2104-10
11. Jenkins KJ, Gauvreau K, Newburger JW, Spray TL, Moller JH, Iezzoni LI. Consensus-based method for risk adjustment for surgery for congenital heart disease. *J Thorac Cardiovasc Surg* 2002;123(1):110-8
12. Lacour-Gayet F, Clarke D, Jacobs J, Comas J, Daebritz S, Daenen W et al. The Aristotle score: a complexity-adjusted method to evaluate surgical results. *European Journal of Cardio-thoracic Surgery* 2004;25:911-24
13. Dibardino DJ, Pasquali SK, Hirsch JC, Benjamin DK, Kleeman KC, Salazar JD et al. Effect of sex and race on outcome in patients undergoing congenital heart surgery: an analysis of the society of thoracic surgeons congenital heart surgery database. *Ann Thorac Surg*. 2012;94:2054-9
14. Chang RK, Chen AY, Klitzner TS. Female sex as a risk factor for in-hospital mortality among children undergoing cardiac surgery. *Circulation*.2002;106:1514-22
15. Klitzner TS, Lee M, Rodriguez S, Chang RK. Sex-related disparity in surgical mortality among pediatric patients. *Congenit Heart Dis*. 2006;1:77-88
16. Marelli A, Gauvreau K, Landzberg M, Jenkins K. Sex differences in mortality in children undergoing congenital heart disease surgery: a United States population-based study. *Circulation*. 2010;122:S234-S40
17. Hassold T, Quillen SD, Yamane JA. Sex ratio in spontaneous abortions. *Ann Hum Genet*. 1983;47(Pt 1):39-47
18. Ingemarsson I. Gender aspects of preterm birth. *BJOG*. 2003;110(suppl 20):34-8
19. Khoury MJ, Marks JS, McCarthy BJ, Zaro SM. Factors affecting the sex differential in neonatal mortality: the role of respiratory distress syndrome. *Am J Obstet Gynecol*. 1985;151:777-82
20. Ingemarsson I, Herbst A, Thorngren-Jerneck K. Long term outcome after umbilical artery acidemia at term birth: influence of gender and duration of fetal heart rate abnormalities. *Br J Obstet Gynaecol*. 1997;104:1123-7
21. Greenough A, Lagercrantz H, Pool J, Dahlin I. Plasma catecholamine levels in preterm infants. Effect of birth asphyxia and Apgar score. *Acta Paediatr Scand*. 1987;76: 54-9
22. Rankin JS, Hammill BG, Ferguson TB Jr, Glower DD, O'Brien SM, De Long ER et al. Determinants of operative mortality in valvular heart surgery. *J Thorac Cardiovasc Surg*. 2006;131:547-57
23. Bondy CA. Aortic coarctation and coronary artery disease: the XY factor. *Circulation*. 2012;126:5-7
24. Engelfriet PM, Duffels MG, Moller T, Boersma E, Tijssen JG, Thaulow E et al. Pulmonary arterial hypertension in adults born with a heart septal defect: the Euro Heart Survey on adult congenital heart disease. *Heart*. 2007;93: 682-7
25. Mc Sweeney JC, Cody M, O'Sullivan P, Elbersson K, Moser DK, Garvin BJ. Women's early warning symptoms of acute myocardial infarction. *Circulation*. 2003;108:26 19 -23
26. Wenger NK. Coronary heart disease: the female heart is vulnerable. *Prog Cardiovasc Dis*. 2003;46:199 -229



27. Samanek M. Boy: girl ratio in children born with different forms of cardiac malformation: a population-based study. *Pediatr Cardiol.* 1994;15:53-7
28. Miller-Hance WC, Tacy TA. Gender differences in pediatric cardiac surgery: the cardiologist's perspective. *J Thorac Cardiovasc Surg.* 2004;128:7-10
29. Chang R-K, Chen AY, Klitzner TS. Female sex as a risk factor for in-hospital mortality among children undergoing cardiac surgery. *Circulation.* 2002;106:1514-22
30. Klitzner TS, Lee M, Rodriguez MS, Chang RK. Sex-related disparity in surgical mortality among pediatric patients. *Congenit Heart Dis.* 2006;1:77-88
31. Harry A. Seifert, MD, MSCE A B, David L. Howard R, Jeffrey H et al. Female gender increases the risk of death during hospitalization for pediatric cardiac surgery. *J Thorac Cardiovasc Surg* 2007;133:668-75
32. Fyler DC. Report of the New England regional infant cardiac program. *Pediatrics.* 1980;65:375- 461
33. Boneva RS, Botto LD, Moore CA, Yang Q, Correa A, Erickson JD. Mortality associated with congenital heart defects in the United States: trends and racial disparities, 1979 –1997. *Circulation.* 2001;103:2376-81



## COMMENTARY

# Workplace Based Assessment: A Step Towards Competency Based Training

Shafaq Sultana

### ABSTRACT:

Assessment of clinical performance is important but challenging and the key features of good assessment include clarity of purpose, formative feedback, transparency, credibility, cost efficiency, use of multiple methods and ongoing quality assurance. In traditional educational paradigm assessment is based on integrating teaching, learning and assessment however complex professional attributes are difficult to assess using standardized assessment methods but can be better assessed in workplace situation. It has been observed that trainees are seldom observed, assessed and given feedback at workplace. Workplace based assessment as a part of an assessment strategy provides an opportunity to incorporate feedback and facilitate integration. It assesses the performance in everyday clinical practice in healthcare setting and tracks the progress in integrating clinical knowledge and skills for clinical decision making. Faculty training is an absolute radical to valid work based assessment and should include feedback training sessions as well as specifics of individual assessment instrument.

