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The prevalence of vitamin D deficiency between Saudis and non-Saudis in Al-Madinah
Al-Munawarah a cross-sectional study

Running headed: Vitamin D deficiency in Saudis and non-Saudis

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34 INTRODUCTION

35

36 Vitamin D is one of the vitamins that are soluble in fat. It is also classified as a
37 prohormone steroid, (1). Vitamin D has important functions in the endocrine, paracrine
38 and autocrine systems, as so, it is regarded as the “sunshine” vitamin (2,3). Naturally, only
39 two forms of vitamin D are available; vitamin D₂, also known as ergocalciferol, and
40 vitamin D₃ (cholecalciferol). Photosynthesis of vitamin D in the skin by the induction of
41 sunlight produces only cholecalciferol, while the dietary sources of vitamin D can provide
42 the vitamin in the two forms (3).

43

44 Globally, Vitamin D deficiency is among the major public health issues (4,5). The
45 deficiency in vitamin D in children leads to a disease called rickets in which bone tissue
46 fails to mineralize properly, causing skeletal deformities and brittle bones (1,6–8). While,
47 the deficiency in vitamin D in adults leads to weakness in the muscles and consequently,
48 increasing the risk of falls and fractures. In adults, vitamin D deficiency causes another
49 disease called osteomalacia, leading to the weakness of bones and exacerbate osteoporosis
50 (1,6).

51

52 Numerous data have been documented on the link between ethnicity, colour of skin and
53 vitamin D serum levels (9,10). In the United States, 90% or more of the non-Hispanic
54 African American were found to have vitamin D levels around 15 ng/mL, the mean
55 vitamin D level of Hispanics was 20 ng/mL, while that of the non-Hispanics white was 26
56 ng/mL, whereas those individuals residing the traditional counties in Central Africa were
57 showed to have mean plasma 25(OH) D levels of about 46 ng/mL (1,11).

58

59 In the Middle East and North Africa (MENA), especially the Kingdom of Saudi Arabia
60 (KSA), high prevalence of vitamin D deficiency has been reported despite the abundance
61 of sunshine (12,13). In 2015, a cross-sectional national multistage survey reported that the
62 prevalence of vitamin D deficiency was 40.6% in male Saudis and 62.65% in female
63 Saudis (14). Similar results were found in other, studies that were conducted in various
64 places in KSA. Two studies that were conducted in Riyadh indicated that vitamin D
65 deficiency among Saudi men was 87.8% (15), and; 78.1% in Saudi females and 72.4% in
66 Saudi males respectively (16). In Damam the prevalence of vitamin D deficiency was
67 found to be greater than 65% (17). Furthermore, Dabbour and colleagues (2016a study)
68 conducted in Makkah estimated that vitamin D levels were very low among the healthy
69 Saudis population (5.26 ± 2.59 ng/ml) (4).

70

71 To our knowledge, up to know no study has yet been conducted to compare the prevalence
72 of vitamin D deficiency between Saudis and non-Saudis living in the same area. This will
73 be the first study to document Vitamin D status in Al- Madinah Al-Munawwarah. Also,
74 this paper will examine predictors of such deficiency.

75

76

77 **Methods**

78

79 **Subjects and Study Design**

80

81 This was a cross-sectional study conducted in Al-Madinah Al-Munawwarah, KSA from
82 October 2017 to May 2018. This period of eight months was chosen to reduce seasonality
83 bias as June, July, August and September are the hottest summer months in KSA. The
84 ethical approval was obtained before the beginning of the study; in October 2017, from
85 Taibah University College of Dentistry Research Ethics Committee (TUCD-REC), which
86 is the ethical review board in Taibah University.

87 Non-diabetic male Saudis and non-Saudis nationalities age between 18 to 65 years old
88 attended Medical Unit, Taibah University were included in this study. Participants were
89 excluded from the study if they consume vitamin D supplementations, diagnosed with
90 renal, liver or cancer, thyroid or parathyroid diseases and receiving any drug interacting
91 with vitamin D such steroids, orlistat, cholestyramine, phenytoin and phenobarbital. Based
92 on the sample size calculation, a sample size of 65 encounters per study arm; 33 in group
93 one (Saudis) and 32 in group two (non-Saudis) would detect the degree of difference with
94 80% power at $\alpha = 0.05$.

95 Prior participants' enrollment commenced, demographic data, a detailed history of
96 diabetes (if present), socio-economic data, information about intake of vitamin D-
97 containing diets, exposure period to sunlight per day were collected. Completion and
98 return of written informed consent before participating in the study indicated voluntary
99 agreement to participate in this study.

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108 **Data Collection**

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110 **Anthropometric Data**

111 The weight was measured in kilograms and rounded to the nearest 100 grams. Standard
112 beam scale was used to measure the weight of participants. The participants weighed
113 wearing light clothes and barefoot

114 A calibrated height board was attached to the scale to measure the height in centimeters.
115 Weight in kilograms were divided by the square of height in meters to calculate the BMI.
116 The body mass index (BMI) was used to evaluate obesity. Participants with BMI between
117 25 and 29.9 kg/m² were considered overweight, participants with BMI between 30 and
118 39.9 kg/m² were considered obese participants. While, when BMI found to be > 40 kg/m²
119 then they were considered morbidly obese. Determination of the waist circumference was
120 done by measuring the broadest area between the edge of lower ribs and the iliac crest.

121

122 **Blood Sample Collection and Laboratory Analysis**

123

124 Five millilitres of blood were collected by trained technicians. This was done under the
125 supervision and guidance of the primary care physicians. Blood tubes were preserved in a
126 cooler or refrigerator immediately. The time of preservation was not less than 30 minutes
127 and did not exceed four hours before the technicians centrifuge them. The centrifugation
128 process was done for about half an hour at 3000 RPM at 4°C. Following that, the
129 technicians immediately separated the serum from the whole blood and freeze them at -
130 20°C. This was done at the biochemistry laboratory at the Medical Unit, Faculty of
131 Medicine, Taibah University, Al-Madinah.

132

133 The 25-hydroxyvitamin D levels were measured by ECLIA assay by Cobas machine e
134 411. The level were considered as deficient (< 20 ng/ml), inadequate (20 –30 ng/ml) and
135 adequate (30 ng/ml) as recommended by the American Endocrine Society Clinical
136 Practice Guidelines (2,18).

