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Surgeries and Cardiovascular System



Evaluation of Blood Elements

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Highlights

Primary Healthcare Units in Quito

Case Report with Review of Literature

Discovering Thoughts, Inventing Future

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Ascending Colon Duplication- A Case Report with Review of Literature

By Guru Prasad Painuly, Col. Ashok Tyagi, Rashk Kaushal & Mini Singhal

Max Super Specialty Hospital

Abstract- Gastro – intestinal duplications are usually detected in children before 2 years of age due symptoms/complications associated with the condition or during surgery in the child for some unrelated condition. In adults colonic duplication is of rare occurrence and often diagnosed during surgery. However, it may be diagnosed pre operatively due to symptoms of obstruction, volvulus or rarely due to perforation. We present an adult having tubular duplication of ascending colon, that presented with perforation. The duplication had a blind end and did not have distal communication. In addition, it had its own blood supply. The management of the case is discussed. Aetiogenesis of the anomaly is enumerated and literature reviewed.

For ascending colon duplication similar meaning word accessory ascending colon has also been used in the manuscript.

Keywords: gastro – intestinal tract duplication, colonic duplication, tubular duplication of colon.

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Abstract- Gastro - intestinal duplications are usually detected in children before 2 years of age due symptoms/complications associated with the condition or during surgery in the child for some unrelated condition. In adults colonic duplication is of rare occurrence and often diagnosed during surgery. However, it may be diagnosed pre operatively due to symptoms of obstruction, volvulus or rarely due to perforation. We present an adult having tubular duplication of ascending colon, that presented with perforation. The duplication had a blind end and did not have distal communication. In addition, it had its own blood supply. The management of the case is discussed. Aetiogenesis of the anomaly is enumerated and literature reviewed.

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Introduction

astro-intestinal duplications are uncommon anomaly. The incidence has been reported as 1 per 4500 births by the various authors¹. Due to complications of acute abdomen or bowel obstruction, most cases (65-80%) are detected in children by the age of 2 years^{2,3,4,5,6,7,8}. This is the reason that most cases have been reported in children and a few number of cases have been reported in adults. Amongst gastro - intestinal duplication colonic duplication represents only 7 - 20% cases^{2,3}. Fotiadis et al stated that most of the time definitive diagnosis of colonic duplication is made during a laparotomy⁶.

The differential diagnosis includes enteric cyst, giant colonic diverticulum, volvulus colon, duplication cyst. If not diagnosed in childhood, then these anomalies may come to notice in the later life as a chance finding (being asymptomatic) during an unrelated surgery. When associated with symptoms of intestinal obstruction, volvulus or compression of normal

adjacent bowel by the expanding duplication^{6,9}. It may also present with symptoms and signs of diverticulitis¹⁰. Rarely in case of presence of ectopic gastric mucosa with ulceration bleeding or perforation may occur^{9,10}. The relative occurrence of digestive duplication has been reported as follows^{2,11}. – jejunum 8%, Ileal 30%, ileocecal valve 30%, colon 6-7%, rectum 2-3%. The complications of colonic duplication include development of adenocarcinoma, squamous carcinoma and carcinoid tumour 12,13,14. Fora true digestive duplication, Rowling has set forth the following criteria¹⁵ –

- 1. The wall of duplication should be in continuity with one of the duplicated organs.
- The cyst is surrounded by a smooth muscle layer.
- 3. A layer of digestive mucosa is present (Typical or Heterotrophic).

Examples include gastric/colonic/pancreatic

CASE REPORT H.

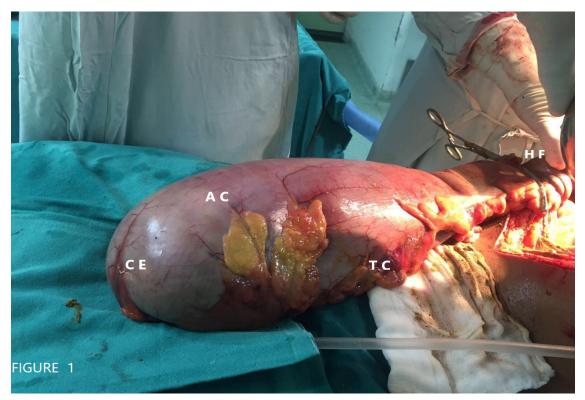
A 45 Year aged male was brought to ER as a case of acute abdomen. He had severe pain in abdomen, abdominal distension and h/o obstipation. He had signs of intestinal perforation. A bed side U/S and Xray abdomen erect (in sitting position) revealed free air under dome of diaphragm and also shadow of colonic lump. The findings were suggestive of intestinal perforation. Patient did not consent for CT abdomen, and hence with a presumptive diagnosis of volvulus sigmoid colon with perforation emergency exploratory laparotomy was done. Exploration revealed a free 40 cm long segment of colon that originated proximal to hepatic flexure. The duplicated ascending colon had a blind end distally, that reached pelvic brim. It was grossly distended and there was a big perforation at its origin from the primary colon (Figure 1, 2 and 3). The ascending colon was excised from proximal to origin of duplication along with right colonic flexure. Both colonic ends were closed and a side-to-side hand sewn anastomosis was done. A covering proximal loop ileostomy was formed for temporary diversion. Patient had postoperative sepsis and had a prolonged hospital stay of 21 days. 4 months later uneventful ileostomy closure was done.

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AC: Accessory ascending colon, TC: Taenia coli

Figure 1: Long duplication of ascending colon with distal free blind end.



VS: Vascular supply, C: Confluence of taenia in accessory ascending colon

Figure 2: Separate arterial feeder of accessory ascending colon.



HF: Hepatic flexure, Transverse colon with colonic band is also visible.

Figure 3: Hepatic flexure with big proximal perforation at the junction of ascending and accessory ascending colon.

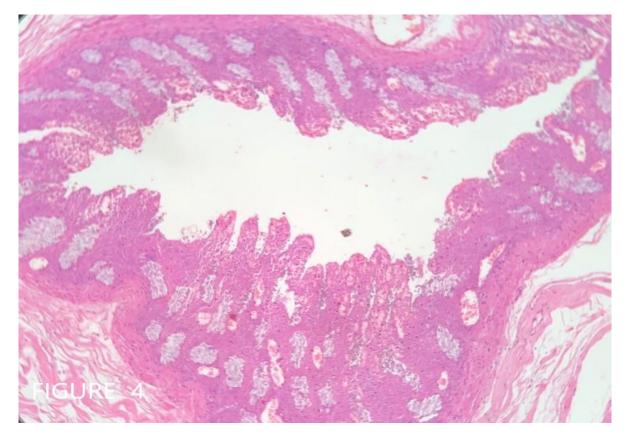


Figure 4: Histo-biopsy of the excised accessory ascending colon

Diagnosis and Management III.

Sonography, CT scan, Contrast enema and Colonoscopy have been suggested for the diagnosis¹⁶. Contrast CT is best modality in diagnosis, however preoperative diagnosis is difficult¹⁷. In addition, MRI and Contrast MRI have also been used in the diagnosis of colonic duplication.

Surgical resection of duplication with attached normal colonic segment to avert the risk of developing cancer in the duplicated colon is recommended^{2,3}. Currently laparoscopic resection of colon duplication has been advocated^{18,19,20}. Laparoscopic surgery is a preferred approach in the management of gastrointestinal duplications²¹. In huge barrel shaped colonic duplication selective mucosal excision with preservation of sero-muscular layer or distal internal drainage by excision of the common wall of duplication may be an effective alternative^{2,3,5,22}.

AETIOGENESIS, CLASSIFICATION AND IV. REVIEW OF LITERATURE

Historically first case of colonic duplication was reported in 1733 by Cadler²³. Another case was described by Suppinger in 1876, and the term 'duplication of alimentary tract' was first coined by Ladd in 1937. The exact cause of colonic duplication remains unknown, however environmental factors like trauma and hypoxia etc. have been implicated in its formation²⁴. Lewis et al proposed the diverticular theory, whereas alterations in closure of embryonic disc have been attributed as the cause of colonic duplication by others. Yet another theory regarding colonic duplication was proposed by Smith. It describes dorsal protrusion of yolk sac caused by its herniation or adherence to ectoderm responsible for the condition. This theory explains the Genito-urinary problems associated with duplication of hindgut. However, the most plausible theory having comprehensive explanation is given by Bremer. It states aberrant lumen recanalization of the gut in the embryo as the cause of colonic duplication. Since the duplication develops within the intestine, the outer wall contains all tissue layers and its counterparts²⁵.

Colonic duplications are difficult to differentiate from Mc Nutt's type 3 or Choony and Frizell's type 2 giant diverticulum^{8,26}. Gross et al described 4 varieties of duplication4 -

- 1. A tubular structure that branches out from intestine and extends for some distance between mesenteric leaves.
- 2. A double barrel structure communicating with the intestine lumen at one or both ends.
- 3. A cystic structure lying in the peritoneal cavity attached by a mesenteric stalk.

4. A spherical lesion contiguous with some part of the bowel particularly along the ileum.

Tubular Duplication can be barrel type or T/Y shaped.

McPherson classified colonic duplication into 3 groups²⁷-

Type I Simple cyst

Type II Diverticulum

Type III Tubular colonic duplication

Stern at al reported that 80% colonic duplications are cystic type, and 20% cases are tubular⁵.

Another classification envisages length/extent of involvement of colon/bowel. 2 Types are described -

Type I is limited to colon or rectum and is usually partial. If these lesions project into the lumen of bowel, patient may have intussusception leading to obstruction^{5,28}.

Type II duplications most of the time involve entire colon and are associated with genital or lower urinary tract anomalies. These may also be associated with intestinal mal rotation, duplication of ileum and appendix, spinal anomalies, omphalocele, exstrophy of urinary bladder and other abdominal wall anomalies. Double barrel duplication is usually associated with distal anomalies, terminal fistula or imperforated anus(type IIb and c). These duplications communicate with the bowel proximally, and in most cases do not have distal communication. Thus, these may become distended with faecal matter and cause obstruction of the adjacent bowel^{29,30}. Patient may have pain due to over distension or inflammation. Due to ulceration of the aberrant mucosa present in the duplicated segment, gastrointestinal bleeding may occur⁴. Unless associated with another congenital anomaly, duplication having a distal communication are usually asymptomatic. Usually, duplication is present along the ante mesenteric border of bowel, whereas it is lateral in position in case of ascending colon. Transverse colon duplications occur along supra colic margin. Rectal duplications occur posterior to rectum and both have common mesentery and blood supply, however loop duplication have a separate mesentery and blood supply. Tubular duplications sometimes can have direct communication with perineum^{3,30}. Tubular type of colonic duplication (T or Y type) have only one communication with the native bowel and the other end forms a blind pouch (as in our present case). Sometimes the distal end may communicate distally forming perianal and Genitourinary fistula and imperforated anus²⁰.

Due to number of cases being small, literature has mostly case reports rather than large series or multiinstitutional series. Rarely complete duplication of colon may occur^{29,30}. Reporting cases from 1950 - 2005, Fotiadis C et al reported total 83 cases⁶.