**Keywords:** Workplace, Based Assessment, Competency, Training

### INTRODUCTION:

It has been very well said that "Assessment drives learning." For just over two decades leading educationists, including medical educators, have highlighted the intimate relationship between learning and assessment. Indeed, in an educational context it is now argued that learning is the key purpose of assessment. Assessment is defined as "any systematic method of obtaining information from tests and other sources, used to draw inferences about characteristics of people, objects, or programs." Good assessment is difficult but critical to effective development of clinical learners.<sup>1</sup> This article provides you some principles of assessment and some professional competencies that are difficult to assess within the traditional assessment system for example multiple choice test and written papers, OSCE. We need to assess the competencies using information directly derived from the working environment.<sup>2</sup>

There is an increasing recognition of the need to include work based assessment as part of an overall assessment strategy; as well as providing an opportunity for authentic assessment incorporating feedback and facilitating integration of assessment and learning it also presents significant challenges.

In 1990 George Miller proposed<sup>3</sup> a frame work to assess clinical competence. It shows that there are different types of competence demonstrated at each stage of the pyramid and that it is vital to record, monitor and assess these in an authentic way. At the lowest level of the pyramid is knowledge (knows), followed by competence (knows how), performance (shows how) and action (does). The 'knows' level of pyramid can be assessed using simple knowledge tests, e.g. multiple choice

questions (MCQs). The 'knows how' level can be assessed using one best MCQs, and unfolding patient management problems (PMPs). Objective structured clinical examination (OSCEs) can assess the 'shows how' level but when we are taking about assessing does level, it refer to assess performance in context<sup>2</sup> (figure 1). The problem is that what doctors do in controlled assessment situations correlates poorly with their actual performance in professional practice. These problems give rise to a need to develop assessment method that focus on top of pyramid that's where workplace assessment comes in.

Utility of any assessment method is a product of its reliability, validity, cost, acceptability and educational impact. It is necessary to set explicit standards and assessment program that should be monitored against these parameters. There will be a balance between the educational impact, acceptability, reliability, validity and feasibility across the suite of WBAs implementation.

### Types of assessment:

1. Formative Assessment: The assessment for learning through observation feedback.
2. Summative Assessment: The assessment of learning for a high stake decision (pass/fail, certification etc.)

### What is work place based assessment?

WPBA is the assessment of competence based on what a trainee actually does in the workplace.<sup>4</sup> In Medical education context it means the assessment that is conducted in the clinical setting. Work place based assessment is usually a competency based assessment. The competencies assessed by workplace based assessment are medical expertise, decision making, communication, team work and collaboration, leader-ships, management and health advocacy, scholarship teaching and professionalism.<sup>6</sup>

The main aim of WPBA is to aid learning (Assessment for learning) by providing trainees with constructive feedback. Trainees can use the same methodology to assess themselves (Reflective practice).<sup>7</sup> The assessments help the supervisor to chart a trainee's progress during a placement.

WPBA is an essential part of an assessment system. It is comprehensive assessment system that collectively

✉ Dr. Shafaq Sultana

Lecturer

Department of Pathology

Bahria University Medical & Dental College

Karachi.

Email: shafaq\_sultana@yahoo.com

Received: 11-02-16

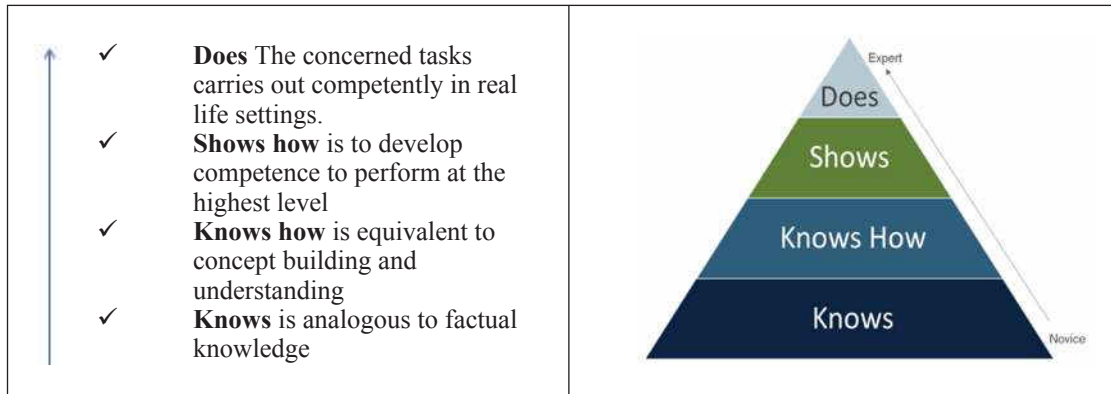
Revised: 25-02-16

Accepted: 27-02-16

forms an overall profile of an individual by testing their skills, knowledge and behaviors.<sup>8</sup> It evaluates trainees in the environment where they will be working upon graduation. The use of WBAs will support the

individual's practice of providing safe patient-centered care. It offers the opportunity of formative assessment and feedback at the same time.

Figure: 1  
Miller's pyramid<sup>3</sup>



**Purpose of work based assessment:**

WPBA provides day to day practice in working environment.<sup>9</sup>It is used to support education and maximize learning impact. It assures patient safety, monitors progression, structures learning plans and provides a transparent policy on assessment for learning and its relationship to assessment of learning.

**Types of WPBA:**

A number of methods of assessment for observation are used in clinical settings.<sup>10</sup>Some of the common methods are Mini - clinical evaluation exercise (mini - CEX), Direct observation of procedural Skills (DOPS), Case - based discussion (CBD), Multi - source feedback (MSF), Mini-PAT peer assessment tool (Figure 2a, 2b).

