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3 **Association between handgrip strength and maximum expiratory**
4 **flow with site-specific bone mineral density of healthy young adults**

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6 **Rimsha Tahir¹, Faateh Ahmad Rauf², Shafaq Ismail³, Huma Saeed Khan⁴,**
7 **Shahid Hasan⁵**

8 **1-3** 3rd Year MBBS Student, CMH Lahore Medical College and Institute of Dentistry, Lahore,
9 Pakistan; **4,5** CMH Lahore Medical College and Institute of Dentistry, Lahore, Pakistan

10 **Correspondence:** Rimsha Tahir. **Email:** Rimsha895@live.com

11
12 **Abstract**

13 **Objective:** To explore the association of handgrip strength and peak expiratory
14 flow rate with site-specific bone mineral density T scores, and to study the
15 correlation of body mass index and physical activity with the three parameters.

16 **Methods:** The cross-sectional study was conducted in November 2018 at the
17 Combined Military Hospital, Lahore Medical College and Institute of Dentistry,
18 Lahore, Pakistan, and comprised healthy young adults. Data was collected using
19 the international physical activity questionnaire. The subjects were grouped into
20 low, moderate and high categories. Handgrip strength, peak expiratory flow rate,
21 bone mineral density T scores and body mass index were measured for each
22 subject. Data was analysed using SPSS 24.

23 **Results:** Of the 102 subjects, 52(50.9%) were males and 50(49%) were females.
24 There was a significant difference between the median values for all parameters
25 of males and females ($p < 0.05$). Bone mineral density and physical activity were
26 moderately associated with handgrip strength and peak expiratory flow rate

27 (p<0.05). Body mass index showed a moderate association with bone mineral
28 density (p<0.05).

29 **Conclusion:** Healthy young adults with better respiratory function and handgrip
30 strength were found to have higher values of site-specific calcaneal bone mineral
31 density. Subjects with higher reported levels of physical activity and a higher
32 body mass index within a healthy range presented with improved values of bone
33 mineral density, handgrip strength and peak expiratory flow rate.

34 **Key Words:** Bone mineral density, Handgrip strength, Peak expiratory flow rate,
35 Body mass index, Physical activity.

36

37 **Introduction**

38 Poor bone health has important implications for individuals, their quality of life
39 and the economics of a society. Presently in Pakistan, low bone health and
40 osteoporosis is not regarded as a national health priority and there are no
41 sufficient health policies on the matter. Bone mineral density (BMD) is used in
42 clinical medicine as an indirect marker of osteoporosis and fracture risk. It is a
43 medical term denoting the amount of mineral matter, or bone mineral content, per
44 square centimetre of bone [1]. BMD results from a dynamic process of
45 osteoclasts resorbing older bone and the periodical deposition of new matrix by
46 osteoblasts, in a process known as remodelling that occurs throughout the
47 lifetime. Bone resorption predominates in the elderly, with each standard
48 deviation (SD) decrease in BMD being linked to 2.6 times increase of hip-fracture
49 risk. Low bone mass is thus an important risk factor for fracture as well as
50 osteoporosis, a condition characterised by skeletal fragility and a T score of <-2.5
51 [2].

52 Common risk factors for osteoporosis include a personal history of fracture,
53 current low bone mass, advanced age, female gender, menopausal status, low
54 lifetime calcium intake, vitamin D deficiency and an inactive lifestyle [3]. There

55 are certain medical conditions from which secondary osteoporosis is known to
56 stem from as well, including endocrine disorders, renal diseases, disorders of the
57 gastrointestinal tract (GIT) as well as manifesting as an adverse effect of
58 medications, like immunosuppressants, anti-seizure medications, corticosteroids
59 etc. It has also been shown in a study that a low haemoglobin (Hb) value
60 significantly affected bone turnover in the tested subjects and has shown to be an
61 independent risk factor for osteoporosis [4]. Moreover, it is imperative to
62 recognise the prevalence of osteoporosis in a first-degree relative as an essential
63 clinical indication for BMD screening. It is important to identify certain pre-
64 emptive measures regarding lifestyle changes, and hormone replacement therapy
65 (HRT) may prove to be beneficial in reducing premenopausal bone loss.

