Abstract
Since the beginning of mankind, man has sought ways to promote and preserve health as well as to prevent disease. Hydration, physical activity and exercise are key factors for enhancing human health. However, either a little dose of them or an excess can be harmful for health maintenance at any age. Water is an essential nutrient for human body and a major key to survival has been to prevent dehydration. However, there is still a general controversy regarding the necessary amount to drink water or other beverages to properly get an adequate level of hydration. In addition, up to now the tools used to measure hydration are controversial. To this end, there are several important groups of variables to take into account such as water balance, hydration biomarkers and total body water. A combination of methods will be the most preferred tool to find out any risk or situation of dehydration at any age range.

On the other hand, physical activity and exercise are being demonstrated to promote health, avoiding or reducing health problems. Therefore, physical activity is also being used as a pill within a therapy to promote health and reduce risk diseases, but as in the case of drugs, dose, intensity, frequency, duration and precautions have to be evaluated and taken into account in order to get the maximum effectiveness and success of a treatment. On the other hand, sedentariness is the opposite concept to physical activity that has been recently recognized as an important factor of lifestyle involved in the obesogenic environment and consequently in the risk of the non-communicable diseases.

In view of the literature consulted and taking into account the expertise of the authors, in this review a Decalogue of global recommendations is included to achieve an adequate hydration and physical activity status to avoid overweight/obesity consequences.

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Key words: Physical Activity, Hydration, Health, Prevention. Decalogue of Recommendations.
Introduction

This article is the result of the presentation and discussion of 10 lectures at the I Symposium on Physical Activity, Hydration and Health, that was held at the Royal and Illustrious Official College of Pharmacists from Sevilla, Spain with the collaboration of Coca-Cola Iberia, on the 7th November, 2013.

All the authors expressed freely their ideas and studies according to their expertise and knowledge. Discussions were held among all of them and the audience.

The Symposium included both the opening and closing lectures, with the respective following titles and speakers:

- **Health maintenance through exercise**, by Pedro Manonelles

Two round tables were included, as follows:

1. “What do we know and what can be improved in healthy subjects? The following lectures were selected (in order of appearance at the scientific programme):
   - *Composition of beverages for sportspeople* by Nieves Palacios,
   - *Evaluation of hydration* by Julia Wärnberg,
   - *Prescription of physical exercise for health* by José Antonio Casajús,
   - *Exercise as medicine* by Margarita Pérez.

2. “Preventing through physical activity and hydration” included the following lectures:
   - *Relationship between physical activity levels and childhood obesity* by Susana Aznar
   - *Myths and realities of the loss weight programmes* by Pedro J. Benito
   - *Physical activity, sedentariness and inflammation in adolescence* by David Martínez-Gómez
   - *Physical exercise and hydration: not too little, not too much* by Francisco B. Ortega.

The coordination of the scientific programme and the discussion of the presentations were led by Ascensión Marcos.

It is well known that since the beginning of mankind, man has sought ways to preserve health and prevent disease. Hydration, physical activity and exercise are key factors for enhancing human health (fig. 1). However, either a little dose of them or an excess can be harmful for people at any age.

Therefore, the aim of this event was to discuss about how these two apparently independent entities such as hydration and physical activity/exercise, have an impor-
tant role both separately and interacting between both of them in the maintenance and improvement of health. Recent findings were discussed and recommendations for appropriate behaviour related to hydration and physical activity have been offered in this overview.

Hydration

Water is an essential nutrient for human body and a major key to survival has been to prevent dehydration. However, there is still a general controversy regarding the necessary amount to drink water or other beverages to properly get an adequate level of hydration and also the best way of measuring hydration in humans in order to know to what extent an individual can be at risk of dehydration and how to prevent any situation of dehydration at any age range.

Measurement of hydration

In clinical nutrition and field studies, commonly used assessment methods of hydration are: estimates of water balance (thirst rating, total water intake and output or body weight changes), hydration markers (plasma or urine osmolarity) and total body water (TBW) measurements by bioelectrical impedance or isotope dilution.

Water balance

Thirst rating

For the majority of healthy population, fluid balance is maintained via thirst, a feedback-controlled variable, acutely regulated by central and peripheral mechanisms. However, voluntary drinking is also a behaviour influenced by other environmental, social, and psychological cues. Indeed, thirst perception is typically assessed by subjective ratings using either categorical or visual analogue scales. Therefore, factors and conditions (e.g., age, disease, temperature) that influence thirst should be also recognized, taking into account that humans may drink also for other reasons, particularly for hedonic ones. For example, during cold exposure, thirst is significantly blunted independently of hydration status or activity.

