Challenges and Opportunities to Scale Up Cardiovascular Disease Secondary Prevention in Latin America and the Caribbean



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Cardiovascular disease (CVD) is the leading cause of death throughout the world; however, a reduction of 21% (age-standardized cardiovascular mortality rates per 100,000 inhabitants) was observed between 1990 and 2010, with more substantial reductions in CVD mortality evident in high-income countries (~42% reduction in CVD deaths) (Table 1) [1,2].

CARDIOVASCULAR DISEASE BURDEN IN LATIN AMERICA AND THE CARIBBEAN

In the Americas, CVD represents about 38% of noncommunicable disease (NCD) deaths and is the leading cause of death, with 1.6 million deaths per year, one-half million occurring before age 70 [3]. Although mortality due to CVD in the Americas declined 19.2% from 2000 to 2007, higher median CVD mortality was found in lowincome (242 per 100,000) and medium-income (186 per 100,000) countries, compared with high-income countries (154 per 100,000) [4]. Between 2000 and 2010, premature CVD mortality (<70 years of age) in the Americas decreased by 21%, with a statistically significant 2.5% average annual decline in the rate of change in the final 5 years. Declines in mortality from ischemic vascular disease (overall, 25%; men, 24%; women, 26%) and cerebrovascular disease (overall, 27%; men, 26%; women, 28%) were observed during the first decade of this century. These declines were not uniform across all countries, with notable disparities in country-specific premature mortality rates (region average, 132.7 per 100,000), including rates above 200 per 100,000 in Guyana, Trinidad and Tobago, the Dominican Republic, Bahamas, and Brazil [5]. Although, substantial decreases in CVD mortality rates have been seen in most regions, including the Americas from 1990 to 2013, the CVD burden, and specifically the absolute number of cases and deaths have risen in some but not all countries of the region [6]. CVD disproportionately affects the poorest population sectors [7], affecting families, communities, and governments due to treatment costs and potential years of life and productivity lost due to premature death and disability [8,9]. Therefore, the direct and indirect costs of these diseases absorb resources that could otherwise be spent in health promotion efforts at the community and clinical levels [7].

In terms of the global economic burden, direct and indirect costs of CVD were estimated to be US\$863 billion in 2010, potentially rising to US\$1.04 trillion by 2030 [10]. Even countries such as Brazil, Chile, Mexico, Argentina, and Peru, which experienced some economic growth in recent years, lack economic assets to efficiently provide resources to prevent and manage CVD [11]. A comprehensive approach to CVD prevention and management is needed for long-term improvements in cardiovascular health. This report focuses on efforts to improve secondary prevention using existing frameworks, tailored to the Latin American and Caribbean (LAC) context.

THE FRAMEWORK OF CVD SECONDARY PREVENTION

Secondary prevention of CVD can be defined as the use of intensive risk-reduction therapies and adoption of healthy lifestyle behaviors in patients with established coronary and/or other atherosclerotic vascular diseases. This would include peripheral artery disease, atherosclerotic aortic disease, and carotid and cerebral artery disease with clinical benefits of 75% estimated efficacy. Efficacy with these outcomes has been assessed by a variety of outcomes, including improved survival, reduced recurrent events, the need for revascularization procedures, and improved quality of life [12-14]. Risk-reduction therapies and healthy lifestyle changes include pharmacological and nonpharmacologic interventions. Examples of pharmacologic interventions for these populations include antiplatelet agents, beta-blockers, angiotensin-converting enzyme (ACE) inhibitors or angiotensin-receptor blockers (ARB), 3-hydroxy-3-methy-glutaryl-coenzymeA reductase inhibitors (e.g., statins), and pharmaceutical therapy for tobacco cessation. Examples of nonpharmacological interventions include smoking cessation counseling, physical activity, and healthy diet. Both pharmacologic and nonpharmacologic interventions aim to reduce the probability

