

Comparative Analysis of American Academy of Pediatrics and European Society of Hypertension Guidelines for the Diagnosis and Treatment of Pediatric Hypertension

Se Jin Park, M.D., Ph.D.¹
Jae Il Shin, M.D., Ph.D.²

¹Department of Pediatrics, Daejeon Eulji Medical Center, Eulji University School of Medicine, Daejeon, Republic of Korea, ²Department of Pediatrics, Severance Children's Hospital, Yonsei University College of Medicine, Seoul, Republic of Korea

Corresponding author:

Jae Il Shin, M.D., Ph.D.
Department of Pediatrics, Severance Children's Hospital, Yonsei University College of Medicine, Seoul 03722, Republic of Korea
Tel: +82-2-2228-2050
Fax: +82-2-393-9118
E-mail: shinji@yuhs.ac

Received: 12 December 2021
Revised: 19 December 2021
Accepted: 24 December 2021

This is an open-access article distributed under the terms of the Creative Commons Attribution Non-Commercial License (<http://creativecommons.org/licenses/by-nc/4.0/>) which permits unrestricted non-commercial use, distribution, and reproduction in any medium, provided the original work is properly cited.

Copyright © 2021 The Korean Society of Pediatric Nephrology

Childhood hypertension (HTN) has become a significant public health issue because of the increased risk of cardiovascular disease in adulthood. However, childhood HTN is underrecognized and underdiagnosed in clinical practice. The European Society of Hypertension in 2016 and the American Academy of Pediatrics (AAP) in 2017 published updated guidelines for the screening, prevention, and management of pediatric HTN. There were notable differences between the two guidelines as well as many similarities. The updated AAP guidelines have clarified and simplified the recommendations for screening, diagnosis, and treatment of childhood HTN based on current evidence. This review highlights the important developments in both guidelines, focusing on recent advances in the classification and treatment of childhood HTN.

Key words: Blood pressure, Child, Guideline, Hypertension

Introduction

Since the “Fourth Report on the Diagnosis, Evaluation, and Treatment of High Blood Pressure in Children and Adolescents” (2004 Fourth Report) was published in 2004, interest in childhood hypertension (HTN) has increased¹. Increased blood pressure (BP) in children and adolescents increases the risk of cardiovascular disease (CVD) in adulthood, which is known as the tracking phenomenon². Mounting evidence has also shown that childhood and adolescent HTN increase the risk of adult cardiovascular morbidity and mortality^{3,4}. The global rise in childhood HTN is likely associated with the growing prevalence of children who are overweight and obese⁵.

In an effort to delay the present trajectory, new studies related to childhood HTN have been conducted, and two updated clinical practice guidelines for the management of BP in children and adolescents have recently been published. In 2016, the European Society of Hypertension (ESH) updated the 2009 guidelines and published new guidelines (2016 the European Society of Hypertension guidelines [2016 ESHG]) to increase efforts toward prevention and management of HTN in the pediatric age, thus also helping relieve the burden of CVD in adults⁶. In 2017, the American Academy of Pediatrics Clinical Practice Guideline (AAP CPG) for Screening and Management of

High Blood Pressure in Children and Adolescents (2017 AAP CPG) was published to update the current guidelines based on a systematic review of evidence published since the 2004 Fourth Report⁷. The aim of the 2017 AAP CPG was not only to provide an update on topics relevant to the diagnosis, evaluation, and management of pediatric HTN, but also to improve early recognition of pediatric HTN by simplifying the diagnosis and evaluation of HTN in children and adolescents. The 2017 AAP CPG includes 30 key action statements and 27 additional recommendations (Appendix 1 in the 2017 AAP CPG).

Both guidelines produced significant reactions and were discussed in the HTN community. Although there are many similarities between these two guidelines, there are also several notable differences. These differences suggest that further studies are needed to prevent and treat HTN in childhood and to improve adult CVD. The goal of this review is to highlight new guidelines for the diagnosis and management of pediatric HTN.

Definition and classification of HTN

Both the 2016 ESHG and 2017 AAP CPG regard a BP value <90th percentile by age, sex, and height as normal BP^{6,7}. HTN is defined in children aged <13 years in the 2017 AAP CPG and <16 years in the 2016 ESHG as a persistent elevation in BP measured clinically at or ≥95th percentile on at least three separate occasions^{6,7}.