Water intake and output

Water balance can be estimated by the assessment of both water intake (input) and water losses (output) during a period of time.

Total water input includes total water intake (from beverages, and food) and to a small extent also from oxidation of macronutrients (metabolic water). Estima-
Total body water (TBW)

Bioelectrical impedance analysis (BIA)

BIA measures the impedance or resistance to a small electrical current as it travels through the body’s water pool. It is a commonly used method to quantitatively estimate body composition and is based on a two-compartment body composition model, namely body fat mass (FM) and fat-free mass (FFM) and assuming that 73% of the body’s FFM is water for TBW estimation. Single-frequency BIA (SF-BIA) is most commonly used for assessing TBW and FFM but cannot distinguish TBW into its intracellular and extracellular compartments. Bioimpedance spectroscopy (BIS) or multifrequency BIA allows for the differentiation of TBW into intracellular and extracellular water compartments. Specific prediction equations may be developed against dilution techniques, e.g. the TBW equation model developed for Spanish children which incorporated height (cm)^2/Reactance and weight: TBW (kg) = 0.495 × height (cm)/Reactance + weight (kg) × 0.107 + 6.08 (R^2 = 0.91; SEE = 2.0 kg)^7.

Dilution and tracer techniques

Dilution techniques are considered the golden standard of the qualitative measurement of FM, FFM and TBW (again, assuming that water in the FFM is constant at about 73%). To this end, a known dose of isotop-labelled water (2H, 18O or 3H) is ingested and allowed to equilibrate within the body water (4-5 h) and urine is collected for later spectrophotometry determinations. The tracer sodium bromide (NaBr) can be used for the measurement of extracellular water space. Administration of these tracers and collection of samples are easy but these methods are expensive and impractical for large-scale studies.

In summary, at the population level one standard method has not been still accepted to adequately assess the hydration status; therefore, a combination of methods is preferred. When body water intake and output are in balance, TBW and P_WE provide objective measurements of volume and concentration. TBW may be assessed by BIA or dilution techniques. At an individual level, maintaining a constant morning body weight, adequate fluid intake, a pale yellow urine colour, and controlled normal urine volume will assist healthy individuals to achieve euhydration.

Interactions between hydration and exercise

The importance of hyponatremia

Childhood and elderly people are groups of population with a high risk of dehydration. Specially, the risks and consequences of dehydration while exercising are well-known. This is the case of hyponatremia that has been defined as plasma sodium concentration below 135 mmol/L, which may cause a pathological situation and might occur when doing exercise and unfortunately it is very common in endurance events. Hyponatremia becomes more frequent as the duration of the exercise increases, specially, in ultra-endurance exercise (events lasting longer than 6 hours). As an example, during the London Marathon in 2006, the incidence of hyponatremia was 12.5%^11. Indeed, it is the first cause of severe disease in ultra-endurance events, such as the popular Ironman. Therefore, in ultra-endurance events (more than 6 hours, but often lasting 10 and 20 hours, such as Ironman) the amount of beverage intake containing carbohydrates and sodium should be smaller, around 0.5 L per hour of event, being the amount of liquid intake per hour smaller as the duration of the sport activity increases in order to avoid any renal or metabolic disfunction.

Recent data support that 95% of the variance in the decrease in sodium concentration (that can lead to hyponatremia) after exercise is explained by increases in body weight as a consequence of over drinking^14,15. These findings may have important implications for guiding athletes participating in ultra-endurance sports (fig. 2).

In addition, since many of the effects of the exercise on the inflammatory cells are mediated by changes in the systemic concentrations of stress hormones and proteins, such as glucocorticoids, catecholamines, and 72 kDa heat shock proteins, an optimal hydration during sport performance must be important in order to avoid potential hemoconcentration of these “stress mediators”.

Beverages for sportspeople

According to Spanish law, beverages for sportspeople are considered among food preparations for dietary and/or special regimes, under the heading on foods adapted to intense muscular wear.

These beverages have a specific composition aimed to achieve rapid absorption of water and electrolytes and to prevent fatigue. In addition, they have specific requirements.

Therefore, the main aims of these beverages are as follows:

1. To provide carbohydrates to maintain an appropriate concentration of glucose in blood and to delay the exhaustion of glycogen deposits.
2. The replacement of electrolytes, particularly of sodium.
3. Hydric replacement to avoid dehydration.

These beverages usually have a particular nice overall perception of flavour, so it is reasonable to bear in mind that they will be more easily consumed than water on its own.