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TABLE 1. Changes in age-standardized all CVD, IHD, and CVD mortality rate and DALY by country group and selected countries (1990 to 2010)

of recurrent cardiovascular events in patients with clinically evident atherosclerotic cardiovascular disease [15]. Moreover, clinical practice guidelines for the management of care with appropriate updates have been periodically published by several medical societies to guide physician prescription of proven, effective treatments in comprehensive CVD care [12,16-23]. Optimal risk factor management through nonpharmacologically based approaches (Table 2) may include tobacco cessation counseling, regular participation in physical activity, achieving a healthy diet, limited alcohol intake, and weight management; with clinical outcome improvements in blood pressure control, lipid management, and diabetes control [12,15,24-28]. Although these interventions have been shown to be effective, their uptake by patients can be challenging, and clinical and public health programs are encouraged to adapt resources to the culture and capability of those they serve. Programs addressing these areas are often composed of a multidisciplinary team of providers and should include individual level interventions using targeted education and behavior modification programs to counterbalance the "continuum" of the atherosclerotic process. Populationwide interventions (e.g., limiting exposure to secondhand tobacco smoke, sodium reduction) to reduce exposures to risk factors should complement individualized interventions that support risk factor management.

CVD SECONDARY PREVENTION IN LATIN AMERICA AND THE CARIBBEAN: A SITUATION ANALYSIS

A large body of robust evidence to guide therapeutic decision making and enhance long-term adherence after an acute episode of CVD, such as ST-segment elevation myocardial infarction, non-ST-segment elevation acute coronary syndromes, or revascularization procedures exists. Despite this, we frequently observe substantial discrepancies between the evidence-based recommendations and what is actually delivered in clinical practice. This can be appropriately documented based on contemporary realworld data on the use of beneficial secondary prevention medications among several geographic regions, including LAC. Examples include clinical as well as communitybased studies. The REACH (REduction of Atherothrombosis for Continued Health) registry included data from North America, Europe, Australia, Asia, Latin America (Brazil, Chile, Mexico, Panama, Costa Rica, Dominican Republic, Ecuador, Guatemala, and Peru), and the Middle East. The use of secondary prevention medications was ascertained among 32,247 patients. Among these patients, a significant gap was detected between clinically indicated and actual rates of use at the first visit assessment and after 3 years of follow-up: aspirin alone (baseline: 56.6% and 3year follow-up: 56.9%); aspirin plus another antiplatelet agent (14.5% and 12.8%); other antiplatelet agents alone (13.6% and 14.2%); and statins (68.3% and 71.9%) [29]. Compared with patients in North America in the REACH registry, patients from Latin America and Asia were less

TABLE 2. Recommendations for secondary prevention as AHA/ACC guideline, 2011

Area for Intervention	Goal
Smoking	Complete cessation. No exposure to environmental tobacco smoke.
Blood pressure control	<140/90 mm Hg
Lipid management	Treatment with statin therapy; use statin therapy to achieve an LDL-C of <100 mg/dl; for very high risk patients an LDL-C <70 mg/dl is reasonable; if triglycerides are \geq 200 mg/dl, non—HDL-C should be <130 mg/dl, whereas non—HDL-C <100 mg/dl for very high-risk patients is reasonable.
Physical activity	At least 30 min, 7 days per week (minimum 5 days per week)
Weight management	Body mass index: 18.5 to 24.9 kg/m ²
	Waist circumference: women $<$ 35 inches ($<$ 89 cm), men $<$ 40 inches ($<$ 102 cm)
Type 2 diabetes mellitus management	Recommendations are for prevention of cardiovascular complications, based on lifestyle modification, weight management, blood pressure control, lipid management, and pharmacotherapy for all diabetes patients.
Medications based on evidence	Must be considered based on clinical condition of the patient, but the combination of them shoul be applied to all patients on secondary prevention: antiplatelet agents/anticoagulants; reninangiotensin-aldosterone system blockers; ACE inhibitors; ARB; aldosterone blockade; betablockers.
Influenza vaccination	Should be applied to all patients with cardiovascular disease an annual influenza vaccination program.
Depression	It is reasonable to screen for depression if patients have access to case management, in collaboration with their primary care physician and a mental health specialist.
Cardiac rehabilitation	All patients with ACS or postintervention procedures should be referred to a cardiac rehabilitation program. Home-based programs have similar results as compared with supervised, center-based program for low-risk patients.