137

138 **Results:**

139 A total of sixty-five healthy participants were recruited in the study, subdivided into two
140 groups, 33 Saudis and 32 non-Saudis. The sample size was estimated to provide 80%

141 power to detect a difference in vitamin D deficiency between the 2 groups at a two-sided
142 0.05 significance level.

143 (25-hydroxyvitamin D) levels less than 20 ng/ml were considered as deficient, while
144 levels between 20–30 ng/ml were considered insufficient, and levels greater than 30 ng/ml
145 were considered adequate vitamin D.

146

147 **Table 1. Baseline Demographic and clinical characteristics of participants**

Indicators	Saudi (n= 33)	Non-Saudi (n= 32)	P-Value
Age (years)			
18-40	30 (91%)	6 (18.75%)	
41-65	3 (9%)	26 (81.25 %)	
Education level (postgraduates)	4 (12.1%)	30 (93.8%)	< 0.001
BMI (kg/m ²)	28.85 ±4.55	27.3± 3.27	
Marital status			
Married	25 (75.8%)	32 (100%)	< 0.003
Unmarried	8 (24.2%)	0 (0%)	
Smoking status (current smoker)	17 (51.5%)	1 (3.1%)	< 0.001
Occupation status (governmental employees)	31 (93.9%)	29 (90.6%)	0.6

148

149 The mean age of the Saudi group was 33.6 ± 7.2 years and the median and mode were
150 found to be 32 years, while the mean age of the non-Saudi group was 46.8 ± 8.1 years,
151 with median 45.5 and mode 43 years respectively.

152

153 *The marital status*

154 All the non-Saudi group were married (100%) versus only 75.8% (n = 25) of the Saudi
155 group were married. A significant difference was found between the two groups in their
156 marital status.

157

158 ***Smoking status***

159 The results of the present study showed that high percentage of the Saudi participants
160 were current and active smokers (n=17, 51.5%) compared to the non-Saudi participants
161 (n=1, 3.1%). Statistical analysis showed that there is a significant variation in the smoking
162 status between Saudi and non-Saudi participants ($p < 0.001$).

163 ***Education status***

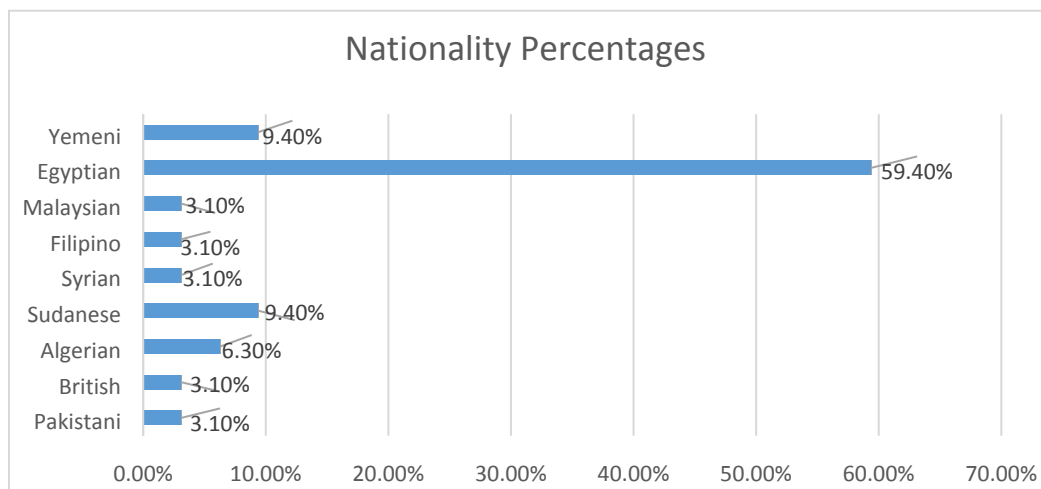
164 Majority of the non-Saudi participants were post graduates, (n = 30, 93.8%) while only
165 12.1% (n = 3) in the Saudi group was postgraduates. Therefore, a significant variation was
166 detected between the two groups in their education status ($p < 0.001$).

167

168 ***The occupation status***

169 The majority of both groups were governmental employees i.e. 93.9% (n = 31) Saudi and
170 90.6% (n = 29) non-Saudi, while only 6.2% (n = 2) of Saudi and non-Saudi participants
171 were students. Besides that, 3.1% (n = 1) of the non-Saudi group were without a job.
172 However, the occupation status did not differ significantly between the two groups ($p =$
173 0.6).

174 **Figure 1. Nationalities of the non-Saudi participants (n=32)**



175

176

177 Majority of the non-Saudi participants were Egyptians (n = 19, 59.40%), followed by the
178 Sudanese and Yemenis were 9.40% (n = 3) each, as shown in figure 1.

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183 **Table 2. Dietary habits for the Saudi and non-Saudi participants**

	Saudi (n=33)		Non-Saudi (n=32)		Total		<i>p</i> -value	
	No.	%	No.	%	No.	%		
Egg yolk	never	1	3.0%	1	3.1%	2	3.1%	0.9
	monthly	1	3.0%	1	3.1%	2	3.1%	
	1 time/week	9	27.3%	7	21.9%	16	24.6%	
	3 times/week	17	51.5%	17	53.1%	34	52.3%	
	Daily	5	15.2%	6	18.8%	11	16.9%	
Oily Fish	never	2	6.1%	6	18.8%	8	12.3%	0.004
	monthly	17	51.5%	4	12.5%	21	32.3%	
	1 time/week	11	33.3%	13	40.6%	24	36.9%	
	3 times/week	3	9.1%	9	28.1%	12	18.5%	
	never	2	6.1%	7	22.6%	9	14.1%	
Liver	monthly	23	69.7%	14	45.2%	37	57.8%	0.1
	1 time/week	8	24.2%	9	29.0%	17	26.6%	
	3 times/week	0	0.0%	1	3.2%	1	1.6%	
	never	1	3.0%	0	0.0%	1	1.5%	
	monthly	7	21.2%	1	3.1%	8	12.3%	
Red Meat	1 time/week	12	36.4%	7	21.9%	19	29.2%	0.04
	3 times/week	11	33.3%	20	62.5%	31	47.7%	

	Daily	2	6.1%	4	12.5%	6	9.2%	
	never	3	9.1%	7	21.9%	10	15.4%	
	monthly	11	33.3%	1	3.1%	12	18.5%	
Fortified food	1 time/week	6	18.2%	8	25.0%	14	21.5%	0.03
	3 times/week	4	12.1%	6	18.8%	10	15.4%	
	Daily	9	27.3%	10	31.2%	19	29.2%	

184

185 Results of this present study showed that egg-yolk consumption in both groups were
 186 almost the same (51.5% versus 53.1%, 3 times/week, 3% versus 3.1% monthly, 15.2%
 187 versus 18.8% daily).