In February, 2001, the European Commission’s Health and Consumer Protection, through the Scientific...
Committee on Food, drew up a report on the composition of food and beverages intended to meet the expenditure of great muscular effort, especially among sportsmen and women. This report points out that specially adapted foods and liquids help solve specific problems in order to achieve an optimum nutritional balance. These beneficial effects are not confined only to sportspeople, who take regular and intensive muscular exercise, but also people who, in their jobs, make major exertions in adverse conditions, and people who during their leisure time do physical exercise and training. It indicates that the sports beverage should supply carbohydrates as a fundamental source of energy and should be efficient in maintaining optimum hydration or in rehydrating recommending the following margins in the composition of the beverages to drink while doing sport:

- Not less than 80 kcal per litre (L) and not less than 350 kcal/L.
- At least 75% of the calories should come from carbohydrates with a high glycemic index (glucose, sucrose, maltodextrins).
- Not less than 460 mg/L of sodium (46 mg per 100 mL/20 mmoL/L) and not more than 1,150 mg/L of sodium (115 mg per 100 mL/50 mmoL/L).
- Osmolality between 200-330 mOsm/kg of water.

**Other components of replacement beverages**

**Antioxidants**

The ingestion of antioxidants to minimize the damage caused by reactive species generated in the electron transport chain has produced different results when assessing an increase in performance, so their presence in beverages for sportsmen and women is not essential.

**Proteins**

The ideal protein concentrate to be added to a beverage for sportspeople would be whey protein from milk. Another alternative is just to provide serum lacto-proteins, i.e. milk serum deprived from lactose, which can be useful in people with lactase deficit.

**General recommendations**

A consensus was reported in view of the needs to get proper beverages for sportspeople, including the definition of the composition and guidelines to healthy liquid replacements.

1. Hydration should be optimized as much as possible.
2. Beverages for sportsmen or women used during training sessions or in competitions should be ranged between 80-350 kcal/1000 mL, of which at least 75%, simple carbohydrates, e.g. glucose.
3. Beverages for sportsmen or women used during training sessions and in competitions should be ranged between 20-50 mmoL/L (460-1,150 mg/L) of sodium ion content according to the heat, intensity and duration of the exertion. The osmolality of such beverages should be ranged between 200-330 mOsm/L of water, and should
not under any circumstances exceed 400 mOsm/L of water.

4. Replacement beverages used after training or competitions should have a calorie content of 300-350 kcal/1000 mL, of which at least 75% should come from a mix of high glycemic load carbohydrates such as glucose, sucrose, maltodextrins and fructose.

5. Beverages for sportsmen used for immediate post-exertion should have sodium ion content in the range of 40-50 mmol/L (920-1,150 mg/L). Likewise, they should provide potassium ion in the range of 2-6 mmol/L. The osmolality of those beverages should be comprised 200-330 mOsm/L of water, not exceeding 400 mOsm/L.

Physical activity/exercise

Not in vain Hippocrates mentioned his famous sentence (IV B.C): “Eating alone will not keep a man well; he must also take exercise. For food and exercise... work together to produce health”.

Physical activity is defined as any bodily movement produced by skeletal muscles that results in energy expenditure beyond resting expenditure (>1.5 METs), as walking to and from work, taking the stairs instead of elevators and escalators, gardening, and doing household chores. Exercise, however, is a type of physical activity that requires planned, structured, and repetitive bodily movement with the intent of improving or maintaining physical fitness level. Exercise can be accomplished through activities such as cycling, dancing, walking, swimming, yoga, working out at the gym, or running, just to name a few. Regular exercise, depending upon the kind, improves aerobic fitness, muscular strength, and flexibility.

In the XXI century the practice of regular physical activity/exercise has been associated with both a healthier and a longer life, being recognized as one of the most powerful health tools for preventing disease and improving the quality of life in developed countries. Indeed, frequent and regular physical exercise boosts the immune system, and can be beneficial in detecting, preventing and managing the “diseases of affluence”, such as hyperlipidaemia, hypertension, type 2 diabetes, obesity, arthritis, dyslipidaemia, depression, chronic obstructive pulmonary disease, nicotine addiction, affective disorders, cancer, osteoporosis, and age-related declines in muscular strength.

The main outcome of regular physical activity, achieving moderate-to-high peak cardiorespiratory fitness (> 8 METs) may reduce the risk of cardiovascular events and all-cause mortality. Those moderate-to-vigorous physical activity (MVPA) (≥450 min/wk) values that are clearly above the minimum international recommendations of 150 min/wk of MVPA are associated with longer life expectancy.

