The use of expensive technologies instead of simple, sound and effective lifestyle interventions: a perpetual delusion

Silvia Carlos,1,2 Jokin de Irala,1,2,3 Matt Hanley,1 Miguel Ángel Martínez-González1,3

ABSTRACT
A dangerous distortion of priorities seems to be currently apparent in the dominant approaches to major public health problems, including cardiovascular disease, diabetes, obesity, cancer and some infectious diseases. Relevant examples suggest an apparently inappropriate tendency to prioritise technocratic, partial solutions rather than confronting their true behavioural and structural determinants. Technically oriented preventive medicine often takes excessive precedence over simpler, more sensible approaches to modify lifestyles, the environment and the social structure. Structural factors (social, cultural, financial, familiar, educational, political or ideological factors) that act as determinants of individual behaviours should be effectively addressed to confront the essential causes of the most prevalent and important health problems. Some consumer-directed commercial forces seem to be increasingly driving many aspects of the current sociocultural environment, and may eventually compromise the main pursuits of public health. Population-wide strategies are needed to create a healthy sociocultural environment and to empower individuals and make themselves resistant to these adverse environmental and structural pressures. Otherwise most public health interventions will most likely end in failures.

LIFESTYLE AND DIET TO PREVENT CARDIOVASCULAR DISEASE INSTEAD OF THE STATIN REDUCTIONISM
At least 80% of premature cardiovascular deaths could be prevented simply through diet, physical activity and smoking avoidance. However, despite an increase in the use of antihypertensive and lipid-lowering drugs, the overall control of classical risk factors in the last years in Europe has been poor. Smoking continued unabated, and overweight/obesity prevalence increased dramatically.1 Cardiovascular prevention might be futile without addressing the actual underlying causes of myocardial infarction and stroke.

The 2013 updated American guidelines for cardiovascular prevention include blood cholesterol levels in all risk prediction algorithms.2 The strongest recommendation in these guidelines is the treatment with cholesterol-lowering drugs, mainly with statins (hydroxymethylglutaryl coenzyme A reductase inhibitors) for wide sectors of the population including subjects with no previous history of cardiovascular disease (CVD, ie, for primary prevention). However, the massive use of statins for primary prevention in healthy individuals is controversial. It is well known that statins do effectively reduce cardiovascular events in selected subgroups of patients with previous CVD, but the causal mechanisms leading from blood lipids to major cardiovascular clinical events need to be more thoroughly reappraised: is blood cholesterol really the major cause of CVD?; is cholesterol reduction the essential and indispensable step in CVD prevention for all kind of patients? Have we replaced the multifactorial origins of CVD by a single risk factor? It is well known that the best way to market a drug is the marketing of a disease (or one proxy of a disease). Are we reductionistically using high blood cholesterol as a commercially profitable proxy for CVD? Today, it is becoming more obvious that it is not LDL-cholesterol that is directly implicated in CVD but small, dense, type B particles generated after a high intake of sugar and processed carbohydrates.3,9 Overnutrition is related to blood lipids, and to blood pressure, resistance to insulin, inflammatory mechanisms, oxidative stress, coagulation factors and many other pathways leading to cardiovascular clinical events. A stronger focus on nutrition instead of the global oversimplification to massive statin use would have been the most sensible approach. The continually widening indication for statin prescription to more and more millions of healthy people does not seem to be scientifically justified.10 11 The US guidelines...
as well as the newly updated UK guidelines,12 seem to assume that the unique solution to the CVD pandemic is the replacement of CVD by a proxy (blood cholesterol) and therefore justify a blanket-wide use of statins. This oversimplification has provoked ample criticisms.8 10 13

Do we need to prescribe statins or ‘polypills’ to more than 30% of the general adult population for CVD prevention when we already know that 80% of CVD events can be prevented with simple dietary and non-dietary lifestyle measures?14–17 As the large PREvención con Dleta MEDiterránea (PREDIMED) primary prevention nutritional trial showed, simple dietary interventions can prevent many different cardiovascular problems. Indeed, only a modest increment in the adherence to the Mediterranean diet supplemented with extra virgin olive oil (100% fat) or with nuts (>50% fat) obtained a 30% relative reduction in major cardiovascular events (stroke, myocardial infarction, cardiovascular death)18 19 and also attained other cardiovascular benefits in primary prevention.20 21 Improved adherence to a high-quality dietary pattern has the additional benefit of enjoying palatable foods and will allow many more people to live longer without any need of recommending low-fat diets or using pharmacological solutions which are not exempt from side effects.13 Yet, even though light to moderate alcohol intake might reduce cardiovascular mortality,22 being physically active and eating a healthier diet can be more effective than drinking a low dose of alcohol, which may have other undesirable effects.22 23