Roberts M et al reported 2 cases of sigmoid duplication that were pre operatively misdiagnosed as carcinoma⁷. Historically a case of colonic triplication has been reported by Gray, A.W.31 Lee KH et al reported laparoscopy for the first time in the management of intestinal duplication in a child³². A case asymptomatic tubular duplication of transverse colon has been reported by Kim YW et al³³. 2 Cases of colonic duplication that presented as rectal bleeding have also been reported by Fotiadis C et al⁶. 7 case reports of colonic duplication in adults were also reported. Of these 4 patients presented as abdominal pain and 3 had intestinal obstruction³⁴. A 'Y' shaped colonic duplication has been reported by Chang et al³⁵. A. Sozutek et al reported a case of perforated caecal duplication cyst presenting as peritonitis³⁶. An adult female had a sigmoid colon duplication, that was preoperatively diagnosed as colonic diverticula⁸. Jung Hi et al have reported a complete tubular duplication of colon in adult female with Colo-vaginal fistula³⁷. Wu X et al also reported a case of tubular colonic duplication in an adult³⁸. Another anomaly presenting with multi segmental asymptomatic duplication of colon has been reported³⁹. Kung-Chuan Cheng et al reported a case of colonic duplication that presented as a huge abdominal mass in an adult female⁴⁰. Yet another case of colonic duplication cyst in adult female has been reported by Shrestha S and Adhikari S⁴¹. Li GB et al managed a case of tubular colonic duplication and published it⁴². Recently Reddyn V et al reported a case of intestinal duplication in an elderly male that presented as sigmoid volvulus⁴³.

TAKE HOME MESSAGE V.

Diagnosis of colonic duplication should be suspected in an adult with chronic colicy pain with constipation as the condition may get unrecognized till adulthood. Patient may also have abdominal lump, distension- usually chronic in nature. However rarely it can also present as acute emergency. The pre-operative diagnosis may be difficult without radiological investigations because of vague clinical and radiological presentation. Treatment is to admit the patient and do open or laparoscopic resection of duplication along with attached normal colonic segment.

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Clinical Pattern and Outcomes of Neck Vascular Injuries at Aden Hospitals

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Abstract- Introduction: Vascular trauma to the neck can result in life-threatening injuries, because it contains a high concreteness of vital organ structures. Therefore, trauma require prompt diagnosis and treatment.

Objectives: To identify the different clinical pattern and outcome of traumatic neck vascular injury in Aden hospitals during 2015 – 2017.

Patients and methods: A retrospective observational study of all patients admitted to hospital with traumatic neck vascular injuries (TNVIs) between 2015 -2017. Data was collected about demographics, pattern of injury, type of vascular injury, associated injuries, hospital stay, and mortality rate.

Keywords: traumatic neck vascular injuries, penetrating injuries, clinical presentation.

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Clinical Pattern and Outcomes of Neck Vascular Injuries at Aden Hospitals

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Patients and methods: A retrospective observational study of all patients admitted to hospital with traumatic neck vascular injuries (TNVIs) between 2015 -2017. Data was collected about demographics, pattern of injury, type of vascular injury, associated injuries, hospital stay, and mortality rate.

Results: Of the total 52 TNVIs patients, male were 98.1 %. The mean age was 28.63 ± 10.07 years. The majority of injuries were penetrating (96.2%); gunshot was the predominant mechanism of injury (75%). Injuries in zone II involved in (57.7%). Of the TNVIs (65.4%) were associated with other injuries most of theme aerodigestive tract injury (38.2%). Patients presented with small non-pulsatile hematoma (28.8%), external bleeding (26.9%), rapid expanding hematoma (25%), neurological deficit (13.5%) and palpable thrill-audible bruit (5.8%). There is significant difference between stability of patients and their presentation (P = 0.033), most of stable patients have small non-pulsatile hematoma (43.3%), whereas an unstable patients have rapid expanding hematoma (36.4%). Neck exploration was performed in (94.2%). Arteries were more prone to injury (72.2%) than veins (28.8%). The common carotid artery is the commonest affected carotid arteries (41.5%), followed by external carotid artery (14.6%) and vertebral artery (12.2%). The internal jugular vein was the commonest injured vein (61.9%). Among those patients managed operatively; (55.9%) of injuries were repaired while ligation was performed on (44.06%). Complications were encountered in (50%) of them (42.3%) neurological deficit, wound infection (3.8%), fistula and hoarseness (1.9% for each). Overall mortality rate was (19.2%). Most of deaths occurred when patients were unstable during presentation (p<0.012).

Conclusion: TNVIs represent an alarming serious entity. Dominant presentation is penetrating injuries, exclusively among young male, mainly exposed to gunshot wound. The need for operative intervention should be based on clinical features. Moreover, further researches and an institutional protocol guidelines management are requested.

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Introduction

he neck is a vulnerable area comprising different vital organs of multiple physiological systems. Due to juxtaposition of these anatomical structures. there is a high predisposition of multi-systemic injuries with potentially life threatening lesions of the major blood vessels that conceder most common injured structure. 1-3

TNVIs represent 5-10% of all severe trauma cases^{.3-5} The neck is a complex anatomical area because in a small space there are abundant critical aerodigestive, neurological, and vascular structures.6 over the neck injury does not usually occur in isolation, therefore trauma are potentially dangerous and require prompt diagnosis and treatment.6

Mostly injured components in cervical trauma related to vascular structures.7 The most common mechanism is penetrating injury which mostly involved the common carotid artery, while the vertebral artery was less common injured due to its anatomical position.^{2,8-10} High velocity penetrating trauma can cause secondary injuries "blunt trauma" by shock wave. 10 The major venous injuries seen after penetrating trauma are the internal jugular vein and external jugular vein, in such cases venous injuries are never recognized due to its lower pressure. 11

Vascular injury is suggested by history and physical examination. Patient clinical presentation and symptoms still hold an important place in the management of penetrating neck injuries (PNIs), patients with signs of significant neck injury, particularly those with "hard" signs of vascular and/or aerodigestive injuries, require immediate surgical exploration.¹²

When an arterial injury is identified during neck exploration, current consensus agrees that primary repair of the artery is preferred than ligation, irrespective of any abnormality in focal preoperative neurologic examination findings. 13 Optimal management of patients with PNIs remains a controversial issue.² The treatment strategy of PNIs acquired from military surgical practice suggested mandatory exploration as a standard treatment to avoid missed injuries. However, civilian adoption of mandatory exploration in PNIs resulted in high negative exploration rate. 12 Therefor in the last three decades, there has been a slow shift towards selective

management for these injuries, which using zones of neck to guide investigations and management a "zonebased" approach or "no zone" approach. 13-15 Injuries are classified by penetration site into the three anatomical zones. 14,16 Those patients should be evaluated using proper physical examination, selective use of investigations 17,18 and managed according to staff experience and resource availability.¹⁴

Ideally, war injuries should be treated by surgeons having military surgery experience. In fact, civilian surgeons may find themselves trapped in wars practicing military surgery without prior training or experience in this field.¹⁹ The purpose of this study was to review our experience with penetrating neck vascular injuries in Aden-Yemen, thereby focusing on surgical management, and early outcomes and to highlight lessons learned from that period.

II. METHOD AND PATIENTS

This retrospectively study was conducted at Department of surgery in four main hospital in Aden city from 1st of April 2015 up to 31st December 2017. All patients of both gender and any age who fulfilling the inclusion criteria were enrolled in this study. During study period, the total number of patients admitted to surgical department in relation to vascular neck injury in Aden Hospitals were 52 patients. Any of the following was considered criteria for exclusion: death before admission, patients with superficial wound (which defined by injuries superficial to the plan of the platesma) or patients with minor neck injury who did not require hospital admission.

All patients were resuscitated in emergency room according to Advanced Trauma Life Support protocols in the hospital field. The clinical diagnosis was done at the time of the admission by history tacking, physical examination and investigations (laboratory, radiological and ultrasound), and/or confirmed during operative intervention.

The variable was collected retrospectively using a questionnaire and interviewing the patients including: age, sex, residence, clinical presentations that include detailed history of the injuries, hemodynamic status, external bleeding, expanding or large haematoma, neurological deficit and palpable thrill / audible bruit, associated injuries, operative finding, treatment option, and outcomes. Types of morbidity included neurological deficit, wound infection, A-V fistula and hoarseness.

Data and statistical analysis: data were presented as a proportion, medians or mean ± standard deviation as a appropriated and the analysis will be covered out using SPSS version 22 (SPSS Inc., Chicago, IL, USA). Quantitative variables like age and duration of symptoms were presented by calculating means ± standard deviation (SD). Qualitative variables like symptoms and operative finding gender,

presented by calculating frequencies and percentages. The statistical significance of differences between categorical variables were calculated by the chi-square test, Fisher's exact test. P-value of <0.05 was considered as statistically significant.

Results III.

During the study period from April 2015 to December 2017, the total number of patients with neck vascular injure was 52 patients. Most of the patients were male (98.1%). The age ranges from 9 to 60 years with a mean age was 28.63 ± 10.07 years. The peak age of patients with neck vascular injury in this study was in second decade (46.2%) followed by the third decade (26.9%) of life.

The majority of injuries was penetrating (96.2%); gunshot was the predominant mechanism of injury (75%), followed by sharp object injury (15.4%). Stab wound and road traffic accidents represent (3.8%) for each, and iatrogenic injury (1.9%). About (76.9%) of patients presented within the first 6 hours to the hospital and (42.3%) of the patients were unstable at the time of presentation. More than half of patients had their injuries on the right side; and the same percentage of injury in zone 2, while 17% of the studied patients had their injury in multiple zones. Concerning presentation of patients, the highest percentage of them presented with small non pulsatile hematoma (28.8%), followed by external bleeding (26.9%) and pulsatile hematomas (25%). Table I

In our study, we found arteries were the commonest injured vessel (71.2%), followed by veins (28.8%). Concerning type of injury, the most common finding was partial cut (55.9%), complete cut (44.1%). The commonest injured vessel is the carotid arteries, the common carotid artery represent (41.5%). followed by external carotid artery (14.6%) and vertebral artery in (12.2%), whereas the internal jugular vein was the most common vein injured (61.9%). Fig.1,2

About two third of our patients had another associated injuries, the most common associated injuries were aerodigestive tract injury (38.2%), followed by brachial plexus injury (23.5%).

The operative intervention was the main therapeutic option for most patients it represent (94.2%) and only (5.8%) of patients managed by non-operative procedures. Among those managed operatively, repair was performed in most of them (55.9%), while the ligation performed in (44.06%). The common type of repair was simple anatomical repair that represent (51.5%) followed by using graft in (33.3%) and end-toend anastomosis in (21.2%) of patients.

Variables				%
	Penetrating		50	96.2
		Gunshot	39	75.0
Machaniam of page yang dar injune	Type of penetrating	Sharp object	8	15.4
Mechanism of neck vascular injury	Type of perietrating	Stab wound	2	3.8
		latrogenic	1	1.9
	Blunt		2	3.8
Time of presentation (hrs)	< 6		40	76.9
Time of presentation (firs)	≥ 6		12	23.1
Hemodynamic state	Stable		30	57.7
Tierriodynamic state	Unstable		22	42.3
	Right		30	57.7
Site of injury	Left		20	38.5
	Bilateral		2	3.8
	zone 1		6	11.5
Zone of injury	zone 2		30	57.7
Zone of injury	zone 3		7	13.5
	Multiple zoon		9	17.3
	Small non pulsatile hematoma		15	28.8
	External bleeding		14	26.9
Presentation	Rapid expanding hematoma		13	25.0
	Neurological deficit		7	13.5
	Palpable thrill / Audible bruit		3	5.8

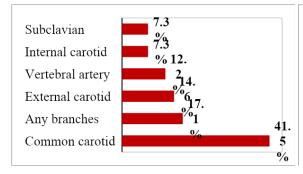


Figure 1: Name of Injured arteries: (n=42)

Our study showed that neck vascular injury in the hospitals are associated with (19.2%) mortality. Complications were found in (50%) patients with vascular neck injury. The most common complications was the neurological deficit (42.3%), while wound infection, A-V fistula, and hoarseness occur in (3.8%), (1.9%) and (1.9%) of patients respectively. The hospital stay ranged from one to more than 15 days. About (50%) of the studied patients stay in hospital 1-7 days, while (40.4%) stay 8-14 days and (9.6%) of them stay more than 15 days, as shown in Table II.