66 Restricted lung function has also been detailed to producing an undesirable effect
67 on BMD. The relationship between maximum expiratory flow with BMD can be
68 understood by analysing the bone mass of patients with chronic obstructive
69 pulmonary disease (COPD) presenting with a reduced peak expiratory flow rate
70 (PEFR). Though the precise mechanism linking diminished lung function to
71 osteoporosis remains unclear, United Kingdom National Institute for Health and
72 Care Excellence (NICE) guidelines on osteoporosis recommend utilising it as a
73 fracture prediction tool, considering COPD as a secondary cause of osteoporosis
74 [5].

75 The current study was planned to explore the association of handgrip strength
76 (HGS) and PEFR with site-specific BMD T scores, and to study the correlation of
77 body mass index (BMI) and physical activity (PA) with the three parameters.

78 **Subjects and Methods**

79 The cross-sectional study was conducted in November 2018 at the Combined
80 Military Hospital (CMH) Lahore Medical College and Institute of Dentistry,
81 Lahore, Pakistan. After approval was obtained from the institutional ethics review
82 committee, the sample size was calculated using the formula [6]:

83
$$\text{Sample size} = \frac{(Z_{1-\alpha/2})^2 SD^2}{d^2}$$

84 In the formula, $Z_{1-\alpha/2}$ is a standard normal variate and at $p < 0.05$ it is 1.96; d
 85 = absolute error calculated via $d^2 = (\mu_1 - \mu_2) = 19.27 - 19.07 = 0.04$. The average
 86 SD of the variable is taken from a previous study [7]: Average $SD^2 = \left\{ \frac{0.94 + 0.74}{2} \right\}^2$
 87 = 0.7056

88 The sample was raised using non-probability convenience sampling and informed
 89 consent was taken from each subject. Those included were normal healthy male
 90 and female volunteers aged 18-25 years. The subjects were without a current
 91 status or any history of smoking, COPD, fractured bones, anaemia, consumption
 92 of nutritional supplements, corticosteroids and HRT within the preceding one
 93 year. Also, the subjects consumed <4 cups of cola per week and <4 cups of
 94 tea/coffee per day. Additional inclusion criteria for female subjects involved
 95 regular menstrual cycle. Those excluded were individuals exhibiting other
 96 recognised causes of osteoporosis, including but not restricted to rheumatic
 97 diseases and endocrine disorders.

98 In order to maintain confidentiality and anonymity, each questionnaire was
 99 assigned a sequential code which was used during data entry and analysis. Socio-
 100 demographic details, including age, gender, year of study, were obtained through
 101 a self-administered questionnaire.

102 The international physical activity questionnaire [8] was administered to the
 103 participants and their level of PA was calculated as low, moderate or high via a
 104 series of 7 pre-set questions. Anthropometric measurements such as height and
 105 weight were measured from the respective subjects according to the National Health and
 106 Nutrition Examination Survey (NHANES) manual [9]. BMI was calculated using the
 107 standard formula BMI = weight in kilograms divided by height in meters squared.

108 Wright's Peak Flow Meter (MicroPeak, Cardinal Health, UK) was used to obtain
 109 PEFr using sterilised mouth-pieces. In a standing position, the participants were

110 instructed to deeply inspire followed by a quick and maximal forceful expiration
111 into the flow meter [10]. The highest of three recordings was selected for
112 assessment.

113 HGS of the dominant hand was measured with the aid of BIOPAC Hand
114 dynamometer (SS25 LA S/N 12013156) and analysed by using the BIOPAC
115 Student Lab software BSL R version 4.0.0). Each subject was seated in a standard
116 position with the forearm flexed at a 90° angle from the elbow, and briefed
117 beforehand on the proceedings. A maximum clench force was exerted followed
118 by a sustained force of contraction till the reading declined to half of the
119 maximum value [11].

120 Site-specific calcaneal BMD T scores was calculated using the Osteosys
121 SONOST 3000 quantitative ultrasound scan (QUS) bone densitometer under the
122 supervision of an experienced technician. By means of the quantitative ultrasound
123 scan, the equipment measured two parameters of bone profile: broadband
124 ultrasound attenuation (BUA) and the speed of sound (SOS). The structural
125 feature of the bone, as a function of different attenuations of the ultrasound wave,
126 was expressed as the BUA (Db/MHz). SOS (m/s) was a counter of the strength,
127 elasticity and fragility by measuring the time taken for the ultrasound to travel
128 through the calcaneus [12]. According to the World health Organisation (WHO)
129 criterion, the obtained T scores were computed and interpreted as; osteoporosis
130 <-2.5 SD, osteopenia -1.1 to -2.4, and normal -1.0 or higher) [2].