188 While, the statistical analysis of oily-fish consumption showed that non-Saudi population
 189 consumed more significantly than Saudi population did. There was not much difference
 190 between the two groups regarding liver intake, but the ingestion of both red meat and
 191 fortified food were significantly lower among the Saudi population compared with the
 192 non-Saudi.

193

194 **Table 3. Duration and pattern of sun-light exposure**

	Saudi (n=33)		Non-Saudi (n=32)		Total		<i>p</i> -value	
	No.	%	No.	%	No.	%		
Duration Of Approximately Sunlight Exposure	15 min	17	51.5%	11	34.4%	28	43.1%	0.5
	30 min	9	27.3%	11	34.4%	20	30.8%	
	60 min	6	18.2%	8	25.0%	14	21.5%	
	More Than 1 hour	1	3.0%	2	6.2%	3	4.6%	
	Yes	1	3.0%	1	3.1%	2	3.1%	

195

196 In the Saudi nationality group, it was observed that 27.3% were exposed to sunlight 15
 197 minutes daily compared to 34.4% in the non-Saudi group, while 18.2% of the Saudi

198 nationality group were exposed to the sunlight 60 minutes daily versus 25% in the non-
199 Saudi group.

200 Noteworthy, the positive exposure to sunlight in this study was defined as; the direct
201 exposure (not indoors via windows) of at least some uncovered body parts, such as the
202 arms and some parts of the legs, during the time period between 10 AM and 3 PM for not
203 less than five minutes.

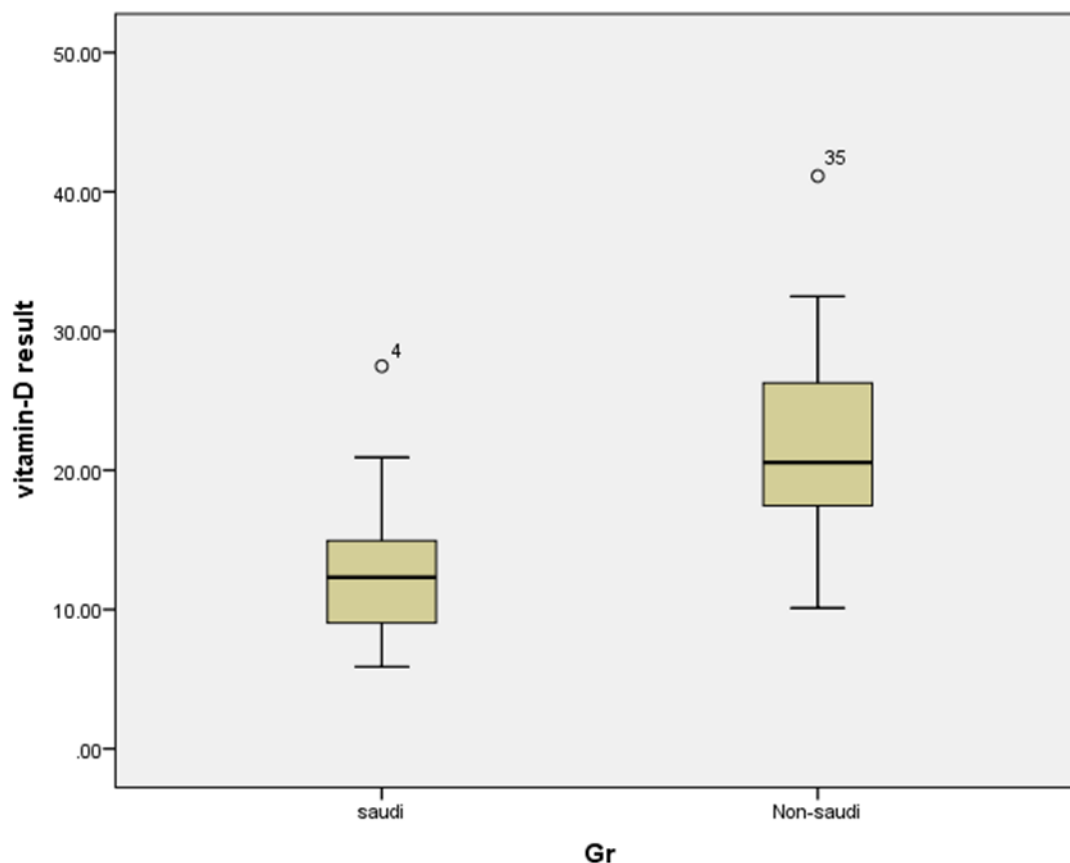
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205 The occupation status was found to be positively correlated with vitamin D deficiency,
206 while the consumption of liver was observed to be negatively correlated with vitamin D
207 deficiency among the Saudi population, as revealed by logistic regression.

208

209 *Vitamin D levels*

210 **Figure 2. Box and whisker plots showing Vitamin D levels**



211

212 Significant percentage of the participants in the Saudi group (n=30, 91%) suffered from
213 deficiency in vitamin D levels [25 (OH) D < 20 ng/ml] 12.57 ± 4.82 (mean \pm SD),
214 compared to only 47% (n=15) in the non-Saudi group [21.56 ± 6.82 (mean \pm SD)].