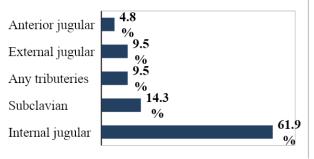


Figure 2: Name of injured vein: (n=17)

Variables			%
Morbidity	No	26	50.0
Worbidity	Yes	26	50.0
	Neurological deficit	22	42.3
Type of morbidity	Wound infection	2	3.8
	A-V fistula	1	1.9
	Hoarseness	1	1.9
	1- 7	26	50.0
Hospital stay (days)	8- 14	21	40.4
	>15	5	9.6
Mortality	No	42	80.8
	Yes	10	19.2

Table II: Outcome of the studied patients with neck vascular injury

IV. Discussion

This study deliberate the first study in Yemen, could be due to the lack of national data. Therefore, we conducted this retrospective descriptive study to establish the frequencies of clinical patterns and the outcomes of TVNIs in population of Aden hospitals within two consecutive years; from 1st April 2015 up to 31st Dec 2017.

Analysis of the personal characteristics of patients in this study showing a high prevalence of TNVIs among young males (98.1%), being similar to the studies reported by Martinakis et al¹ in Athens and Reva et al² in Russia.

The peak age for TNVIs in this study was in the second decade (46.2%) follow by third decade of life (26.9%). Similar observation was reported in other studies. 1,3

More than half of the patients in this study were solider (65.4%) while the incidence of traumatic injuries in Civilian (34.6%), showing a similarity to studies performed by White et al²⁰ and Rasmussen et al²¹ reports that civilian traumatic injuries of neck vessels are relatively uncommon.

The predominant mechanism of injury to the neck in this study is PNIs: (75% gunshot, 15.4% sharp objects and 3.8% stab wound) with a percentage of (96.2%), which is consistent with almost all other studies. 1,6,13,22 The blunt injuries are rare and consist (3.8 %) of studied patient as demonstrated by other studies.3,15

Findings at emergency department are small non pulsatile hematoma (28.8%), external bleeding (26.9%), rapid expanding hematoma (25.0%),neurological deficit (13.5%) and palpable thrill-audible bruit (5.8%), nearly similar result found by Reva et al² report cases on Physical examination reviled non pulsatile hematoma (36%), active bleeding from neck wound (19.6%), expanding or large hematoma in the injured area (19.6%), neurological deficit (17.4%), and asymptomatic injury (15.2%), while Teixeira et al¹⁹ reports external Bleeding (64%), expanding hematoma

(22%), and Nasr et al²³ reviled (65.5%) of the sample had active bleeding and (17.2%) hematomas.

In these study, commonest injured vessel were arteries it represent (71,2%), veins injury reviled (28.81%) this is in agreement with previous reports, where Prichayudh et al¹⁷ report 36 patients underwent immediate neck exploration, identified (55.6%) arterial injuries, (19.4%) venous injury, Bodanapally et al¹⁷ report 51 patient, (76.5%) patients had arterial injuries, (116) in contrast to another study by Ghnnam et al⁶ that report commonly affected anatomical structures were the neck veins (24.5%), followed by neck arteries (10.2%). However other studies were reported venous injury more than arteries injury.^{24,25} These differences may because majority of jugular venous injuries are probably unrecognized without exploration owing to the lowpressure venous system.²²

In current study, a higher proportion of TNVIs had associated injuries (65.4%), while (34.6 %) of patients had isolated neck trauma. Biffl et al²⁶ demonstrated similar incidence of associated injuries (33.1%). The most common associated injuries were aerodigestive tract injury (38.2%) similar to Pirrelli et al²⁷, in this study brachial plexus represent (23.5%), spinal injury represent (14.7%), and facial injury (11.8%).

Our study shows that, the operative intervention was the therapeutic option for most patients it represent (94.2%), only (5.8 %) were managed by conservative management. One study report neck exploration was performed in (88.2%) in his series while other demonstrated (84.4%) surgical intervention. 3,22

In this series 49 patients, underwent exploration, for positive physical examination findings and workup, this is supported by a study by Thomas et al²⁸ published a prospective study in which complementary diagnostic studies were used based on physical examination findings and no CTA was performed, showing good results. Furthermore, Mahmoodie et al²² and his group demonstrated that physical examination did not miss any major vascular or esophageal injuries that required intervention.

Among those managed operatively, repair was perform in most of them (55.9%) whereas the ligation performed in (44.06%). The common type of repair was simple repair that represent (51.5%) follow by using graft (33.3%) and end-to-end anastomosis (21.2%). One study established surgical interventions mainly included simple repair (53%).3

In the studied hospitals patients are treated with primary repair for carotid artery regardless of the degree of consciousness depression and preoperative condition similar to was found in Pirrelli et al²⁷ study. Other author recommend repair for CAI only in the absence of severe neurologic deficit. The overall mortality rate was (19.2%) this is high compared to other studies, in which mortality rates ranged from (11.8%) as Al Thani et al⁴ as well Ghnnam et al⁶ (12.2%).

The hospital stay ranged from one to more than fifteen days. About (50.5%) of the studied patients stay 1 -7 days, while 8-14 days represent (40.4%) and (9.6%) they stay more than 15 days. Ghnnam et al⁶ proved the mean duration of hospital stay 6.6 days; it is similar to this series.

Complications found in (50%) of the studied patient's. The common complications were the neurological deficit (42.3%), wound infection (3.8%), while A-V fistula and hoarseness (1.9%) for each, similar finding was reported by Reva et al.2

Conclusion

TNVIs represent an alarming serious entity. Dominant presentation is penetrating injuries, exclusively among young male, mainly exposed to gunshot wound. The need for operative intervention should be based on clinical features. Moreover, further researches and an institutional protocol guidelines management are requested.

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Evaluation of Blood Elements and Red Blood Cell Indices among Sudanese Cannabis and Cigarette Smokers in Khartoum State

By Dr. Abdelgadir Ahmed Abdelgadir, Alaa Ibrahim Mohammed, Abeer Saifaldeen Basher & Dr. Amged Hussein Abdelrahman

Abstract- Background: Cannabis, also known as (marihuana), is a psychoactive drug from the cannabis plant used for medical or recreational purposes. It is one of the most commonly used psychoactive drug worldwide, and it is the most popular illegal drugs. Long-term use of cannabis has acute effects on hemostasis of the body and hematological parameters of addicted individuals. Cigarette smoking is one of the major leading causes of death throughout the world: smoking has both acute and chronic effects on hematological parameters; many studies proved its harmful effects on many organ systems, mainly respiratory, reticuloendothelial system, and cardiovascular systems. Tobacco cigarette smoking is one of the main leading causes of death worldwide. Continuous cigarette smoking has severe adverse effects on hematological parameters (e.g., hemoglobin, white blood cells count, mean corpuscular volume, mean corpuscular hemoglobin concentration, red blood cells count, hematocrit).

Keywords: cannaibs; cigarette cannaibs smokers, hematological parameters.

GJMR-I Classification: NLMC Code: QW 300



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Dr. Abdelgadir Ahmed Abdelgadir a, Alaa Ibrahim Mohammed, Abeer Saifaldeen Basher & Dr. Amged Hussein Abdelrahman^ω

Abstract- Background: Cannabis, also known as (marihuana), is a psychoactive drug from the cannabis plant used for medical or recreational purposes. It is one of the most commonly used psychoactive drug worldwide, and it is the most popular illegal drugs. Long-term use of cannabis has acute effects on hemostasis of the body and hematological parameters of addicted individuals. Cigarette smoking is one of the major leading causes of death throughout the world: smoking has both acute and chronic effects on hematological parameters; many studies proved its harmful effects on many organ systems, mainly respiratory, reticuloendothelial system, and cardiovascular systems. Tobacco cigarette smoking is one of the main leading causes of death worldwide. Continuous cigarette smoking has severe adverse effects on hematological parameters (e.g., hemoglobin, white blood cells count, mean corpuscular volume, mean corpuscular hemoglobin concentration, red blood cells count, hematocrit). These represented a predisposing factor for the development of various pathological conditions and diseases such as atherosclerosis, polycythemia vera, chronic obstructive pulmonary disease and cardiovascular diseases.

Objectives: This work aimed to study the effect of Cannabis abuse and cigarette smoking on some hematological parameters in Sudanese smokers.

Methodology: This study was an observational comparative cross-sectional community-based study that enrolled a total of hundred Sudanese participants, divided into fifty (50) Sudanese individuals smoking cannabis plants as a case group and fifty (50) non-smoker healthy Sudanese individuals as a control group. Venous blood samples was collected from each participant, and a Complete Blood Count (CBC) was carried out using an automated hematology analyser to measure the hematological parameters for both groups.

Results: The result stated that the levels of total leukocyte count (WBCs), hematocrit (HCT), Neutrophils absolute count, and Red Cell Distribution Width (RDW) were significantly higher in cannabis smokers as compared to non-smokers with P.values equal (0. 001), (0. 005), (0.04) and (0. 000), respectively. The study revealed a significant decrease in platelet count and lymphocyte percentage count with P.values equal (0.04) and (0.02), respectively. The study also show significant effect of long-term smoking of cannabis among the case group in reducing the level of (HCT) with P.value (0.04).

This study concludes that the rates of examinations among cigarette smokers compared to non-smokers state a significantly higher values of Mean Corpuscular Volume (MCV) (p<0.0362) Platelet Distribution Width (PDW) (P<0.025) and its significance in WBCs depending on duration (p<0.0419) and depending on the dose (p<0.0473).

Conclusion: This study indicated that cannabis abusing could change the hematological parameters; therefore, the indiscriminate use of them should be discouraged as it is detrimental to the body results of our study conclude that there is a significant increase in MCV and PDW among smokers. But there was no change in other CBC parameters. Also, results conclude that smoking duration and frequency significantly increase white blood cell count.

Keywords: cannaibs; cigarette cannaibs hematological parameters.

Introduction I.

arijuana also known as (Cannabis) or Cannabis Sativa, is a psychoactive drug from the cannabis plant used for medical recreational purposes (1). It is one of the most commonly used psychoactive drugs worldwide, and it is one of the most popular illegal drugs (2). Main psychoactive part of cannabis is tetrahydrocannabinol, which has a scientific name called "Cannabinoids and Cannabidiol. Cannabinoids are usually classify as endocannabinoids, phytocannabinoids, and synthetic cannabinoids. Synthetic cannabinoids are in the gathering of drugs called new psychoactive substances and these technically synthetic cannabinoid receptor agonists are designer drugs that mimic psychoactive effects of cannabis (3). There is a long tradition of cannabis use for culinary, medicinal and ,ceremonial purposes in many developing countries. Various intake routes of tetrahydrocannabinol THC (intravenous, smoke, inhalation and, oral) so can be used by smoking, vaporizing, in food, or an extract. The plasma levels are related to onset, degree and duration of clinical effects. The degree of response and plasma cannabinoid levels attain edvary in a dose-related manner depending upon the potency of smoked marihuana(4). Over the last decades, there have been considerable researches involving cannabinoids and their importance in regulating a variety of physiological

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behavior, lipid metabolism, pleasure sensation, and immune system (4). The physical harm caused by cannabis is less well-known. In adults, chronic bronchitis, lung cancer, myocardial infarction, hepatotoxicity, decreased sperm count and motility, gynecomastia in males, suppression of ovulation among females, low birth weight and delayed visual system, and development among the newborns of cannabis using females (3). Plant-derived cannabinoids include delta-9-tetrahydrocannabinol (THC), the psychoactive component of cannabis. Cannabinoids mediate their effects through binding specific receptors, which are members of the G protein-coupled receptor superfamily. Two cannabinoid receptors was identified: Cannabinoid-1 receptor (CB1) and cannabinoid-2 receptor (CB2). CB1 is expressed primarily in the central nervous system (CNS) and is responsible for the psychoactive effects of cannabinoids by modulating neurotransmitter release). In contrast, CB2 is localized primarily in immune cells such as lymphocytes, macrophages, and neutrophils and is responsible for the immunomodulatory effects of cannabinoids (5). Smoking is the most important health problem in the world. Many studies proved its harmful effects on many organ systems like respiratory, reticuloendothelial system, and cardiovascular systems. (6) Tobacco cigarette smoking is one of the main leading causes of death worldwide. Continuous cigarette smoking has severe adverse effects on hematological parameters (e.g., hemoglobin, white blood cells count, mean corpuscular volume, mean corpuscular hemoglobin concentration, red blood cells count, hematocrit). These represented a predisposing factor for the development of various pathological conditions, and diseases such atherosclerosis, polycythemia vera, obstructive pulmonary disease and cardiovascular diseases. (7) Smoking effects on hematological indices observed on routine complete blood count testing (CBC). Smoking- induced increased in red blood cell count (RBC) was described. Current smoking has been reported as an associative factor with leukocytosis (TLC), thrombocytosis (PLTS) in some reports. Nicotineinduced JAK-STAT &NF-Kb signaling pathways are thought to mediate the increase in RBC. (8)

and psychological processes such as pain, feeding

Materials and Methods

Study population: Sudanese voluntary cannabis abusers and Cigarette smokers in Khartoum state, Sudan.