Areas of competence assessed through WPBA

Figure 2a<sup>3</sup>

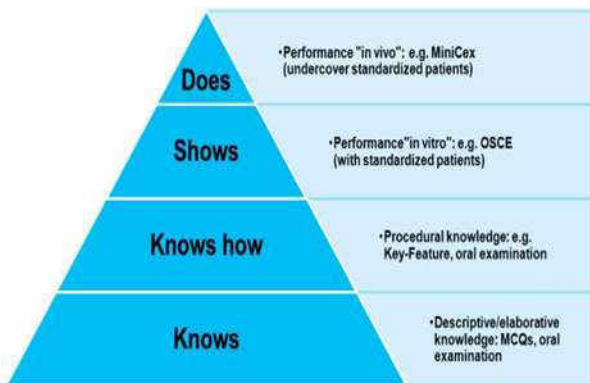
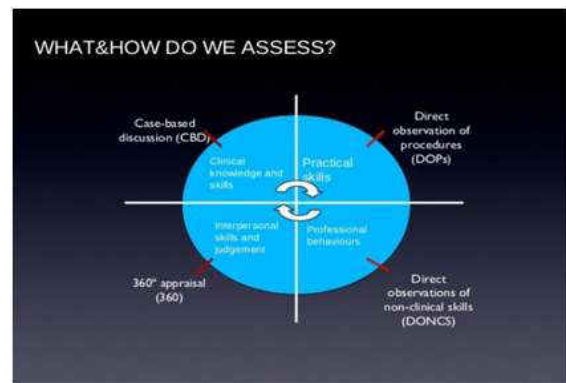


Figure 2b<sup>15</sup>



**Effective feedback:**

The work place based assessment is a useful exercise to get personalized feedback to create, enhance and support learning, thus strengthening the formative impact of this assessment.<sup>11</sup>It provides an educational supervision to learners about progress and encourages the practice of reflection. It fosters an environment where assessment for learning along with assessment of learning is seen. Thus effective feedback informs trainees of their progress, facilitates learning, and motivates them to engage in appropriate learning activities.<sup>11</sup>

**Strengths and limitations:**

WPBA is potentially highly valid assessment tool.<sup>12</sup> It

can assess 'does' (what the doctor actually does in practice) and contribute to an understanding of whether the trainee can apply the skills and knowledge in particular situation. It focuses on Trainee experience and maps achievement in a competency framework.<sup>13</sup> Moreover it helps to identify those who might need particular educational support early in training and creates a nurturing culture and provides feedback. It samples widely in the workplace across the curriculum and utilizes a range of judges and assessors.<sup>14</sup>Not yet robust in terms of reliability. Other assessments of 'show how' and 'know how' are needed to provide reassurance in terms of reliability. WPBA does not assess knowledge

directly. If educational supervision is not working appropriately trainees are more likely to try to delay or avoid assessments, or ignore feedback.<sup>15</sup> WPBA is learner dependent and vulnerable.

#### CONCLUSION:

Workplace Based Assessment is definitely a step towards Competency Based Training. The assessment tools should be designed and continuously refined to maximize their validity in competency frame work. They also ensure what they assess? and what they are intended to assess? WPBA have positive impact on learning and performance and creates a nurturing culture.

Several good methods are available that have major influence on learning and should be utilized. The opportunity for educational feedback is an important contribution to the assessment process. At the end of each assessment session the experts should provide a comprehensive evaluation to trainee based on strengths and weaknesses. Faculty members then should be encouraged for self - assessment and develop action plans, which will enable the trainees to address any deficiencies. In a sense, these methods bring summative and formative assessment closer to each other. However faculty need to be trained on 'how to give effective feedback' as it is one of the most important success factor in workplace based assessment.

#### REFERENCES:

1. Miller MD, Linn RL, Gronlund NE. The Role of Measurement and Assessment in Teaching. In: Robb C. Measurement and Assessment in Teaching. 10th Ed. USA: Kevin M. Davis; 2009: 26-46
2. Overeem K, Faber MJ, Arah OA, Elwyn G, Lombarts KM, Wollersheim HC, et al. Doctor performance assessment in daily practise: does it help doctors or not? A systematic review. *Med Educ* 2007;41:1039-49
3. Miller GE. The assessment of clinical skills/competence/performance. *Acad Med* 1990; 65(9):S63-S7
4. Norcini J. Work place assessment. In: Swanwick T (eds.) *Understanding Medical Education: Evidence, Theory and Practice*. London: Wiley-Blackwell; 2010: p 232-45
5. Norcini J, Burch V. Workplace-based assessment as an educational tool: AMEE guide no.31. *Med Teach* 2007; 29:855-71
6. Van der Vleuten CPM, Schuwirth, LWT. Assessing professional competence *Medical Education* 2005;39:309-17
7. Rethans JJ, Norcini JJ, Baron-Maldonado M, Blackmore D, Jolly BC, LaDuca T et al. The relationship between competence and performance: implications for assessing practice performance. *Med Educ* 2002;36:901-9
8. Wilkinson J, Crossley J, Wragg A, Mills P, Cowan G, Wade W. Implementing workplace-based assessment across the medical specialties in the United Kingdom. *Medical Education* 2008; 42: 364-73
9. Norcini JJ Current perspectives in assessment: the assessment of performance at work. *Medical Education*. 2005 39:880- 9
10. Wass V, Van der Vleuten CPM, Shatzer J, Jones R. Assessment of clinical competence. *The Lancet* 2001;357:945-9
11. Cantillon P, Wood D. Direct observation tools for workplace-based assessment. In: Cantillon P, Wood D, (Eds). *ABC of learning and teaching in medicine*. 2nd ed. London: Wiley-Blackwell; 2010:P. 52-9
12. Miller A, Archer J. Impact of workplace based assessment on doctors' education and performance: a systematic review. *BMJ* 2010;341:Cite this as: *BMJ* 2010;341:c5064
13. Van der Vleuten CPM. The assessment of professional competence: development, research and practical implications. *Adv Health Sci Educ* 1996;1:41-67
14. Postgraduate Medical Education and Training Board Workplace Based Assessment Subcommittee. Workplace based assessment. Postgraduate Medical Education and Training Board, 2005
15. General Medical Council, Academy of Medical Royal Colleges. Workplace based assessment: a guide for implementation. 2010



