



## Hypertension Among Patients Younger than 40 Years: An Endocrine Point of View

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### Abstract

**Background and objectives:** Hypertension is common among older adults and the elderly, but some studies also reported data on young adults. The aim of this study is to diagnose the endocrine causes of secondary hypertension among younger adults.

**Methods:** This cross-sectional study enrolled 100 outpatients with hypertension from Endocrine and Internal Medicine Consultation Rooms, Shar Teaching Hospital, Sulaimaniyah, Iraq, from January 2023 to December 2023. Data was collected from patients with secondary hypertension using a questionnaire by face-to-face interviews during their hospital visits. Then, blood samples were collected to determine hematological parameters, kidney function tests, blood electrolytes, lipid profiles, hormones, and vitamin D levels.

**Results:** Most patients (68%) aged >30 years, females (56%), overweight (51%), visited the hospital as outpatient due to a headache (58%), had no clinical finding (81%), smoking history (83%), family history of hypertension (53%), positive past medical history (90%) and positive past surgical history (92%). The mean systolic/diastolic blood pressures were high ( $99.64 \pm 10.82/158.12 \pm 20.74$  mmHg). Mean hematological parameters, kidney function tests, electrolytes, lipid profiles, hormones, and vitamin D3 levels were observed. Most patients had negative dexamethasone suppression test (19%), essential hypertension (54%) with normal general urine examination (96%), abdominal ultrasound (72.05%), abdominal Computed Tomography (41%), Doppler investigations (90%), and Echocardiogram (57%).

**Conclusions:** Hypertension is a primary health issue among young adults that is mainly caused by endocrine disorders such as thyroid problems and primary hyperaldosteronism.

**Keywords:** Endocrine cause, Hypertension, Young individuals

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## Introduction

Hypertension is among the most typical (common) disorder and main risk factors for sudden death, especially among aged and elderly individuals. It is one of the non-communicable diseases that burden people worldwide, leading to coronary artery disease, myocardial infarction, ischemia, stroke, congestive heart failure, and increased cardiovascular risk. Hypertension also caused by primary aldosteronism (PA) among patients aged <40 years old.<sup>1, 2</sup> Hypertension increases the risk of emerging many ailments several times, such as coronary artery ailment twice, congestive heart failure four times, and cerebrovascular illness seven times. It is the 4<sup>th</sup> disease that causes premature death in developing nations and 7<sup>th</sup> in developed ones.<sup>3</sup> Younger adults' regular exercise training and healthy lifestyles are anticipated to exist as a subordinate occurrence of HPT in the general community. These outcomes describe a comforting clinical situation in that the rate of HPT in a large population of modest athletes (aged 18 to 40 years) is less. However, a deceptive inconsistency occurs between the expected and the reported prevalence of systemic HPT in athletic individuals.<sup>4, 5</sup> Currently, unwholesome lifestyles have increasingly complicated children and adolescents worldwide, thus causing a further rise in the risk of HPT in young populations, especially isolated systolic HPT.<sup>1</sup> Generally, high body mass index (BMI) and a family history of HPT were known as the sturdiest interpreters of HPT in multivariate analysis. However, many other factors are associated with developing HPT in young adults. Thus, the epidemiology, clinical manifestation, outcome, and management of HPT in young people should be considered seriously.<sup>6</sup> Essential hypertension (EHPT) is a significant threat to young adults in most societies, and most hypertensive cases are

diagnosed accidentally. Thus, detecting such cases as early as possible is very important. Excellence Hypertension Centers displayed that the prevalence of secondary HPT in young hypertensive adults is 30%, while it is 10% in hypertensive adults.<sup>7</sup> Consequently, the need arises to conduct a study on this aspect to find the prevalence of HPT in a young people and the correlation of accompanying factors. Thus, this study aimed to assess the incidence of endocrine diseases leading to secondary HPT among young adults in Sulaymaniyah City, Iraq.

