1	Diabetes in the Shadow of Conflict: Understanding and Addressing the Crisis
2	in Gaza
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10	Abstract
11	The conflict in Gaza has inflicted severe damage on an already fragile healthcare system, leading
12	to significant challenges in providing adequate care for individuals, particularly those with
13	chronic diseases like diabetes. The conflict has caused immense strain on healthcare facilities,
14	resulting in shortages of essential supplies and medications, including insulin, and disrupting the
15	provision of proper nutrition and medical care. This situation has led to an increase in
16	communicable diseases and a significant impact on individuals with chronic conditions, such as
17	diabetes.
18	Patients with diabetes in Gaza face numerous challenges, including the inability to access
19	necessary medications, adhere to special diets, and receive proper surveillance and care for their
20	condition. The lack of appropriate management can lead to a deterioration in the quality of life

21 and an acceleration of the disease processes, increasing the risk of complications such as

22 cardiovascular disease, nephropathy, retinopathy, and neuropathy.

23 To address these challenges, short-term strategies must focus on promptly identifying diabetic 24 patients and ensuring their continuous access to essential medications, particularly insulin. Long-25 term strategies should include strengthening health information systems, ensuring continuous 26 access to medication and adequate supplies, and providing comprehensive healthcare services targeting diabetes within primary health centers. Additionally, training healthcare professionals 27 28 in diabetes care, developing national emergency response plans, and fostering effective 29 communication between governments are crucial steps in managing diabetes during crises like 30 the conflict in Gaza.

31 Key words:

32 Gaza conflict, Diabetic patients, Diabetes, Challenges, Recommendations

33 Introduction

Diabetes Mellitus (DM), once considered prevalent primarily in affluent nations, has now become a pervasive global phenomenon. Presently, it afflicts 537 million adults worldwide, and projections suggest this number will increase to 783 million within the next two decades (1). The implications and complications associated with this disease are profound, affecting not only individual health but also exerting significant strain on national economies and stability. This burden is especially pronounced in underrepresented communities, including low-income and 40 Indigenous subgroups, where the impact is disproportionately felt (2), particularly in regions
41 characterized by high conflict and limited access to healthcare facilities (3).

The healthcare sector in occupied territories faces significant challenges, with physicians
enduring constant threats and Palestinians not guaranteed access to hospitals (4). With the global
rate of undiagnosed DM at one individual for every two cases, the lack of testing centers and
consistent healthcare provision exacerbates this disparity (5).

Gazans, comprising 64% of whom are descendants of refugees, find themselves on the 46 47 disadvantaged side of an expanding health disparity (6). They have endured the historical trauma 48 of forced displacement, known as "Al Nakba" (the great catastrophe), which paved the way for 49 Israel's control over the West Bank and Gaza. This control has perpetuated further displacement 50 and instability among the local populations. A significant ongoing issue exacerbating the broader 51 conflict faced by Gazans is the siege imposed on Gaza in 2007, intensifying humanitarian 52 concerns and severely restricting access to essential resources. The persistent state of war renders life and basic amenities uncertain for many (7). 53

The healthcare system routinely faces compromise, either directly through the destruction of medical facilities or through the withholding of medical supplies. Many life-saving devices and medications, crucial for various medical interventions, are subject to arbitrary rules, leaving many civil structures vulnerable during the war. The few remaining institutions struggle to meet the growing demand and cope with limited capabilities, heavily reliant on foreign aid, rendering them largely inadequate (7).

60	As the need to address the growing epidemic grows, many nations have made it a routine to
61	allocate funding appropriately. Healthcare funding is expected to increase significantly to meet
62	the projected surge, with global diabetes-related health expenditures estimated to reach \$966
63	billion USD in 2021 and projected to rise to \$1.054 trillion USD by 2045 (8). In countries with
64	high GDP, funding is often barely sufficient to meet the demand for improved chronic healthcare
65	management centers (9). Chronic conditions like DM require specialized care and high
66	maintenance. Tackling increasing co-morbidity requires constant surveillance due to various
67	factors (1). Sub-optimal diets play a significant role in the predictive aspects of DM (10).
68	The growing number of adverse health outcomes related to DM in Gaza is influenced by the lack
69	of healthcare access due to geopolitical instability and limited resources (6). There is an apparent
70	link between the development of DM and economic development (10), raising concerns
71	regarding the healthcare situation in Gaza.