215 Vitamin D deficiency was found to be significantly higher in the Saudi group than the
216 non-Saudi group with $P = 0.001$

217

218

219 **Discussion**

220 **Anthropometric Features and Socio-demographic data**

221 *Age of the participants*

222 In this study, the ages of the Saudis were 33.63 ± 7.17 years (mean \pm SD) with median and
223 mode of 32 years. Meanwhile, the ages of the non-Saudis were 46.78 ± 8.1 years (mean \pm
224 SD), with median 45.5 and mode 43 years. Noticeably, the non-Saudis were in higher age
225 group than the Saudi population in phase one of this study. That seemed reasonable
226 because the majority of the non-Saudis (93.85%, $n = 30$) had postgraduate degrees; this
227 process took a long time – around six to eight years after graduation. In addition, the main
228 reason for the presence of the non-Saudis in KSA is for work, which they would not have
229 been eligible for without at least three years of experience. All these justified why the non-
230 Saudis were found to be in a higher age group than the Saudis.

231 On the contrary, the majority of the Saudi participants were without postgraduate degrees.
232 All of them were local and working in their own country, which made it easier for them to
233 be offered jobs in a short time just after graduation. In addition, most of the Saudi
234 participants were administrative employees, for which postgraduate studies were not a
235 must, so opportunities for governmental jobs were easy in their country. So, the ages of
236 the Saudis were less than that of the non-Saudis.

237 This finding was consistent with that of Naugler et al. (2013) who observed that vitamin D
238 deficiency was less prevalent in the older age group (19). However more than one study
239 has demonstrated the high prevalence of vitamin D deficiency in different age groups
240 among the Saudi population (20–22). The documented high prevalence of vitamin D
241 deficiency among the Saudi population with wide age groups was mainly the result of the
242 recent huge socioeconomic change that led to urbanization of lifestyles and westernization
243 of dietary habits of most of the Saudi population.

244 *BMI of participants*

245 The BMI of the Saudis was 28.85 ± 4.55 (mean \pm SD), while that of the non-Saudis was
246 27.3 ± 3.27 (mean \pm SD). No statistical difference was detected between the two groups.
247 The present study has shown that the Saudi population was characterized by higher BMI.
248 Alqarni (2016) showed in a qualitative review an increasing pattern in the overweight and
249 obesity prevalence in Saudi Arabia, which was expected to further increase in the future.
250 Unbalanced diets – high consumption of carbohydrates in the form of rice, bread, pizza
251 and dates with low vegetables and fruit intake –, in addition to the lack of physical

252 exercise, contributed to the higher BMI and overweight status of the Saudi population
253 (23–25).

254 In agreement with this finding, AlHumaidi, Adha, and Dewish in 2013 reported that obese
255 Saudis with BMI 32.6 ± 6.7 (mean + SD) were found to have very low vitamin D levels
256 (11.1 ± 5.9 ng/mL). Similarly, Al-Daghri et al. (2016) showed that overweight participants
257 with BMI 28.0 ± 6.6 suffered from either vitamin D insufficiency (51.5%) or deficiency
258 (29.9%) (26,27).

259 Also, the Saudi group had a greater deficiency in vitamin D than the non-Saudis. High
260 BMI or obesity and vitamin D deficiency in this research was consistent with other
261 researches that have evaluated the link between obesity and vitamin D status in different
262 populations.

263 Bettencourt et al. (2018) performed a research in Portugal to assess the vitamin D status
264 and its related factors. Their results showed that vitamin D levels were negatively
265 correlated with obesity (i.e. BMI). This was mainly because of the sequestration of
266 vitamin D in the body fat compartment, which could lead to the reduction of the
267 bioavailability of vitamin D that was either photosynthesized through the skin by direct
268 exposure to sunlight or from dietary sources (28). Other researchers have revealed the
269 correlation of obesity with hypovitaminosis D with enhanced catabolism of the said
270 vitamin in fat tissues and their conversion into inactive metabolites (29).

271 Kaddam et al. in 2017 conducted a cross-sectional study to assess the prevalence of the
272 aforementioned condition and its associated factors among students and employees in
273 three regions of KSA. The authors observed that obesity was associated with vitamin D
274 deficiency (30). Tønnesen and colleagues in (2016) observed that obese participants had
275 the highest relative risk for vitamin D deficiency in their study (31). Vimalleswaran et al
276 (2014) showed that for every 1 kg/m^2 higher in BMI, vitamin D deficiency was increased
277 by 1.15% (32). Furthermore, Barchetta and colleagues in 2011 found that higher vitamin
278 D deficiency was accompanied by increasing BMI and waist circumference (33).
279 Chakhtoura and colleagues (2018) documented in their review on the prevalence of
280 vitamin D deficiency in the Middle East and North Africa that higher BMI was one of the
281 predictors of reduced serum vitamin D levels (13).

282 Therefore, it is important to consider much higher doses (2.5 times) of vitamin D
283 supplementation in the obese vitamin D-deficient individuals due to their reduced
284 bioavailability of vitamin D and/or the enhanced catabolism of vitamin D, as well as their
285 conversion into inactive metabolites (1,18).

286 ***Marital status***

287 The results of phase one in the present study have shown that all of the non-Saudi group
288 and most of (75.8%, $n = 25$) the Saudi group were married ($P = 0.003$). As described
289 before, the non-Saudi participants were of the higher age group than the Saudi

290 participants. So, not surprisingly, the percentage of married non-Saudis was higher than
291 that of the Saudis.

292 This finding could point towards the impact of marital status and social stability on the
293 type of food, which might be healthier and containing a higher content of vitamin D. On
294 the contrary, the unmarried individuals usually consume more fast and junk food that
295 contained very small amounts of the vitamin.

296 *Education status*

297 The results have showed that the education status of the non-Saudi group was significantly
298 higher than that of the Saudi group, where 93.85% (n = 30) of the non-Saudis had
299 postgraduate degrees as compared to only 12.1% (n = 4) in the Saudi group ($P = 0.001$).
300 This is because the administrative professions were restricted to the local residents (Saudi
301 nationality) in KSA while the academic professions were mostly occupied by non-Saudis.

302 It is elicited in this present study that participants with higher education levels had higher
303 vitamin D levels. The same observation was seen in a number of previous studies. In the
304 Emirates, Bani-Issa and colleagues performed a quantitative cross-sectional study in 2017
305 to assess vitamin D deficiency and the factors associated with this condition. The authors
306 have concluded that the less-educated, employed Emirati participants had a significantly
307 higher percentage of vitamin D deficiency than the higher-educated participants (12).

