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# RESISTANT ARTERIAL HYPERTENSION: DIAGNOSTIC APPROACHES AND MULTIMODAL TREATMENT STRATEGIES

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#### **Annotation**

In this article, resistant hypertension is discussed as a condition affecting around 10 to 15 percent of individuals with high blood pressure, carrying a high risk of heart and vascular complications. Diagnostic attention is given to confirming that patients take their medications, excluding elevated readings due to clinical settings, and identifying secondary causes. Treatment includes changes in lifestyle, the use of scientifically supported drug combinations, and new procedures such as nerve-related kidney treatments. The updated 2024 recommendations by the European Society of Cardiology highlight personalized care and the use of blood pressure monitoring outside clinical settings. The article presents practical guidance for healthcare professionals based on the latest diagnostic and treatment approaches.

**Keywords:** resistant hypertension, ambulatory blood pressure monitoring, antihypertensive therapy, renal denervation, secondary hypertension, cardiovascular risk, medication adherence, lifestyle interventions

Today's, the growing clinical importance of resistant arterial hypertension is addressed as a major challenge in modern cardiovascular care. This condition is defined by the failure to control blood pressure despite adherence to a full regimen of at least three antihypertensive medications, including diuretics, at optimal doses. Patients with resistant hypertension face significantly higher risks of heart attack, stroke, heart failure, and death compared to those with controlled hypertension. Epidemiological data indicate that 10 to 15 percent of hypertensive individuals globally-nearly 100 million people-are affected. Recent advances such as ambulatory and home blood pressure monitoring have helped distinguish true resistance from pseudo-resistance due to white coat effects or nonadherence. Additionally, new treatment options, including device-based therapies like renal denervation and innovative medications, offer promising solutions. The 2024 guidelines from the European Society of Cardiology emphasize a patient-centered approach, improved diagnostic methods, and the

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routine use of out-of-office monitoring, representing a shift toward more personalized and comprehensive hypertension management.

### **MAIN PART**

Accurate diagnosis of resistant arterial hypertension requires thorough evaluation of contributing factors, including confirming medication adherence and excluding white-coat hypertension. Proper blood pressure measurement techniques with calibrated devices and standardized protocols are essential, involving multiple readings taken after the patient rests seated for five minutes. The 2024 European Society of Cardiology guidelines strongly recommend out-of-office blood pressure monitoring, with ambulatory monitoring considered the gold standard. This method helps confirm true resistance by eliminating white-coat effects and provides valuable information on blood pressure patterns such as nocturnal hypertension and morning surges, important for cardiovascular risk assessment.

Home blood pressure monitoring serves as an accessible alternative to ambulatory monitoring, providing valuable information about blood pressure control in the patient's natural environment. Patients should be trained in proper measurement techniques and encouraged to maintain detailed blood pressure logs that include timing of measurements, concurrent medications, and relevant clinical symptoms. Medication adherence assessment represents a critical component of resistant hypertension diagnosis, as poor adherence accounts for a significant proportion of apparent treatment resistance. Clinical evaluation should include direct patient interviews regarding medication-taking behaviors, pill counting, pharmacy refill records, and consideration of biochemical adherence monitoring when available. Healthcare providers must approach adherence assessment in a non-judgmental manner, acknowledging the multiple barriers that patients may face in maintaining optimal medication regimens. Investigation of secondary causes of hypertension becomes paramount in patients with confirmed resistant hypertension, as the prevalence of secondary hypertension approaches 30 percent in this population compared to 5-10 percent in the general hypertensive population. Systematic screening should include assessment for primary aldosteronism through aldosterone-to-renin ratio measurement, evaluation for renovascular disease using duplex ultrasonography or magnetic resonance angiography, investigation of sleep-disordered breathing through sleep studies, and assessment for pheochromocytoma or paraganglioma in appropriate clinical contexts. Laboratory evaluation should encompass comprehensive metabolic panels including serum electrolytes, kidney function markers, lipid profiles, and glucose metabolism assessment. Electrocardiographic examination and echocardiography

provide valuable information regarding target organ damage, particularly left ventricular hypertrophy and cardiac function assessment. Additional imaging studies, including renal ultrasonography and fundoscopic examination, help identify hypertensive complications and guide therapeutic decision-making.