This review aims to delve into the complexities of this conflict's impact while also examining its 72 73 limitations and challenges to propose unique recommendations.

Impact of the Conflict 74

The conflict in Gaza has put an enormous strain on the already stretched healthcare sector in the 75

- strip, a region with around 2.2 million residents, of which almost 1.7 million have been 76
- 77 displaced. As of the 27th of March 2024, more than 74,000 have been injured, and 32,000 have
- 78 died. The displaced population took refuge in religious centers, United Nations Relief and Works

79	Agency for Palestine	Refugees in th	he Near East	(UNRWA)	facilities, schools,	and hospitals,
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80 which has led to further degradation of healthcare services provided (11,12).

According to the World Health Organization (WHO), the conflict has led to the degradation of 81 82 healthcare facilities, with only 2 out of 3 field hospitals fully functioning. Additionally, 12 out of 83 36 hospitals in the strip are partially functioning, and the average bed occupancy in the hospitals is approximately 327%. Furthermore, only 20 out of 82 primary healthcare facilities are 84 functional, and 104 ambulances have been affected since the onset of the war. Fuel, electricity, 85 86 food, water, medicine, and medical equipment shortages have resulted in inadequate treatment of the patient population in the Gaza Strip. These shortages have also led to a significant increase in 87 Communicable Diseases, with more than 524,363 cases of acute respiratory infections recorded 88 since mid-October 2023 and the number of recorded cases of diarrhea reaching 296,152 (11,13-89 90 15).

91 Gaza's healthcare system provides for 350,000 individuals suffering from chronic diseases, 1000 patients on hemodialysis, 50,000 pregnant women who result in 183 births per day, and 130 92 premature infants dependent on incubators, in addition to numerous other chronic diseases that 93 94 have gone untreated (13). According to the World Health Organization (WHO), individuals 95 suffering from long-term health conditions in Gaza are numbered to be more than 485,000 96 individuals with mental health disorders, 225,000 with hypertension, 45,000 with cardiovascular 97 disease (CVD), and more than 60,000 patients suffering from DM as of the 31st of October 2023 98 (14).

Patients with chronic diseases in ordinary conditions face challenges in terms of access to 99 100 healthcare, including the ability to acquire the medication they need, medication side effects, 101 healthcare facilities, compliance, and the psychosocial impact of their disease conditions. More 102 than 60,000 individuals with DM need special diets to control blood glucose levels, slow down 103 disease progression, manage their weight, and handle the comorbidities that typically accompany 104 DM. These diets are low in carbohydrates, sodium, saturated, and trans fats and provide large amounts of proteins, low-fat dairy, fruits, and vegetables. Considering the events in Gaza, 105 106 providing such a diet consistently would be difficult due to the high prices—up to fifty times the 107 prices before the conflict—and the lack of fresh supplies (50). The World Food Programme 108 (WFP) had warned of the ongoing famine in the northern governorates, which, according to the 109 warnings, was projected to occur anytime between mid-March and May 2024 (16). Diabetic patients, like other patients with chronic diseases in Gaza, suffer from the scarcity of 110 medications. In the United States, DM patients typically use about 5.9 different medications. 111 112 Patients may require statins, Angiotensin Converting Enzyme (ACE) inhibitors, sulfonylureas, 113 alpha-glucosidase inhibitors, and insulin, in addition to other medications needed to manage 114 concurrent comorbidities (17). The entry of medicines, including insulin pens, into the Gaza 115 Strip, has proven to be a difficult topic due to circumstances surrounding the matter. 116 A parallel can be drawn regarding the war in Ukraine. The availability of medications varies by 117 the affected area, being worse in contested areas, which is not a privilege Gaza has as the effects

of the conflict have been far-reaching. The issue of cost has emerged in Gaza, just as it has in