308 In Alberta, Canada, an ecological study has been conducted by Naugler et al. (2013) to
309 evaluate the association between vitamin D deficiency and a number of socio-
310 demographic factors. The authors have observed that vitamin D level was associated with
311 education status, and they attributed their finding to the increased consumption of vitamin
312 D supplements by the higher-educated individuals (19).

313 Moreover, Ardawi and colleagues in 2012 studied the high prevalence of vitamin D
314 deficiency among the Saudi males. The authors attributed the significant deficiency in
315 vitamin D among Saudis to a number of factors, among which was the lack of education
316 (34). Similarly, a Finnish study performed by Jääskeläinen et al. (2012), has documented
317 the association between vitamin D levels with the education status (35). Furthermore, in
318 China, Song et al. (2013), who studied the prevalence of vitamin D deficiency in pregnant
319 women, have observed that the long duration of sunlight exposure, and the subsequently
320 alleviation of vitamin D deficiency was in favour of the higher-educated women (36).

321 Notably, high education level rendered the individuals to be more knowledgeable and
322 have greater awareness of at least the basic role of vitamin D in body-functioning and the
323 negative impact of its deficiency.

324 *Smoking status*

325 There was a significant variation in the smoking status between the two groups with $P =$
326 0.001, whereby 17 out of 33 (51.5%) from the Saudi group were current smokers, while
327 only 1 out of 32 (3.1%) from the non-Saudi was a current smoker.

328 According to Al-Nozha and colleagues, smoking rates were high among Saudis who were
329 living in urban areas (Al-Nozha et al., 2009). This was consistent with the finding of this
330 present study, as Al-Madinah is one of the urban cities in KSA.

331 (37).

332 Smoking status of the Saudi population could help explain the deficiency in vitamin D
333 levels detected in this population. Chakhtoura and colleagues, in their systematic review in
334 2018, have reviewed trials that studied the vitamin D deficiency in the MENA region.
335 They indicated that smoking status was one of the possible consistent predictors of
336 vitamin D deficiency (13).

337 In addition, several studies that have been conducted in different countries and used
338 different methodologies indicated similar findings regarding the link between smoking
339 status and vitamin D levels (38–40).

340 Hoteit et al. (2014) who studied the prevalence of vitamin D deficiency in 9147 Lebanese
341 subjects reported smoking status was a robust predictor of vitamin D deficiency (38).
342 Jiang et al. (2016), via a cross-sectional study that was performed in China, observed that
343 the current smokers were associated with more severe vitamin D deficiency than non-
344 smokers (39). In addition, Tønnesen et al. (2016) considered smoking as one of the
345 modifiable factors for vitamin D deficiency as they showed that smoker participants were
346 associated with a higher relative risk of vitamin D deficiency than non-smokers (31).

347 Furthermore, in 2014, Mulligan and colleagues have assessed the impact of current
348 smoking on vitamin D-serum levels. Their results showed that smoking caused a
349 significant reduction in vitamin D serum levels (40). Ardawi and colleagues (2012)
350 reported a high prevalence of vitamin D deficiency (87.8%) among Saudis living in
351 Jeddah. The authors have shown that smoking was one of the attributable factors and was
352 associated with the low vitamin D status found in the Saudi population (34). Notably, Brot
353 and colleagues (1999) have conducted a cross-sectional study to investigate the effects of
354 smoking on the levels of vitamin D and calcium metabolism. The authors have observed
355 that smoking adversely affected calcium and vitamin D metabolism, mainly by depressing
356 vitamin D-parathyroid hormone pathway (41).

357 Not surprisingly, that the results of all these studies were in agreement with this present
358 finding regarding the link between smoking and low vitamin D levels, as it has been
359 documented that smoking negatively affected various body functions, harmed organs, and
360 had a lot of general adverse consequences throughout all the body (42).

361 ***Occupation status***

362 There was no significant difference in the occupation status between the two groups ($P =$
363 0.60), as 93.9% ($n = 31$) from the Saudi group and 90.6% ($n = 29$) from the non-Saudi
364 group were governmental employees. The participants in the two groups were employees
365 in Taibah Univeristy, which is a governmental university in Al-Madinah Al-Munawarah.

366 Being office-bound employees had a negative impact on their lifestyle, rendering them to
367 have more sedentary lifestyles and decrease their exposure to sunlight – which is the
368 major source for the photosynthesis of vitamin D. Tønnesen et al. (2016) have
369 documented that the more the time spent in physical exercise, the more reduction in the
370 relative risk of vitamin D deficiency. The authors have also added that fast food
371 consumption was associated with a high relative risk of vitamin D deficiency (31).
372 Furthermore, Dabbour and colleagues performed a research in Makkah region in Saudi
373 Arabia in 2016 to evaluate the association of vitamin D deficiency with T2DM. The
374 authors have observed that vitamin D deficiency – which is highly prevalent in KSA – was
375 associated with the lack of physical activity and decreased exposure to sunlight. They
376 further added that physical activity could aid in the maintenance of vitamin D levels,
377 hence concluding that duration of physical activity was positively correlated with serum
378 vitamin D levels (4).

379 **Dietary habits of participants**

380 Detection of dietary habits of the Saudi population could be considered as a very
381 important point, because there is still scarcity in the information on vitamin D status and
382 its association with the Saudi population's dietary habits. Such information can help
383 ministries in charge (e.g. ministry of health, ministry of education, and ministry of media)
384 guide the Saudi population to overcome this medical problem (i.e. hypovitaminosis D),
385 either by guiding the general population to increase exposure to sunlight or by focusing on
386 the fortification of food products (43). The answers to these two points have been
387 mentioned by this present study. Such valuable information could significantly help
388 develop a guideline for the Saudi population.

389 The dietary habits between the two groups were found to be as follows:

390 **Oily fish:** the consumption of oily-fish was significantly higher in the non-Saudi group
391 than in the Saudi group ($P = 0.004$).