**MEDICATION VERSUS LIFESTYLE IN DIABETES PREVENTION**

Type 2 diabetes (T2D) is likely to become the next major epidemic largely as a result of the current obesity pandemic. T2D will affect more than 10% of the adult population in many countries during the next two decades, with a projected increase of 55% by 2035.24 25 T2D is also increasing in the youth.26 However, T2D is largely preventable.27 Up to 91% of T2D cases could be prevented by relatively modest lifestyle changes.28 Therefore global growing rates of T2D represent a profound humiliation for public health. Randomised controlled trials and longitudinal observational studies have confirmed that

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**Table 1  Public health problems and their respective technological, behavioural or structural solutions**

<table>
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</table>

A mark (✓) indicates an appropriate evidence-based preventive intervention.

*Learning process that enables to understand, care about and act, on core values such as respect, justice, civic virtue and citizenship, and responsibility for self and others, which are the hallmark of safe, healthy and informed communities that serve as the foundation of our society.

CT, computed tomography; HPV, human papillomavirus.
dietary changes combined with non-dietary lifestyle modifications can have a long-term effect on T2D prevention.\textsuperscript{29, 30} The large Diabetes Prevention Program trial (figure 1) in 2002 showed that a lifestyle intervention based on dietary changes and on physical activity performed better than the metformin-medicalised option: 58\% (lifestyle) versus 31\% (metformin) relative reduction in risk over 3.2 years of follow-up, and 34\% versus 18\%, respectively, at 10 years of follow-up.\textsuperscript{31, 32} Furthermore, metformin efficacy was only limited to subjects with a body mass index $>$35 kg/m$^2$.\textsuperscript{31} The outstanding results of this landmark trial continue today to support the superiority of lifestyle interventions in comparison with pharmacological approaches to prevention. This is also true for realistic improvements on different components of the metabolic syndrome and on the atherogenic dyslipidaemia frequently observed in patients with diabetes or prediabetes. These nutritional improvements help to reduce T2D and cardiovascular risk in people with impaired glucose tolerance.\textsuperscript{31, 34} Moreover, recent evidence supports that participants in nutrition intervention studies appear to sustain the dietary changes over time.\textsuperscript{35} Additionally, pharmacological approaches to prevention may have adverse effects. Metformin has a good record for effectiveness and safety, but the potential adverse effects of other medications used against diabetes (eg, thiazolidinediones)\textsuperscript{36–38} add reasons for caution. The alarming worldwide prevalence of undiagnosed T2D, with almost half of all cases still not detected\textsuperscript{25} is a good reason to use preventive population-wide dietary and lifestyles strategies. Thus, lower-cost healthy lifestyles should be the priority for the global prevention of T2D.\textsuperscript{27, 28, 29} The Mediterranean diet, with a millennia-long history of no harm, is beneficial for primary cardiovascular prevention,\textsuperscript{40} and to reduce T2D risk.\textsuperscript{41} Indeed, the large PREDIMED randomised trial has also recently shown that changes in the overall dietary pattern with a long-term (up to 7-year follow-up) adoption of the Mediterranean diet can reduce the risk of T2D among persons at high cardiovascular risk.\textsuperscript{30}

Another essential strategy for T2D prevention is physical activity, because physical inactivity is responsible of around 27\% of T2D cases. The promotion of physical activity has been included in many international clinical guidelines for the prevention of major chronic diseases.\textsuperscript{42, 43} The new fact sheet of the WHO recommends that adults should do at least 150 min/wk of physical activity of moderate intensity.\textsuperscript{44} However, we are still far from reaching this recommendation and global levels of physical activity keep falling.\textsuperscript{45, 46} Public health policies need to implement effective promotion programmes to make physical activity accessible and attractive for whole populations. An active lifestyle can be promoted with correct and specific instructions to be included in the routines of the general population at large. Personal, social and environmental contexts have to be considered to achieve this purpose.\textsuperscript{43, 47}

The current absence of large randomised trials to assess the benefits of physical activity using hard clinical end points is somewhat surprising. Large randomised trials to assess the long-term effects of physical activity are badly needed.