Inclusion criteria: This study included Sudanese voluntary cannabis abusers and Cigarette smokers in Khartoum state. Sudan.

Exclusion Criteria: The Participant with any disease or smoking any other type of smoking was excluded.

Data collection: Collected using self administrated per coded questionnaire, which was specifically designed to obtain information to this study.

Blood sampling: Venous blood was collected using sterile disposable plastic syringes after cleaning the venous puncture area with 70% ethanol, the blood 2.5 ml was added to the anticoagulant container EDTA.

Methods: The result was calculated by CBC analyzer. Whole blood is passed between two electrodes through apertures so narrow that only one cell can pass through at a time.

Statistical Analysis: Statistical assessment was carried out with statistical package for social sciences (SPSS) version 17.0 for windows statistical software.

Ethical Considerations: All participants were voluntarily submitted written informed consent before the commencement of the study. Neither the participant name nor situation or any other information was used in this study.

III. RESULTS

A total of 100 Sudanese participants were enrolled in our research, divided into 50 Sudanese individuals smoking cannabis plants as a case group and 50 healthy Sudanese individuals as a control group. (Fig.1). All study participants were males with ages ranged from (17) to (35) years old in both groups. (Fig.2). Concerning some other Cannabis smoking characteristics, the majority of cases group 38 (76%) reported smoking duration more than three years and almost nearly all of them, 49 98%, reported smoking frequencies of more than five times per week as detailed in figure (3), and Table (1). In regards to the effect of cannabis smoking on the results of complete blood count test results, our study showed that measures of cases group were significantly higher among cases group compared to the control group in white blood cells count (p = 0.0121), HCT (p = 0.0055), neutrophil count (p = 0.0428) and in RDW - SD (p = 0.004). The study showed that measures of the cases group were significantly lower among case group compared to control group in platelets count (p = 0.0477), and in the Lymphocytes count (p = 0.0238) as detailed in the table (2). Lastly, the study did not find a significant difference in the complete blood count measures according to the duration of cannabis smoking among the case group except in hematocrit (p = 0.041) as detailed in Table (3) the study was not able to assess the effect of cannabis smoking frequency/week of the measures of complete blood count because of the lack of variation in the relevant data; because all most all 49 (98%) of the case group had a similar frequency of weekly cannabis smokers. Among the cigarette smokers this study showed that the majority, 36 72% of the cases group (smokers) were within the age group 20 - 25, as detailed in Table (4). Concerning some other smoking characteristics, more than half of cases group 27 (54%) reported smoking cigarette duration less than five years, and almost the majority of them, 40 80% reported smoking frequencies less than ten times per day as detailed in Tables 5 and 6. In regards to the effect of cigarette smoking on the results of complete blood count measures, our study showed that results of cases group were significantly higher among cases group compared to control group in mean cell volume (p =

0.0362), in PWD (p = 0.0259) as detailed in Table (7) Moreover, the study did not find a significant difference in complete blood count measures according to the duration of smoking among case group except in white blood cells count (p = 0.0419) as detailed in Table 8. This study did not find a significant difference in complete blood count measures according to the frequency of smoking among case group except in white blood cell count (p = 0.0473) as detailed in Table (9).

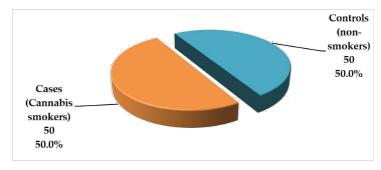


Figure. (1): The distribution of the study participants according to the study groups.

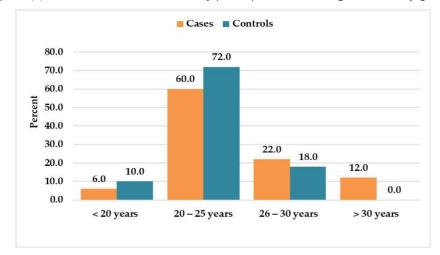


Figure. (2): The distribution of the study participants according to their age in years.

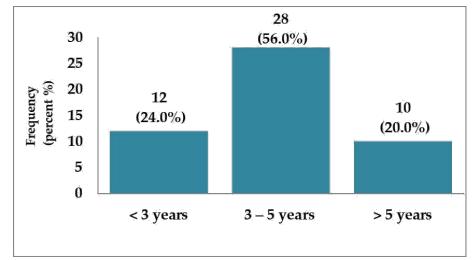


Figure. (3): The distribution of the study participants according to their cannabis smoking duration in years (n = 50 cases).

Table. (1): The distribution of the study participants according to their cannabis smoking frequency per week (n = 50 cases).

Cannabis smoking frequency – week	Frequency	Percent (%)
< 5 times / week	1	2.0
> 5 times / week	49	98.0
Total	50	100.0

Table (2): The difference in complete blood count results between the study groups (n = 100, 50 cases + 50 controls).

	Study groups			
Complete blood count	Mean in Case	Mean in Control	Difference	P value
WBC (× 10³)	6.26	5.20	1.07	<u>0.0121</u>
RBC (× 10³)	5.45	5.24	0.21	0.3539
HGB (g/dL)	14.36	14.04	0.32	0.4234
HCT (%)	46.62	43.29	3.33	<u>0.0055</u>
MCV (fL)	84.48	82.15	2.33	0.2855
мсн	27.78	26.95	0.83	0.1231
MCHC (g/L)	35.19	32.68	2.51	0.4774
PLT (× 10³)	237.06	269.44	-32.38	0.0477
LYM (× 10³)	2.22	2.15	0.07	0.7814
NEUT(× 10³)	2.50	1.88	0.63	0.0428
MIX (× 10 ³)	1.16	1.13	0.03	0.8482
LYM (%)	35.88	42.08	-6.20	0.0238
NEUT (%)	39.18	36.35	2.83	0.4033
MIXD (%)	18.91	20.20	-1.29	0.6402

RDW-SD (%)	44.73	42.41	2.31	0.0004
RDW-CV (%)	13.58	14.11	-0.53	0.0833
PDW (%)	16.85	15.80	1.05	0.0819
MPV (fL)	11.20	10.95	0.26	0.6012
P-LCR (fL)	27.14	27.33	-0.19	0.8557
PCT (ng/ml)	0.25	0.29	-0.03	0.0777

Table (3): The effect of Cannabis smoking duration on the measures of complete blood count (n = 50 cases)

Complete Blood Count	Duration of Cannabis smoking (years)			Overall mean	P value
(CBC)	< 3	3 - 5	> 5		
WBC (× 10³)	6.36	6.20	6.33	6.26	0.967
RBC (× 10³)	5.35	5.23	6.17	5.45	0.241
HGB (g/dL)	14.75	14.19	14.38	14.36	0.692
HCT (%)	50.41	45.61	44.91	46.62	<u>0.041</u>
MCV (fL)	81.68	84.24	88.51	84.48	0.236
мсн	27.67	27.64	28.32	27.78	0.577
MCHC (g/L)	46.02	31.72	31.91	35.19	0.163
PLT (× 10³)	243.17	228.04	255.00	237.06	0.792
LYM (× 10³)	2.32	2.02	2.65	2.22	0.676
NEUT(× 10³)	2.04	2.59	2.82	2.50	0.327
MIX (× 10³)	1.14	1.14	1.25	1.16	0.776
LYM (%)	36.23	33.11	43.21	35.88	0.374
NEUT (%)	35.08	40.25	41.12	39.18	0.391

MIXD (%)	20.18	18.31	19.07	18.91	0.811
RDW-SD (%)	45.10	44.77	44.15	44.73	0.525
RDW-CV (%)	14.26	13.19	13.88	13.58	0.582
PDW (%)	16.31	16.85	17.51	16.85	0.309
MPV (fL)	12.83	10.55	11.08	11.20	0.177
P-LCR (fL)	25.95	27.39	27.86	27.14	0.379
PCT (ng/ml)	0.28	0.23	0.28	0.25	0.923

Table (4): The distribution of the study participants according to their age - years (n = 50 cases)

Age – years	Frequency	Percent (%)
< 20 years	5	10.0
20 – 25 years	36	72.0
26 – 30 years	9	18.0
Total	50	100.0

Table (5): The distribution of the study participants according to their smoking duration - years (n = 50 cases)

Smoking duration – years	Frequency	Percent (%)
< 5 years	27	54.0
5 – 10 years	19	38.0
> 10 years	4	8.0
Total	50	100.0

Table (6): The distribution of the study participants according to their smoking frequency - day (n = 50 cases)

Smoking frequency – week	Frequency	Percent (%)
< 10 cigarettes / day	40	80.0
≥ 10 cigarettes / day	10	20.0
Total	50	100.0

Table (7): The difference in complete blood count results between the study groups (n = 100, 50 cases + 50 controls)

Complete blood count	Study group		Difference	P value
	Case	Control		
WBC (× 10³)	5.38	5.10	0.28	0.3340
RBC (× 10³)	5.24	5.22	0.02	0.7970
HGB (g/dL)	14.06	13.91	0.15	0.5760
HCT (%)	43.08	42.48	0.60	0.3547
MCV (fL)	82.61	77.20	5.41	0.0362
MCH (pg)	27.04	26.84	0.20	0.6957
MCHC (g/L)	32.69	32.67	0.02	0.9353
PLT (× 10³)	263.86	278.52	-14.66	0.3274
LYM (× 10³)	2.20	2.10	0.09	0.4726
NEUT (× 10³)	0.73	0.75	-0.02	0.7940
MIX (× 10³)	2.49	2.25	0.24	0.3236
LYM (%)	41.64	41.68	-0.04	0.9854
MIXD (%)	13.71	15.11	-1.40	0.4214
NEUT (%)	44.43	42.37	2.06	0.4397
RDW-CV (%)	14.08	16.26	-2.18	0.3119
RDW-SD (%)	42.78	42.34	0.44	0.3973
PDW (%)	16.78	15.59	1.19	0.0259
MPV (fL)	11.23	12.71	-1.48	0.4458
P-LCR (fL)	28.30	26.81	1.49	0.1019
PCT (ng/ml)	0.29	0.29	0.01	0.7018

Table (8): The effect of smoking duration on the measures of complete blood count (n = 50 cases)

