

# Hypertension: JNC-6 (Archive)

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Physician Assistant Program

Management and Treatment of Hypertension  
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## HYPERTENSION (NIH Study)

### I. Introduction

A. NIH Study: since its inception (1972) progress in the detection, treatment, and control of HTN has been remarkable

- in the last 2 decades, the following observations were reported:
  - (1) number of patients aware of their hypertensive condition has increased dramatically
  - (2) the percentage of hypertensive patients taking medication and controlling their HTN improved significantly
  - (3) mortality from coronary heart disease (CHD) decreased by approx. 50%
  - (4) mortality from stroke decreased by 57%
- progress in the detection, treatment, and control of HTN has significantly contributed to the reported decreases in mortality rates for CHD and stroke since HTN is a major risk factor for CHD and the most important risk factor for stroke

### B. Clinical Evaluation

- 50 million Americans have elevated BP:
  - SBP  $\geq$  140 mm Hg and/or
  - DBP  $\geq$  90 mm Hg
- the prevalence of HTN:
  - increases with age
  - is greater for blacks than for whites
  - is greater in less educated people (both races, whites and blacks)
  - is much greater in lower socioeconomic groups
  - is greater in men than women (in young adulthood and early middle age)
  - is greater in women than men (after middle age)
  - is greater in people living in the southeastern U.S. than people living in other areas of the country

## B. Clinical Evaluation (continued)

- risks for cardiovascular diseases (CVDs), including CHD and stroke, are lowest for adults with average SBP < 120 mm Hg and average DBP < 80 mm Hg
- at every level of DBP, risk of CVDs are greater with higher levels of SBP
- in middle-aged and older people, elevated SBP with normal DBP is associated with increased risk of CVD → "isolated systolic hypertension"

## C. New Classification of High Blood Pressure

(1) high-normal BP is a new category (SBP=130-139 / DBP=85-89)

- people with SBP and/or DBP in this range are at increased risk of progressing to high BP → increased risk of experiencing non-fatal and fatal CVDs
- people with high-normal BP should be:
  - monitored frequently
  - counseled about life-style measures → reduce BP
- pharmacologic treatment is rarely needed in people with high normal BP

(2) table 2: Classification of BP for Adults (18 Years and Older)

### Classification of Blood Pressure for Adults Aged 18 Years & Older

Category	Systolic (mm Hg)	Diastolic (mm Hg)
Normal	< 130	< 85
High Normal	130-139	85-89
Hypertension		
Stage 1 (mild)	140-159	90-99
Stage 2 (moderate)	160-179	100-109
Stage 3 (severe)	180-209	110-119
Stage 4 (very severe)	> or = 210	> or = 120

## (2) table 2: Classification of BP for Adults (continued)

- all stages of HTN are associated with increased risk of nonfatal & fatal CVDs and renal disease
- the higher the BP, the greater the risk of CVDs
- stage 1: (previously termed "mild" HTN)
  - most common form of HTN
  - responsible for a major proportion of the incidence of morbidity, disability, and mortality associated with HTN
- all stages of HTN require long-term therapy

## D. Cost of Drugs

- generic drugs → reduced drug costs
- new drugs used in HTN are expensive (up to 30 times more than generic diuretics and beta blockers)
- drug dosages may be reduced with patient education and counseling for diet and weight control → decreased drug costs
- dividing a scored tablet in half for the prescribed dose → cost savings

## E. Treatment of HTN

## (1) Goal

- the ultimate goal in treating HTN is to prevent morbidity and mortality
  - objectives:
    - a. control BP by the least intrusive method possible → life-style modification
    - b. maintain:                    SBP < 140    and    DBP < 90

**or**

SBP < 135    and    DBP < 85 (if possible)

## (2) Life-Style Modification (previously termed "nonpharmacologic therapy")

## a. lose wt if overweight

- wt reduction decreases BP in most pts who are 10% above ideal wt

$$\text{IBW} = (2.3 \times \text{ht in inches above 5'}) + 45.5 \text{ kg} \quad (\text{female})$$

$$\text{IBW} = (2.3 \times \text{ht in inches above 5'}) + 50 \text{ kg} \quad (\text{male})$$

## b. limit alcohol intake to &lt; or = 1 ounce of alcohol (24 oz of beer, 8 oz of wine, or 2 oz of 100 proof whiskey)

## c. exercise

## d. reduce sodium intake &lt; 100 mmol/d (&lt; 6 gm of NaCl)

## e. maintain adequate dietary K, Ca, and Mg

- hypokalemia → may increase BP and induce ventricular ectopy

→ important in patients on diuretics

- hypocalcemia → associated with increased BP (in some studies)

- there is no rationale for increasing Ca intake in excess of the recommended daily allowance (800-1200 mg)