According to the 2016 ESHG, children with an average systolic BP (SBP) and/or diastolic BP (DBP) between the 90th and 95th percentile are classified as having high-normal BP⁶. The diagnostic criteria for elevated BP in children depend on the fact that BP in children increases with age and body size. Therefore, it is inappropriate to use a single BP level to define HTN, as is done in adults. For adolescents aged ≥16 years, the same single, static thresholds (≥140/90 mmHg) defining BP categories used in adults were adopted, allowing a gentle transition from pediatric to adult providers⁶. This is quite different from the Fourth Report that used percentiles to define the BP category for adolescents, while maintaining the Fourth Report criteria in young people <16 years of age.

However, in 2017, the term “elevated BP” was used to replace “prehypertension”^{7,8}. New thresholds for elevated BP, stage 1 HTN, and stage 2 HTN were also provided for adolescents aged ≥13 years^{7,8}. The term “high-normal BP” in 2016 ESHG was converted to “elevated BP” in the 2017 AAP CPG. The new BP cutoff values for screening and diagnosing pediatric HTN, proposed by the 2017 AAP CPG, differ from those by the ESHG⁷. The cutoffs of SBP and DBP were lowered, whereas a single cutoff of 130/80 mmHg and 140/90 mmHg was recommended to define stage 1 and stage 2 HTN, respectively, in adolescents aged ≥13 years (Table 1). This decision has provoked a large debate not only for the adults’ criteria, as the ESH was determined to maintain the cutoff of 140/90 mmHg, but also for the pediatric criteria, as the ESHG has not been renewed

Table 1. Diagnostic Criteria in Pediatric Hypertension

2017 AAP CPG		2016 ESHG
1–13 years	Children	0–15 years
<90th p	Normal BP	<90th p
≥90th p to <95th p or 120/80 (whichever is lower)	Elevated BP (High-normal BP)	≥90th p to <95th p
≥95th p to <95th p+12 mmHg or 130/80 to 139/89 mmHg (whichever is lower)	Stage 1 HTN	95th p to 99th p+5 mmHg (ISH: SBP ≥95th p & DBP <90th p)
≥95th p+12 mmHg or ≥140/90 mmHg (whichever is lower)	Stage 2 HTN	>99th p+5 mmHg
≥13 years	Adolescents	≥16 years
<120/<80 mmHg	Normal BP	<130/85 mmHg
120/<80 to 129/<80 mmHg	Elevated BP (High-normal BP)	130–139/85–89 mmHg
130/80 to 139/89 mmHg	Stage 1 HTN	140–159/90–99 mmHg (ISH: ≥140/<90 mmHg)
≥140/90 mmHg	Stage 2 HTN	160–179/100–109 mmHg

Abbreviations: AAP CPG, American Academy of Pediatrics Clinical Practice Guideline; ESHG, European Society of Hypertension Guideline; BP, blood pressure; HTN, hypertension; ISH, isolated systolic hypertension; SBP, systolic blood pressure; DBP, diastolic blood pressure; p, percentile.

yet⁹). The difference in the definition of HTN in children and adolescents may cause confusion and contrasting results in both epidemiological research and clinical practice.

In the previous guidelines from 2004, children who are overweight and obese were included. In 2017, the AAP CPG renewed the normative tables for pediatric BP values by excluding overweight and obese children who had a body mass index (BMI) \geq 85th percentile^{10,11}. This resulted in a decrease of 2–3 mmHg in BP values compared to those in the 2004 Fourth Report. Stage 2 HTN for children <13 years was defined as BP values \geq 95th percentile+12 mmHg (or \geq 140/90 mmHg) instead of the 99th percentile+5 mmHg (Table 1). The 2017 AAP CPG applied an adult threshold of 130/80 mmHg to adolescents, both male and female, aged \geq 13 years¹⁰. The absolute BP cutoff points were introduced to follow the updated adult HTN guidelines and to simplify the process of identifying and classifying HTN in adolescents¹². In particular, in view of Korean pediatricians, the new percentile tables are easy to use at office, as height values were added to the table in centimeters, which are used as the unit⁷. In addition to applying a static threshold to adolescents, the 2017 AAP CPG has a new simplified screening table with BP cutoffs based on the 90th percentile BP at the 5th percentile of height for every age for males and females <13 years of age⁷. The goal of this table is not to diagnose elevated BP or HTN by itself but to identify children and adolescents who need further evaluation of their BP with repeat BP measurements.