likely to be fully adherent to guideline-based treatments at 4 years [30]. Data from the large community-based PURE (Prospective Urban Rural Epidemiology) study have provided relevant assessment on the use of secondary prevention therapies across broad geographic and income levels. The use of effective secondary preventive medications (antiplatelet agents, beta-blockers, ACE inhibitors or ARB, and statins) in 153,996 adults ages 35 to 70 years with a history of coronary heart disease or stroke, from rural and urban communities in countries classified at various income levels by the World Bank has been evaluated. Few patients in the study were taking antiplatelet agents (25.3%), beta-blockers (17.4%), ACE inhibitors or ARB (19.5%), or statins (14.6%). Furthermore, use was lowest in low-income countries (8.8%, 9.7%, 5.2%, and 3.3%, respectively) [31].

Recent data from the PURE study have also shown poor adoption of healthy lifestyle behaviors among those with known CVD, emphasizing the urgent need of efforts to implement the nonpharmacologically based interventions. For example, among 7,519 individuals with self-reported coronary heart disease or stroke, a substantial proportion continued some level of tobacco use following their event (18.5%), only 35.1% participated in high levels of physical activity and only about 40% had diets classified as healthy. Changes in dietary habits may be a significant barrier in many environments, particularly low-income

countries. In addition, 14% did not undertake any of these 3 key healthy lifestyle behaviors and only 4.3% were classified as achieving all 3 recommended healthy behaviors. Variations in participation in healthy lifestyle behaviors were seen among different subgroups. For example, whereas over 50% reported smoking cessation during the period assessed, differences were noted among highincome countries (74.9%), upper-middle-income countries (56.5%), low-middle-income countries (42.6%), and low-income countries (38.1%) [32]. The existing evidence shows that lifestyle modifications to reduce the risk of recurrent CVD events are not being maximized, and these could be implemented alongside long-term pharmacological management of CVD. However, current approaches to modifying lifestyle behaviors, based on individual counseling, are expensive and only modestly effective, particularly among lower income countries, with significant barriers in health system and clinical capacity, broader approaches should be considered [31].

Overall, each year, millions of individuals survive ischemic heart disease or stroke events. This creates substantial clinical and public health challenges in the provision of CVD risk factor management, availability and affordability of effective therapies, adherence to prescribed medications, and available financial and human resources to care for individuals. Secondary prevention of CVD requires a comprehensive system that includes increased

patient awareness, prescription of evidenced-based treatments using standardized approaches, patient adherence, and functional health systems; all of which are influenced by local socioeconomic factors. A fundamental aspect of improving CVD care is the concept of individual and system level factors being strongly influenced by the existence of effective national policies aimed to control NCD. Many efforts had been made by international organizations, such as the World Health Organization (WHO) and the Pan American Health Organization (PAHO), along with national organizations and medical societies, to prevent recurrence of CVD events in LAC by means of national and regional policies. Countries in the region have engaged in implementing those policies supported by WHO and PAHO; however, there is much to be done to reduce the high burden of disease associated with CVD in LAC [33] Due to the existing economic diversity within and across the region, large geographic density and countries in different stages of epidemiological transition, LAC constitutes a vast field to build efficient and complementary models for secondary prevention of CVD.