Indeed, regular exercise is probably the lifestyle intervention with the most profound up-regulating effect on hundreds of genes involved in tissue maintenance and homeostasis, implying a complex cross talk between muscles and other tissues, which has been selected for optimizing aerobic metabolism to conserve energy in an environment of food scarcity resulting in numerous beneficial adaptations and the benefits of exercise on metabolic, psychological and physiological health (Table I).

| Table I |
| Benefits of regular physical activity on metabolic, psychological and physiological health |
| **Effects on performance / fitness** |
| Improved endurance |
| Improved strength |
| Improved balance |
| Improved flexibility |
| **Reduction of cardiovascular risk factors** |
| Sedentary lifestyle |
| Fibrinogen |
| Insulin sensitivity |
| **Reduction of social misbehaviours** |
| Violence, smoking, consumption of alcohol, drugs and unhealthy diets (especially in children and young people) |
| **Prevention of chronic diseases** |
| Obesity / overweight |
| Hypertension |
| Ischemic cardiopathy |
| Stroke |
| Hypercholesterolemia |
| Type 2 diabetes |
| Osteoporosis |
| Musculoskeletal disorders |
| Lumbar pain |
| Cancer: colon, breast, prostate |
| **Prevention of psychological, psychiatric and behavioural disorders** |
| Stress |
| Increased psychological balance |
| Improved cognitive function |
| Anxiety |
| Depression |
| Self-confidence |
| Self-esteem |
| Attenuation of CV and neuroendocrine responses to mental stress |
| Reduction of some type A behaviours |
| **Aging** |
| Prevention of muscle loss |
| Prevention of bone loss |
| Reduction of functional limitation |
| Reduction of falls |
| Reduction of fractures |
| **Reduction of mortality** |
| Global |
| Cardiovascular |
However, it is important to highlight that nowadays we are facing with a paradox: technology development and certain social behaviours have caused increased physical inactivity rates that have altered the configuration of the human biological machine; when we forget our biological evolutionary process we become hypactive, sedentary, the contrary to what our evolution has been developed over thousands of years. The result is decoupling numerous biological functions involving loss of health and rise of the so-called “diseases of civilization”. Both physical inactivity and sitting behaviour in contemporary obesogenic environments initiate a situation of maladaptation that may lead to chronic diseases, and therefore it has become one of the most important health problems across all over the world.

Sedentary behaviour refers to activities that do not increase energy expenditure substantially above the resting level (1-1.5 METs) and includes activities such as sitting, lying down, and watching television, and other types of screen-based entertainment. Hundreds of thousands of people die each year as a result of developing diseases highly related to inactivity. The direct economic costs as consequence of inadequate physical activity behaviours are huge.

Unfortunately, there are no reliable estimates of the economic costs, advantages and savings that may produce weight loss programmes based not only on dietary restrictions, but especially on promotion of physical activity and the avoidance of sedentariness. Only few data are known regarding some specific diseases, but certainly very far from reality, since the investment in all kinds of slimming products is much higher than the official pharmaceutical costs.

The EEUU is the country where the most health care resources are associated with obesity and the estimated amount is around 5.5 to 9.4% of their health expenditure (more than 100,000 million dollars). In other countries, such as Canada, Switzerland, New Zealand, Australia, France and Portugal, obesity has been reported to cause between 2 and 3.5% of health expenditures. In Spain, the cost of obesity could reach 7% of health expenditure of the Spanish National Health System, although these figures are still controversial. Indeed, due to the fact that the figures in Spain are surprising, being more than 2500 million euros a year only spent in public health on healthy problems related to overweight and obesity, rationalizing resources is crucial.

Nevertheless, before interventions can be properly designed and their findings correctly interpreted, it is important to understand how physical activity levels can be changed across different periods of life.

**Physician prescription**

The US Office of Disease Prevention and Health Promotion pointed out that at that time over 8000 articles reported the benefits of exercise. According to this report, two important conclusions must be drawn:

- Important health benefits can be obtained by performing 30 minutes of moderate physical activity on most, if not all, days of the week.
- Additional health benefits result from greater amounts of physical activity.

Evidence supports the inverse relationship between physical activity intensity and premature mortality, cardiovascular diseases, stroke, type 2 diabetes, metabolic syndrome, colon cancer, breast cancer or depression, vigorous physical activity being more effective (around 30-40%) than moderate physical activity levels (fig. 3).
The way to perform physical activity has consequences and should be prescribed but it is recommended in moderate or moderate-to-vigorous intensity levels for overall health maintenance, since strenuous exercise, especially in training sportspeople, can lead to risks and harms in some occasions (fig. 4)\(^39\).