Complete blood count	Duration of smoking (years)			Overall mean	P value
	< 5	5–10	> 10		, , , , , , ,
WBC (× 10³)	4.11	5.22	6.58	5.38	0.0419
RBC (× 10³)	5.29	5.17	5.25	5.24	0.5110
HGB (g/dL)	14.10	14.13	13.45	14.06	0.4020
HCT (%)	43.46	42.83	41.73	43.08	0.1730
MCV (fL)	82.67	83.14	79.70	82.61	0.5610
MCH (pg)	26.96	27.43	25.73	27.04	0.7550
MCHC (g/L)	32.54	32.99	32.28	32.69	0.6360
PLT (× 10³)	266.74	262.37	251.50	263.86	0.6480
LYM (× 10 ³)	2.12	2.41	1.68	2.20	0.9810
MIX (× 10³)	0.67	0.84	0.57	0.73	0.5760
NEUT (× 10³)	2.59	1.96	4.33	2.49	0.4640
LYM (%)	40.08	46.57	28.78	41.64	0.8560
MIXD (%)	12.57	16.33	8.98	13.71	0.6810
NEUT (%)	46.61	37.37	63.25	44.43	0.7870
RDW-CV (%)	14.11	14.02	14.13	14.08	0.6190
RDW-SD (%)	43.06	42.69	41.28	42.78	0.1140
PDW (%)	16.84	16.56	17.38	16.78	0.9280
MPV (fL)	11.41	11.01	11.05	11.23	0.1730
P-LCR (fL)	29.04	27.38	27.65	28.30	0.0840
PCT (ng/ml)	0.30	0.29	0.27	0.29	0.2690

Table (9): The effect of smoking frequency on the measures of complete blood count (n = 50 cases)

Complete blood count	Frequency of s	smoking (daily)	Overall mean	P value
	< 10	≥ 10		
WBC (× 10³)	4.27	6.01	5.38	0.0473
RBC (× 10³)	5.27	5.13	5.24	0.3669
HGB (g/dL)	14.12	13.80	14.06	0.3188
HCT (%)	43.31	42.18	43.08	0.2127
MCV (fL)	82.65	82.45	82.61	0.9104
MCH (pg)	27.06	26.99	27.04	0.9342
MCHC (g/L)	32.69	32.73	32.69	0.9085
PLT (× 10³)	268.55	245.10	263.86	0.2790
LYM (× 10³)	2.23	2.07	2.20	0.4955
MIX (× 10³)	0.71	0.82	0.73	0.3690
NEUT(× 10³)	2.54	2.28	2.49	0.5985
LYM (%)	42.10	39.82	41.64	0.5710
MIXD (%)	13.11	16.11	13.71	0.1775
NEUT (%)	44.52	44.08	44.43	0.9267
RDW-CV (%)	14.12	13.91	14.08	0.0932
RDW-SD (%)	42.98	41.99	42.78	0.1453
PDW (%)	16.74	16.94	16.78	0.7147
MPV (fL)	11.23	11.21	11.23	0.9414
P-LCR (fL)	28.34	28.13	28.30	0.8401
PCT (ng/ml)	0.30	0.27	0.29	0.1809

IV. DISCUSSION

effect of Cannabis "marihuana" The hematological parameters has been discussed by many authors as it is the most popular illegal drug used worldwide. This study had demonstrated the effects of cannabis abusing and Cigarette smoking on some hematological parameters, including 100 Sudanese participants, divided into 50 Sudanese individuals smoking cannabis plants as the case group and 50 healthy Sudanese individuals as the control group. Our study revealed a significant higher increase in the WBCs, HCT, absolute neutrophil count and RDW-SD, with P.values equal= (0.012), (0.005), (0.04) and (0.0004) respectively. The WBCs count was higher in case group (mean=6.26) cell/µl compared with control group (mean=5.20) cell/µl. This result was agreed with Dervas study (8) and disagreed with Amaechi and his colleagues study (6). Also, HCT showed a statistically significant higher difference between the case group and the control groups, in which the mean in case group was (46.6) % and (43.2) % in control group. Our result disagreed with the study carried in Nigeria by Amaechi and his colleague which showed lower HCT (6). The mean of Neutrophils absolute count in the case group was (2.5) cell/μl compare with (1.88) cell/μl in the control group. Our result was agreed with stud carried by Derya (8) and disagreed with the studies carried by Amaechi (6) and Salma studies (7). While the brilliant highly significant difference in the mean of RDW-SD in case group, which equal (44.7) fl was higher compare with control group (42.4) fl. This result was agreed with findings of the Derya study that their study showed a significantly increase in RDW-SD (8). On the other hand, our study revealed a significant lower difference in platelete count and lymphocyte percentage count with Values equal (0.04) and (0.02), respectively. Which agreed with the studies carried out by Amaechi (6) and salma (7). While the insignificant differences of RBCs (P.Value=0.35) were agreed with Salma, Amna and, Bashiri studies (7, 9, 10). Also, Hb concentration revealed no significant difference between both groups with P. Value (0.42) which agreed with Amna, Bashiri and, Derya studies (10). All studies agreed with our findings revealed no significant difference on RBCs indices; MCV, MCH, MCHC, MPV, PDW and P-LCR. Differences in our findings from previous studies may be attributed to the racial, genetic, geographic, nutritional status, duration of cannabis abusing and differences in sample size included. (8, 9). Among Sudanese cigarette smokers, our result showed that measures of CBC were significantly higher among the cases group compared to the control group in mean cell volume (p = 0.0362), and in PDW (p = 0.0259) and the duration of smoking and its frequency significantly increase white blood cells count (p = 0.0419), (p = 0.0473) respectively. Our result revealed that there was a significant increase in MCV and this finding disagree with Naser M Ergiah, et al, Rawia O. A Mustafa and Muhammad Asif, et al finding that showed there was no significant difference in MCV.(14) (15) (17) Similarly, we found that there was a significant increase in PDW in among smokers compared to non-smokers and these findings disagree with Muhammad Asif, et al. Findings which show that did not show any significant difference in PDW. (17) Furthermore, our result stated that there was no significant difference in MCH, RDW, PLT, MPV and this result agree with Muhammad Asif, et al findings which show that it did not show any significant difference in MCH, RDW, PLT, MPV.(17). On the other hand, our result showed no significant difference in WBCs, RBCs, Hb and ,MCHC and this disagree with Naser M Ergiah, et al findings that stated there was a significant increase in these parameters.(14) Also, our result reported that there was no statistically significant difference in total WBCs and, platelets count and this result disagrees with Dinesh et al which their results show there was a slight increase in white blood cells (WBCs) and platelets observed in smokers compared with non-smokers.(13). We found that there was no statistically significant difference in MCH and PLT and this findings agreed with Naser M Ergiah, et al result.(14). Similarly, we found that there was no statistically significant difference in RBCs, HB, HCT, MCH, RDW-SD, RDW-CV, Neutrophils, Monocytes and, Eosinophils. And this result disagrees with Rawia O. A Mustafa finding that stated a increase in these parameters and significantly Muhammad I. Khan, et al result which reported that Hemoglobin, RBC Count, HCT and MCH were significantly increased.(15) (16). Furthermore, we found that there was no significant difference in PLT count and these finding disagrees with Rawia O. A Mustafa finding that showed there was a significant decrease observed in PLT count. (15). Finally, we found no significant difference in WBC, RBC, Hb, HCT, MCHC and PCT and, this result disagrees with Muhammad Asif, et al findings which show that WBC, RBC, Hb, and HCT were significant high, at the same time MCHC and PCT were significantly low. (17)

Conclusion

Our study concluded that some hematological parameters in cannabis smokers differ significantly from non-smoker ones. The most likely consequences are an increase in TWBCs count, HCT, Neutrophils absolute count and, RDW. Also, the study revealed that cannabis abusing results in a low level of platelet count and lymphocyte percentage count., there was no change in RBCs count, Hb concentration, MCV, MCH, MCHC, MPV, PDW and ,P-LCR. Among Sudanese cigarette smokers, the study concluded that smoking might result an increase of MCV and PDW and, long smoking duration and high frequency per day may lead to high

total white blood cells count. Our result concluded that there was no significant difference in other CBC

parameters.

List of Abbreviation

RBCs	Red blood cells	
Hb	Hemoglobin	
PCV	Packed cell volume	
MCV	Mean cell volume	
MCH	Mean cell hemoglobin	
MCHC	Mean cell hemoglobin concentration	
RDW	Red cell distribution width	
WBCs	White blood cells	
plts	Platelets	
PCT	Plateletcrit	
MPV	Mean Platelet Volume	
PDW	Platelet Distribution Width	

Declarations

Ethical approval and consent to participant: Approval of This study was obtained from the hematology department of medical laboratory science MLS, Alfajir College, and the ministry of health issued by the local ethical committee Khartoum State Sudan. Written consent was taken from each member of the study.

Consent for publication Not applicable.

Availability of data and materials

The datasets generated during and / or analyzed in this study are not publicly available due to ethical policy in order to protect participant confidentiality.

Competing interest

The authors declare that they have no competing interests.

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Authors contributions

AA, AI, AS and, AH contributed in literature search and manuscript writing. Al and AS had the main idea of the study and contributed to manuscript writing, AA contributed to clinic work, AH contributed to statistical analysis. AA supervised the study and, critically reviewed the manuscript. All authors read and approved the final draft of the manuscript.

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Factors Conditioning the Adherence to Pharmacological Prescription in Patients with Hypertension Attending Primary Healthcare Units in Quito – Ecuador

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Keywords: medication adherence; hypertension; risk factors; primary health care.

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Introduction I.

ypertension (HTN) constitutes a public health problem and represents the core risk factor for premature death worldwide (Chow & Gupta, 2019; GBD 2017 Risk Factor Collaborators, 2018). Its control depends directly on the quality and timeliness of the medical diagnosis, treatment, and follow-up of the individual patient and effective health systems (Marrugat et al., 2003). Given that cheap and effective antihypertensive treatments are available, hypertension control and prevention of the morbidity and mortality that

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it generates should be achievable (Chow et al., 2013; McAlister et al., 2011). However, even in high-income countries, a significant proportion of people with hypertension do not know their diagnosis, and among those who do, a minority have been treated and have appropriate control (Maimaris et al., 2013; Pereira et al., 2009). Worldwide, it is estimated that only one in seven people have their pressure under control, that is, a blood pressure (BP) less than 140/90 mm Hg. Some of the obstacles that have been described to achieve BP control are the absence of comprehensive health services at the primary care level, limited access to drugs, and failure of health systems to effectively manage the prevention and treatment of the disease (MacMahon et al., 2008; Patel et al., 2016).

Available literature shows a clear correlation between a low level of adherence and less blood pressure control. Additionally, it has been associated with adverse outcomes such as cerebrovascular accidents, heart failure, myocardial infarction, and death. Impaired adherence constitutes a critical barrier for the reduction of blood pressure (Gosmanova & Kovesdy, 2015; Peacock & Krousel-Wood, 2017). However, the rates of adherence to medical prescriptions by patients with chronic conditions are very low. It is estimated that between 20-50% of patients suffering from these diseases do not take their medication as prescribed. An even larger number do not comply with the indications on changes in lifestyle (Kripalani et al., 2007). In the case of HTN, evidence shows that it affects 30 to 65% of patients and that 87.3% of uncontrolled patients presented failures in adherence to medical prescription (Abegaz et al., 2017). Because to its consequences on the health status, such as therapeutic failures, increased hospitalization rates, and increased healthcare costs, the WHO considers low adherence a priority public health issue (World Health Organization, 2004).

Lack of adherence is multidimensional. The factors that contribute to its existence are categorized into those related to the patient, such as socioeconomic status, inadequate knowledge of the disease, and costs of medication, and those related to the health system and equipment. Additionally, there are factors related to the treatment itself and those related to the disease (World Health Organization, 2004). It is necessary to study the adherence to prescription, as well as the main factors that condition compliance with medical indications, in particular those related to deficiencies in the quality of care provided by health services to hypertensive patients and their consequences on the morbidity and mortality caused by HTN.

The objective of this study was to determine the factors related to the patient, their clinical status, and the health services that facilitate or limit the adequate adherence to pharmacological treatment of patients who attend first-level health units in the South of Quito.