ROADMAP FOR SECONDARY PREVENTION IN LATIN AMERICA AND THE CARIBBEAN

The World Heart Federation (WHF) Roadmaps are aligned with the WHO goal of reducing premature mortality from CVD by at least 25% by 2025, and it may be viewed as a model to be locally customized for countries into currently established CVD prevention programs or to build implementation strategies where an efficient program is not yet implemented. The Secondary Prevention Roadmap in LAC aims to optimize actions to prevent and control CVD. Based on clear evidence of effectiveness, including feasibility and cost effectiveness, a practical roadmap identifies potential barriers (roadblocks) to implementation of prevention and control efforts and potential solutions for bypassing those barriers [15]. In some circumstances, proven therapies are neither available, nor affordable, and increased availability of proven therapies by means of expanded health coverage would potentially address that roadblock [34]. Finally, implementation of information and communication technology concerning the use of key therapies from simplified recommendations would maximize use of available resources for better treatment and patient adherence. The ehealth technology has the potential to be incorporated into daily clinical practice due to its low cost and being simple and widely available. About Latin America, around 150 million individuals use smartphones, and there are predictions that this number will increase up to 245 million users in 2019. Therefore, there is a good opportunity and potential utility of e-health technologies, particularly in terms of secondary prevention strategies in Latin America.

Two main actions are necessary to achieve the desired target for secondary prevention. First, priority interventions provided in a standardized algorithm (pharmacologic availability and use: ACE inhibitors, aspirin, statins, beta-

blockers, tobacco cessation agents, and nonpharmacologic strategies that facilitate lifestyle modification, particularly smoking cessation) should be used broadly in a standardized manner by multidisciplinary teams in diverse settings. Second, systems of care should be structured to improve the likelihood of patient adherence to these interventions [15]. Table 3 identifies roadblocks and potential solutions for LAC that were the result of a joint consultation meeting between several international organizations (PAHO, WHF, the Latin American Society of Hypertension, the Inter-American Society of Cardiology [IASC], the Caribbean Public Health Agency, the Centers for Disease Control and Prevention), and representatives of Ministries of Health of selected countries (Brazil, Chile, Cuba) held in São Paulo. Brazil, at the end of 2015. This joint consultation, which reviewed the applicability of the WHF Global Roadmap on Secondary Prevention for Latin America and explored opportunities to improve secondary prevention, concluded that WHF Roadmap framework is well aligned with the WHO strategy and priorities for NCD/CVD management as well as with the priorities of many other relevant global and regional organizations in this particular matter. In addition, this consultation highlighted that the WHF Roadmap offers an important opportunity to work in the coordinated manner at local and regional levels to coordinate efforts among between partners to scale up secondary prevention of CVD in the Americas.

The following actions should be prioritized to obtain the desirable outcomes (Figure 1): 1) integration of CVD secondary prevention interventions into the general framework of CVD/NCD management; 2) increasing the availability of health care professionals to provide recommendations to patients with known CVD; 3) simplifying the treatment of patients with known CVD; 4) ensuring that health care professionals are aware of and follow recommendations; 5) increasing the availability and affordability of key priority interventions; 6) helping patients to adhere to recommendations; 7) strengthening the governance of health care systems to enhance prevention; and 8) strengthening health information systems [15].

LATIN AMERICAN CVD PREVENTION AND CONTROL PRIORITIES

In 2006, in response to the regional and global concern about NCD, the Directing Council of PAHO approved a comprehensive regional plan for the prevention and control of NCD [35]. This program was operationalized through biannual plans aimed at achieving specific objectives, which included 4 lines of action: integrated management of NCD and risk factors; health promotion and disease prevention; surveillance; and policy and advocacy [33]. Since then, significant progress in implementing national chronic disease programs has been reported in the majority of countries [33].