131 Data was analysed using SPSS 24. Shapiro-Wilk's test was used to assess the
132 normality of continuous variables. The non-normally distributed quantitative
133 variables were presented as medians and interquartile ranges (IQRs: Q3-Q1).

134 Intergroup comparison of means was done using non-parametric Mann Whitney
135 U test to verify the differences in gender. Spearman's rank correlation analysis
136 was used to assess the association involving PEFr, HGS, BMD (T score), BMI
137 and PA. In all cases, $p < 0.05$ was taken as statistically significant.

138 **Results**

139 Of the 102 subjects, 52(50.9%) were males and 50(49%) were females. Based on
140 gender, there was a significant difference in height, weight, BMI, PEFr, HGS
141 and BMD (Table 1).

142 Correlation between PEFr, HGS, BMD, BMI and PA was significant (Table 2).

143 Based on low, moderate and high categories of PA, PEFr, HGS and BMD were
144 higher in subjects performing more exercise (Figure).

145

146 **Discussion**

147 To our knowledge, this is the first study exploring the association of HGS, PEFr
148 and BMD in Pakistani population.

149 The results indicate that HGS and PEFr were moderately positively correlated
150 with BMD (T score). This is in agreement with a study [7]. A higher PEFr was
151 found in individuals with higher BMD. One study showed that kyphosis and
152 vertebral fractures secondary to osteoporosis had a direct relationship with
153 decline in vital capacity (VC) [13]. Another longitudinal study found that a
154 decrease in the forced expiratory volume (FEV1) was associated with increased
155 prevalence of osteoporosis during a 3-year follow up [14].

156 Likewise, the current study observed a correlation of PA with HGS and PEFr.
157 This can be proposed to individuals engaging in a high level of PA having
158 markedly increased levels of lung function evident from the higher peak flow
159 rate. A similar positive correlation of PA was found with HGS. The mechano-
160 responsive cells of the bone are known to be the osteocytes and osteoblasts. These
161 cells respond to dynamic stress placed on the bone via strain-adaptive
162 remodelling. Alterations in exercise levels due to occupational factors during
163 early adulthood have a significant effect on the maintenance and generation of
164 peak bone mass and thus compensatory exercise is highly relevant for bone health
165 in young adults [15]. Peak bone mass is defined as the amount of bony tissue

166 present at the end of the skeletal maturation. Physical exercise has been associated
167 with higher peak bone mass in young adults independent of gender and vitamin
168 D levels for both males and females [16]. There are primarily two methods to
169 measure BMD: dual ray X-ray absorptiometry (DEXA) and QUS. While DEXA
170 has long since been the gold standard for diagnosis of osteoporosis, it is an
171 expensive test and not readily available everywhere. BMD T scores measured at
172 the calcaneus using QUS have been proven to be highly correlated with BMD T
173 scores measured using DEXA at the femoral neck, total femur and lumbar spine
174 [17].

175 The Osteosys SONOST 3000 QUS bone densitometer was used to measure BMD
176 in the current study. This particular machine has been used in various researches
177 and, thus, its results are reliable and reproducible [18]. This particular device was
178 used to investigate BMD in pre- and post-menopausal Pakistani women [19].

179 In a country where the healthcare system is absorbed in tackling infectious
180 diseases and the population branding osteoporosis as a foreseeable disease of old
181 age [20], the urgency of dealing with osteoporosis is not ranked as a priority. An
182 ambulatory study setting in Lahore in 2013 revealed the prevalence of
183 osteoporosis in 18.6% of the study population and osteopenia in 64.1% [21]. The
184 mean age of the participants was 34 ± 11.8 years, which was different than the age
185 group included in the current study. These figures disclose that an imperative
186 intervention is required. The development of osteoporosis is characterised by a
187 micro-architectural deterioration of bone tissue as well as contribution from
188 extra-skeletal factors. Consequently, this proposes the shift of osteoporosis
189 management from a reactive to a predictive field. To attain better management, it
190 is important to comprehend both the physiological and lifestyle factors that
191 contribute to a low bone mass as well as to identify the individuals at risk for
192 osteoporosis. It is estimated that approximately one in two women and one in four
193 men will suffer from osteoporotic fractures in their lifetime [22].