II. Methods and Procedures

a) Study Design

This research is a cross-sectional study carried out between February and November 2012 by the Faculty of Medicine of the Pontifical Catholic University of Ecuador (PUCE) and the University of Michigan. The Research Bioethics Committee of the Pontifical Catholic University of Ecuador (PUCE) and the Central University of Ecuador (Universidad Central del Ecuador) approved the study protocol. The Ministry of Public Health of Ecuador also approved the project.

b) Research scope and subjects

Study universe corresponded to all patients with diagnosed hypertension who regularly attended the health services belonging to the Ministry of Public Health of the District of Guamaní. This District is located in a suburban area south of the city of Quito; it serves 51,986 direct beneficiaries, 99,143 indirect beneficiaries with a total of 151,129 beneficiaries: Health care is covered by a type C Health Center and by type A and B health services. Three units were intentionally selected: the Guamaní Health Center, which, as the main center of the area, brings together the biggest number of patients with HTN. The Social Assistance Health Sub center that has the highest number of hypertensive patients registered in the health units attached to the area; and the Pueblo Unido health sub-center; where a family doctor works and provides care and follow-up to patients and their families with a comprehensive health approach.

Sampling was not carried out since all patients registered in these three units were included in the research, given they could be located and voluntarily agreed to participate in the study. The total number of patients with HTN registered in the units was 298, of which 188 (63.1%) participated in the study; one survey was eliminated due to incomplete information, thus n = 187.

c) Data collection

A questionary that included sociodemographic information and information on the diagnosis, control, and follow-up by the health services of each patient was applied. A clinical evaluation including taking, recording, and classification of blood pressure values was carried out following the recommendations of the JNC-7 (DEPARTMENT OF HEALTH AND HUMAN, 2003); as well as anthropometric measurements (weight, height, abdominal circumference), assessment cardiovascular risk and cognitive ability through the application of the Minimental test. Medical students from PUCE and the University of Michigan who received training and supervision from a faculty professor performed these procedures.

A subsample of the 130 patients who accepted the procedure had their blood glucose, cholesterol and creatinine determined. The samples were collected and analyzed by technicians from the PUCE clinical laboratory (DISERLAB) following international recommendations on biosafety and blood transport.

The criteria for evaluating the clinical status of the patient were established based on the European HTN Clinical Practice Guide (CPG) (ESH / ESC, 2013), which considers controlled hypertension when blood pressure values are ≤ 140/90 mg / mm and overweight or obesity when BMI ≥ 25. Fasting blood glucose <101 mg/dl was classified as normal, 102 - 125 mg/dl as altered and ≥ 126 mg/dl as diabetes; creatinine values to normal, altered renal function were ≥ 1.3 mg/dl in men and 1.2 mg/dl in women; and the total cholesterol level was considered elevated when the values were 190 mg/dl.

Treatment adherence defined as compliance with the pharmacological indications, both in dose and schedule and complete consumption of the medication, was self-reported, based on the Morinsky test (Morisky et al., 1986), considering as "adherent" those patients who fully complied with all the indications in the two evaluated dimensions.

d) Information Analysis

For the statistical analysis, a database was generated in Excel, and quality control of the information was carried out. It was then imported into the SPSS v 18.0 program. The statistical analysis initially included descriptive statistics: the mean, median, mode, and standard deviation values of the quantitative variables such as Glucose, Creatinine, Cholesterol, BMI, and Creatinine Clearance values were determined. Percentages were calculated for the qualitative variables. Subsequently, a bivariate analysis was performed, using the OR as association measures and the Chi-square as a significance test for qualitative variables. For quantitative variables, the Kolmogorov normality test was applied to establish the test to be used for the difference in means, be it the Student's t or the Mann-Whitney U; finally, a multiple regression model was performed.

RESULTS III.

The demographic and clinical characteristics of the study population are described in Table 1. The mean age was 63.8 years (SD 12.7) with a range of 35 to 89 years; the proportion of women was higher (83.4%) than that of men. Patients reported a low level of education with an average of 4.27 (SD 3.38) years completed. Regarding the clinical status of the patients who underwent laboratory tests (n = 130), it should be noted that more than 60% had hypercholesterolemia, 8.5% diabetes, 2.3% altered kidney function; and 86.5% were overweight or obese. 57.1% of patients had an uncontrolled blood pressure value.

a) Medication Adherence and its conditioning factors

28.9% (95% CI: 22.4% -35.4%) of the patients reported good adherence to the drug prescription, while 71.1% (95% CI: 64.6% -77, 6%) were non-adherent.

Table 2 shows the sociodemographic characteristics of the patients and their relationship with low adherence. Age (OR = 0.98); sex (OR = 0.69), living alone (OR = 0.79), having a partner (OR = 1.36) or working (OR = 1.28), were not statistically significantly associated with adherence pharmacological treatment. In contrast, the average number of years of formal education completed was significantly lower in non-adherent patients (3.75 \pm 3.05 years) than in adherent patients (5.65 \pm 3.7 years). Factors related to self-care, such as referring to carrying out actions and the need for another person's assistance for this purpose, were not significantly associated with adherence. However, the proportion of non-adherence was higher in patients who reported performing these actions (69.2%) or perceiving the need to do so 74.8%, with ORs of 1.25 and 1.48, respectively.

Table 3 presents the factors related to the patient's clinical status, showing that there are no significant differences between adherents and nonadherents in terms of the presence or absence of hypercholesterolemia, diabetes, being overweight or obese, or with the cognitive status of the patient. However, people who presented moderate or severe depression showed a higher risk of non-adherence than those without this health problem (OR = 2,054; CI 1,064 -3,964).

b) Conditioning Factors for Adherence Related to Health Services

Table 4 presents the conditioning factors for adherence related to the organization and operation of health services.

The need for help in reading medical indications was greater among the non-adherent, but the difference with the adherent subjects was not statistically significant. That doctors inquire about adherence and patient understands the prescribed medications, disrespect for patients' opinions by health professionals, and not shared goal setting with patients, were also not associated in a statistically significant degree with adherence. Regarding the indications written by the doctor or nurse, it was found that patients who reported having little or no understanding of these had a higher risk of non-adherence (OR = 2.3 Cl 1.188 -4.477) than those who did understand them. The irregular and incomplete supply of drugs by health units also constituted a risk factor for low adherence (OR = 1.93 Cl 1.022-3.74). Difficulty in accessing prescribed drugs due to lack of economic means increased the risk of not complying with treatment (OR = 5.715; CI 2.85-11.43). The perception of patients about disrespect for their beliefs (OR = 1.97 Cl 1.08 - 3.817) and the little or no satisfaction of patients with health services (OR = 2.45; Cl 1.202-5.005) were also related to low adherence to pharmacological indications.

The logistic regression model (Table 5) shows that factors that constitute facilitators of adequate adherence were higher education level, having economic means to access the prescribed medications. respect for patient's beliefs, and patient satisfaction with health services.

Even though non-adherents had a higher proportion of inadequate blood pressure control, the difference with adherents was not statistically significant (p = 0.081); see table 6.

Discussion IV.

present study contributes to the understanding of the obstacles to adherence to their pharmacological indications in patients with HTN seen at the first level of care. Collected data showed that seven out of ten hypertensive patients were nonadherent to the prescribed drug treatment, and more than half of the patients had inadequate blood pressure control. In addition, about three-quarters of the patients were obese overweight, 60% or hypercholesterolemia, and 21% had moderate to severe cognitive impairment. However, no association was found between low adherence and inadequate control of hypertension or an impaired clinical status, except that patients with moderate or severe depression were less compliant. Regarding patient-related factors linked to poor adherence, the low level of education (average years of formal education completed) stands out. Regarding health services, it was found that poor understanding of medical indications, irregular supply of medicines, reduced access to affordable drugs, and low level of satisfaction with health services increased the possibility of poor adherence.

Reduced adherence to medical indications in general, and pharmacological prescription in particular,

is a highly prevalent problem in the care of patients with chronic diseases and represents a priority for public health (Patel et al., 2016; Peacock & Krousel-Wood, 2017). In patients with hypertension poor adherence has been associated with lesser blood pressure control, cardiovascular complications, and death (Burnier & Egan, 2019; Gosmanova & Kovesdy, 2015). In the United States, an estimated 125,000 annual deaths and 33% to 69% of hospital admissions are due to poor adherence (Bosworth et al., 2011). Regarding health systems treatment failure increases hospitalization rates and health costs (World Health Organization, 2004). Therefore, timely detection and correction of low adherence should be part of the care and follow-up of patients with chronic diseases.

This study did not show a statistically significant association between non-adherence and poor blood pressure control. However, the proportion of nonadherence was higher in uncontrolled patients (61%). Other studies have shown that non-adherence is the key factor for uncontrolled blood pressure (Burnier & Egan, 2019; van der Laan et al., 2017); non-adherence increases significantly in uncontrolled patients (from 45.2% to 87.3%) (Abegaz et al., 2017). Lor Maichou et al. (Lor et al., 2019) asserts that adherence above 80% to antihypertensive medication is essential for optimal blood pressure control.

WHO estimates that the prevalence of nonadherence to antihypertensive drug treatment worldwide ranges from 30 to 50% (World Health Organization, 2004). In the study by Zullig (Zullig et al., 2015), nonadherence was 58.1%, and in that of Lor (Lor et al., 2019), non-adherence was 88.4%. A systematic review of 28 studies carried out in 15 countries shows a prevalence of non-adherence of 45.2% (Abegaz et al., 2017), and the research by Steiner (2009) found a mean therapeutic adherence of 70%, where only 36% obtained more than 80% of the prescription drugs. Adherence rates in patients with hypertension are very diverse, and the magnitudes reported in the different studies cannot be compared due to the use of various indicators and measurement instruments. However, the default value found in this study (71.1%) may indicate of the gap that exists between high and low and middle-income countries.

Lack of adherence is a complex, multifactorial problem; its conditioning factors can be classified according to their relations with the pathology, the patient, the health system, the doctor-patient relationship, or with the treatment itself (Khatib et al., 2014; van der Laan et al., 2017). In this study, there was no association between demographic factors such as sex, ethnicity, or age of the patient. While some research has found that men are less adherent than women, that evidence is conflicting. The study "Cohort Study of Medication Adherence among Older Adults (CoSMO)" (M. Krousel-Wood et al., 2011) reported very low

adherence to medication (14%) in patients older than 65 years of age; these values vary according to age group, sex, and ethnicity (MA Krousel-Wood et al., 2009). In the present study, patients with lower education had lower adherence, which corroborates what was reported by other studies in which a low level of education and reduced knowledge about the disease and treatment were significantly associated with low adherence (Parra et al. al., 2019).

International evidence suggests that clinical conditions, and in particularly cognitive dysfunction and mental health problems, constitute relevance obstacles for adherence (Steiner et al., 2009; Zullig et al., 2015). In this study, no significant association was found between hypercholesterolemia, excess weight or obesity, and low adherence. However, the coexistence hypercholesterolemia and hypertension is common: Tadick & Cuspide found that 69.7% of hypertensive patients presented hypercholesterolemia.

This study shows that the presence of moderate/severe depression constitutes a risk factor for non-adherence; these results are similar to those reported by (M. Krousel-Wood et al., 2011; Lor et al., 2019), who found that depressed people with chronic diseases are more likely to not comply with the prescribed treatment. Therefore, doctors should carry out a timely detection and adequate treatment of depression, especially in patients with poor adherence and poor blood pressure control.