It is important to highlight that physicians also need more training in how to make best use of a powerful therapy, such as the physical activity. Physicians can successfully encourage activity by giving patients a written exercise prescription along with printed advice on how to design a safe and enjoyable routine\(^26\).

For physicians, the prescription pad is a familiar and easy way to transmit the recommendations to maintain or recover health. Similarly, the exercise prescription directs patients to initiate, maintain or increase their physical activity levels. Unfortunately at the end of the XX century primary care physicians, family doctors, etc. were limited to recommend exercise only in certain clinical situations without specifying the type of physical activity performed and no guarantee adhering to it. The academic subject “exercise prescription” is unknown in undergraduate studies and the physician must update as a postgraduate. Thus, the initiative “Exercise is Medicine”\(^40\) promoted by the American College of Sports Medicine could help physicians and other health professionals to understand the exercise prescription process. The performance of prescription requires a multidisciplinary team that will need to have an expert on physical activity. The success of prescription depends on the ability of the health team to prepare and develop a specific exercise prescription for each subject\(^41\). In this context, the concept of a *polypill* is receiving growing attention to prevent cardiovascular disease. Indeed, compared with drugs, exercise is available at low cost and relatively free of adverse effects\(^42\).

The components of a prescription for medicaments include the name of the medication, strength, route and frequency of administration, as well as duration and precautions related to doses. The components of an exercise prescription follow a similar format, defining type of exercise, intensity, frequency, duration, precautions and progression of the doses. The prescription of exercise for sedentary people should begin with a minimal effective “dose”, focusing first on the preliminary aspects of the regular exercise programme. From this “small dose” of exercise, the patient, with the professional encouragement and guidance should hopefully progress to the final optimal dose (table II).

**Physical activity and sedentariness inadequacy in early ages**

Due to the limitations of self-reported methods to assess physical activity in young people of different ages, the information available on this regard has been very limited.

The European Youth Heart Study (EYHS) is a school-based, cross-sectional study designed to examine the interactions between personal, environmental, and lifestyle influences on the risk factor for future cardiovascular diseases in several European countries. Within the EYHS, in the assessment of physical activity in a 6 to 10 year-follow up conducted in Sweden and Estonia, a decline in MVPA (overall change = 30 min/d) was observed together with an increase in sedentary time (overall change = 2 h 45 min/d) from childhood to adolescence\(^43\). These authors observed that MVPA decreased from childhood to adolescence (-1 to -2.5 min/d per year of follow-up, \(P = 0.01\) and < 0.001, for girls and boys, respectively) and also from adolescence to young adulthood (-0.8 to -2.2 min/d per year of follow-up, \(P < 0.01\) for both genders). The authors of the EYHS suggested that the increase in sedentary time was associated with the decrease in physical activity. In general, the sedentary time increased from childhood to adolescence and further to young adulthood for both genders. The decrease in physical activity was associated with an increase in body mass index. In this study, the authors observed that the children who engaged in moderate-to-vigorous physical activity had a lower body mass index. They concluded that physical activity should be encouraged in early ages to prevent obesity and other chronic diseases.
year, \( P = 0.02 \) and < 0.001 for girls and boys, respectively). Sedentary time increased from childhood to adolescence (+15 and +20 min/d per year, for girls and boys, respectively, \( P < 0.001 \)), with no substantial change from adolescence to young adulthood. The magnitude of the change observed in sedentary time was 3-6 times higher than the change observed in MVPA. Overall, these results support that MVPA declined 30 min/d while sedentary time increased 2 h 45 min/d from childhood to adolescence. These findings are of concern and might increase the risk of developing obesity and other chronic diseases later in life. In addition, these findings suggest that if a long-term intervention conducted in individuals of this age obtains no change in physical activity or sedentary time, it would actually be a very successful intervention, since the observed trend above reported has substantially modified43.