## Patients and methods

In this cross-sectional study, 100 outpatients with HPT were recruited at Shar Teaching Hospitals from January 2023 to December 2023. Patients with sustained HPT aged <40 years old were included, while pregnant and postpartum women with those had long-standing HPT and aged >40 years were excluded. A validated questionnaire was utilized to collect hypertensive patients' sociodemographic and clinical data, including age, gender, height and weight, to determine BMI, smoking status, family history of HPT, clinical findings, systolic blood pressure (SBP), diastolic blood pressure (DBP), past medical history (PMH), and past surgical history (PSH). Diagnosis of the patients with HPT was confirmed using either ambulatory BP monitoring or manual BP measurement using a mercury sphygmomanometer on different occasions. Then, 5.0 mL blood was taken from each patient to perform hematological parameters, kidney function tests, protein levels, blood electrolytes, lipid profiles, hormones, and vitamin D levels. On the other hand, diagnostic tests were done for each patient, including general urine examination (GUE), dexamethasone suppression test (DST), electrocardiogram (ECHO), abdominal ultrasound (U/S), abdominal computed tomography (CT), and Doppler U/S. The Scientific and Ethical Committees of Shar





Teaching Hospital in Sulaymaniyah and Kurdistan Higher Council of Medical Specialties (no.1569 21/9/2023) (KHCMS), Iraq, revised and accepted the study protocol. Written informed consent was gained from patients/their guardians before starting the study, and their data was kept confidential. The data obtained were analyzed using the Statistical Package for Social Science (Chicago, USA, version 26). Age, systolic BP, diastolic BP, BMI, Na, K, Hb, etc. were given as mean and standard deviation (SD), while gender, smoking, complaints, etc. were shown as numbers and percentages. This is not a comparable study to find the level of significance between groups.

## Results

The mean age of the participants was  $32.37 \pm 6.5$  years, ranging from 10-40 years. Most patients (68%) were aged  $>30$  years and were females (56%). The mean BMI was  $28.49 \pm 6.18$  kg/m<sup>2</sup>, and most patients (51%) were overweight, followed by obese (28%), then normal weight (20%). The mean SBP/DBP was  $99.64 \pm 10.82/158.12 \pm 20.74$  mmHg. Moreover, 58% were admitted outpatients to the hospital with a complaint of headache, 22% with dyspnea, 14% with chest palpitation, 2.0% with edema/weakness and 2.0% with no complaint. Furthermore, 81% had no clinical findings, 83% had no smoking history, 53% had no family history of HPT, 90% had no PMH, and 92% had no PSH, as found in Table (1).

**Table (1):** Sociodemographic and clinical data of the participants.

Variables	Number (%)
Age (Years)	
10-20	8 (8.0)
20-30	24 (24)
30-40	68 (68)
Gender	
Male	44 (44)
Female	56 (56)
BMI (Kg/m <sup>2</sup> )	
Underweight	1 (1.0)
Normal weight	20 (20)

Overweight	51 (51)
Obese	28 (28)
Blood pressure (Mean±SD)	
SBP (mmHg)	$99.64 \pm 10.82$
DBP (mmHg)	$158.12 \pm 20.74$
Complain	
No	2 (2.0)
Yes	98 (98)
Clinical finding	
No	81 (81)
Yes	19 (19)
Smoking	
No	83 (83)
Yes	17 (17)
Family history	
No	53 (53)
Yes	47 (47)
PMH	
No	90 (90)
Yes	10 (10)
PSH	
No	92 (92)
Yes	18 (18)

DBP: Diastolic blood pressure, PMH: Past medical history, PSH: Past surgical history, SBP: Systolic blood pressure