**THE MIRAGE OF ANTIOBESITY DRUGS**

Worldwide, around 35\% of the adult population are overweight and 12\% are obese.\textsuperscript{47, 48} The prevalence of overweight and obese children and adolescents is also rapidly increasing,\textsuperscript{49–51} with an expected 9.1\% worldwide prevalence in 2020.\textsuperscript{52–54} One temptation is to seek ‘technical’ or pharmaceutical solutions which allow to eat calorie-dense foods \textit{ad libitum} and still not become obese. But the story of what has happened with several seemingly ‘magic’ antiobesity drugs is instructive: some were rejected by the food and drug administration (FDA), others were withdrawn from the market or are still awaiting decisions because of safety concerns.\textsuperscript{55–57} Although it is easier for doctors and patients to turn to a pill than to address behaviour, we likely do not need more antiobesity drugs trials but a change of paradigm.\textsuperscript{58, 59} The problem is that our culture, which advertises how to eat more, resists the simple message ‘eat less’.\textsuperscript{60} An obesogenic environment is a strong determinant of the current obesity epidemic. However, not all subjects exposed to this environment become obese. Susceptible individuals become obese, whereas resistant subjects remain lean. A strong hedonistic attraction to palatable foods and to overeating has proven to be a decisive factor to explain weight gain in an obesogenic environment.\textsuperscript{60} Given the actual prevalence of obesity and its worrisome health consequences, the population would benefit more from being turned towards attractive and high-quality lighter meals, with an adequate proportion of the evidence-based recommended foods.

**CANCER: COMPUTED TOMOGRAPHY SCREENING**

The development of the ‘omics’ sciences (genomics, transcriptomics, proteomics, metabolomics) has contributed to over-shadowing the fact that it is not genetic predisposition but social, environmental and behavioural factors which cause most cancers.\textsuperscript{2, 61–63} The good news is that these environmental and behavioural factors are modifiable.\textsuperscript{64}

Twelve forms of cancer are the major global contributors to deaths by neoplasia (figure 2). The cancer which causes most deaths—lung cancer—is the one that could be most easily and cost-effectively prevented, just by altering behavioural (active smoking) and environmental (passive smoking) factors.\textsuperscript{65–67} Expense screening of lung cancer with low-dose CT has been recently endorsed by the United States Preventive Services Task Force (USPSTF) in high-risk subjects because of their tobacco smoking history.\textsuperscript{68} However, this approach, which has important harms associated with false positive results, overdoses and radiation, cannot prevent most lung cancer-related deaths, as it results in only a 16\% reduction in lung cancer mortality.\textsuperscript{69} In comparison, a near 40\% reduction in lung cancer

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**Figure 1** Cumulative incidence of type 2 diabetes in the Diabetes Prevention Program trial (n=3234). p<0.001 for all comparisons. Among participants with BMI $<$35 kg/m$^2$ (n=2040), no significant differences were found between metformin and placebo. From (Kowler et al\textsuperscript{25})Copyright (2002) Massachusetts Medical Society. Reprinted with permission from Massachusetts Medical Society.)
Deaths was already attributed to decreases in smoking in the past century,69 and a 90% risk reduction for overall smoking-associated mortality has been recently estimated.70 The lure of a high tech screening device can overshadow the fact that billions of cigarettes are still smoked worldwide every day and that the number of smokers is increasing in many countries.69 71 Nineteen types of cancer are attributable to tobacco smoking.63 Is it not worthy to tackle, as the first priority, this common cause of a largely lifestyle-associated disease projected to keep increasing?69 Tobacco retail prices remain too low in some countries and although tobacco has become less affordable in some countries it is now more affordable in many others (figure 3). Sadly, the global burden of disease attributable to tobacco smoking (including secondhand smoke) has changed little.2

Screening costs were not considered in the USPSTF and other guidelines67 68 and screening is not an alternative to smoking cessation but an additional intervention. Therefore, governmental actions must focus first on tobacco-avoiding policies.72–79

The priority in lung cancer prevention is not screening...
(secondary prevention) but complete avoidance of smoking (primary prevention) and even *primordial* prevention to completely eradicate the existence of smoking in our societies and prevent the initiation of smoking for all members of future generations.61 69 80