In 2010, a regional consultation on cardiovascular health, led by PAHO, proposed a core set of priority

TABLE 3. Roadblocks and potential solutions to achieve appropriate cardiovascular secondary prevention targets in LAC

Referral to rehabilitation E- services; improve schedule of follow-up visits at outpatient clinics	Education provided to the patient and family: the disease and possible complications,
	importanceof the use of specific medications, adoption of healthy lifestyle
expand clinic operating hours; schedule first and follow-up visit in collaboration with patients; diversify locations of health care units, especially in rural communities.	
Jtilizing e-health to its maximum capacity.	
Clinical decision support systems (e-health) to increase awareness.	
Clinical decision support systems (e-health); implement recognition programs to drive positive change.	
Provide financial and social support for patients to purchase priority interventions.	
rc	support for patients to purchase priority

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TABLE 3. Continued

Roadblock		Potential	Solutions	
Patients are not aware of the importance of and need for long-term treatment.	Engage diverse health care staff (e.g., pharmacy, nurses) to education patients and family about the health condition; promote continuous education on consequences, daily activities of living, and impact on survival.	Public campaigns; performance measures as metrics to inform patients and family; provider point of care decision prompts in risk factor review and medication adherence.	Empower patients to have a key role in the decision-making process (patient advocacy), sharing comments, and thoughts with health care providers.	
Patients do notremember to follow recommendations.	Use information and communication technology to remind patients about key and simple recommendations (mHealth).	Shift conversation away from directly blaming patient to one of mutual understanding of need for lifestyle changes to manage and improve chronic condition.	Use patient-nominated, nonprofessional treatment supporters (family and close friends to come to the first visit at the outpatient clinic after the CV event).	Encourage community engagement and ownership of health promotion activities through integration of new technologies.
Patients with low health literacy who do not know what medication to take.	Develop easy to interpret medication labels (e.g., symbols for dosing) and educational materials.	Include education of broader social support network, including family members.		

interventions to improve cardiovascular health in the region and, since then, much progress has been reported. Policies related to improvements in smoke-free spaces were most likely to be implemented. Progress was apparent in surveillance of risk factors, actions to improve the quality of mortality data, and access to medication for hypertension among disadvantaged populations. However, significant challenges persist in many priority areas. For instance, on the prevention side, less than one-half of countries reported more focus and resources for cardiovascular health components of the NCD plans. What was also evident in that report was the limited progress on the clinical management of CVD. For example, less than 50% of the countries reported clear progress on the following: emphasis of therapeutic control of hypertension; CVD risk assessment in primary care; public health programs for early recognition of CVD; institutional incentives for doctors to use clinical practice guidelines in hypertension management and for quality improvement of acute coronary syndrome [36].

PROPOSAL FOR A REALISTIC AND EFFICIENT PREVENTION COALITION PLAN IN LATIN AMERICA

Taking into account the heterogeneity of health systems, management strategies for CVD prevention, and patient adherence in LAC, a proposal for a coalition toward an efficient and integrated prevention plan will not be an easy task; however, this is indeed feasible. A common, comprehensive, regional, and country-applicable agenda is urgently needed and highly desirable for secondary

prevention of CVD in LAC. To address the high burden of CVD in the region, a coalition plan should be developed. This would include the integrated efforts of clinical medicine, public health and health professional societies and organizations, government at all levels (federal, state, and municipal), patient organizations, universities, private initiatives, and international organizations such as PAHO and WHF. The guarantee of availability and accessibility of an essential set of medications is critical to moving forward with the secondary prevention of the CVD agenda. In this regard, the role of PAHO Strategic Fund would be relevant to support the efforts of countries to improve the access to high-quality and affordable, low-cost medications [37].

It would be very desirable that following this joint consultation meeting, the LAC region can move toward the construction of a regional plan to scale up the implementation of the CVD secondary prevention. Such a plan should start with a situational analysis in each country to properly and reliably assess the current landscape of secondary CVD prevention. The plan should focus on clear and measurable objectives, contain straightforward implementation actions, and include a core set of indicators for monitoring and evaluation.

SIMPLE AND PRACTICAL GUIDELINES FOR HEALTH CARE PROVIDERS

Guideline-based recommendations continue to be underused, and some observational studies have shown that low implementation in daily practice is associated with the high degree of complexity of the recommendations [28,29,38].