The mean WBC, Hb and platelets were  $10020.2 \pm 11845.78$  cells/ $\mu$ L,  $13.72 \pm 1.61$  g/dL and  $278470.0 \pm 83469.40$  cells/mL, respectively. Blood urea and serum creatinine were  $29.05 \pm 8.25$  mmol/L and  $0.76 \pm 0.2$  mg/dL, respectively. In respect of the minerals, the mean value of  $140.05 \pm 2.30$ ,  $4.31 \pm 13.92$ ,  $1.97 \pm 0.13$ ,  $9.04 \pm 0.59$ ,  $3.77 \pm 0.59$  and  $103.54 \pm 3.68$  mmol/L was found for sodium, potassium, magnesium, calcium, phosphate, and chloride, subsequently. The mean lipid profile was  $189.53 \pm 41.18$ ,  $119.31 \pm 29.49$ ,  $43.31 \pm 9.53$  and  $180.16 \pm 90.63$  mg/dL for total cholesterol, low-density lipoprotein, high-density lipoprotein, and triglyceride, respectively. On the other hand, the levels of free thyroxine, thyroid stimulating hormone, parathyroid hormone, aldosterone, plasma renin and plasma aldosterone renin ratio was  $17.96 \pm 12.50$  ng/dL,  $4.03 \pm 8.47$  mIU/mL,  $52.15 \pm 27.76$  pg/mL,  $144.97 \pm 83.69$  ng/dL,  $26.73 \pm 36.72$  pg/dL and  $20.18 \pm 29.96$  ng/dL, respectively. Patients had a low





vitamin D3 ( $25.13 \pm 8.98 \mu\text{g}$ ) with normal serum albumin ( $4.42 \pm 0.43 \text{ g/dL}$ ). The mean level of RBS was  $112.34 \pm 41.30 \text{ mg/dL}$ . Lastly, the plasma level of Metanephrine was evaluated for 24 patients; five patients had an average level, while others had a mean level of  $105.35 \pm 120.54 \mu\text{g}$ , as shown in Table (2).

**Table (2):** Laboratory findings of the hypertensive patients.

Laboratory finding	No.	Min.	Max.	Mean $\pm$ SD
<b>Hematology</b>				
WBC (cells/ $\mu\text{L}$ )	100	3500	89000	$10020.2 \pm 11845.78$
Hb (g/dL)	100	9.8	18	$13.72 \pm 1.61$
Platelet (cells/mL)	100	117000	650000	$278470 \pm 83469.4$
<b>Renal function</b>				
Urea (mmol/L)	100	14.6	61	$29.05 \pm 8.25$
Creatinine (mg/dL)	100	0.2	1.4	$0.76 \pm 0.2$
<b>Mineral (mmol/L)</b>				
Sodium	100	134	145	$140.05 \pm 2.3$
Potassium	100	2.4	143	$4.31 \pm 13.92$
Magnesium	57	1.6	2.22	$1.97 \pm 0.13$
Calcium	100	7.9	10.3	$9.04 \pm 0.59$
Phosphate	99	2.5	5.5	$3.77 \pm 0.59$
Chloride	100	96	111.3	$103.54 \pm 3.68$
<b>Lipid profile (mg/dL)</b>				
Cholesterol	100	93	336	$189.53 \pm 41.18$
LDL	100	50	234	$119.31 \pm 29.49$
HDL	13	29	63	$43.31 \pm 9.53$
TG	100	10	499	$180.16 \pm 90.63$
<b>Hormone</b>				
Free T4 (ng/dL)	100	6.8	128	$17.96 \pm 12.5$
TSH (mIU/mL)	99	0.0	62	$4.03 \pm 8.47$
PTH (pg/mL)	29	16	141	$52.15 \pm 27.76$
Aldosterone (ng/dL)	31	12.74	360	$144.97 \pm 83.69$
P. Renin (pg/mL)	32	1.64	134	$26.73 \pm 36.72$
PARR (ng/dL)	31	0.10	144	$20.18 \pm 29.96$
<b>Vitamin</b>				

Vitamin D3 (ng/mL)	97	4.0	55	$25.13 \pm 8.98$
<b>Protein</b>				
Albumin (g/dL)	98	3.77	5.6	$4.42 \pm 0.43$
<b>Miscellaneous</b>				
RBS (mg/dL)	100	74	314	$112.34 \pm 41.3$
Plasma Metanephrine ( $\mu\text{g}$ )	24	2.9	456	$105.35 \pm 120.54$

WBC: White blood cell; Hb: Hemoglobin; LDL: Low density lipoprotein; TG: Triglyceride; T4: Thyroxine; TSH: Thyroid stimulating hormone; PTH: Parathyroid hormone; PARR: Plasma aldosterone renin ratio; RBS: Random blood sugar; Min.: minimum; max.: maximum