392 Salmon reportedly contains 988 IU of vitamin D per 100 grams; this amount accounts for
393 about 247% of the Reference Daily Intake (RDI) of vitamin D. Herring and sardines: the
394 fresh Atlantic herring contains around 1628 IU of vitamin D per 100-grams; this amount
395 accounts for around 4 times the RDI. Pickled herring contains about 680 IU/100 g that
396 accounts for 170% of RDI. Sardines provide 272 IU per serving. Mackerel provides 360
397 IU/serving. In addition, cod liver oil is a good source of our concerned vitamin as it
398 contains around 450 IU/teaspoon. Furthermore, canned tuna provides up to 236 IU/100 g
399 (44).

400 Oily fish contains omega-3 fatty acids which have a well-known protective effect against
401 dyslipidemia, hypertension, stroke, and cardiovascular diseases, in addition to the fair
402 contents of protein, vitamins, and other essential nutrients for good health (Burger et al.,
403 2014). The consumption of oily fish by the Saudi community remained poor. This could
404 have been due to cultural and traditional reasons as the main food types in Saudi Arabia
405 are mainly rich in carbohydrates and fibers like rice, potatoes, bread, and dates. In

406 addition, of the diet types and habits have increasingly westernized in the last few years
407 due to the introduction of multi-national restaurants in the Saudi market. Furthermore, the
408 lack of health and nutritional education played a crucial role in the dietary habits of the
409 Saudi population. These observations were consistent with those of Burger et al. (2014)
410 who have studied the behaviour and rates of fish consumption among the natives and non-
411 natives living in KSA. The authors have shown that the percentage of the male Saudis
412 who consumed fish was 3.7% and that of the male non-Saudis was 6.6% (45). Kaddam et
413 al. in 2017 conducted a cross-sectional study to assess the occurrence of hypovitaminosis
414 D among students and employees in three regions of KSA and its causative factors. The
415 authors observed that lack of omega-3 in the diet of the students was associated with
416 vitamin D deficiency (30).

417 **Red meat:** the non-Saudi group consumed red meat significantly more than the Saudi
418 group ($P = 0.04$), as 62.5% ($n = 20$) of the non- Saudis consumed red meat three times
419 weekly, and 12.5% ($n = 4$) ingested red meat on a daily basis, as compared to 33.3% ($n =$
420 11) and 6.1% ($n = 2$) of the Saudi group who consumed red meat in the same pattern.

421 A study has documented the levels of vitamin D₃ in types of meat, which were found to be
422 9.0 µg/kg for beef, 1.0–23.0 µg/kg for pork, 1.0–61.0 µg/kg for lamb, 0.0–50.0 µg/kg for
423 veal, 0.0–14.0 µg/kg for poultry, and 0.0–23.0 µg/kg for various meat product (46).

424 As mentioned before, the inherited traditional dietary habits in the Saudi community led
425 them to prefer foods that were rich in carbohydrates and fibre, and low in fats and
426 cholesterol. Moreover, the socioeconomic conditions, which have changed in the last
427 decade in Saudi Arabia, caused the migration of significantly large proportions of the
428 Saudi population from the rural areas to settle in the urban and large cities. It has been
429 forecasted by the Ministry of Municipal and Rural Affairs (MMRA) that by the year 2025,
430 88% of KSA residents will settle in the urban areas (47). Hence, the modification and
431 urbanization of their lifestyles have led to the alarming increase in the consumption of fast
432 and junk foods such as pizzas, pastas, and sandwiches. This explained the Saudis' low
433 consumption of red meat.

434 This present study's finding was consistent with a nationally representative survey done in
435 2016 by Moradi-Lakeh and colleagues to evaluate the dietary habits in KSA. The authors
436 have discovered that the type of food least consumed by the Saudi population was red
437 meat; 4.8 ± 0.2 g (mean \pm SE). They added that only a small proportion of the Saudis met
438 the dietary recommendations (48).

439 **Fortified food:** Although, fortified food may be considered as the ideal and the easiest
440 alternative to sunlight exposure in the Saudi lifestyle to guard against vitamin D
441 deficiency, yet the consumption of fortified food was significantly higher in the non-Saudi
442 group than in the Saudi group ($P = 0.03$).

443 The known sources of fortified food are cow's milk which usually contains around 130 IU
444 vitamin D/cup (237 ml), soy milk which can contain up to 119 IU vitamin D/cup, orange
445 juice which can also be fortified with vitamin D (usually 142 IU/cup), and cereal and

446 oatmeal which can be fortified with about 154 IU vitamin D per serving (49). This finding
447 is consistent with two recent studies which were conducted in Saudi Arabia. Al-Daghri
448 (2015) has studied the correlation between vitamin D levels and the consumption of dairy
449 and fortified products in Saudi Arabia. The author has observed the poor consumption of
450 fortified and dairy products in the overall Saudi population, apart from showed that the
451 vitamin D status was significantly associated with the consumption of dairy and fortified
452 products (20). In addition, in 2012, Ardawi and colleagues considered poor consumption
453 of dietary and fortified products as one of the contributory factors to vitamin D deficiency
454 in the Saudi population (34).

455 **Egg-yolk:** in contrast, there was no significant difference between the two groups in the
456 consumption of egg-yolk ($P = 0.9$). Only a small proportion of the two groups ate egg-
457 yolk as per the dietary recommendations. This could be justified by the dramatic
458 socioeconomic change and hence, urbanization of lifestyle and westernization of food
459 habits which pointed towards the large consumption of fast-food. Egg-yolk in general is
460 rich in fats, vitamins, and minerals. Egg-yolk contains around 39 IU of vitamin D, while
461 egg-yolk which comes from chickens that spent more time under the sun produced 3 to 4
462 times higher vitamin D. Chickens fed with vitamin D-enriched feed produced egg-yolks
463 with around 6000 IU vitamin D (49).

464 **Liver consumption** did not vary significantly between the two groups ($P = 0.1$). Even
465 though liver is considered as one of the rich animal sources of vitamin D that reaches as
466 high as 140 $\mu\text{g}/\text{kg}$, the consumption of liver was quite poor in both groups (46).

467 **The duration of sunlight exposure** did not significantly vary between the two groups (P
468 $= 0.5$). As observed, both groups were poorly exposed to sunlight (18.2% from the Saudi
469 group and 25% from the non-Saudi group were exposed to sunlight for one hour daily).