## STUDENTS CORNER

# Oral Health Education Poster Competition 2015 by Department of Community Dentistry at BUMDC

Raima Bashir<sup>1</sup>, Kulsoom Fatima Rizvi<sup>2</sup>

The Department of Community Dentistry of Bahria University Medical & Dental College, organized a students' "Oral Health Education Poster competition" on Wednesday 7<sup>th</sup> Oct 2015 from 9am to 2pm at the Ibn-e-Sina auditorium of BUMDC. The program was headed by Dr. Kulsoom Fatima Rizvi (Head of Department) and hosted by Dr. Raima Bashir and Dr. Anum Sami (lecturers of the department). The event commenced by display of scientific posters on a variety of topics relating to Community Dentistry and pertaining to oral conditions like xerostomia, baby bottle tooth decay, dental anxiety, prevention strategies and various other topics like cosmetic dentistry, which were prepared and presented by the 2<sup>nd</sup> Year BDS students. The whole Class had been divided into 14 groups which were supervised by Dr. Raima and Dr. Anum Sami. The Director General of BUMDC Vice Admiral (Retd) Tehseenullah Khan and Dean & Principal Health Sciences, Brig (Retd) Dr. Shaheen Moin were invited as chief guests. Respectable heads and faculty of entire medical & dental sections of BUMDC as well as guests from other Medical & Dental Colleges were also invited to provide support and encouragement to the participants. The Jury comprised of Dr Ashar Afaq (HOD Community Dentistry & Vice Principal of DUHS OJHA campus), Dr Mariam Azfar (HOD Community Dentistry, JSMU), Dr Ambreen Usmani (HOD Anatomy BUMDC) and Dr Mushtaq Memon (HOD Periodontology/Oral Medicine BUMDC) who judged each poster on their concept, knowledge, and presentation skills and scored them accordingly. The results were then compiled by commutating the marks allotted by each jury member. Every one appreciated the efforts of the students and shared their expert opinions and knowledge with them.

The event then proceeded with an introductory presentation on departmental achievements and a featured

video on outcomes and accomplishments of a "Community Support Program" running successfully in BUMDC under the dynamic supervision of Dr. Kulsoom Fatima.

Dr. Kulsoom Fatima welcomed the respected guest, faculty and students of all BDS batches and congratulated all the participants and the organizers for conducting a successful event. Two students of 3<sup>rd</sup> year BDS, Affaf Fatima and Safia Anwar presented their researches before the audience that have been presented earlier at both national and international conferences. The motive was to encourage the upcoming batches and students towards research work, extracurricular participation and enhancing their knowledge and confidence. Dr. Ashar Afaq and Dr. Mariam Azfar also addressed the audience sharing their experience and delivered presentations on oral health promotion.

The ceremony was concluded with prize distribution of shields and certificates to meritorious students. 1<sup>st</sup> prize was awarded to poster titled "Xerostomia" by Anika Choudary, Hafsa Saeed, Kulsoom Zaidi and Anum Malik, 2<sup>nd</sup> prize was bagged by poster on "Importance Of Salivary Diagnostic Tools In Oral Disease" by Bibi Hafsa, Tooba Taj, Saba Gul and Moneezay Jaffer and poster titled "Dental care for mother and child" by Ramsha Iqbal, Madiha Parveen and Rehab Tahir aquired 3<sup>rd</sup> prize. Honorary shields were also presented to respected judges and certificates of appreciation were also awarded to the organizers namely Dr Raima Bashir, Dr Anum Sami and Assistant Qadeer Ahmed for working day in and out in making this event a complete success. The Director General Vice Admiral (Retd) Tehseenullah Khan and Dean health Sciences Brig (Retd) Dr Shaheen Moin congratulated the participants and the department with words of tremendous appreciation for working wholeheartedly as a team in the execution and completion of the event and emphasized that more of such activities should be conducted on regular basis.

Lastly, Dr Kulsoom gave her vote of thanks to the chief guests for sparing their precious time, entire medical and dental faculty for joining them, her team and the entire 2<sup>nd</sup> year BDS student for their tremendous hard work. BUMDC encourages such healthy extracurricular activities for students as they set a platform for all the young nurturing dentists and researchers to present their real-time work and get appreciations which are their real achievements.

### ✉ Dr. Raima Bashir

Lecturer

Department of Community Dentistry  
Bahria University Medical & Dental College  
Email: raimabashir@gmail.com

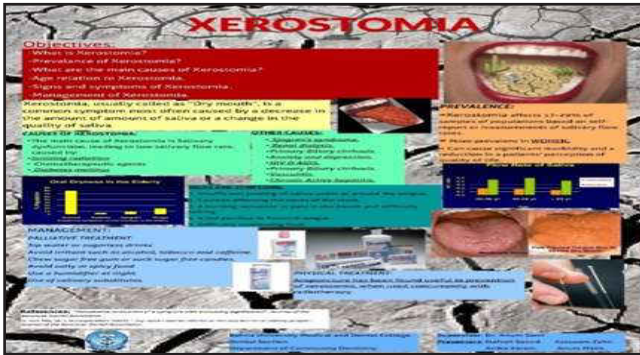
### ✉ Dr. Kulsoom Fatima Rizvi

Assistant Professor and Head  
Department of Community Dentistry  
Bahria University Medical & Dental College  
Karachi

Received: 27-12-2015

Revised: 02-01-2016

Accepted: 04-01-2016



1<sup>st</sup> position Poster  
Titled: Xerostromia



2<sup>nd</sup> Position Poster  
Titled: Importance of Salivary Diagnostics in Oral Diseases



3<sup>rd</sup> Position Poster  
Titled: Dental Care for Mother and Child



1<sup>st</sup> Position Holder Group  
Group members  
Hafsa Saeed, Anika Akram, Anum Malik, Kulsum Zaidi



2<sup>nd</sup> Position Holder Group  
Group Members  
Bibi Hafsa, Moneezay Jaffar, Saba Gul, Tooba Taj



3<sup>rd</sup> Position Holder Group  
Group Members  
Ramsha Iqbal, Madiha Perveen, Rehab Tahir



Students Presenting before a Judge for Poster Evaluation



Group Photo with Director General, Judges and Departmental Faculty

## CASE REPORT

# Endodontic Therapy of Mandibular Canines with Two Canals in a Single Root

Shama Asghar<sup>1</sup>, Mahwesh Hasan<sup>2</sup>, Asghar Ali<sup>3</sup>

### ABSTRACT:

Mandibular canines have less anatomical diversities than other teeth. Mandibular canine is generally a single rooted tooth with one wide root canal. This case describes the root canal treatment of a mandibular canine with two completely separate root canals in a single root. Clinical and radiographic examination revealed a mandibular canine with carious lesion and pulp exposure, tender to percussion. The precise understanding of the dental endocanalicular system's anatomy is critical in the success of the root canal management.