The above-mentioned non-communicable diseases (NCDs) are largely preventable by addressing common lifestyle determinants (unhealthy diet, physical inactivity and tobacco use).81 82 A recent modelling study has analysed the potential impacts in NCD mortality of reducing six selected preventable risk factors (tobacco, alcohol, salt intake, obesity, raised blood pressure and raised blood glucose). It has concluded that achieving the reduction of these six risk factors will decrease the probability of premature deaths from 2010 to 2025 by 33% from CVDs, 12% from lung cancer and 5% from diabetes.83 Therefore, sociological and cultural issues need to be fully addressed and require educational and structural responses.84–89 From an equity perspective, environments should make the healthier choice the easiest choice so that prevention becomes accessible, available and affordable to all sectors of the population.57 58 90 On the other hand, energy-dense fast foods, sugar sweetened beverages, smoking and sedentary lifestyles should be made the *harder* options. These are not *sweet* truths for some food or beverage corporations or for the technocratic business,59 95 but they are indeed the priorities for public health.

**HPV Vaccine: The Glamour of Technoprevention**

Also, some approaches to the prevention of communicable diseases have apparently forgotten the need to give appropriate priority to lifestyle interventions and heavily relied on technocratic solutions. Universal policies to vaccinate girls against human papillomavirus (HPV) were adopted very rapidly during 2006–2009. The technical merits of this vaccine should be acknowledged and appreciated. However, many issues on this publicly funded mass vaccination remain unresolved: its coverage and affordability for the most vulnerable and affected countries, its long-term effectiveness and the uncertain—though potential—replacement of the vaccine-targeted strains with other high-risk types.96–98 Strong conflicts of interest are another reason for concern as available trials were sponsored by the manufacturers.

Affordability and an adequate coverage with complete vaccine regimens are two essential components for HPV vaccination programmes success. The vaccination programmes have mostly been implemented and publicly funded in higher-income countries although most of them have very low rates of HPV-related diseases.99 The rapid introduction of this publicly funded vaccine in Spain does not match the epidemiological needs of the country because there is no epidemic of cervical cancer in Spain whatsoever. On the contrary, mortality and incidence of cervical cancer are very low and show a decreasing trend in Spanish women (figure 4).

In developed countries the coverage and effectiveness are lower for girls at higher risk.100 In addition, very few developing countries have introduced HPV vaccine into their national programmes although the burden of HPV-related diseases is highest there. At the current cost, mass vaccination is unaffordable in low-income and middle-income countries. In Africa, although several countries are implementing HPV vaccine demonstration projects, only Rwanda initiated a countrywide vaccination, after receiving a donation of several million vaccine doses from the manufacturer.101 102 The Global Alliance for Vaccines and Immunisation has offered some of the world’s poorest countries the quadrivalent vaccine at $4.50/dose. But overall, vaccination programmes will still be prohibitively expensive and their widespread implementation will not probably be sustainable.

Moreover, vaccine efficiency heavily depends on an adequate coverage with the complete regimen103 and we do not yet know the long-term efficacy even when a high coverage is reached. Today, HPV vaccine coverage is still quite low in many countries, including economically developed countries such as the USA, where coverage for adolescent girls with all three doses was 33.4% in 2012.

Is the vaccination supplanting the need to combat the major risk factors? Why has an expensive and only partially preventive measure been rapidly introduced while other well-known and sufficiently proven measures been downgraded?

The rates of cervical cancer in a country are highly dependent on the availability of and women compliance with screening programmes. HPV infection strongly depends on partners’ number and other sexual risk behaviours. Long-term primary prevention strategies are needed, which include structural interventions and sexual education with correct and complete information to increasing awareness about the basic facts of HPV infection. These simple, affordable and sustainable measures are absolutely essential for cancer prevention, especially in developing countries.

**Risk Compensation**

If preventive strategies focus preferentially on glamorous technological options, apart from the problems herein described, another important unexpected consequence is risk compensation. People may consider themselves healthy enough when they eat foods labelled as low-energy/low-fat, take the prescribed pills and vaccines or attend the recommended screening programmes no matter if they keep sedentary, adhere to low-quality food patterns, smoke or have many other dietary and non-dietary risk behaviours. For example, the new USPSTF guidelines for lung cancer screening notice that people may keep on smoking thinking that screening programmes can reduce their smoking-related risks.104 Believing in a new preventive technology may lead to a dangerous sense of immunity. The available evidence is not fully consistent, but risk compensation has also been associated with HPV vaccine acceptability and this affects vaccine coverage and consequently, effectiveness and efficiency.96 97 99 103 Therefore, it is important that preventive messages address the specific target population groups in which these biomedical strategies are to be recommended so that no erroneous beliefs turn up. Always, apart from the high-risk subjects, the general population needs to be correctly informed because population-wide strategies to promote healthy lifestyles are frequently the most cost-effective option.57