Recent cross-sectional studies have reported positive associations between physical activity and obesity prevention as well as with cardiovascular risk facts in European children44-46. The majority of Spanish adults and children do not engage in enough physical activity to achieve beneficial effects for health47. There is a need for objective and accurate assessment of the proportion of children meeting the recommendation for healthy levels of physical activity. Current health-related physical activity recommendations for children and adolescents have been established in at least 60 minutes of MVPA for 5 days a week48, 49 and they were updated later32. Assessing the patterns of physical activity both between days (during weekdays and weekend days) and within days is of interest to improve our understanding of the variation in physical activity in Spanish children and to provide more efficient intervention programmes to prevent obesity. Therefore, some studies have been aimed to find out the patterns of physical activity in children and to compare them between overweight/obese and non-overweight/obese children50 and also to assess the association be-

### Table II

A comparison between several factors involved for medication and exercise prescriptions

<table>
<thead>
<tr>
<th>Medication Prescription</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Drug</td>
<td>Acetaminophen with codeine</td>
</tr>
<tr>
<td>Strength</td>
<td>300 mg / 30 mg, Tablet</td>
</tr>
<tr>
<td>Route</td>
<td>By mouth</td>
</tr>
<tr>
<td>Frequency</td>
<td>1 tablet every 8 h</td>
</tr>
<tr>
<td>Duration</td>
<td>1 week</td>
</tr>
<tr>
<td>Precautions</td>
<td>Avoid drinking alcohol</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Exercise prescription</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Type of exercise</td>
<td>Walk</td>
</tr>
<tr>
<td>Intensity</td>
<td>4-6 km·h⁻¹ RPE 3-5</td>
</tr>
<tr>
<td>Frequency</td>
<td>Three days a week, Target 5/w</td>
</tr>
<tr>
<td>Duration</td>
<td>Forever</td>
</tr>
<tr>
<td>Precautions</td>
<td>Sprain in the right ankle two months ago</td>
</tr>
<tr>
<td>Progression</td>
<td>Start at 4 km·h⁻¹. Increase gradually every 4 weeks</td>
</tr>
</tbody>
</table>

MVPA weekday and weekend day patterns in non-overweight/obese and overweight/obese 8-10 yr-old children. The EYHS study.
between current physical activity guidelines and obesity in children (fig. 5). As an example, one of the studies performed in Spain was a part of the EYHS and involved a total of 439 children (233 girls, 216 boys) aged 8-10 y, with the main outcomes as follows: a) children’s MVPA levels are more closely associated to gender than to obesity status, b) children tend to be more active during school periods, however they achieved a low amount of MVPA and c) 60 minutes of MVPA is inversely associated with overweight and obesity risk; however, vigorous physical activity (VPA) is a more important component and it should be specifically included into the children physical activity recommendations.

These results emphasize the relevance of physical activity in children to prevent obesity. This outcome showed the need to focus on VPA opportunities and these efforts should be carried out particularly outside school time (afternoon-evening time).

Sedentariness: on the threshold of inflammatory processes

The New England Journal of Medicine 2008 has pointed out the importance of skeletal-muscle homeostasis. Skeletal-muscle fibres can produce several hundred secreted factors, including proteins, growth factors, cytokines, with such secretory capacity increasing during muscle contractions, myogenesis or after exercise training. Several studies conducted have considered the potential impact of sarcopenia on metabolic function, chronic disease, and mortality. Acutely, i.e. in response to a single bout of exercise, a pro-inflammatory response is generated (increase in circulating number and function of leukocytes and in systemic concentrations of pro-inflammatory cytokines and chemokines), whose damaging potential is limited by simultaneous activation of anti-inflammatory cytokines and chemokines, whose damaging potential is limited by simultaneous activation of anti-inflammatory mechanisms. Conversely, repeated exercise training results in significant reduction of the systemic inflammatory state. The overall health effects of exercise are therefore induced by the correct balance between these apparently opposed pro- and anti-inflammatory effects. The physiological relevance of pro- and anti-inflammatory effects of exercise will be also depending on the inflammatory status of people performing exercise, especially in individuals with low-grade inflammatory diseases.

Cardiovascular diseases (CVD) are the principal causes of death in developed countries. It is well documented that the genesis of CVD occurs in early ages, although the clinical symptoms are not clearly observed until adulthood. The trigger of CVD is the atherosclerosis, which involves an inflammatory process during the atherogenesis with a continued and substantial increase of inflammatory cytokines and acute-phase reactant levels within the arterial wall. Since CVD risk factors usually track from childhood to adulthood, a healthy lifestyle beginning in early ages is the focus of public health strategies.

Sedentary lifestyle has officially been recognized as a major risk factor for CVD, being responsible for 6% of the major CVD mortality, which makes inactivity comparable to well-known risk factors such as smoking and obesity. Information on the detrimental effect of sedentariness (mainly sitting) on health is relatively new, but there is compelling evidence for its key role on CVD mortality, regardless of physical activity. Since both physical activity and sedentary behaviours have been identified as crucial risk factors for CVD mortality, public health interventions to (i) increase physical activity and (ii) decrease sedentary behaviour in children and adolescents might have the potential to provide health protection against future CVD.