119 Ukraine, alongside the logistical challenges of importing or manufacturing medications. In

120	Ukraine, prices of medications have increased by up to 60%. The World Health Organization
121	(WHO) has predicted that disruptions in the supply of medications would lead to a rise in
122	morbidity and mortality in Ukraine. Mirroring the situation, a worse outcome would be expected
123	to occur in the Gaza Strip (18,19).
124	The lack of medication, appropriate diet, proper surveillance of DM complications, and
125	circumstances conducive to adequate glycemic control would result in a deteriorating quality of
126	life and the acceleration of disease progression, leading to DM complications. Patient
127	noncompliance has been found to increase overall mortality in both Type 1 DM and Type 2 DM.
128	Type 1 DM is an autoimmune disease that results in the destruction of pancreatic beta cells that
129	release insulin (20). On the other hand, Type 2 DM is a chronic disease that presents as a
130	resistance to the peripheral actions of insulin or an impairment in the secretion of insulin (21).
131	Medication non-adherence has also been associated with worsened quality of life, poorer
132	outcomes, increased complications, and greater resource utilization by patients (22-24).
133	Not only death but CVD, nephropathy, retinopathy, and neuropathy are all outcomes that may
134	become more prevalent or be accelerated because of the lack of appropriate management
135	systems. Bourne et al.'s systemic analysis between 1990 and 2010 concluded that blindness
136	caused by DM-associated retinopathy accounted for 2.6% of blindness worldwide (25). With
137	80% of end-stage renal disease caused by either DM, hypertension, or a combi in a nation of
138	both (25). Patients suffering from DM are 10 to 20 times more likely to undergo amputation of
139	the lower limbs (26).

140 The psychological aspect of DM affecting adherence has been documented in the literature. 141 Patients with DM tend to score lower on EQ-5D quality of life assessments than similarly aged 142 peers, and their scores are even lower with the onset of macrovascular and microvascular 143 complications of DM. The presence of both complications leads to even lower scores (27). DM 144 has also been found to be highly associated with depression (28). The psychological impact of 145 DM, leading patients to suffer from depression, stress, and anxiety, is exacerbated by the worsened circumstances in the Gaza Strip, which would lead to decreased adherence to self-care 146 147 behaviors, including medication adherence (29,30).

148 Limitations and Challenges

149 Based on previous conflicts, such as the conflict with Nicaragua, it has been evident that warfare significantly impacts health and access to essential services (31). It has been noted that access to 150 151 medications may be constrained during conflicts due to insufficient medical supplies and 152 logistical challenges such as limitations in shipping and electricity infrastructure. Maintaining the integrity of insulin, particularly in extreme environments with extreme temperatures, becomes 153 challenging due to electricity shortages, which may result in the degradation and loss of potency 154 155 of the medications (32). The cessation of electrical power disrupts cooling systems and 156 refrigeration, resulting in a temperature fluctuation between 25° and 37°, leading to the loss of 157 insulin effectiveness for short-acting, intermediate-acting, and mixed insulin (33). Furthermore, 158 the breakdown of data systems in conflict zones poses difficulties in accurately assessing 159 mortality and morbidity rates of diseases (34), impeding the identification of needs for managing 160 non-communicable diseases such as DM (35). Additionally, prioritizing infectious disease

161	management due to concerns about transmission often results in decreased attention to the care
162	of non-communicable diseases (35). It is well-established that specific dietary requirements and
163	proper nutrition are crucial in managing and controlling DM. However, during times of conflict,
164	the scarcity of food resources can profoundly affect patient outcomes (36). Moreover, the
165	reallocation of financial resources from social services to military endeavors exacerbates the
166	scarcity of resources such as medications and food (37). Despite increased physical activity and
167	weight loss during wartime, no improvement in serum lipid levels has been observed. This is
168	attributed to the limitations in antioxidant vitamins found in fruits and vegetables and the
169	heightened levels of smoking prevalent during times of conflict (38). Additionally, poor mental
170	health, stress, and anxiety serve as significant risk factors for the development of type 2 DM
171	(39). The prevalence of post-traumatic stress disorder, depression, and severe emotional
172	disorders during and after wartime exacerbates health challenges in conflict-affected areas
173	(40). Table 1 shows the limitations and challenges DM patients face in the conflict.