470 It was expected that 80% of the daily requirement of vitamin D in the majority of
471 individuals would be generated through sunlight exposure, as the major source of vitamin
472 D was photosynthesis. However, this was not the reality nowadays, presumably due to the
473 fear of erythema and some types of skin cancer, or that the increasing use of technology in
474 the 21st century kept most of the people spending most of their time indoors (1). In Saudi
475 Arabia, apart from the abovementioned possible causes of the avoidance of exposure to
476 sunlight, this could be attributed to the extreme hot temperature in KSA due to its
477 geographical location, abundant use of cars, and the presence of more highways.

478 Consistently with our results, Ardawi and colleagues in 2012 assessed the vitamin D
479 deficiency prevalence among healthy Saudi males. They have observed the high and
480 significant prevalence of vitamin D deficiency in this population was because of a number
481 of factors, among which included poor sunlight exposure (34). Almeahdi et al. (2016)
482 studied the vitamin D levels among children in Jeddah, KSA. Their result – poor sunlight
483 exposure in Saudi Arabia – was consistent with the finding of this present study (50). The
484 discrepancy between both studies was in the study population, where this present research
485 focused on adults and not children.

486 **Vitamin D levels among Saudi and non-Saudi populations**

487 In phase one, 30 out of 33 participants in the Saudi group (91%) were vitamin D-deficient
488 (calcidiol levels < 20 ng/ml) as compared to 15 out of 32 participants (47%) in the non-
489 Saudi population. Subsequently, vitamin D deficiency was found to be significantly higher
490 in the Saudis (12.57 ± 4.82) (mean \pm SD) than the non-Saudis (21.56 ± 6.82) (mean \pm SD),
491 with $P = 0.001$.

492 In terms of the socio-demographic data, occupation status was positively correlated with
493 the vitamin D deficiency in the Saudi group and was shown to be the only predictor of
494 vitamin D deficiency, as revealed by logistic regression and expressed by exponential
495 beta. This finding could be explained by the fact that the majority of the Saudi population
496 in this present study were governmental employees, which rendered them more office-
497 bound, and consequently reducing their duration and the time of sunlight exposure (51).
498 Among the dietary habits, the logistic regression analysis revealed that the consumption of
499 liver was negatively correlated with vitamin D deficiency and was considered as a
500 predictor of vitamin D deficiency.

501 Previously, there was a controversy regarding the exact level of vitamin D which could be
502 considered as deficiency or even insufficiency. In the renal and extra-renal tissues, the
503 enzyme 1 α -hydroxylase (CYP27B1) targets its substrate calcidiol and converts it into the
504 active form, calcitriol. It has been documented that not more than 50% of maximal 25-
505 (OH) D-1 α -hydroxylase activity (Km) was reached when 25-(OH) D level reached 40
506 ng/mL (100 nmol/L), which in turn relied on having adequate amounts of vitamin D (18).
507 On the kinetics of 25-(OH) D-1 α -hydroxylase, the most recent studies were consistent
508 with the USA Endocrine Society's recommendations that the targeted level for optimal
509 vitamin D effects was 30–50 ng/mL (75–125 nmol/L), preferably achieving the level of
510 40–60 ng/mL (100–150 nmol/L). Meanwhile, vitamin D deficiency was denoted by serum
511 vitamin D levels of < 20 ng/mL (< 50 nmol/L), and insufficient or suboptimal status 20–
512 30 ng/mL (50–75 nmol/L) (18,52,53).

513 Notably, 13 out of 33 Saudi non-diabetics were overweight (BMI > 25), and 5 out of 33
514 were obese (BMI > 30). So, it was observed from the finding of present study that both
515 conditions – overweight and vitamin D deficiency – were higher in the Saudi population.
516 Furthermore, it is well known that T2DM became an increasingly widespread chronic
517 disease in the Saudi community (as mentioned in the Introduction Chapter). Thus,
518 overweight or obesity and vitamin D deficiency may be considered indicators or
519 predictors for the development of T2DM among the Saudi population. Likewise, Bani-Issa
520 and colleagues (2017) have observed obesity and T2DM to be bivariate correlators (which
521 were independent of each other) with vitamin D deficiency in Emiratis (12). In addition,
522 Sadiya et al. (2014) documented in their cross-sectional study which was performed in
523 UAE the coexistence of these three cofactors – obesity, vitamin D deficiency and T2DM
524 (54). Therefore, the present study recommends early screening and detection of T2DM in
525 the presence of obesity and vitamin D deficiency, with rapid and aggressive corrections of
526 these two predisposing factors.

527 As long as vitamin D deficiency is highly prevalent in the Saudi population relative to the
528 non-Saudis, there might be some genetic factors playing a role in this scenario. Al Kadi
529 and Sonbol studied the high prevalence of vitamin D deficiency among Saudi population
530 in 2015. They have stated that the increased rate of consanguineous marriages in the Saudi
531 population have led to some types of genetic mutations, such as calcium-sensing receptor
532 gene A986S polymorphisms (55). In addition, Sonbol and colleagues have reported the
533 association between low levels of vitamin D and the ‘‘S’’ of the calcium-sensing receptor
534 gene (56).

535 Furthermore, in 2016, Nayak and Ramnanansingh in their study which investigated the
536 deficiency of vitamin D among the Trinidadian population have attributed this deficiency
537 to VDR genetic polymorphism. This polymorphism might lead to the dimensional
538 conformation of the vitamin D receptors which showed variations in their affinities
539 towards vitamin D (57). This hypothesis gave rise to the need for other studies to confirm
540 this hypothesis. A possible explanation for the rampant hypovitaminosis D among Saudis
541 could have been due to the dark skin, which could reduce the Ultraviolet B (UVB) rays’
542 penetration into the skin, thus minimizing the synthesis of vitamin D (58).

543 However, all the various factors that have been discussed previously in this study or in
544 other studies (poor consumption of vitamin D-containing food, skin pigmentation, sunlight
545 exposure, obesity, smoking, interacting drugs, etc.) which are implicated in the incidence
546 of vitamin D deficiency could lead to a decrease in the synthesis and bioavailability, and
547 elevations in catabolism and urinary excretion of vitamin D (1,59).

548 In addition, the cultural clothes used by the male Saudi population which covered the
549 whole body (even their heads and necks) reduced their body surface exposure to sunlight.
550 Consequently, their cutaneous vitamin D synthesis was decreased.