**Keywords:** Mandibular canine, Endodontics, Two canals, Anatomical variations

### INTRODUCTION:

The objective of endodontic therapy is the eradication of infection from the root canal and the prevention of reinfection.<sup>1</sup> Abnormal root and root canal morphology can be found associated with any tooth with varying degree and frequency and affects endodontic management.<sup>2</sup> Knowledge of the root canal anatomy is the basic pre-requisite for successful completion and outcome of endodontic treatment.<sup>3</sup> Mandibular canines' anatomy usually presents just one wide canal associated with a single root.<sup>2</sup> In mandibular canine, the occurrence of two roots and more than two root canals is rare, ranging from 1% to 5%. Most of the lower canines' studies (98.3%) presented a single root, with three internal variations, one canal and one foramen (92.2%), two canals and one foramen (4.9%), two canals and two foramina (1.2%). This paper reports the case of a patient with mandibular canines with a single root and two entirely separated root canals.

### CASE REPORT:

A 47-year-old woman patient reported to Department

✉ **Dr. Shama Asghar**  
Assistant Professor & Head  
Department of Operative Dentistry  
Dental Section  
Bahria University Medical and Dental College  
Karachi  
E-mail: shama.asghar@yahoo.com

✉ **Dr. Mahwesh Hasan**  
Registrar  
Department of Operative Dentistry  
Dental Section  
Bahria University Medical and Dental College  
Karachi

✉ **Dr. Asghar Ali**  
Associate Professor and Head  
Department of Community Dentistry  
Baqai Dental College  
Karachi  
Received: 13-12-2015  
Revised: 05-01-2016  
Accepted: 07-01-2016

of Operative and Endodontics of Bahria Dental College with the chief complaint of pain in lower left anterior region for the last one week. The clinical examination revealed that the mandibular left canine had proximal caries involving pulp. Periapical radiograph showed pulp exposure and patient was diagnosed with irreversible pulpitis. Any unusual medical histories were not revealed. Informed consent was obtained and root canal procedure was started after administering local anesthesia, 2% lidocaine. Rubber dam was placed and endodontic access cavity was created through the center area of the lingual surface using a long-shank, round bur on a high-speed handpiece and an Endo Z tapered safe-end bur. Only one root canal orifice was found in the center of the tooth after access cavity opening. The pulp was extirpated completely using a K-hand file size No. 10 and 15. After removal of infected pulp, another canal toward labial side was visible (Figure 1a). The pulp was removed from this canal by using a K-hand file size No. 10 and 15. Working length of both canals was taken with electronic apex locator. The length of the buccal canal was 22 mm and the lingual was also 22 mm and No. 25 K-files were replaced in both canals (Figure 1b) and radiograph was taken at two different angulations to confirm the presence of extra canals (Figure 2a). Radiograph revealed the presence of two canals and one root. In the next stage, the mechanical treatment was performed. The canals were prepared, using a step back instrumentation technique with a hand file with master apical filling up to No. 30. Rinsing of the endodontic space was done with plenty of antiseptic solution, using a 2.5% of sodium hypochlorite as irrigant, at every change of instruments. The canals were dried with sterile paper points.

The root canals were filled with intra-canal medicament with the help of Lentulo-spiral and cavity was sealed with temporary filling material and analgesic was given. After a week, patient was recalled for obturation, she had no pain. Temporary filling was removed, and canals were irrigated with normal saline and dried with sterile paper points. The canals were filled with gutta-percha cones and root canal sealer (Figure 2b).

Figure: 1a  
Two canals in left lower mandibular canine  
(one Lingual and Labial)



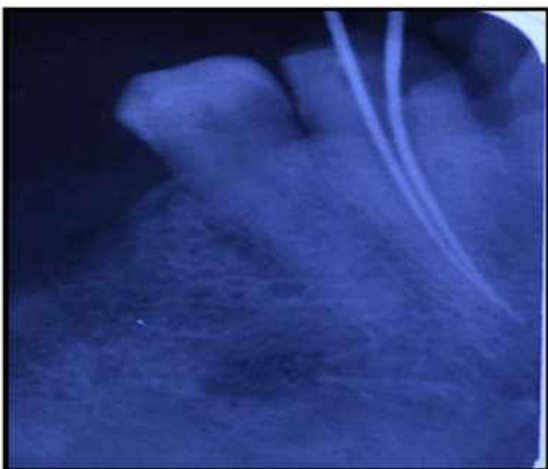
Figure: 2b  
Obturation of both canals  
(filled with GuttaPurcha points)



Figure : 1b  
Left lower mandibular canine with two K-file No. 25 in  
both canals



Figure: 2a  
Radiograph of working length of left lower mandibular  
canine with two K-file No. 25 in both canals



### DISCUSSION:

Knowledge of anatomic variations is essential because endodontic success is related to a thorough debridement of the root canal system.<sup>2</sup> The mandibular canine is the second longest tooth in dentition.<sup>5</sup> It is only 1-2 mm shorter than the upper canine.<sup>6</sup>