**Common Building Blocks: Empowerment and Structural Changes**

Screening, diagnosis and treatment represent real breakthroughs and have accounted for tremendous advances in health status. In fact, the technological preventive interventions mentioned in this paper can surely have some role and should be applied when they are appropriately justified. However, the unprecedented and continuous increase in the use of pharmacological/technological approaches, confirms that these approaches might not be adequate to manage the complex pathogenesis and consequences of chronic diseases. In addition, the partially successful pharmacological/technological approaches are responsible for inevitable and unsustainable increases in health expenditures; hence, alternative lifestyle and dietary strategies should be given a sufficiently high priority. A body of sound epidemiological evidence exists to support the effectiveness of diet and
lifestyles in the prevention of NCD. The technologically driven mentality might lead us further away from, rather than closer to, simpler, sensible approaches to the main behaviour-related public health problems. Only the societal and cultural approaches confront the root causes.

A common denominator for all the examples reviewed is that our culture fosters behaviours mainly driven by pleasure-seeking which are known to be associated with a clustering of unhealthy lifestyles. Some global commercial powers have fostered a pleasure-seeking cultural environment which rich and the poor sectors of society. In this context, cultural deprivation seems more important than monetary poverty and needs to be counterbalanced by the cultural empowerment of individuals. This empowerment of individuals is needed to make them resistant to an environmental pressure favouring unhealthy behaviours.88

However, the success of individually focused interventions might be substantially improved when wider structural factors (social, cultural, familiar, educational, political or ideological factors) that act as determinants of individual behaviours are also addressed.69 As Geoffrey Rose already stated, population-based approaches addressing these structural determinants of health should be prioritised.104 We need to incorporate interventions that modify the environment through all public and private policies, in line with the idea of ‘Health in All Policies’. They will facilitate healthy choices that depend strongly on circumstances beyond individual control. Today people cannot make choices about behaviours in the circumstances of their own choosing. On the contrary, the promotion, price, availability and marketing of products affect their perceived costs and benefits, and consequently their choices. Decisions about lifestyles depend now on choices/interests of governments, producers and multinational companies and consequently, on the incentives of financial institutions. A few powerful institutions and decision-makers who set the agenda and decide what is fashionable and what is not, are making unhealthy choices the easiest choices, shaping the preferences of the population at large and encouraging them to make unhealthy choices. Consequently, as a result of the conflict with the dominant economic interests, wealth is being placed before health and an overemphasis on expensive technologies has led to underfunded prevention strategies. Public health intervention policies are often failing to overcome these powerful forces.105

We acknowledge that the needed cultural change is not easy. Cultivating an environment where healthier options are encouraged requires entering in decision arenas and counteracting commercially driven forces investing millions in advertising and opinion-making. Therefore, it does not always conform with the ‘politically correct’ positions and it is far from cheap and simple to achieve. However, if sociocultural and normative changes are attained, they will lead to dramatic reversals of epidemics. We do not need to shape individuals’ minds to fit our increasingly unhealthy societies but we need to redesign the environment to fit our populations’ health.

The implementation of all the strategies proposed here should be considered a priority for public health.

**What is already known on this subject?**

▸ Reducing some of the main preventable risk factors described here (tobacco, alcohol, salt intake, high blood pressure and blood glucose and diabetes) will lead to higher reductions in probability of death compared with current trends with no additional action (22% in men and 19% in women) and even more in case of early action (24% and 21%).83

**What this study adds?**

▸ The new guidelines for cardiovascular prevention can be misinterpreted and lead to an excessive use of statins; statin use should be decided depending on individual circumstances.

▸ The growing trends in type 2 diabetes (T2D) prevalence represent a humiliation for public health because T2D is easily prevented with diet and lifestyle.

▸ The key message for obesity prevention is “to eat less”.

▸ Screening for lung cancer is probably inefficient and not affordable and may distort the true priority: smoking avoidance.

▸ Technoprevention with HPV vaccination can backfire without addressing the true underlying behavioural determinants of risk.
Contributors All authors have: Made substantial contributions to the conception or design of the work; Drafted the work or revised it critically for important intellectual content; Approved the final version to be published; and Agreed to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

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