Recent evidence revealed that the use of the 2017 AAP CPG identified a higher proportion of hypertensive young people with abnormal cardiometabolic risk¹³ or target organ damage (TOD)¹⁴, resulting in an overall increase in the prevalence of HTN^{8,15,16}. The simplified screening table in the 2017 AAP CPG better contributes to recognizing and classifying those with obesity and other CVD risk factors as hypertensive when compared to the 2016 ESHG. According to the screening table, children who have increased BP should be evaluated using the extended percentile table or a cutoff BP value of 120/80 mmHg should be used for children aged \geq 13 years⁷.

Treatment strategies

1. Target blood pressure values

The main goals of treatment for both primary and secondary HTN are to reduce the risk of TOD in childhood and decrease the risk of HTN and CVD in adulthood. The 2017 AAP CPG recommends that the treatment goal of non-pharmacologic and pharmacologic therapy is to reduce SBP and DBP to <90th percentile and <130/80 mmHg in adolescents \geq 13 years old⁷. The mean arterial pressure should be maintained <50th percentile in children with chronic kidney disease (CKD)⁷.

In the 2016 ESHG, a BP <95th percentile for age, sex, and height is recommended for the general hypertensive population. However, if the goal can be accomplished by well-tolerated treatment, a BP <90th percentile should be considered⁶. For children with CKD, strict BP control leads to better long-term renal survival¹⁷. A BP goal of <50th percentile is recommended in children with proteinuric CKD, as proteinuria is an important modifier of the renoprotective effect of intensified BP control⁶. In patients aged \geq 16 years, the cutoff values for office BP were 130/80 mmHg or 125/75 mmHg with proteinuric CKD⁶.

Children with type 1 or 2 diabetes mellitus (DM) are considered to have increased long-term risk for HTN and renal damage^{18,19}. Therefore, it is appropriate for target BP values to be <75th percentile in children with non-proteinuric CKD and <50th percentile in children with proteinuric CKD⁶.

2. Lifestyle interventions

Both the 2017 AAP CPG and the 2016 ESHG initially recommended lifestyle modifications to control BP. Particularly, in overweight and obese children, it has been demonstrated that lifestyle modifications such as a healthy diet, exercise, and behavioral therapy lead to improvements in both weight reduction and BP control²⁰. The 2017 AAP CPG seems to emphasize the Dietary Approach to Stop Hypertension (DASH)-type diet, which consists of a high daily intake of fruits and vegetables, low-fat milk products, and whole grains, and a low intake of added sugar and sweets (including sweetened beverages) and dietary sodium. A reduction in salt (<3 g per day) decreases SBP and DBP by 1.2 and 1.3 mmHg, respectively (Table 2)⁶.

Moderate to vigorous exercise at least 3 to 5 days per week (30–60 minutes per session) helps to reduce BP^{6,7}. Of note, the 2016 ESHG recommends that BMI should be maintained at <85th percentile to prevent becoming overweight⁶. Children with a BMI >95th percentile require gradual weight loss (1–2 kg/month) to achieve a value <85th percentile.

3. Pharmacologic treatment

Despite lifestyle modifications, children who remain hypertensive or who have symptomatic HTN, stage 2 HTN without modifiable risk factors (e.g., obesity), or any stage of HTN associated with CKD or DM therapy should be initiated with pharmacologic treatment⁷. An individual approach can be made in children with high-normal BP if hypertensive TOD is already present⁶.

It is logical to initiate monotherapy that can be administered once a day due to the benefits of simplicity and adherence to administration. After treatment with the lowest dose, BP should be measured at 2–4-week intervals. If the target BP goal is not achieved, the dose should be increased. Adverse effects should also be monitored during dose increments.