To date, there are a few studies that examined whether regular physical activity influences the inflammatory process in children and adolescents. The main findings in such studies suggest that the total amount and intensity of physical activity (i.e. light, moderate and vigorous intensities) are not directly associated with inflammatory markers in youth, but it would have a crucial indirect role through increasing cardiorespiratory fitness and decreasing body fat. The majority of Spanish adults and children do not engage in enough physical activity to be beneficial for health. Regarding the influence of sedentary behaviour on inflammation in these ages, nowadays there is limited evidence to draw any conclusion. Overall, some studies that used accelerometers for assessing sedentary time found null associations with inflammatory biomarkers in youth. Also, there is some evidence that high levels of television viewing might have an indirect role on the inflammatory process in youth because this behaviour is associated with unhealthy dietary patterns. Further longitudinal or clinical trials will provide insights into the role of physical activity and sedentary behaviour on inflammation in young people.

The interrelationship between physical activity and the immune system

It is well known that, together with skeletal muscle, metabolic and cardiovascular systems, physical activity also strongly modulates the immune system. As a consequence, people who perform some type of sport regularly have been associated with less susceptibility to infection compared with sedentary people. However, while regular moderate exercise is very likely to be associated with decreased susceptibility to infection, intense exercise has been associated with symptoms of transient immunodepression, leading to increased susceptibility to infection, especially in high competition athletes. Thus, as on the other physiological systems, the effects of exercise on the immune system also depend on the frequency, intensity, and duration. Nevertheless, there has been an excessive generalization of the idea that, while moderate exercise is beneficial, intense exercise is harmful for the immune system. This general finding cannot be extended to the
innate/inflammatory response mediated by phagocytes, which can be stimulated even after intense sessions of exercise. The reduction of the functional capacity of lymphocytes in situations of excessive intense exercise can induce a temporary immunosuppression that allows microorganisms time to evade immunological recognition and become established, giving rise to infections in athletes. Innate immune defences may play an important role in the defence against infection of sportspeople, probably preventing the entry and maintenance of the antigen in situations where the adaptive immune response is depressed. Indeed, the stimulation of the innate/inflammatory responses during strenuous physical activity might counterbalance the decreased lymphocyte activity, and this may be regarded as an adaptation of this response to exercise-stress situations, in which stress hormones and mediators are involved (fig. 6).

It is also necessary to take into account that while innate and/or inflammatory responses are crucial in host defence for healthy people, uncontrolled inflammatory reactions may be responsible for the initiation and progression of autoimmune and inflammatory diseases. Thus, many of the benefits induced by exercise have been proposed to be mediated by the induction of an anti-inflammatory response. However, it is not formally proven whether an induced anti-inflammatory effect of exercise in healthy people, with an optimal inflammatory regulation could be beneficial for an optimal regulation of homeostasis. Then, the potential anti-inflammatory effects of exercise would be positive only for those people with unhealthy high inflammatory status. In addition, while cellular oxidative stress is dangerous for most of the cells, this process is necessary for phagocytes in order to destroy pathogens, and phagocytic cells also need a good balance between oxidative and anti-oxidative mechanisms. From an immunophysiological perspective, anti-oxidant and anti-inflammatory supplements make sense especially when one physical activity induces excessive oxidative/inflammatory response.

In this context, physical exercise has been found to improve the overall health of people suffering from certain autoimmune diseases such as the case of children with cystic fibrosis, who often have to be hospitalized because of acute exacerbation of their immune system exhibiting respiratory symptoms. Therefore, this is a powerful prognostic factor since aerobic fitness is associated with lower risk of hospitalization in children with cystic fibrosis, given that physical exercise can reduce the decline in VO₂ peak that occurs in these patients.

**Physical activity/exercise as a therapy**

**Cardiovascular diseases**

Exercise training has been shown to exert a restoring/improving effect on endothelial function, which is shown to be a risk factor for CVD. Longitudinal studies have shown that increased levels of physical activity reduce thrombosis-related cardiovascular events.

**Diabetes**

A meta-analysis has demonstrated that exercise training is associated with an overall 0.67% decline in glycated hemoglobin levels. This reduction, derived from the exercise, is similar to that reached by treatment of oral antidiabetic drugs.

A more recent meta-analysis of randomized control trials has shown a significant decrease in triglycerides...
after exercise interventions but no effects were found on total cholesterol, high-density lipoprotein cholesterol or low-density lipoprotein cholesterol. A meta-analysis was recently conducted on both the efficacy and tolerability of polypills in 2,218 subjects (fig. 7). The authors concluded that the polypills can reduce blood pressure and lipids in comparison with a placebo group and the differences found in both groups about the tolerability of polypills were moderate. Therefore, further studies are necessary to elucidate the status of polypills in primary care and prevention strategies.