On studying the literature it becomes seeming clear that there is disagreement of opinion as to the structure of root canal of human permanent teeth. The occurrence of two canals in single rooted tooth has been reported to be as low as 0.0% and as high as 6.25%<sup>7</sup>. Incidence of one root with two canals in mandibular canines was detected by several authors.<sup>5,6,7</sup> Hess in 1921, Barrett in 1925, Pucci and Rei in 1944, Madeira in 1973, De Deuss in 1982, all have demonstrated the case of single root and two canals.<sup>6,7,8</sup> In 1972, Pineda F and Kuttler Y found that 18.5% of the mandibular canines had two canals through a study on 187 radiological images.<sup>9</sup> Green D reported 13% in 1973 following the analysis of 100 teeth.<sup>10</sup> Hession reported 11% in 1977.<sup>11</sup> Similar results were obtained by Kaffe I et al. in 1985, in a radiological study on 400 mandibular canines, in vivo, which showed a percentage of 13.75%.<sup>12</sup> Laurichesse et al in 1986 informed that 2% of mandibular canines presented with single root and two canals and 1% had two roots and two canals.<sup>13</sup> Our case report demonstrated two canals in a single root. In 2006, Bakianian studied 100 canines by using the stereomicroscope; he noticed the occurrence of two radicular canals in 12% of the cases.<sup>14</sup> Another study conducted on internal anatomy of mandibular canine, showed that 4.9% had two canals and one foramen, 1.2% had two canals and two foramina.<sup>15</sup> According to Vertucci, in single-rooted mandibular canines, type II and type III configurations may be found in 14% and 3% of the cases, respectively.<sup>16</sup> Other researchers have performed in-vitro studies using sectioning<sup>9</sup> or radiographic<sup>10</sup> techniques: they also reported that about 15% of single-rooted lower canines showed two canals with one or two foramina. The anatomy of root canal system dictate the condition under which root



canal therapy is carried out and can directly affect this prognosis.<sup>16</sup> Extra root or rootcanals if not detected, remain a major reason for failure of treatment. Incomplete removal of all the irritants from the pulp space may increase the possibility of treatment failure.

#### CONCLUSION:

Clinicians should be aware of anatomical variations in the teeth they are managing, and should never assume that canal systems are simple. Even though the most common anatomy of mandibular canines comprises a single root and a single root canal, clinicians should consider the possible variations and always search for the second root canal in teeth with either one or two roots.

**Conflicts of interest:** The authors have no conflicts of interest relevant to this article.

#### REFERENCES:

1. Cleghorn B, Christie W, Dong C. Anomalous mandibular premolars: a mandibular first premolar with three roots and a mandibular second premolar with a C-shaped canal system. *International Endodontic Journal*. 2008;41(11):1005-14
2. Irodi S, Farook AZ. Three Rooted Mandibular Molar; Radix Entomolaris and Paramolaris. *International Journal of Dental Clinics*. 2011;3(1):102-4
3. Khandelwal A MB P. An Endodontic management of mandibular incisor with bifurcated root canal. *International Journal of Dental Clinics*. 2011;3(2):87-8
4. Salgar AR, Chandak MG, Manwar NU. Surgical endodontic management of external root resorption. *International Journal of Dental Clinics*. 2011;3(2):93-4
5. Gopikrishna V, Bhargavi N, Kandaswamy D. Endodontic management of a maxillary first molar with a single root and a single canal diagnosed with the aid of spiral CT: a case report. *Journal of Endodontics*. 2006;32(7):687-91
6. Victorino FR, Bernardes RA, Baldi JV, Moraes IG, Bernardinelli N, Garcia RB, et al. Bilateral mandibular canines with two roots and two separate canals: case report. *Brazilian Dental Journal*. 2009;20(1):84-6
7. Ouellet R. Mandibular permanent cuspids with two roots. *Journal of Canadian Dental Association*. 1995;61(2):159-61
8. Shanna R, Pécora JD, Lumley P, Walmsley A. The external and internal anatomy of human mandibular canine teeth with two roots. *Dental Traumatology*. 1998;14(2):88-92
9. Green D. Double canals in single roots. *Oral Surgery, Oral Medicine, Oral Pathology*. 1973;35(5):689-96
10. Gpineda F, Kuttler Y. Mesiodistal and buccolingual roentgen graphic investigation of 7,275 root canals. *Oral Surg Oral Med Oral Pathol*. 1972; 33:101-10
11. Hession RW. Endodontic morphology. II. A radiographic analysis. *Oral Surg Oral Med Oral Pathol*. 1977;44:610-20
12. Kaffe I, Kaufman A, Littner MM, Lazaron A. Radiographic study of the root canal system of mandibular anterior teeth. *Int Endod J*. 1985;18:253-9
13. Laurichesse JM, Maestroni J, Breillat J. *Endodontie Clinique*, 1986; 1stedn. Paris, France: Edition CdP, 64-6
14. Bakianian Vaziri P, Kasraee S, Reza Abdolsamadi H, Abdollahzadeh S, Esmaeili F, Nazari S, Vahedi M. Root canal configuration of one-rooted mandibular canine in an Iranian population: an in vitro study. *J Dent Res Dent Clin Dent Prospects*. 2008, 2 :28-32
15. Vertucci FJ. Root canal anatomy of the mandibular anterior teeth. *J Am Dent Assoc*. 1974; 89:369-71
16. D'Arcangelo C, Varvara G, De Fazio P. Root canal treatment in mandibular canines with two roots: a report of two cases. *International Endodontic Journal*. 2001;34:331-4



## LETTER TO EDITOR

# Medical Brain Drain – An Increasing Social Stigma

Maria Shoaib

To,  
The editor,

Migration of talented and educated professionals from their native country, in search of higher salaries, advanced technology, stable political conditions, and a better living, defines the term brain drain. This is not only concerned with people but is also concerned with migration of education, intellect, talent and resources as well. Currently in our country, brain drain of doctors is a rising social stigma as Pakistan ranks as 3<sup>rd</sup> leading country for International Medical Graduates working abroad.<sup>1</sup>The reason may be financial, social stressors, job satisfaction and better learning opportunities but this mobility is very asymmetrical. The immigrants are usually from developing nations to countries like USA, UK, Canada, Australia which are marked 1<sup>st</sup> world and beneficiaries of large scale physician immigration over last fifty years.<sup>2</sup>

These professional émigrés are competent, profound and skilled people, who are moreover an asset to that society, their contributions at home would have been valuable to their health care and socioeconomic sectors. Further, they could be role models and a source of great academic inspiration. It is of great interest that the recipient countries and the migrants are not at loss, in fact the donor country is decreasing doctor-patient ratio and in short loss of resources and human capital there.<sup>3</sup> Although at some point there are remittances that immigrant physicians be sent home, however the stated disadvantages are not compensated by the clinical and educational link they establish or their economical support.