- hypomagnesemia → "suggested" association with increased BP

- there is no conclusive data to justify Mg supplementation to lower BP

## f. stop smoking

## g. reduced dietary saturated fat and cholesterol intake

## F. Pharmacologic Treatment of HTN

### (1) General Considerations

- the decision to initiate drug therapy in patients with HTN requires the following considerations:
  - a. severity of the HTN
  - b. TOD (target-organ disease)
  - c. presence of other conditions and risk factors

table 3: **Manifestation of Target-Organ Disease**

<b>Organ System</b>	<b>Manifestations</b>
Cardiac	Clinical, electrocardiographic, or radiologic evidence of coronary artery disease; left ventricular hypertrophy or "strain" by EKG or left ventricular hypertrophy by EKG; left ventricular dysfunction or cardiac failure
Cerebrovascular	Transient ischemic attack or stroke
Peripheral Vascular	Absence of 1 or more major pulses in extremities (except for dorsalis pedis) with or without intermittent claudication; aneurysm
Renal	Serum creatinine $\geq$ or = 1.5 mg/dL; proteinuria (1+ or greater); microalbuminuria
Retinopathy	Hemorrhages or exudates, with or without papilledema

### (2) Pharmacologic Efficacy

- reducing BP with drugs  $\rightarrow$  decreases cardiovascular mortality and morbidity  $\rightarrow$  decreases incidence of stroke, coronary events, CHF, more severe HTN, and all-cause mortality
- studies indicate that:
  - 42% reduction in stroke from a 5 to 6 mm Hg drop in DBP
  - 20 - 25 % reduction in the rate of CHDs

### (3) Stage 1 and Stage 2 Hypertension

- if BP remains at 140/90 during a 3 - 4 month period → pharmacologic tx (after encouragement of life-style modifications)
- if DBP = 90 - 94 and SBP = 140 - 149 → withhold drug tx in the absence of TOD & risk factors → follow-up at 3 to 6 month intervals (since BP may increase)

### (4) Initial Drug Therapy

- initial drug therapy (stage 1 and stage 2) → monotherapy
  - a. diuretics and beta blockers → preferred for initial drug therapy
    - clinical trials have shown diuretics and beta blockers reduce cardiovascular M/M
  - b. Alternative Drugs
    - calcium channel blockers, ACE inhibitors, alpha-1 blockers, and alpha-beta blockers
      - equally effective in reducing BP
      - have not been shown in long-term clinical trials to reduce M/M
      - should be used when diuretics and beta blockers have been ineffective or unacceptable (d/t side effects, etc...)
  - c. Supplemental Antihypertensives (not for initial monotherapy)
    - direct-acting smooth muscle vasodilators (hydralazine, minoxidil)
      - induce reflex sympathetic stimulation of the cardiovascular system
      - cause fluid retention
    - alpha-2 agonist
      - cause annoying side effects in many patients
    - peripheral adrenergic neuron antagonists
      - cause annoying side effects in many patients

## d. Special Considerations

### i. Demographic Characteristics

- blacks → more responsive to diuretics & Ca channel blockers than beta blockers or ACE inhibitors
- geriatrics → responsive to all classes of drugs
- gender → no special determinations
- NOTE: issues concerning race, age, or gender are not reasons for avoiding any drug class especially if a particular drug offers other therapeutic benefits since efficacy differences may be accomplished by the addition of a diuretic or another drug

### ii. Concomitant Diseases and Therapies

- anti-HTN drugs may exacerbate some disease states while improving others:
  - beta blockers improve
    - angina
    - cardiac dysrhythmias
    - migraine headaches
    - prolong life after a MI
  - beta blockers worsen
    - asthma
    - diabetes
    - peripheral ischemia
- selecting an anti-HTN drug that treats a coexisting disease state
  - simplifies drug regimen
  - reduces side effects
  - reduces drug costs

### iii. Quality of Life → Side Effects

- anti-HTN drugs may cause undesirable side effects:
  - central acting drugs → drowsiness & sedation
  - beta blockers → reduce exercise tolerance