194 According to the WHO, globally, 23% of adults and 81% of adolescents do not
195 meet the global recommendations on PA for health. The 2015 Insufficient
196 Physical Activity country factsheet for Pakistan put these figures at 26% for
197 adults and 84.5% for adolescents, and they were bound to increase if an
198 intervention was not done [23]. The clinical implication of this work lies in the
199 fact that patients with low HGS or expiratory flow rates should be investigated
200 for loss of BMD due to limitations on PA imposed by their co-morbid conditions.
201 QUS can be used to screen for low BMD in high-risk populations in lieu of DEXA
202 in low-income or remote areas, such as basic health units (BHUs). Furthermore,
203 QUS machines are more portable and reduce patient exposure to ionizing
204 radiation [24]. It has been proven to be an effective tool to screen for osteoporosis.
205 Though not assessed in the current study, the effect of nutritional supplements,
206 including vitamin D, vitamin E and calcium, has been reported in literature to
207 have a marked positive effect on BMD. Likewise, individuals who underwent
208 HRT showed increased levels [25]. Exercise serves to maintain BMD whereas
209 dietary intake of calcium and vitamin D helps reverse bone loss. Previous BMD
210 studies in Pakistani population revealed similar results regarding both genders
211 having a low BMI being significantly susceptible to having a low BMD as well
212 [26]. Nevertheless, a few variables affect BMD negatively too and are vital to be
213 understood in order to establish a proactive approach against osteoporosis. These
214 include consumption of alcohol and carbonated drinks, specifically cola. At the
215 same time, individuals who are habitual smokers are susceptible to low BMD
216 levels [25].

217 The WHO's Global Action Plan on Physical Activity 2018-2030 [27] aims at
218 reducing physical inactivity by 10% by the year 2025 and 15% by 2030. As a
219 member state of the World Health Assembly (WHA), the 4 objectives and 20
220 policy actions highlighted in the plan should be put into practice in Pakistan. Key
221 among these are governmental support and enabling policies to promote physical
222 activity with national targets that should be met. Physical education should be

223 mandatory in the curriculum of schools and colleges at all levels for both genders
224 in order to form a healthy habit from a very early age. Governmental programmes
225 to create public parks and recreation spaces for the purpose of promoting PA
226 should be enacted. Population-based interventions should be carried out, such as
227 national awareness campaigns, aimed at educating the public about the
228 importance of PA and positive lifestyle modifications to reduce the risk of non-
229 communicable diseases. These measures will save the national exchequer billions
230 by decreasing the disease burden of preventable problems.

231 The current study has limitations as the sample size was too small to validate the
232 results for generalisation. Further studies are recommended with a larger sample,
233 concurrently using a more extensive dietary assessment regarding intake levels
234 of calcium and vitamin D.

235

236 **Conclusion**

237 High PEFR and HGS proved to be beneficial for improving the quality of BMD,
238 thereby reducing fracture risk. PA had a role in enhancing PEFR and HGS in
239 young adults. A protective effect of increasing BMI on the BMD of healthy adults
240 was established.

241

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245

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Table 1: Comparison of variables based on gender

	Males (n=52) Median (IQR)	Females (n=50) Median (IQR)	p-value*
Age (years)	21.00 (1)	21.00 (1)	0.399
Height (m)	1.74 (0.09)	1.61 (0.08)	0.0001*
Weight (kg)	71.50 (13.8)	56.50 (13.8)	0.0001*
BMI (kg/m²)	23.95 (4.4)	21.70 (5.4)	0.002*
PEFR (L/min)	505.00 (100)	302.50 (100)	0.0001*
HGS (kg)	29.85 (13.1)	13.85 (5.3)	0.0001*
T score	-0.50 (1.4)	-1.30 (1.0)	0.0001*

346 BMI: Body mass index, PEFR: Peak expiratory flow rate, HGS: Handgrip
 347 strength. P-value calculated by Mann Whitney U Test, *p<0.05 = statistically
 348 significant.

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 352

Table 2: Spearman correlation between PEFR, HGS, BMD, BMI, PA.