There are eight classes of antihypertensive drugs that may potentially be first-line agents, including angiotensin-converting enzyme inhibitors (ACEi), angiotensin receptor blockers (ARBs), beta-blockers, calcium channel blockers (CCBs), and diuretics⁶. Drug choice should be targeted to the child's underlying pathophysiology and the presence of concurrent disorders⁶. Although there are limited

evidence and trials related to antihypertensives used in childhood, and their adverse effects and long-term cardiovascular outcomes, the use of ACEi and ARBs are recommended as first-line agents in populations with obesity-associated primary HTN²¹. CCBs are a reasonable alternative in the absence of treatment effects with ACEi and ARBs, whereas beta-blockers are avoided in terms of safety in children without vasodilatory capacity and thiazide diuretics²². In children with CKD and HTN, proteinuria, or DM, an ACEi or ARB is also recommended as the initial antihypertensive agent, unless there is an absolute contraindication such as pregnancy in adolescence⁷.

The use of combination therapy is recommended if a single agent does not successfully achieve target BP⁶. The second drug to be chosen is diuretics because antihypertensive agents can generally cause water and salt retention. Increased consumption of vegetables and fruits, restricted salt intake, and physical activity should be maintained. No two drugs (ACEi and ARBs) that act separately on the renin-angiotensin system should be used in combination because of the risks of hyperkalemia, acute kidney injury, and hypotension⁶. It is preferred to combine agents from different drug classes and complementary modes of action⁶.

4. Treatment-resistant or refractory HTN

When a therapeutic plan that includes lifestyle modifications and intake of three or more antihypertensive drugs (one diuretic) at maximally effective doses has failed to lower SBP and DBP sufficiently^{23,24}, HTN may be defined as resistant to treatment or refractory^{6,7}. This condition

Table 2. Non-pharmacologic and Pharmacologic Treatment of Pediatric Hypertension

	2017 AAP CPG	2016 ESHG
Lifestyle modifications	DASH-type diet	3 months of exercise training → lowering SBP 7–12 mmHg and DBP 2–7 mmHg
	Moderate to vigorous physical activity: at least 3 to 5 days per week (30–60 min per session)	More than two servings of fruits and vegetables/day → lowering the risk of HTN by 35%
	Motivational interviewing, Stress reduction	3 g/day reduction in salt intake → lowering SBP and DBP of 1.2/1.3 mmHg
Pharmacologic treatment	Indications: lifestyle modifications failure, LVH on echocardiography, symptomatic HTN, stage 2 HTN without a clearly modifiable factor (obesity etc.)	Indications: symptomatic, secondary, organ damage, diabetes, hypertensive emergency and urgency
	ACEi, ARBs, long-acting CCBs, thiazide diuretics	ACEi, ARBs, calcium antagonist, beta blockers, thiazide diuretics
Remarks	Target BP: SBP and DBP to <90th percentile and <130/80 mmHg in younger children	ACEi should not be used with ARBs

Abbreviations: AAP CPG, American Academy of Pediatrics Clinical Practice Guideline; ESHG, European Society of Hypertension Guideline; DASH, dietary approaches to stop hypertension; BP, blood pressure; LVH, left ventricular hypertrophy; HTN, hypertension; SBP, systolic blood pressure; DBP, diastolic blood pressure; ACEi, angiotensin-converting enzyme inhibitors; ARBs, angiotensin receptor blockers; CCBs, calcium channel blockers.

requires precise investigation of the secondary causes of resistant HTN and prompt intervention. Primary glomerulopathies, renovascular disease, coarctation of the aorta, and neurological tumors are the most common causes of secondary HTN in children <6 years of age (Table 3). According to the clinical status, renal Doppler ultrasonography, magnetic resonance imaging, and computed tomography angiography may be helpful in the differential diagnosis. In addition, renin-aldosterone values and serum potassium levels provide important information to help rule out the genetic causes of HTN. Ambulatory BP monitoring may be performed to identify true resistant HTN and confirm the diagnosis⁷⁾.

Treatment includes dietary sodium restriction, avoidance of substances and drugs known to elevate BP, the identification of previously undiagnosed secondary causes of HTN, the optimization of current therapy, and the addition of extended-release drugs at the highest dose as needed^{7,25)}. As there are limited data related to treatment-resistant HTN in children, evaluation and management strategies similar to those effective in adults should be taken⁷⁾.