Pakistan has a greater burden of diseases and increasing mortality, with a growing disease load of cancers, cardio vascular diseases, disabilities and an increasing rate of infectious diseases and nutritional deficiencies.<sup>4</sup> It is indeed a socioeconomic need to identify the reasons

which push towards efflux of the talented young graduates or trained professionals to the other countries at national level. Disparities in working conditions and demotivating factors should be resolved. The young graduates and students should be given some incentives, leadership and better chances to progress. The government should design policies, introduce teaching research programs and take responsibility to train, retain and sustain its youthful and important work force. Many countries intensify their efforts to attract and retain foreign students, which increases the risk of brain drain in the developing countries. In poor countries, this transfer can change the skill structure of the labor force, cause labor shortages, and affect fiscal policy. It can be a boon or a curse for developing countries.<sup>5</sup> Finding opportunities and seeking better choices is a basic human right. But a promising future with good learning and work environment, improvised salaries, optimism, peace and equity can certainly help to reduce and control the social malaise of the medical brain drain.

### REFERENCES:

1. Dodani S, LaPorte RE. Brain drain from developing countries: how can brain drain be converted into wisdom gain? *JRSM*. 2005;98(11):487-91
2. Mullan F. The metrics of the physician brain drain. *New England Journal of Medicine*. 2005;353(17):1810-8
3. Chen LC, Boufford JI. Fatal flows-doctors on the move. *New England Journal of Medicine*. 2005;353(17):1850-2
4. Talati JJ, Pappas G. Migration, medical education, and health care: a view from Pakistan. *Academic Medicine*. 2006;81(12):S55
5. The brain drain from developing countries. *IZA World of Labor* 2014: 31 doi: 10.15185/izawol.31 | Frédéric Docquier © | May 2014 | wol.iza.org



✉ **Dr. Maria Shoaib**  
Email: syedamariashoaib@gmail.com  
Received: 13-02-2016  
Revised: 16-02-2016  
Accepted: 18-02-2016

# JBUMDC INSTRUCTION TO AUTHORS

The Journal Of Bahria University Medical and Dental College abbreviated as JBUMDC is a peer reviewed biannual multidisciplinary biomedical journal of basic and clinical health sciences. It accepts manuscripts prepared in accordance with the "Uniform Requirements for Submission of Manuscripts for Biomedical Journals, updated October 2008", adopted by International Committee of Medical Journal Editors (ICMJE). & PMDC guidelines for medical & Dental journals. The Journal will encompass manuscripts from all fields of biomedical sciences in the form of Editorial (Invited), Original Article, Review Article, Short Communication, (Commentary), Case report and Letter to editor.

## Peer Review Policy:

Every paper will be read by the editor. Selected papers will be sent to two reviewers. If statistical analysis is included examination by the staff statistician will be carried out.

## Plagiarism:

JBUMDC follows the ICMJE, PMDC and HEC guidelines for plagiarism.

## Preparation of Manuscript:

Type the manuscript on ISO A4 (212 × 297 mm), with margins of at least 25 mm (1 inch). Type or print on only one side of the paper. Use double spacing throughout the manuscript. Start each section on new page. Number pages consecutively, beginning with the title page. Put the page number in the lower right-hand corner of each page.

## Contents of Manuscript for submission:

Submission items include a Covering letter, Letter of undertaking duly signed by all authors, Title page and the Manuscript [Abstract, Key words, Introduction, Materials & Methods, Results, discussion, conclusion, acknowledgement, Authorship, Conflict of interest, References, Tables, Figures]. Title page should have complete title of the manuscript, the names of all authors with qualifications, their department, affiliation, telephone number, e-mail, corresponding author, address for correspondence, short running title, source of funding (grant/equipment/drugs), number of figures and tables, total word count, total number of pages.

### 1. Abstract

It should have no more than 150 words for unstructured abstracts or 250 words for structured abstracts. The abstract should state the purpose of the study (objective), basic procedures (materials & methods with study design, subjects/animals, place & duration of study, drug/chemical/equipment, procedure or protocol), main findings (results) and conclusion. It should emphasize new and important aspects of the study. Below the abstract provide, 3-10 key words that will assist indexers in cross-indexing the article and may be published with the abstract.

### 2. Introduction

State the purpose of the article and summarize the rationale for the study. Give only strictly pertinent references and do not include data or conclusions from the work being reported.

### 3. Materials & Methods

Describe your selection of the observational or experimental subjects (patients or laboratory animals, including controls) clearly. Identify the age, sex, and other important characteristics of the subjects. Identify the methods, apparatus (give the manufacturer's name and address in parentheses), and procedures in sufficient detail to allow other workers to reproduce the results. Identify precisely all drugs and chemicals used, including generic name(s), dose(s), and route(s) of

administration. For randomized clinical trials provide information on all major study elements, including the protocol (study population, interventions or exposures, outcomes, and the rationale for statistical analysis), assignment of interventions (methods of randomization, concealment of allocation to treatment groups), and the method of masking (blinding). Authors submitting review manuscripts should include a section describing the methods used for locating, selecting, extracting, and synthesizing data. These methods should also be summarized in the abstract. All studies must be approved by the relevant Ethics Committee/Institution Review Board of the respective institutions.

### 4. Results

Present your results in logical sequence in the text, tables, and illustrations. Do not repeat in the text all the data in the tables or illustrations; emphasize or summarize only important observations. Describe appropriate indicators of measurement error or uncertainty such as confidence intervals, P values. Report complications of treatment & dropouts from a clinical trial. Specify any general-use computer programs employed for analysis.