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Table 1. *The limitations and challenges for DM patients through the conflict.*

Limitations and Challenges

- 1. Limited access to medications and medical supplies due to logistical constraints in wartime conditions (31,32)
- 2. Difficulty in preserving insulin due to electricity scarcity, leading to loss of insulin activity (32,33)
- 3. Impaired data systems hinder accurate assessment of disease mortality and morbidity, affecting non-communicable disease care (34,35)
- 4. Prioritization of infectious disease care reduces attention to non-communicable disease management (35)
- 5. Food limitations during wartime affect diabetes management outcomes (36)
- 6. Allocation of budget from social services to military services leads to resource constraints (37)
- 7. Lack of improvement in serum lipid levels during war attributed to limitations in

antioxidant vitamins and increased smoking rates (38)

8. Poor mental health, stress, and anxiety increase the risk of developing type 2 diabetes mellitus, posing challenges in conflict areas (39,40)

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177

178 **Recommendations**

179 Several recommendations were made to address the proper management of DM during conflicts. 180 They discussed both short- and long-term plans (41). In terms of short-term management, the study emphasizes the critical need for local and international healthcare systems to promptly 181 182 identify diabetic patients and ensure their continuous access to essential medications (41). Notably, the study underscores the importance of identifying type 1 diabetic patients and 183 providing them with necessary insulin. Additionally, it highlights the importance of providing 184 suitable diabetic care for vulnerable populations such as pregnant women, the elderly, and 185 patients with prior complications from DM, such as foot ulcers and end-stage renal disease (41). 186 It is imperative for functioning healthcare systems in Gaza to maintain sufficient data on diabetic 187 patients despite the challenges in obtaining such information. Developing a plan to understand 188 189 the epidemiology of diabetes, including standardized tools, methods, and diagnostic tests, is 190 deemed beneficial.

191 Providing adequate support to diabetic patients in ongoing crises through continuous

surveillance, the use of health information systems, and planning for medical emergencies

associated with diabetes is essential for the long-term management of DM and limiting

194 complications (42). As per the "Boston Declaration," one of its four primary objectives is to

195 ensure universal access to insulin, essential medications, and diagnostics for managing glycemic 196 and blood pressure levels during humanitarian crises. Issued in 2019 after a conference in 197 Boston, this declaration aimed to address the global health and humanitarian dimensions of DM 198 during crises. The conference deliberated on pressing issues and barriers necessitating attention, 199 culminating in formulating a coordinated action plan. Moreover, it underscored the heightened 200 vulnerability of individuals with type 1 DM to mortality (43). Thus, ensuring the provision of essential medications and insulin to Gaza emerges as an imperative mandate. Evidence-Based 201 202 Clinical Guidelines for Diabetes Management in Crisis provide recommendations regarding glucose testing methods, glycemic goals, and the role of Electronic Medical Records (EMR) and 203 204 mobile health (mHealth) technologies in clinical decision-making. Further research is needed to improve the diagnosis and treatment of DM in humanitarian crises (42). Moreover, within the 205 framework of short-term management strategies for addressing DM amidst humanitarian crises, 206 207 an essential aspect involves training humanitarian healthcare workers. This training regimen 208 should encompass comprehensive instruction on insulin administration and managing DM-209 related emergencies (41). Furthermore, insights gleaned from a case study in Mali underscore the 210 importance of equipping healthcare workers with essential resources. These include rapid-acting 211 insulin, insulin syringes, antibiotics, dressings for foot infections, blood glucose meters, urine 212 test strips, and intravenous fluids, particularly in conflict zones (44). Patient education is an 213 additional critical facet emphasized in managing DM during crises. This entails focusing on key 214 aspects such as guiding in managing medications during illness episodes and recognizing 215 symptoms of dehydration, hypoglycemia, and Diabetic Ketoacidosis (DKA). DKA is a life-216 threatening complication of DM that is usually seen in type 1 DM patients rather than type 2.