Obesity

With the aim of losing weight there are more and more alternative programmes and therapies that include different types of exercise along with dietary food products, specific diets, besides of surgical methods, certain drugs and finally “miracle” type products.

However, it is important to highlight that thousands of these supposed miracle products and exercises very frequently show up in developed countries, claiming any number of healthy advantages, although there is always a lack of scientific precision. Facing to the difficulties found to lose weight the consumers still wonder about the reasons and about the short period of success if any. In this scene, the consumer should be helped by health professionals to dispel any doubt about what may be valid or under which circumstances these alternatives can be applied.

The most agreed explanation is the multifactorial component of overweight/obesity, such it can be seen at the following website: http://www.shiftn.com/obesity/Full-Map.html where many factors are involved, such as: energy balance, individual, social or collective

![Fig. 7.—Comparison on the effects of the polypill vs. exercise interventions on outcomes related to CDV risk using data from meta-analyses.](image-url)
psychological factors, individual and social factors of physical activity, energy consumption, access to food products, individual or collective physiological factors (genetic, epigenetic, etc.). This is the reason why those interventions available to combine a large group of factors are more sensible to achieve success.

Certainly, in the area of physical activity it is necessary to further fight for achieving successful goals by avoiding myths such as the use of abdominal exercise or vibration platforms to lose weight. Even the apparently best classic cardiovascular exercise has been evaluated and discredited in an interesting meta-analysis showing that isolated cardiovascular exercise only causes an average weight loss of 1.7 kg in 12 months.

Aging

Moreover, regarding aging, physical activity has been shown as a protective agent of healthy mental and organic improvement, leading to an enhanced autonomy situation and thus, contributes to attenuate aging autonomic dysfunction and to the reduction of morbidity and mortality.

Final remarks

Hydration status and physical activity/exercise are very important concepts to work on to tackle inadequate nutritional situations and promote well-being. Further research studies are necessary regarding the measurement of the hydration status and the binomial physical activity/sedentariness as well as the doses of water intake supplied both by foods and beverages and the ideal proportion of physical activity/sedentariness to achieve the healthiest status by reducing disease risks.

In view of all the opinions expressed by the speakers according to their knowledge, there was a consensus about a guideline as a Decaologue with several advices related both to hydration and physical activity/exercise to achieve an adequate health status.

Decaologue: global recommendations to achieve an adequate hydration and physical activity status to avoid overweight/obesity consequences

Hydration

1. For the general population, intake of 2-2.5 litres (L)/day of fluids, including water, infusions, natural juices, soups and beverages, all of them in moderation and variety.
2. For sportspersons, intake of adequate amounts of liquids according to the training periods or during competitions (80-350 kcal/1,000 mL, of which at least 75% should come from a mix of high glycemic load carbohydrates such as glucose, sucrose, maltodextrins and fructose).
3. In order to avoid hyponatremia, sportspersons should drink beverages with appropriate amounts of sodium (460-1,150 mg/L) between 0.6-1.2 L/hour in long-term sport activities.

Food consumption

4. The dietary daily intake has to be balanced: 55-60% of total carbohydrates from which 5-10% should be simple carbohydrates; 30-35% fats as long as 10% are monounsaturated fatty acids, especially supplied by olive oil, 10% saturated fatty acids and 10% polyunsaturated fatty acids; and 10-12% protein, providing a high variety of different foods but in moderate amounts (less is more).
5. Consumption of 20-30 g/day fibre (including legumes, natural cereals and 3 portions/day of each vegetables and fruits)
6. An appropriate food behaviour consisting of 4-6 meals during the day (breakfast, mid-day snack, lunch, afternoon snack, dinner, after dinner snack) in adequate amounts and at the appropriate times.

Physical activity, exercise and sedentary behaviours

7. 420 minutes of moderate to vigorous physical activity per week for children and adolescents, and at least 3 days per week of vigorous-intensity physical activity.
8. 150 minutes of moderate physical activity per week or 75 minutes at vigorous intensity for adults. Every minute of physical activity really does count and intensity plays an important role for health.
9. Sitting breaks every 30 minutes to avoid the negative consequences for health associated with prolonged sedentary behaviours.

Global remark

10. Globally, each person should find his/her own energy balance to achieve a healthy and well-being status through the establishment of an adequate nutritional, hydration and physical activity/inactivity status.

References