	PEFR (L/m)	HGS (kg)	T score	BMI (kg/m²)	PA
PEFR (L/m) Correlation Coefficient Sig.	1.000 . .	0.659 0.001	0.417 0.001	0.351 0.001	0.426 0.001

HGS (kg) Correlation Coefficient Sig	0.659 0.001	1.000 .	0.364 0.001	0.383 0.001	0.364 0.001
T score Correlation Coefficient Sig	0.417 0.001	0.364 0.001	1.000 .	0.462 0.001	0.238 0.016
BMI (kg/m²) Correlation Coefficient Sig	0.351 0.001	0.383 0.001	0.462 0.001	1.000 .	0.151 0.130
PA Correlation Coefficient Sig	0.426 0.001	0.364 0.001	0.238 0.016	0.151 0.130	1.000 .

353 p-value <0.05 = statistically significant. PEFR: Peak expiratory flow rate, HGS:
354 Handgrip strength, BMI: Body mass index, PA: Physical activity.

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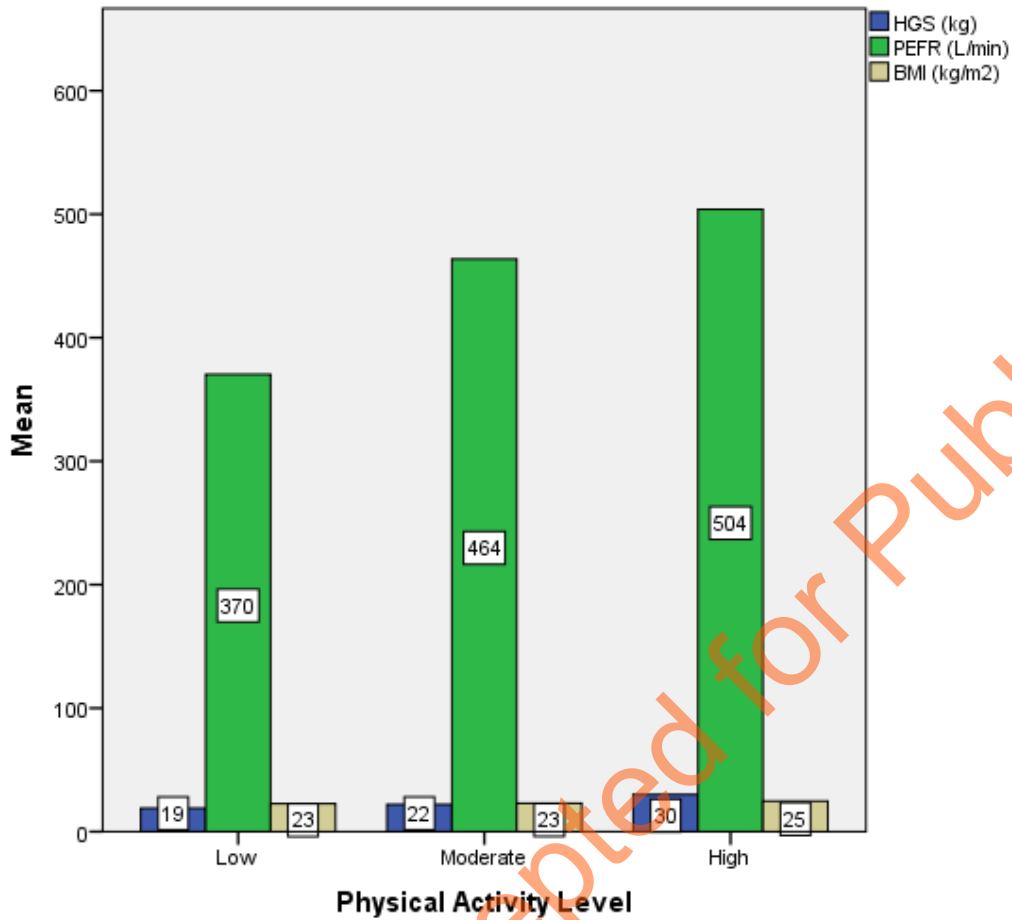
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364 **Figure:** Peak expiratory flow rate (PEFR), Handgrip strength (HGS) and body
365 mass index (BMI) values for young adults based on physical activity (PA)
366 expressed in categories.

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