### 5. Discussion & Conclusion

Emphasize the new and important aspects of the study and the conclusions that follow from them. Do not repeat in detail data or other material given in the Introduction or the Results section. Include in the Discussion section the implications of the findings and their limitations, including implications for future research. Relate the observations to other relevant studies. Link the conclusions with the goals of the study.

### 6. Acknowledgment

List all contributors who do not meet the criteria for authorship, such as a person who provided purely technical help, writing assistance, or a department chair who provided only general support. Financial and material support should also be acknowledged.

### 7. Authorship

Authorship credit is based only on 1) substantial contributions to conception and design, or acquisition of data, or analysis and interpretation of data; 2) drafting the article or revising it critically for important intellectual content; and 3) final approval of the version to be published. Conditions 1, 2, and 3 must all be met. Authors should provide a description of what each contributed.

### 8. Conflict of interest

All authors have to disclose and submit any financial /personnel relationship that might bias and inappropriately influence their work.

### 9. References

Majority of the references must be from last five years. Local references must also be included. Vancouver style should be followed. Examples are:

#### a) Standard journal article

List the first six authors followed by et al.

I) Less than 6 authors:

Vega KJ, Pina I, Krevsky B. Heart transplantation is associated with an increased risk for pancreato-biliary disease. *Ann Intern Med* 1996 Jun 1;124 (11):980-3.

II) More than six authors:

Parkin DM, Clayton D, Black RJ, Masuyer E, Friedl HP, Ivanov E, et al. Childhood leukaemia in Europe after Chernobyl: 5 year follow-up. *Br J Cancer* 1996;73:1006-12.

#### b) Organization as author

The Cardiac Society of Australia and New Zealand. Clinical exercise stress testing. Safety and performance guidelines.

Med J Aust 1996; 164: 282-4.

**e) No author given**

Cancer in South Africa [editorial]. S Afr Med J 1994;84:15.

**d) Chapter in a book**

Phillips SJ, Whisnant JP. Hypertension and stroke. In: Laragh JH, Brenner BM, editors. Hypertension: pathophysiology, diagnosis, and management. 2nd ed. New York: Raven Press; 1995. p. 465-78.

**e) Newspaper**

HasanMansoor. Excessive use of drugs creating resistance to antibiotics. The Dawn 2013, 24 June; sect. Metropolitan (col.1-4)

**10. Tables**

Type or print out each table with double spacing on a separate sheet of paper. Number tables consecutively in the order of their first citation in the text and supply a brief title for each. Give each column a short or abbreviated heading. Place explanatory matter in footnotes. Explain in footnotes all nonstandard abbreviations that are used in each table. Identify statistical measures of variations, such as standard deviation and standard error of the mean. Do not use internal horizontal and vertical rules.

**11. Illustrations (Figures)**

Figures should be professionally drawn and photographed. Photographic prints 127 × 173 mm (5 × 7 inches). Photomicrographs should have internal scale markers. Symbols, arrows, or letters used in photomicrographs should contrast with the background. If photographs of people are used, either the subjects must not be identifiable or their pictures must be accompanied by written permission to use

the photograph. Figures should be numbered consecutively according to the order in which they have been first cited in the text. If a figure has been published, acknowledge the original source and submit written permission from the copyright holder to reproduce the material.

**Legends for Illustrations**

Type or print out legends for illustrations using double spacing, starting on a separate page, with Arabic numerals corresponding to the illustrations. When symbols, arrows, numbers, or letters are used to identify parts of the illustrations, identify and explain each one clearly in the legend. Explain the internal scale and identify the method of staining in photomicrographs.

**Units of Measurement**

Measurements of length, height, weight, and volume should be reported in metric units. Temperatures in degrees Celsius, Blood pressure in millimeters of mercury & all hematologic and clinical chemistry measurements in the metric system in terms of the International System of Units (SI).

**Abbreviations and Symbols**

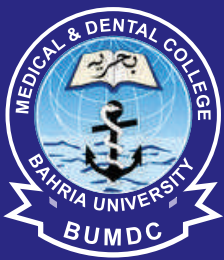
Use only standard abbreviations. Avoid abbreviations in the title and abstract. The full term for which an abbreviation stands should precede its first use in the text unless it is a standard unit of measurement.

**Sending the Manuscript to the Journal**

Submit manuscript by e-mail: editor.bumdc@bahria.edu.pk or by post on CD with two hard copies to: Editor, JBUMDC, Bahria University Medical & Dental College, DHA Phase-II, Adjacent PNS Shifa, Karachi. All correspondence regarding submitted manuscripts will be via e-mail.



S #	Type of Article	Abstract type & word count	Key words	Total word count	References	Tables (Max)	Figures (Max)
1	Editorial	-	-	1000-1500	10-12	-	-
2	Review Article	Unstructured (150)	3-6	3000-3500	40-60	4	2
3	Original Article	Structured (250)	3-10	2500-3000	25-35	4	3
4	Medical Education	1. Original Structured (250)	3-10	2500-3000	25-35	4	3
		2. Review Unstructured (150)	3-6	3000-3500	40-60	4	2
		3. Reproducible work (guide lines, questionnaire)	Mention Source, Accessed on, Retrieval date				
5	Short Communication /Commentary/ Opinions/ Perspective	Unstructured (150)	3-6	1200-1500	15-20	2	1
6	Student Corner	1. Original article Structured (250)	3-10	2500-3000	25-35	4	3
		2. Views/Perspectives/ Opinions Unstructured (150) Students Activity Report (BUMDC)	3-6	1200-1500	8-10	1	1
7	Case Report	Unstructured (150)	3-5	1200-1300	10-12	1	2
8	Letter to Editor	-	-	400-500	5	-	-
9	Instruction to Author	Please See the Text Detail					



## **Bahria University Medical and Dental College, Karachi**

Published by: Bahria University Medical & Dental College

Adjacent PNS Shifa DHA Phase II Karachi, Pakistan.

Ph: +92-21-35319491-9

Website: <http://jbumdc.bahria.edu.pk>

JBUMDC Web Mail: [editor.bumdc@bahria.edu.pk](mailto:editor.bumdc@bahria.edu.pk)