As a result, patients fail to benefit optimally from advances in health care and are exposed to unnecessary risks, and health care systems are exposed to unnecessary expenditures. The utilization of clinical practice guidelines is influenced by local barriers or facilitators for the individual clinician and the professional context. There are gaps between knowledge and application, and these could be linked to: 1) guidelines characteristics (e.g., format, external validity, and scientific basis); 2) physician factors that include knowledge, training, and agreement with guidelines; and 3) both physician and organizational environments [38]. Francke et al. [39] suggested that guidelines that are easy to understand and simple to use in daily practice are more likely to be fully implemented. Moreover, various characteristics of patients, health care professionals, and the environment may influence guideline implementation. It seems clear that effective strategies to improve guideline adherence that are based on multiple domains and broader approaches are more likely to be successful. WHO has promoted the use of a Package of Essential Noncommunicable disease interventions for primary health care in low-resource settings. In the context of CVD secondary prevention, the WHO Package of Essential Noncommunicable [40] should be considered as a simple and practical tool for health care providers that could be customized to address local needs. Technical guidance packages from WHO should also be considered to guide interventions, including those focused on CVD prevention and management (e.g., Global HEARTS [Technical Package For Cardiovascular Disease Management In Primary Health Care]), complemented by activities focused on sodium reduction (e.g., SHAKE [Technical Package For Salt Reduction]) and tobacco cessation (MPOWER [Technical Package To Defeat The Global Tobacco Epidemic]) [41].

FINAL CONSIDERATION

CVD imposes a heavy and lifelong burden in LAC. Availability of scientific evidence confirming efficacy of pharmacological and nonpharmacological interventions requires appropriate translation and implementation to fully recognize their benefits. The WHF 25 × 25 Roadmap aiming at reducing premature cardiovascular mortality, globally, includes 3 major components: smoking; hypertension; and cardiovascular secondary prevention. To achieve efficiently the proposed target, a coalition including academic, government, health organizations, private stakeholders, and civil society, including nongovernment organizations, must work together toward a better and more efficient system for cardiovascular care delivery in LAC. Therefore, there is a clear role for PAHO, WHF, and organizations such as the Inter-American Society of Cardiology to be responsible for organizing and harmonizing the CVD secondary prevention in Latin America. In that context, the IASC would have a crucial role leading the National Cardiology Societies in Latin America.

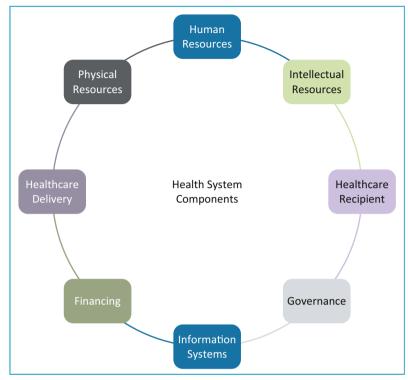


FIGURE 1. Health system requirements to achieve secondary prevention targets. Reproduced with permission from Perel et al. [15].

Regarding the role of primary care providers on the care delivery in substantial proportion of secondary prevention, especially outside the large cities and in poorer regions where there are fewer cardiologists, understanding and proposing how they might be reached and included in these efforts represents a key challenge. In fact, we would like to suggest that based on the 25 \times 25 Roadmap launched in Brazil, the Family Health Strategy, linked to the Universal Health Coverage of the Brazilian Health System, should definitely include health workers in order to identify patients with secondary prevention eligibility and to reassure the best possible adherence rates for evidence-based medications. Consequently, the inclusion of the health system component, ideally through health workers for the implementation of strategies for CVD secondary prevention is highly recommended for a successful nationwide preventive program.

Finally, nongovernmental organizations serve as strategic partners and can support and deliver robust technical assistance. Such technical assistance must be evidence-based and action-oriented and include the CVD prevention and management. The customization of the WHF Roadmaps are complementary to the WHO Global Action Plan for NCD prevention and control, and their implementation in LAC will allow us to positively influence the significant burden of CVD over the next 10 years.

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