Ecuador has made great efforts to implement comprehensive reforms and a public policy aimed to regulate health care (Espinosa, 2017), this does not guarantee compliance at the local level. Free access to care in public health services and the formulation of plans and programs aimed at providing permanent and timely access to comprehensive health care, based on established in the Constitution principles (Constitution of the Republic of Ecuador, 2008), have not always crystallized in benefits provided to patients with chronic diseases who are cared for at the first level. Different authors report an association between low adherence with the following factors of the health care system: not being able to read the written instructions regarding the management of the disease, not receiving information about hypertension, no availability of out-ofpocket expenses to access care and medications, deficiencies in the relationship and communication between patient and their healthcare provider, in addition to a general patient's perception of receiving poor quality of care (Parra et al., 2019; van der Laan et al., 2017).

To reduce the risk and burden of morbidity and mortality from cardiovascular diseases in low or medium per capita income countries WHO proposes different strategies to improve control, monitoring, and follow-up of hypertensive patients. The main recommendation is to strengthen actions to improve the levels of

knowledge, treatment, and control of hypertension, emphasizing the timely detection, selection, and appropriate use of medications and therapeutic adherence (World Health Organization, 2013). The challenge for many health systems is to seek strategies that reduce detected deficiencies, such as failures in the follow-up and the coordination in care, and patients receiving insufficient information that leads them to have difficulties in self-managing their disease (Samb et al., 2010). It is essential to improve patient follow-up and provide quality care in medical consultation, as it constitutes an opportunity to make clinical decisions based on blood pressure values and at the same time, provide counseling and evaluate adherence to treatment (Zuo et al., 2019).

In the clinical management of hypertension, patients are in charge of 95% of their daily care. Although this self-care goes beyond taking the prescribed medication, this aspect being the least complicated, it is vital to guarantee control of blood pressure levels (Comarca, 2011). Ultimately, adherence to the pharmacological prescription corresponds to decision and autonomy; however, health patients' services have a relevant role to play to help them have better compliance to medical prescription. Establishing effective channels of communication between health providers and their patients, which can help to understand the disease and the treatment by the latter, as well as share decision-making regarding the clinical management, are key conditioning factors to achieve this goal (Martínez C. et al., 2016: World Health Organization, 2013).

This study has the following limitations: it was carried out in a specific area of the Metropolitan District of Quito and one Type- C and two Type-A healthcare units belonging to the Ministry of Health; therefore, it cannot be generalized to all health services in Quito. Self-reporting was used in several of the indicators, especially in adherence. Additional questions were included in the Morinsky test to have more information on adherence, but this limits its international comparability. People who refused to take one of the tests represented up to 6% of the non-adherent subsample.

V. Conclusions

The efficacy of antihypertensive medications in controlling blood pressure and reducing adverse outcomes in morbidity and mortality from cardiovascular diseases is well documented. However, patient's adherence to the pharmacological treatment prescribed by the health professional plays a central role in the objectives for adequate management of this disease. The factors that determine the low adherence are multidimensional; in the present study, the logistic regression model showed that higher educational level and having the economic means to access drugs represent factors that facilitate compliance with the pharmacological prescription. Regarding the healthcare services and team, the main conditioning factors for adherence are respect for the beliefs of patients and patient satisfaction with health services. Comprehensive strategies should implemented for the management of hypertension, improving patient follow-up, not only about blood pressure control but also through the implementation of locally adapted strategies aimed at improving the clinical condition of patients and their quality of life.

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Table 1: General Characteristics of the Study Population

Sociodemographic	Frequency or Mean	Percentage or SD
Age	63.8 (x̃)	12.7 (SD)
Level of education		
Years approved	4,27 (x*)	3,38 (SD)
Sex		
Female	156	83.4
Male	31	16.6
Marital Status		
No partner	79	42.0
With partner	108	57.8
Employment		
Yes	51	27.4
No	135	72.6
Clinical Status		
Cholesterol		
Normal	46	35,4
Elevated	84	64,6
Blood Glucose		
Normal	109	83,8
Altered	10	7,7
Diabetes	11	8,5
Creatinine (renal dysfunction)		
Yes	3	2,3
No	127	97,7
BMI		
Normal	25	13,5
Overweight/Obesity	160	86,5
Cognitive impairment		
Mild	143	79
Moderate	38	21
Depression		
Minimal/mild	61	67,2
Moderate/severe	125	32,8
Hypertension		
Uncontrolled	78	42,9
Controlled	104	57,1

Table 2: Pacient'S Factors Associated with Medication Adherence

Demographic	Non ac	herents	s (n=133)	Adheren	ts (n=54)	OR	CI 95%	p-Value
characteristics	Frequer	псу	Percentage	Frequency	Percentage			
Age								
≤ 65 years	69		71,9	27	28,1	0,98	0,43 -	0,816
> 65 years	64		70,3	27	29,7		1,74	
Sex						0,692	0,306 -	0,374
Male	20		64.5	11	35.5		1,56	
Female	113		72,4	43	27,6			
Living with								
Alone	10		66,7	5	33,3	0,797	0,259 -	0,768*
Family	123		71,5	49	28,5	,	2,45	,
Marital status			,		•		,	
No partner	59		74,7	20	25,3	1,355	0,708 -	0,358
With partner	74		68,5	34	31,5	*	2,59	,
Employment			,		,		,	
Yes	38		74,5	13	25,5	1,275	0,615 -	0,513
No	94		69,6	41	30,4	,	2,64	•
Education			·		•		·	
(years								
approved)	3,75			5,65				0,001**
Mean	3,05			3,7				,
SD	ŕ			·				
Impaired family								
economy								
Yes	93		76,2	29	23,8	1,924	0,99 -	0,049
No	40		62,5%	24	37,5%	,	3,70	•
Selfcare			•					
Routine care								
Few or nothing	56	73,7	20		26,3	1,249	0,64 -	0,50
Often/always	74	69,2	33		30,8	•	2,40	•
Support care								
Yes	80		74,8	27	25,2	1,481	0,780 -	0,22
No	52		66,7	26	33,3		2,815	

Note: Some questions were not answered, they were considered as missing data. Thus, frequencies not always total the number of subjects included in the sample

Table 3: Factors Related to Patient'S Clinical Status Associated Whit Adherence to Medication

Clinical features	Non-ad	Non-adherents		erents	OR	Cl95%	p-value
	Frequency	Percentage	Frequency	Percentage			
Cholesterol				•			
Elevated	56	66,7	28	33,3	0,875	0,403 - 1,899	0,735
Normal	32	69,6	14	30,4			
Blood glucose							
Normal	70	64,2	39	35,8			
Altered	9	90,0	1	10,0			0,144
Diabetes	9	81,8	2	18,2			
BMI							
Normal	17	68	8	32	0,832	0,335 -	0,690
Overweight/Obesity	115	71,9	45	28,1		2,062	
Cognitive impairment							
Mild	99	69,2	44	30,8			
Moderate	29	76,3	9	23,7	0,698	0,305 - 1,598	0,431
Depression							
Moderate/severe	95	76,0	30	24,0	2,054	1,064 - 3,964	0,030
Mild	37	60,7	24	39,3			

Note: Some questions were not answered, they were considered as missing data. Thus, frequencies not always total the number of subjects included in the sample.

Table 4: Factors Related to Health Services Associated Whit Medication Adherence

Medical care	Non-adhe (n=13		Adherents (n=54)		OR	Cl95%	p- value
	Frequency	Percentage	Frequency	Percentage			
Understanding							
written prescription							
Few or nothing	98	76,6	30	23,4	2,306	1,188 - 4,477	0,013
Very good /excellent	34	58,6	24	41,4			
Help to read							
instructions	74	67,3	36	32,7	0,638	0,329 - 1,237	0,182
Never/mild	58	76,3	18	23,7			
Always/often							
Health services							
providing		7					
prescripted							
pharmaceuticals	84	5,7	27	24,3	1,93	1,022 - 3,74	0,041
Occasionally/never	43	61,4	27	38,6			
Always							
Money availability to							
compliance with							
medication	106	82	23	17,8	5,715	2,857 - 11,43	0,000
No	25	44,6	31	55,4			
Yes							
Physician Inquiries							
Adherence							
Yes	123	70,3	52	29,7	0,788	0,154 - 4,036	0,775
No	6	75,0	2	25,0			
Understands about							
medicines	00	70.0	20	00.4	4 400	0.700 0.70	0.000
Few/nothing	92	73,6	33	26,4	1,428	0,738 – 2,76	0,289
Excellent	41		31				
Respect to patient's	07	70.4	0.4	04.0	4.07	0.070 0.40	0.440
opinion	67	78,1	21	21,9	1,67	0,876 - 3,18	0,118
Never/eventually	63	65,8	33	34,2			
Always							
Respect for	60	70.4	00	00.0	4.07	1.00 0.017	0.040
patient's beliefs	69 56	76,1	20	23,9	1,97	1,08 – 3,817	0,042
Never/eventually	56	65,6	32	34,4			
Always							
Shared goal setting	65	75.0	04	04.4	1 001	0.0400000	0.440
Never/eventually	65	75,6	21	24,4	1,621	0,848 -3,098	0,142
Always	63	65,6	33	34,4			
Satisfaction with		0					
Healthcare	F O	8	10	10.4	0.450	1 000 5 005	0.010
Sometimes/never	59	1,9	13	18,1	2,453	1,202 - 5,005	0,012
Always	74	64,9	40	35,1			

Note: Some questions were not answered, they were considered as missing data. Thus, frequencies not always total the number of subjects included in the sample.

Table 5: Logistic Regression Model of Enabling Factors for Medication Adherence

Facilitators fa	ctors	OR AJUSTED	Confidenc	es Intervals	p-value
Level of Educa	tion	1,1	1,034	1,294	0,011
Access medicines	to	4,725	2,121	10,527	0,000
Respect patient's beliefs	for s	2,326	1,022	5,294	0,044
Satisfaction health services	with	2,342	0,956	5,738	0,063

Table 6: Association of Adherence to Prescribed Medication And Hypertension Control

Adherence	Control (n=78) Unc		Control (n=78) Uncontrolled (n=104)		OR	lc95%	p-value
	Frequency	Percentage	Frequency	Percentage			
Non-adherents	50	38,8	79	61,2	0,565	0,296 - 1,077	0,081
Adherents	28	52,8	25	57,2			

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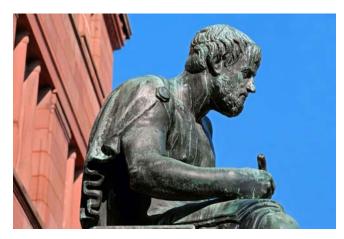
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Acknowledgments

Contributors to the research other than authors credited should be mentioned in Acknowledgments. The source of funding for the research can be included. Suppliers of resources may be mentioned along with their addresses.

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The following is the official style and template developed for publication of a research paper. Authors are not required to follow this style during the submission of the paper. It is just for reference purposes.



Manuscript Style Instruction (Optional)

- Microsoft Word Document Setting Instructions.
- Font type of all text should be Swis721 Lt BT.
- Page size: 8.27" x 11'", left margin: 0.65, right margin: 0.65, bottom margin: 0.75.
- Paper title should be in one column of font size 24.
- Author name in font size of 11 in one column.
- Abstract: font size 9 with the word "Abstract" in bold italics.
- Main text: font size 10 with two justified columns.
- Two columns with equal column width of 3.38 and spacing of 0.2.
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- The paragraph before spacing of 1 pt and after of 0 pt.
- Line spacing of 1 pt.
- Large images must be in one column.
- The names of first main headings (Heading 1) must be in Roman font, capital letters, and font size of 10.
- The names of second main headings (Heading 2) must not include numbers and must be in italics with a font size of 10.