5. Treatment of monogenic HTN

Monogenic HTN syndromes refer to hypertensive disorders caused by a single gene mutation that follows Mendelian inheritance patterns²⁶⁾. According to the 2017 AAP CPG, monogenic forms of HTN are uncommon because of the lack of exact incidence data⁷⁾. In the 2016 ESHG, the Working Group also stated that monogenic causes of HTN are rare but should be discovered during the pediatric age for successful treatment and avoidance of HTN-related

morbidity and mortality in adulthood⁶⁾.

Although genetic testing is a confirmative tool for diagnosis, monogenic causes of HTN may be suspected in low renin HTN, family history of early onset HTN, death from cerebral vascular accidents and heart failure, and refractory HTN⁶⁾. Early recognition and proper management are crucial because specific antihypertensive agents should be targeted to defective tubular function. Based on the results of genetic testing, treatment may consist of thiazides, amiloride, triamterene, dexamethasone, spironolactone, or eplerenone⁶⁾.

Conclusions

HTN in childhood and adolescence progresses to adulthood HTN, resulting in increased cardiovascular events such as heart disease and stroke. Therefore, efforts should be made to detect children who are at risk of HTN to prevent progression to sustained HTN and to avoid the development of hypertensive CVD. Nonpharmacological and pharmacological treatments should be used in the management of children and adolescents with HTN. Prospective reassessment and ongoing revision are needed to improve the recommendations made in both guidelines with regular updates based on new evidence as it is generated.

Funding

This research did not receive any specific grant from fun-

Table 3. Etiology of Secondary Hypertension

2017 AAP CPG	2016 ESHG
Renal parenchymal disease and renal structural abnormalities	Chronic kidney disease
Renovascular disease	Renovascular HTN
Cardiac, including aortic coarctation	Pheochromocytoma and paraganglioma
Endocrine HTN: catecholamine excess, mineralocorticoid excess, glucocorticoid excess, hyperthyroidism, hyperparathyroidism etc.	Primary aldosteronism
Environmental exposures: lead, cadmium, mercury, phthalates etc.	Cushing's syndrome
Neurofibromatosis	Obstructive sleep apnea
Pheochromocytoma	Coarctation of aorta
Medication-related HTN	Drug-induced HTN
Monogenic HTN	Monogenic causes of HTN Hyperthyroidism and congenital adrenal hyperplasia

Abbreviations: AAP CPG, American Academy of Pediatrics Clinical Practice Guideline; ESHG, European Society of Hypertension Guideline; HTN, hypertension.

ding agencies in the public, commercial, or not-for-profit sectors.

Conflicts of interest

No potential conflict of interest relevant to this article was reported.