217 DKA presents with uncontrolled hyperglycemia, metabolic acidosis, and an increase in the 218 concentration of ketones (41,45,46). However, it is imperative to acknowledge the potential 219 obstacles posed by language barriers, translation issues, and the complexity of medical 220 terminology, which may impede effective communication between patients and healthcare 221 providers. Various approaches can be employed to mitigate these challenges, including using 222 simple and visual communication methods. Additionally, adopting mobile applications, such as Cardmedic, accessible via app stores, can facilitate communication and comprehension in 223 224 healthcare settings (47). In the context of long-term management, it is imperative to ensure 225 consistent access to indispensable medications and to guarantee that healthcare facilities are 226 equipped with essential supplies and laboratory testing capabilities for parameters including HbA1C, total cholesterol, HDL, LDL, triglycerides, urine albumin to creatinine ratio (41). 227 Additionally, adopting additional cost-effective approaches for the management of type 2 DM is 228 229 advisable, such as the utilization of metformin, statins, and ACE inhibitors, coupled with the 230 implementation of systematic screenings for conditions like gestational diabetes, microalbuminuria, retinopathy, and diabetic foot complications (48). Addressing the necessity 231 232 for maintaining specific temperature conditions for medical equipment, the deployment of 233 thermoelectric coolers emerges as a viable solution. Additionally, employing drones for 234 medication transportation presents itself as a feasible option (49). These strategies collectively 235 contribute to developing and implementing comprehensive, continuous DM care protocols. 236 Suppose the conflict persists for an extended duration spanning several weeks or months. In that 237 case, providing comprehensive healthcare services targeting DM, hypertension, and CVD within 238 primary health centers is paramount (41). Additionally, there is a critical need to prioritize

training healthcare professionals (HCPs) in DM care, as this enhances local capacities and

240 resilience. National emergency response plans should be formulated with a dedicated strategy for

addressing DM care and ensuring the availability of insulin during crises Moreover, leveraging

online consultations facilitated by volunteers comprising both DM and mental health specialists

243 presents a promising avenue to alleviate the inevitable shortage in healthcare services resulting

- from military conflict (47). While utilizing technology, medical equipment, and specialized
- 245 expertise holds significant value in achieving these recommendations, paramount importance lies
- in fostering effective communication between all parties involved in the conflict. This requires
- accommodations to ensure the safe transport of people and critical supplies to areas in need (47).
- **Table 2** summarizes the short- and long-term recommendations.
- 249 **Table 2.** The short-term and long-term recommendations.

Short-Term Management

1. Prioritize recognition of diabetic patients and ensure continuous access to vital medications, particularly insulin, in both local and international healthcare systems (41).

2. Provide suitable diabetic care for vulnerable populations, including pregnant women, the elderly, and patients with previous complications from diabetes (41).

3. Enhance data collection on diabetic patients and develop standardized tools and methods to understand the epidemiology of diabetes locally (41).

4. Strengthen health information systems and surveillance for epidemiological purposes and ensure sufficient medication and diagnostic supplies in healthcare centers (41,42).

5. Train humanitarian healthcare workers in insulin administration and diabetes emergencies and equip them with necessary supplies for managing diabetes in conflict zones (41,44).

Long-Term Management

1. Ensure continuous access to essential medication and supplies and conduct laboratory testing for diabetes-related parameters (41).

2. Implement cost-effective approaches such as employing metformin, statins, and ACE inhibitors, and conducting screenings for various diabetic complications (41,48).

3. Utilize thermoelectric coolers for maintaining medical equipment temperatures and consider using drones for medication transportation (49).

4. Provide comprehensive care for diabetes, hypertension, and cardiovascular diseases in primary health centers, along with culturally appropriate diabetes education for patients and families (41).

5. Incorporate strategies for diabetes care into national emergency response plans, including the provision of insulin and the development of emergency kits containing essential medications and diagnostic tools (41,50).

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253 Conclusion

- 254 The conflict in Gaza has had a devastating impact on healthcare, particularly for individuals with
- chronic diseases like DM. Immediate action is needed to ensure the provision of essential
- 256 medications and care for DM patients, both in the short and long term, to mitigate the worsening
- 257 health outcomes caused by the conflict.

258 Submission statement

- 259 This work has not been submitted for publication elsewhere, and all the authors listed have
- approved the enclosed manuscript.

261 Availability of data and materials

- 262 The datasets used and/or analyzed during the current study are available from the corresponding
- author upon reasonable request.

264 **Conflict of interest**

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- 269 None
- 270 List of abbreviations
- 271 ACE: Angiotensin-Converting Enzyme, CVD: Cardiovascular Diseases, DKA: Diabetic
- 272 Ketoacidosis, DM: Diabetes Mellitus, EMR: Electronic Medical Records, GDP: Gross Domestic
- 273 Product, HCPs: Healthcare Professionals, mHealth: Mobile Health, UNRWA: United Nations
- 274 Relief and Works Agency for Palestine Refugees in the Near East, USD: United States Dollar,
- 275 WHO: World Health Organization, and WFP: World Food Programme.

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