**Table (3):** Diagnostic results of study patients.

Variable	Number (%)
<b>GUE</b>	
Normal	96 (96)
RBC cast	4.0 (4.0)
<b>DST</b>	
Negative	19 (19)
Positive	7.0 (7.0)
<b>Abdominal U/S</b>	
Normal	72 (72)
Fatty liver	6.0 (6.0)
Others	17 (17)
<b>Abdominal CT</b>	
Normal	41 (41)
Adenoma	3.0 (3.0)
Others	5.0 (5.0)
<b>Doppler US</b>	
Normal	90 (90)
Left renal artery stenosis	2.0 (2.0)
<b>ECHO</b>	
Normal	57 (57)
LVH	38 (38)
<b>Diagnosis</b>	
EHPT	54 (54)
Cushing syndrome	8.0 (8.0)
Hyperaldosteronism	11 (11)
Metabolic syndrome	3.0 (3.0)
Others	24 (24)

GUE: General urine examination; DST: Dexamethasone suppression test; U/S: Ultrasound; CT: Computed tomography; ECHO: Echocardiogram; RBC: Red blood cell; LVH: Left ventricular hypertrophy; EHPT: Essential hypertension





The results of patients associated with metabolic syndrome were high levels of BMI ( $43.53 \pm 5.05$  kg/m<sup>2</sup>), TG ( $298.67 \pm 173.75$  mg/dL), and SBP/DBP ( $101.67 \pm 7.64$  /  $153.33 \pm 11.55$  mmHg), but not RBS ( $107.66 \pm 5.51$  mg/dL), as displayed in Table (4)

**Table 4:** Clinical and laboratory results of the patients associated with metabolic syndrome.

Clinical and laboratory test	Mean $\pm$ SD
Body mass index (kg/m <sup>2</sup> )	43.53 $\pm$ 5.05
Triglyceride (mg/dL)	298.67 $\pm$ 173.75
Random blood sugar (mg/dL)	107.66 $\pm$ 5.51
Systolic/Diastolic blood pressure (mmHg)	101.67 $\pm$ 7.64/153.33 $\pm$ 11.55

## Discussion

In this study, most hypertensive patients aged >30 years, females (56%), overweight (51%), and admitted the hospital due to a headache (58%). In contrast, most patients had no clinical findings (81%), no smoking history (83%), no family history of HPT (53%), no past medical history (90%), and no past surgical history (92%). In this regard, Sidenur et al. reported that HPT among young adults aged 20–40 years was directly related to a family history of HPT, smoking, age, gender, and BMI.<sup>2</sup> Thus, the first two variables are against our findings. Whereas Caselli et al. identified that the rate of HPT in young adults is low (3.0%) and mainly related to family history and overweight.<sup>4</sup> These variations among studies might be associated with quick urbanization and lifestyle alterations.<sup>2</sup> Generally, high BMI, smoking, and having a family history of HPT were found to be the toughest predictors of HPT.<sup>4,8</sup> Moreover, in this study, the HPT rate was higher among patients aged >30 years and lowest among those <20 years. Similarly, another study found the incidence of HPT was more in those 35–40 years (27%) and the less in those 20–24 years (5.3%).<sup>2</sup> Simultaneously, Panesar et al. showed that