Contemporary management of resistant arterial hypertension implementation of comprehensive, multimodal treatment approaches that address the complex pathophysiological mechanisms underlying therapeutic resistance. Effective treatment of resistant hypertension includes ongoing lifestyle modifications and collaboration with patients to detect and address barriers to optimal medication adherence. This holistic approach recognizes that successful blood pressure control requires integration of multiple therapeutic modalities rather than reliance on pharmacological interventions alone. Lifestyle modification strategies form the cornerstone of resistant hypertension management and must be emphasized throughout the treatment continuum. Dietary interventions, particularly sodium restriction to less than 2.3 grams daily and adoption of the Dietary Approaches to Stop Hypertension eating pattern, demonstrate significant blood pressure lowering effects that complement pharmacological therapy. Weight reduction through caloric restriction and increased physical activity provides substantial cardiovascular benefits, with each kilogram of weight loss associated with approximately 1 millimeter of mercury reduction in systolic blood pressure. Regular aerobic exercise, performed for at least 150 minutes weekly at moderate intensity, produces both acute and chronic blood pressure lowering effects through multiple mechanisms including improved endothelial function, reduced sympathetic nervous system activity, and enhanced insulin sensitivity. Resistance training, when performed safely under appropriate supervision, provides complementary cardiovascular benefits and may be particularly beneficial for patients with concurrent metabolic disorders. Pharmacological optimization represents the primary therapeutic intervention for resistant hypertension management, requiring systematic approach to drug selection, dosing, and combination therapy. Resistant hypertension is defined as blood pressure above goal despite confirmed adherence to 3 firstline antihypertensive agents or when blood pressure is controlled with 4 or more medications at maximal or maximally tolerated doses. The foundation of pharmacological therapy typically includes an angiotensin-converting enzyme inhibitor or angiotensin receptor blocker, a calcium channel blocker, and a thiazide or thiazide-like diuretic, each administered at maximum tolerated doses. When initial three-drug combinations fail to achieve target blood pressure levels, fourth-line agents should be considered based on individual patient characteristics and

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comorbidities. Spironolactone, a mineralocorticoid receptor antagonist, represents the preferred fourth-line agent for most patients due to its proven efficacy in resistant hypertension and relatively favorable side effect profile. Alternative fourth-line options include beta-blockers, particularly in patients with concurrent coronary artery disease or heart failure, and alpha-blockers for patients with benign prostatic hyperplasia.

Novel pharmacological approaches continue to emerge, with particular interest in dual combination tablets that improve medication adherence through reduced pill burden. Fixed-dose combinations containing complementary agents with different mechanisms of action provide convenient dosing regimens while maintaining therapeutic efficacy. The selection of specific combination therapies should be individualized based on patient characteristics, comorbidities, and potential drug interactions. Treatment of secondary causes of hypertension requires specialized therapeutic approaches tailored to the underlying pathophysiology. Primary aldosteronism management may include surgical intervention for unilateral adenomas or medical management with mineralocorticoid receptor antagonists for bilateral hyperplasia. Renovascular hypertension treatment considerations include percutaneous revascularization for appropriate anatomical lesions, while sleep-disordered breathing requires continuous positive airway pressure therapy or alternative respiratory support modalities. The 2024 European Society of Cardiology guidelines introduce renal denervation as a validated option for resistant hypertension, showing promising results in recent trials with blood pressure lowering by an average of 10-12 millimeters of mercury. This minimally invasive procedure involves catheter-based radiofrequency or ultrasound energy delivery to ablate sympathetic nerve fibers surrounding the renal arteries, resulting in sustained blood pressure reduction through modulation of sympathetic nervous system activity. Devicebased therapies represent an emerging frontier in resistant hypertension management, with renal denervation demonstrating the most robust clinical evidence to date. Patient selection for renal denervation requires careful consideration of anatomical suitability, absence of significant renal artery stenosis, and comprehensive evaluation of alternative therapeutic options. The procedure should be performed by experienced interventional specialists in centers with appropriate expertise and infrastructure.

The management of resistant hypertension has evolved significantly with the 2024 European Society of Cardiology guidelines, which prioritize patient-centered, comprehensive strategies. These guidelines address diagnostic and therapeutic gaps by emphasizing early detection, lifestyle modification, and the consideration of socioeconomic factors. Out-of-office

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blood pressure monitoring now holds a Class I recommendation, marking a shift from reliance on office-based readings. Personalized risk assessment and shared decision-making are central to updated treatment algorithms. Technological advances, such as telemedicine, remote monitoring, and mobile health applications, are increasingly recognized as supportive tools to improve adherence, education, and clinical monitoring without increasing healthcare costs. In pediatrics, resistant hypertension requires tailored diagnostic and treatment approaches. The 2023 European Society of Hypertension guidelines apply percentile-based criteria for children under 16, transitioning to adult criteria afterward. Most pediatric cases stem from secondary causes and require specialist evaluation. Lifestyle changes remain the first-line treatment, with pharmacotherapy used cautiously, considering developmental impacts. The therapeutic landscape continues to expand with novel pharmacological agents, such as dual angiotensin receptor-neprilysin inhibitors and sodium-glucose cotransporter-2 inhibitors, showing promise for resistant cases, especially in patients with diabetes or heart failure. Device-based treatments Genetic like carotid baroreceptor stimulation are also under investigation. pharmacogenomic approaches are paving the way for personalized therapy by identifying patient-specific drug responses. Additionally, artificial intelligence and machine learning may enhance care through predictive analytics and individualized treatment planning.

In conclusion, resistant arterial hypertension is a complex condition requiring accurate diagnosis, systematic evaluation, and evidence-based, personalized treatment. Effective management includes lifestyle changes, optimized medication use, and, when appropriate, advanced device-based therapies. The 2024 European guidelines emphasize out-of-office blood pressure monitoring and patient-centered care. Collaboration between healthcare teams and patients is essential for long-term success. Future progress will rely on technological innovation, personalized medicine, and continued research to reduce cardiovascular risks and improve patient outcomes.