#### iv. Physiologic Factors

- body wt, heart rate, plasma renin activity, hemodynamic measures → helpful in choosing a specific agent

high renin levels → ACE inhibitor

#### v. Economic Considerations

- cost of drugs → may be a barrier to controlling HTN

#### e. Dosage and Follow-up

- i. the lowest dosage of initial drug should be used even though it may not immediately control BP → avoidance of side effects
- ii. the lowest dose should be given for several weeks before increasing the dosage of the drug → time required to assess efficacy and side effects
- iii. a drug given once or twice daily → improves patient compliance → controls BP
- iv. after 1 to 3 months, if response to initial therapy is (1) adequate, (2) pt is not experiencing significant side effects, and (3) patient compliance is adequate, then 3 options may be considered:
  1. increase the dose of the first drug
  2. substitute an agent from another class
  3. add a second drug from another class
    - combining antihypertensives with different mechanisms of action → lower doses of drugs used to achieve BP control → decreases dose-dependent side effects
    - addition of a 2<sup>nd</sup> agent usually enhances the effects of the 1<sup>st</sup> drug
      - example: adding a diuretic enhances the effects of other agents
    - if addition of a 2<sup>nd</sup> drug results in adequate BP control, an attempt to discontinue the 1<sup>st</sup> drug may be worthwhile since monotherapy will control BP for at least 50% of all patients
    - after achieving a level of BP control, substituting comparable combination tablets may simplify patients' drug regimens and promote compliance

## v. Causes of Lack of Responsiveness to Therapy

- clinicians should assess the possible reasons for lack of responsiveness to drug therapy before proceeding to the next treatment step

### (5) Stage 3 and Stage 4 Hypertension

(patients with DBP  $\geq$  or = 110 and/or SBP  $\geq$  or = 180)

- patients in stage 3 and 4 may respond adequately to monotherapy; however, a 2<sup>nd</sup> or 3<sup>rd</sup> agent is usually necessary for BP control
  - intervals between regimen changes may be decreased for BP control
  - maximum doses of some drugs may be increased for BP control
- some patients may require initial treatment with more than one drug for adequate antihypertensive effects
- patients with DBP  $\geq$  120 with significant TOD may require hospitalization

### (6) Isolated Systolic Hypertension (ISH)

#### a. Adolescents and Young Adults

- ISH  $\rightarrow$  often indicates hyperdynamic circulation which may be predictive of future elevation of DBP
- if life style-modifications ineffective, drug therapy should be considered

#### b. Older Persons

- ISH  $\rightarrow$  SBP  $\geq$  or = 160 and DBP  $<$  90  $\rightarrow$  drug therapy if life-style modifications ineffective

### (7) Step-Down Therapy

- after BP has been controlled for 1 year (with at least 4 office visits), an attempt to reduce antihypertensive drug therapy in a gradual, progressive manner is recommended
- step-down therapy is particularly successful in patients who are also practicing life-style modifications
- regular follow-up should be arranged after discontinuing antihypertensive drugs since BP usually increases to hypertensive levels months or years after discontinuation especially if life-style modifications were discontinued

### (8) J-Curve Hypothesis

- concerns about lowering DBP with antihypertensive drugs has raised the J-curve hypothesis:
  - low DBP ( $< 90$ )  $\rightarrow$  low diastolic perfusion pressure in coronary circulation  
 $\rightarrow$  risk of coronary disease
- this hypothesis may be more important in patients with preexisting CAD
- currently, there is no evidence to support the J-curve hypothesis and clinical trials have supported the reduction of DBP to less than 90 mm Hg

### (9) Resistant Hypertension

a. HTN is considered resistant if a patient's BP:

- i. cannot be reduced to less than 160/100 by a triple-drug regimen given in maximal doses
- ii. pretreatment BP was  $> 180 / 115$

**or**

- i. cannot be reduced to less than 140 / 90 by a triple-drug regimen given in maximal doses
- ii. pretreatment BP was  $< 180 / 115$

b. isolated systolic HTN is considered resistant if an older patient's SBP:

- i. cannot be reduced to less than 170 mm Hg
- ii. pretreatment SBP was  $> 200$  mm Hg

**or**

- i. cannot be reduced to less than 160 mm Hg & by at least 10 mm Hg
- ii. pretreatment SBP was 160 - 200 mm Hg

### (10) Follow-up Visits

- a. goal of drug therapy: achieving and maintaining target BP with the lowest possible drug doses → requires ongoing follow-up for dosage adjustments
- b. patients with stage 1 HTN without TOD
  - i. should be seen within 1 to 2 months after start of therapy to assess BP control, patient compliance, and presence of side effects
  - ii. once BP is controlled, follow-up at 3- to 6- month intervals

### (11) Strategies to Improve Adherence to Therapy

- poor patient compliance to long-term tx (life-style modifications and pharmacologic therapy) → inadequate control of BP → increased HTN-related M/M
- guidelines to improve adherence to treatment:
  - a. simplify the drug regimen
  - b. include patient in the decision making process
  - c. incorporate the treatment into the patient's daily life -style
  - d. set realistic short-term objective of the treatment plan with the patient
  - e. encourage discussion of side effects
  - f. minimize drug costs
  - g. indicate that you will ask about compliance at the next office visit

## Pharmacologic Treatment of Hypertension