551 The findings explored in this current study were consistent with one systematic review and
552 six studies conducted in different regions in Saudi Arabia. However, to our best
553 knowledge until now our study was the first to compare the prevalence of vitamin D
554 deficiency between the Saudi population and participants from other nationalities living in
555 the same environment. In addition, it was the first to assess the prevalence of vitamin D
556 deficiency in Al-Madinah Al-Munawarah region.

557 The systematic review and a meta-analysis was performed by Al-Dagrhi and published in
558 2018. The author has reviewed the literature for the most recent epidemiological clinical
559 trials that studied the prevalence of hypovitaminosis D in the Saudi population from 2011
560 till 2016. He included 13 local trials (performed in KSA), in which 24 399 healthy Saudi
561 participants were included. This systematic review-cum-meta-analysis has shown that
562 81.0% of the Saudi population were deficient in this micronutrient, and that the deficiency
563 in vitamin D in the Saudi population was accompanied with several diseases, among
564 which were asthma, cancer, cardiac diseases, chronic kidney disease, obesity, metabolic
565 syndrome, type 1 diabetes, type 2 diabetes, and skeletal effects. The author has concluded

566 in line with our results in which there were widespread hypovitaminosis D that was
567 associated with insulin resistance and related comorbidities in the Saudi populations (43).

568 Mogahed (2018) has conducted a study in Riyadh city among 100 Saudi patients to detect
569 their vitamin D status. The results showed that 80% of the participants suffered from
570 vitamin D deficiency (< 20 ng/ml). The author has concluded that prevalence of vitamin D
571 deficiency in the Saudi population was very high (60).

572 Dabbour et al. performed a case-control study on 200 participants in Makkah Region in
573 Saudi Arabia in 2016 to evaluate the levels of vitamin D and to assess the associated
574 factors in healthy and diabetic Saudi participants. This study has concluded that there was
575 a very significant deficiency in vitamin D among the healthy and diabetic Saudi
576 populations (4).

577 However, this present trial differed from Dabbour et al. (2016) and Mogahed (2018) in the
578 checking and comparison of vitamin D levels among Saudi and non-Saudi population, and
579 in the evaluation of the dietary habits of the participants in an attempt to assess their
580 correlations with vitamin D deficiency.

581 A retrospective study has been conducted by Alfawaz et al. in Riyadh city to determine
582 the prevalence of vitamin D deficiency and its association with cardiovascular disease and
583 other factors in the Saudi population. Their results have showed that the overall
584 prevalence of vitamin D deficiency in their study was 78.1% in Saudi females and 72.4%
585 in Saudi males. Moreover, the prevalence of vitamin D deficiency was associated with
586 age, and obesity, and positively correlated with calcium, albumin, and phosphorus. It was
587 negatively correlated with alkaline phosphatase and PTH levels (16). This present study
588 showed a higher prevalence of vitamin D deficiency among the Saudi population in Al-
589 Madinah Al-Munawarah (91%) than what was reported by Alfawaz et al. (2014) in
590 Riyadh (72.4.1% in males). Furthermore, this present study has not only detected the
591 prevalence of vitamin D deficiency among the Saudi population, but also compared the
592 high prevalence with those of the non-Saudi population living in the same environment via
593 a prospective study design.

594 A multi-centered case-control study has been done by Alhumaidi and colleagues in the
595 Southern Region of Saudi Arabia (mainly Khamis Mushyt and Abha) in 2013. Their aim
596 was to assess the vitamin D status among the Saudi population. They have included 172
597 diabetic Saudi patients and 173 non-diabetic Saudi participants in their trial. They have
598 showed a very significant vitamin D deficiency in the healthy and diabetic Saudi
599 population (98.5%). The authors have suggested the need for larger studies for better
600 detection of vitamin D deficiency in the Saudi population as a whole, especially among
601 the diabetic patients (26). The lifestyle, socio-demographic data and dietary patterns of
602 their study population have not been studied by Alhumaidi and colleagues, in contrast to
603 this present study. In addition, participants with thyroid diseases were included in their
604 study; 23.5% were found to have euthyroid multinodular goiter and 33.8% were with
605 hypothyroidism. These could have influenced their results (61). Furthermore, 17.4% of the

606 non-diabetic Saudi participants were lost to follow-up. In the contrary, this present study
607 has excluded participants with any thyroid and parathyroid disease and the compliance
608 with the study protocol in this particular phase was 100%.

609 Also, a study has been done in Saudi Arabia in 2012 by Ardawi et al. This study design
610 was cross-sectional and was aimed to assess the prevalence of deficiency in vitamin D in
611 Saudi men in Jeddah. The authors have concluded that the deficiency in vitamin D was
612 highly prevalent (87.8%) among Saudi men living in Jeddah (62). Although this present
613 study was similar to that of Ardawi et al. (2012) in detecting the prevalence of vitamin D
614 deficiency in only males and both have reported the high prevalence of vitamin D
615 deficiency in Saudi men. Ardawi and colleagues neither compared this high prevalence in
616 Saudi men with other nationalities residing in the same environment, nor evaluated the
617 association of socio-demographic factors, sunlight exposure, and dietary habits with the
618 high prevalence of vitamin D deficiency in the Saudi and non-Saudi population.

619

620 In Norway, a study has been performed in 2013 to compare the serum vitamin D levels
621 among different populations living in Norway. The results of this study have revealed that
622 immigrants from South Asia, the Middle East, and Africa were found to be more vitamin
623 D-deficient (25-(OH) D < 20 ng/mL) than East Asians (63). Notably, the authors have
624 recommended the routine measurement of vitamin D levels and early detection of vitamin
625 D deficiency in these populations (Middle East and Africa).

626

627 **Conclusion**

628 Vitamin D deficiency was significantly higher in the Saudi population than the non-Saudis
629 population. The occupation status was found to be the only factor positively correlated
630 with vitamin D deficiency.

631 Early screening for vitamin D serum-level is recommended for the Saudi population and
632 rapid correction of vitamin D deficiency with vitamin D supplements should be
633 considered.

634

635 **Limitations**

636 Multi-centred study might have provided more generalizable results.

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