the occurrence of HPT was significantly higher in 30–40 years (29.6%) than those aged 20–30 years (14.8%).<sup>9</sup> Moreover, the mean age of our patients was  $32.37 \pm 6.5$  years, similar to that found by another study ( $32 \pm 7.0$  years).<sup>7</sup> This is probably due to an environmental factors and body physiological senescence.<sup>2</sup> Concerning gender, we found that HPT is more predominant among females (56%), which does not agree with another study that stated male prevalence (53%) among hypertensive patients rather than females.<sup>7</sup> Most studies agreed that smoking is directly correlated with the frequency of HPT. Sidenur et al. reported that the frequency of HPT was higher in smokers (23.1%) than non-smokers (17.1%).<sup>2</sup> Singh et al. also established that the prevalence of HPT among smokers was highly significantly higher (53.06%) compared to non-smokers (18.11%) ( $p < 0.001$ ).<sup>10</sup> Furthermore, the mean BMI in this study was  $28.49 \pm 6.18$  kg/m<sup>2</sup>. Hence, the prevalence of HPT was higher among overweight individuals (51%) rather than obese and non-weighted patients. Most studies agreed that HPT is more predominant among obese patients, such as that found by Sidenur et al., which mentioned that obese patients with moderate risk (class 2) (100%) had the highest prevalence than those with normal BMI (14.1%).<sup>2</sup> Similarly, Menaga et al. found that the prevalence of HPT was more significant among obese (48.3%), that might be associated to obesity-related HPT.<sup>11</sup> Urea, creatinine, and platelet measurements could be indicators of renal function and endothelial impairment in patients with HPT.<sup>12</sup> In this study, WBC, Hb and platelets were not related to the prevalence of HPT among young adults and serum creatinine; however, blood urea was very high, which supports a significant linear relationship between blood urea and HPT among young adults. Thus, blood urea levels can be used non-invasively to detect hypertensive cases.<sup>13</sup>





Nutrition is undeniably known to impact BP, especially NaCl, but potassium plays a prominent role. The current study showed average Na, K, Ca, Cl, and Mg levels, with higher phosphate levels. Alonso et al. do not support this outcome, which reported that phosphorus was allied with lower SBP and DBP and reduced risk of HPT.<sup>14</sup> The TC, LDL, and HDL were standard, while TG was high among our patients. These findings are not coincidental with the outcomes of Chen et al., who found elevated TC, LDL, and non-HDL about the incidence of HPT in adult Chinese males, while TG was not significantly associated.<sup>15</sup> They displayed that age, BMI, fasting plasma glucose, and TC were strongly associated with HPT incidence. In the present study, high levels of T4 and TSH with normal PTH were found among hypertensive patients. This was in line with Mehran et al., who stated that thyroid hormone resistance was significantly linked to increased BP in euthyroid.<sup>16</sup> Also, Laclaustra et al. showed that high T4, obesity and diabetes were common in those with a higher resistance thyroid hormone index.<sup>17</sup> We found high levels of aldosterone, plasma renin, and PARR among patients, which aligned with Trenkel et al., who found PARR is a valuable parameter for primary aldosteronism in hypertensive cases.<sup>18</sup> Also, high level of Metanephrine were found among our patients which agreed with that of Wang et al., who found high levels of Metanephrine among patients with EHPT that might be associated with a higher risk of cardiovascular and cerebrovascular events.<sup>19</sup> Regarding vitamin D correlation and HPT incidence, studies have shown a controversial relationship between vitamin D and lowering BP among various populations.<sup>20</sup> In this study, most patients had normal GUE, DST, abdominal ultrasound/CT, Doppler and ECHO results, which means these parameters are not potential enough to be used as a diagnostic tool for HPT among young people.

However, most patients had EHPT (54%), hyperaldosteronism (11%), Cushing syndrome (8.0%), and metabolic syndrome (3.0%). In this respect, Noilhan et al. reported that primary aldosteronism and fibromuscular dysplasia are the leading organic etiologies among men and women younger than 40 with HPT.<sup>7</sup> Consequently, metabolic syndrome was found among patients (3.0%) due to high levels of BMI, TG, and SBP/DBP. In this regard, Yano et al. revealed that young and middle-aged individuals with isolated systolic HPT had more risks for cardiovascular/coronary heart diseases mortality.<sup>21</sup>

### Conclusions

Hypertension is one of the leading health problems among young adults in our locality (Sulaymaniyah /Iraq); that might not be related to the medical history of HPT or smoking status but is mainly correlated to endocrine disorders, such as thyroid issues and primary hyperaldosteronism. Additionally, BMI and TG play a significant role in the prevalence of this disease.

### Conflict of interest

Not declared.

### Acknowledgements

The authors would like to thank the healthcare staff at Shar Teaching Hospital, Sulaymaniyah, Iraq.

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