ORCID

Se Jin Park, <https://orcid.org/0000-0002-7650-5393>

Jae Il Shin, <https://orcid.org/0000-0003-2326-1820>

References

- National High Blood Pressure Education Program Working Group on High Blood Pressure in Children and Adolescents. The fourth report on the diagnosis, evaluation, and treatment of high blood pressure in children and adolescents. *Pediatrics* 2004;114:555-76.
- Rademacher ER, Jacobs DR Jr, Moran A, Steinberger J, Prineas RJ, Sinaiko A. Relation of blood pressure and body mass index during childhood to cardiovascular risk factor levels in young adults. *J Hypertens* 2009;27:1766-74.
- Falkner B, Lurbe E. Primary hypertension beginning in childhood and risk for future cardiovascular disease. *J Pediatr* 2021;238:16-25.
- Falkner B, Gidding S. Life-course implications of pediatric risk factors for cardiovascular disease. *Can J Cardiol* 2021;37:766-75.
- Wühl E. Hypertension in childhood obesity. *Acta Paediatr* 2019;108:37-43.
- Lurbe E, Agabiti-Rosei E, Cruickshank JK, Dominiczak A, Erdine S, Hirth A, et al. 2016 European Society of Hypertension guidelines for the management of high blood pressure in children and adolescents. *J Hypertens* 2016;34:1887-920.
- Flynn JT, Kaelber DC, Baker-Smith CM, Blowey D, Carroll AE, Daniels SR, et al. Clinical practice guideline for screening and management of high blood pressure in children and adolescents. *Pediatrics* 2017;140:e20171904.
- Tran AH, Urbina EM. Hypertension in children. *Curr Opin Cardiol* 2020;35:376-80.
- Bonito PD, Licenziati MR, Baroni MG, Maffei C, Morandi A, Manco M, et al. The American Academy of Pediatrics hypertension guidelines identify obese youth at high cardiovascular risk among individuals non-hypertensive by the European Society of Hypertension guidelines. *Eur J Prev Cardiol* 2020;27:8-15.
- Blanchette E, Flynn JT. Implications of the 2017 AAP clinical practice guidelines for management of hypertension in children and adolescents: a review. *Curr Hypertens Rep* 2019;21:35.
- Rosner B, Cook N, Portman R, Daniels S, Falkner B. Determination of blood pressure percentiles in normal-weight children: some methodological issues. *Am J Epidemiol* 2008;167:653-66.
- Whelton PK, Carey RM, Aronow WS, Casey DE Jr, Collins KJ, Himmelfarb CD, et al. 2017 ACC/AHA/AAPA/ABC/ACPM/AGS/APhA/ASH/ASPC/NMA/PCNA guideline for the prevention, detection, evaluation, and management of high blood pressure in adults: executive summary: a report of the American College of Cardiology/American Heart Association task force on clinical practice guidelines. *Circulation* 2018;138:e426-83.
- Sharma AK, Metzger DL, Rodd CJ. Prevalence and severity of high blood pressure among children based on the 2017 American Academy of Pediatrics guidelines. *JAMA Pediatr* 2018;172:557-65.
- Khoury M, Khoury PR, Dolan LM, Kimball TR, Urbina EM. Clinical implications of the revised AAP pediatric hypertension guidelines. *Pediatrics* 2018;142:e20180245.
- Dong Y, Song Y, Zou Z, Ma J, Dong B, Prochaska JJ. Updates to pediatric hypertension guidelines: influence on classification of high blood pressure in children and adolescents. *J Hypertens* 2019;37:297-306.
- Al Kibria GM, Swasey K, Sharmeen A, Day B. Estimated change in prevalence and trends of childhood blood pressure levels in the United States after application of the 2017 AAP guideline. *Prev Chronic Dis* 2019;16:E12.
- ESCAPE Trial Group, Wühl E, Trivelli A, Picca S, Litwin M, Peco-Antic A, et al. Strict blood-pressure control and progression of renal failure in children. *N Engl J Med* 2009;361:1639-50.
- Raile K, Galler A, Hofer S, Herbst A, Dunstheimer D, Busch P, et al. Diabetic nephropathy in 27,805 children, adolescents, and adults with type 1 diabetes: effect of diabetes duration, A1C, hypertension, dyslipidemia, diabetes onset, and sex. *Diabetes Care* 2007;30:2523-8.
- Lurbe E, Redon J, Kesani A, Pascual JM, Tacons J, Alvarez V, et al. Increase in nocturnal blood pressure and progression to microalbuminuria in type 1 diabetes. *N Engl J Med* 2002;347:797-805.
- Ho M, Garnett SP, Baur L, Burrows T, Stewart L, Neve M, et al. Effectiveness of lifestyle interventions in child obesity: systematic review with meta-analysis. *Pediatrics* 2012;130:e1647-71.
- Elliott WJ, Meyer PM. Incident diabetes in clinical trials of antihypertensive drugs: a network meta-analysis. *Lancet* 2007;369:201-7.
- Mancia G, Grassi G, Zanchetti A. New-onset diabetes and antihypertensive drugs. *J Hypertens* 2006;24:3-10.
- Mancia G, Fagard R, Narkiewicz K, Redón J, Zanchetti A, Böhm M, et al. 2013 ESH/ESC guidelines for the management of arterial hypertension: the task force for the management of arterial hypertension of the European Society of Hypertension (ESH) and of the European Society of Cardiology (ESC). *J Hypertens* 2013;31:1281-357.

24. Macumber I, Flynn JT. Does treatment-resistant hypertension exist in children? A review of the evidence. *Pediatr Nephrol* 2020; 35:969-76.
25. White WB, Turner JR, Sica DA, Bisognano JD, Calhoun DA, Townsend RR, et al. Detection, evaluation, and treatment of severe and resistant hypertension: proceedings from an American Society of Hypertension Interactive Forum held in Bethesda, MD, U.S.A., October 10th 2013. *J Am Soc Hypertens* 2014;8:743-57.
26. Ahn SY, Gupta C. Genetic programming of hypertension. *Front Pediatr* 2018;5:285.