Structure and Format of Manuscript

The recommended size of an original research paper is under 15,000 words and review papers under 7,000 words. Research articles should be less than 10,000 words. Research papers are usually longer than review papers. Review papers are reports of significant research (typically less than 7,000 words, including tables, figures, and references)

A research paper must include:

- a) A title which should be relevant to the theme of the paper.
- b) A summary, known as an abstract (less than 150 words), containing the major results and conclusions.
- c) Up to 10 keywords that precisely identify the paper's subject, purpose, and focus.
- d) An introduction, giving fundamental background objectives.
- e) Resources and techniques with sufficient complete experimental details (wherever possible by reference) to permit repetition, sources of information must be given, and numerical methods must be specified by reference.
- Results which should be presented concisely by well-designed tables and figures.
- g) Suitable statistical data should also be given.
- h) All data must have been gathered with attention to numerical detail in the planning stage.

Design has been recognized to be essential to experiments for a considerable time, and the editor has decided that any paper that appears not to have adequate numerical treatments of the data will be returned unrefereed.

- i) Discussion should cover implications and consequences and not just recapitulate the results; conclusions should also be summarized.
- j) There should be brief acknowledgments.
- k) There ought to be references in the conventional format. Global Journals recommends APA format.

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Abstract

The abstract is the foundation of the research paper. It should be clear and concise and must contain the objective of the paper and inferences drawn. It is advised to not include big mathematical equations or complicated jargon.

Many researchers searching for information online will use search engines such as Google, Yahoo or others. By optimizing your paper for search engines, you will amplify the chance of someone finding it. In turn, this will make it more likely to be viewed and cited in further works. Global Journals has compiled these guidelines to facilitate you to maximize the webfriendliness of the most public part of your paper.

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One must be persistent and creative in using keywords. An effective keyword search requires a strategy: planning of a list of possible keywords and phrases to try.

Choice of the main keywords is the first tool of writing a research paper. Research paper writing is an art. Keyword search should be as strategic as possible.

One should start brainstorming lists of potential keywords before even beginning searching. Think about the most important concepts related to research work. Ask, "What words would a source have to include to be truly valuable in a research paper?" Then consider synonyms for the important words.

It may take the discovery of only one important paper to steer in the right keyword direction because, in most databases, the keywords under which a research paper is abstracted are listed with the paper.

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Numerical methods used should be transparent and, where appropriate, supported by references.

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Authors are advised to submit any mathematical equation using either MathJax, KaTeX, or LaTeX, or in a very high-quality image.

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Figures are supposed to be submitted as separate files. Always include a citation in the text for each figure using Arabic numbers, e.g., Fig. 4. Artwork must be submitted online in vector electronic form or by emailing it.

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- 1. Choosing the topic: In most cases, the topic is selected by the interests of the author, but it can also be suggested by the guides. You can have several topics, and then judge which you are most comfortable with. This may be done by asking several questions of yourself, like "Will I be able to carry out a search in this area? Will I find all necessary resources to accomplish the search? Will I be able to find all information in this field area?" If the answer to this type of question is "yes," then you ought to choose that topic. In most cases, you may have to conduct surveys and visit several places. Also, you might have to do a lot of work to find all the rises and falls of the various data on that subject. Sometimes, detailed information plays a vital role, instead of short information. Evaluators are human: The first thing to remember is that evaluators are also human beings. They are not only meant for rejecting a paper. They are here to evaluate your paper. So present your best aspect.
- 2. Think like evaluators: If you are in confusion or getting demotivated because your paper may not be accepted by the evaluators, then think, and try to evaluate your paper like an evaluator. Try to understand what an evaluator wants in your research paper, and you will automatically have your answer. Make blueprints of paper: The outline is the plan or framework that will help you to arrange your thoughts. It will make your paper logical. But remember that all points of your outline must be related to the topic you have chosen.
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Verbs have to be in agreement with their subjects. In a research paper, do not start sentences with conjunctions or finish them with prepositions. When writing formally, it is advisable to never split an infinitive because someone will (wrongly) complain. Avoid clichés like a disease. Always shun irritating alliteration. Use language which is simple and straightforward. Put together a neat summary.

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- **15. Never start at the last minute:** Always allow enough time for research work. Leaving everything to the last minute will degrade your paper and spoil your work.
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- 17. Never copy others' work: Never copy others' work and give it your name because if the evaluator has seen it anywhere, you will be in trouble. Take proper rest and food: No matter how many hours you spend on your research activity, if you are not taking care of your health, then all your efforts will have been in vain. For quality research, take proper rest and food.
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- **20.** Think technically: Always think technically. If anything happens, search for its reasons, benefits, and demerits. Think and then print: When you go to print your paper, check that tables are not split, headings are not detached from their descriptions, and page sequence is maintained.
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- **23. Upon conclusion:** Once you have concluded your research, the next most important step is to present your findings. Presentation is extremely important as it is the definite medium though which your research is going to be in print for the rest of the crowd. Care should be taken to categorize your thoughts well and present them in a logical and neat manner. A good quality research paper format is essential because it serves to highlight your research paper and bring to light all necessary aspects of your research.

INFORMAL GUIDELINES OF RESEARCH PAPER WRITING

Key points to remember:

- Submit all work in its final form.
- Write your paper in the form which is presented in the guidelines using the template.
- Please note the criteria peer reviewers will use for grading the final paper.

Final points:

One purpose of organizing a research paper is to let people interpret your efforts selectively. The journal requires the following sections, submitted in the order listed, with each section starting on a new page:

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The discussion section:

This will provide understanding of the data and projections as to the implications of the results. The use of good quality references throughout the paper will give the effort trustworthiness by representing an alertness to prior workings.

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Specific editorial column necessities for compliance of a manuscript will always take over from directions in these general guidelines.

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- Submitting a manuscript with pages out of sequence.
- In every section of your document, use standard writing style, including articles ("a" and "the").
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- Use paragraphs to split each significant point (excluding the abstract).
- Align the primary line of each section.
- Present your points in sound order.
- Use present tense to report well-accepted matters.
- Use past tense to describe specific results.
- Do not use familiar wording; don't address the reviewer directly. Don't use slang or superlatives.
- Avoid use of extra pictures—include only those figures essential to presenting results.

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Choose a revealing title. It should be short and include the name(s) and address(es) of all authors. It should not have acronyms or abbreviations or exceed two printed lines.

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Reason for writing the article—theory, overall issue, purpose.

- Fundamental goal.
- To-the-point depiction of the research.
- Consequences, including definite statistics—if the consequences are quantitative in nature, account for this; results of any numerical analysis should be reported. Significant conclusions or questions that emerge from the research.

Approach:

- Single section and succinct.
- An outline of the job done is always written in past tense.
- o Concentrate on shortening results—limit background information to a verdict or two.
- Exact spelling, clarity of sentences and phrases, and appropriate reporting of quantities (proper units, important statistics) are just as significant in an abstract as they are anywhere else.

Introduction:

The introduction should "introduce" the manuscript. The reviewer should be presented with sufficient background information to be capable of comprehending and calculating the purpose of your study without having to refer to other works. The basis for the study should be offered. Give the most important references, but avoid making a comprehensive appraisal of the topic. Describe the problem visibly. If the problem is not acknowledged in a logical, reasonable way, the reviewer will give no attention to your results. Speak in common terms about techniques used to explain the problem, if needed, but do not present any particulars about the protocols here.



The following approach can create a valuable beginning:

- o Explain the value (significance) of the study.
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When a technique is used that has been well-described in another section, mention the specific item describing the way, but draw the basic principle while stating the situation. The purpose is to show all particular resources and broad procedures so that another person may use some or all of the methods in one more study or referee the scientific value of your work. It is not to be a step-by-step report of the whole thing you did, nor is a methods section a set of orders.

Materials:

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- Report the method and not the particulars of each process that engaged the same methodology.
- Describe the method entirely.
- o To be succinct, present methods under headings dedicated to specific dealings or groups of measures.
- Simplify—detail how procedures were completed, not how they were performed on a particular day.
- o If well-known procedures were used, account for the procedure by name, possibly with a reference, and that's all.

Approach:

It is embarrassing to use vigorous voice when documenting methods without using first person, which would focus the reviewer's interest on the researcher rather than the job. As a result, when writing up the methods, most authors use third person passive voice.

Use standard style in this and every other part of the paper—avoid familiar lists, and use full sentences.

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- o Resources and methods are not a set of information.
- o Skip all descriptive information and surroundings—save it for the argument.
- o Leave out information that is immaterial to a third party.



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Results:

The principle of a results segment is to present and demonstrate your conclusion. Create this part as entirely objective details of the outcome, and save all understanding for the discussion.

The page length of this segment is set by the sum and types of data to be reported. Use statistics and tables, if suitable, to present consequences most efficiently.

You must clearly differentiate material which would usually be incorporated in a study editorial from any unprocessed data or additional appendix matter that would not be available. In fact, such matters should not be submitted at all except if requested by the instructor.

Content:

- Sum up your conclusions in text and demonstrate them, if suitable, with figures and tables.
- o In the manuscript, explain each of your consequences, and point the reader to remarks that are most appropriate.
- o Present a background, such as by describing the question that was addressed by creation of an exacting study.
- Explain results of control experiments and give remarks that are not accessible in a prescribed figure or table, if appropriate.
- Examine your data, then prepare the analyzed (transformed) data in the form of a figure (graph), table, or manuscript.

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- Do not discuss or infer your outcome, report surrounding information, or try to explain anything.
- Do not include raw data or intermediate calculations in a research manuscript.
- o Do not present similar data more than once.
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- Never confuse figures with tables—there is a difference.

Approach:

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Figures and tables:

If you put figures and tables at the end of some details, make certain that they are visibly distinguished from any attached appendix materials, such as raw facts. Whatever the position, each table must be titled, numbered one after the other, and include a heading. All figures and tables must be divided from the text.

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Infer your data in the conversation in suitable depth. This means that when you clarify an observable fact, you must explain mechanisms that may account for the observation. If your results vary from your prospect, make clear why that may have happened. If your results agree, then explain the theory that the proof supported. It is never suitable to just state that the data approved the prospect, and let it drop at that. Make a decision as to whether each premise is supported or discarded or if you cannot make a conclusion with assurance. Do not just dismiss a study or part of a study as "uncertain."



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- o You may propose future guidelines, such as how an experiment might be personalized to accomplish a new idea.
- o Give details of all of your remarks as much as possible, focusing on mechanisms.
- o Make a decision as to whether the tentative design sufficiently addressed the theory and whether or not it was correctly restricted. Try to present substitute explanations if they are sensible alternatives.
- One piece of research will not counter an overall question, so maintain the large picture in mind. Where do you go next? The best studies unlock new avenues of study. What questions remain?
- o Recommendations for detailed papers will offer supplementary suggestions.

Approach:

When you refer to information, differentiate data generated by your own studies from other available information. Present work done by specific persons (including you) in past tense.

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Topics	Grades		
	A-B	C-D	E-F
Abstract	Clear and concise with appropriate content, Correct format. 200 words or below	Unclear summary and no specific data, Incorrect form Above 200 words	No specific data with ambiguous information Above 250 words
Introduction	Containing all background details with clear goal and appropriate details, flow specification, no grammar and spelling mistake, well organized sentence and paragraph, reference cited	Unclear and confusing data, appropriate format, grammar and spelling errors with unorganized matter	Out of place depth and content, hazy format
Methods and Procedures	Clear and to the point with well arranged paragraph, precision and accuracy of facts and figures, well organized subheads	Difficult to comprehend with embarrassed text, too much explanation but completed	Incorrect and unorganized structure with hazy meaning
Result	Well organized, Clear and specific, Correct units with precision, correct data, well structuring of paragraph, no grammar and spelling mistake	Complete and embarrassed text, difficult to comprehend	Irregular format with wrong facts and figures
Discussion	Well organized, meaningful specification, sound conclusion, logical and concise explanation, highly structured paragraph reference cited	Wordy, unclear conclusion, spurious	Conclusion is not cited, unorganized, difficult to comprehend
References	Complete and correct format, well organized	Beside the point, Incomplete	Wrong format and structuring



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