Low calorie sweeteners: Role and benefits
This booklet has been developed for healthcare professionals and is designed to provide factual information on low calorie sweeteners, their characteristics, the evidence supporting their safety and how they can help manage calorie intake. It is based on publicly available science, with references and contributions from internationally recognised experts.
Introduction

With food plentiful in developed countries and more people leading sedentary lifestyles, the innate human preference for sweet taste is something we need to manage more effectively than ever before. High obesity rates show that more people need to focus on active, healthy lifestyles and energy balance – that is, balancing the calories consumed with the calories burned through physical activity.

Low calorie sweeteners provide a simple way of reducing the amount of calories in our diet without affecting the enjoyment of sweet tasting foods and drinks. As such, low calorie sweeteners can play a helpful role in assisting the achievement of weight maintenance or weight loss, as part of a balanced diet.

In recent years there has been a steady and significant increase in consumer demand for low calorie products. As a result there is growing interest among healthcare professionals and the general public to learn more about low calorie sweeteners, the foods and drinks in which they are found, how they help to reduce calorie intake and contribute to weight management and improved overall health.

Low Calorie Sweeteners: Role and benefits is supported by contributions from a group of eminent scientists and doctors who have undertaken a significant amount of research in the area of low calorie sweeteners, toxicity, epidemiology, appetite/satiety and weight management.

We hope you find this booklet useful and that it will serve as a valuable reference tool in your daily work.
Leading scientists and researchers working in the areas of toxicology, epidemiology, satiety and weight management have reviewed the content of this booklet and provided answers to the most frequently asked questions about low calorie sweeteners from their expertise:

**Professor Andrew Renwick OBE, PhD, DSc, Emeritus Professor, School of Medicine, University of Southampton (Southampton, UK)**

Professor Renwick’s work on species differences and human variability in metabolism and kinetics in relation to the safety factors used in risk assessment led to World Health Organisation (WHO) initiatives to develop chemical-specific adjustment factors. In 2002 he received the George H. Scott Memorial Award from the Toxicology Forum. He retired from the University of Southampton in September 2004.

He has published over 160 original research papers and 35 book chapters and other contributions on the metabolic fate of medicines and other foreign chemicals, on what happens to chemicals in the body, on food chemical safety and on low calorie sweeteners. He has served as a member of a number of UK Government Advisory Committees, and he was awarded an Officer of the Order of the British Empire (OBE) in the New Year Honours List in 2000. He was a member of the European Food Safety Authority’s (EFSA) Contaminants Panel for 2 years and has attended The Joint Expert Committee on Food Additives (JECFA) as a WHO temporary advisor for the past decade.

**Dr Adam Drewnowski, PhD, Professor of Epidemiology and Director of the Nutritional Science Program, University of Washington (Seattle, US)**

Dr Drewnowski is the Director of the Center for Public Health Nutrition and the University of Washington Center for Obesity Research and a joint member of the Fred Hutchinson Cancer Research Center in Seattle.

Dr Drewnowski develops new methods and measures to explore links between food and diet quality, price, and sustainability. He is the author of the Nutrient Rich Foods Index, which ranks foods based on their nutritional value and helps to identify affordable healthy foods. The Seattle Obesity Study (S.O.S.), led by Dr Drewnowski, applies spatial analyses to survey research in helping to determine who buys what foods, where, why and for how much. Dr Drewnowski has conducted numerous studies on hunger, appetite and satiety to determine how different nutrients and food ingredients can help in the management of body weight.

**Dr Carlo La Vecchia, Chief of Epidemiology, Mario Negri Institute (Milan, Italy)**

Dr La Vecchia received his medical degree from the University of Milan and a Master of Science degree in clinical epidemiology from the University of Oxford, UK. He is recognised worldwide as a leading authority in cancer aetiology and epidemiology with over 1,470 peer-reviewed papers published. Dr La Vecchia serves as an editor for numerous clinical and epidemiologic journals.

Dr La Vecchia is also an Adjunct Professor of Medicine at Vanderbilt Medical Center and the Vanderbilt-Ingram Cancer Center and Adjunct Professor of Epidemiology at the University of Lausanne, Switzerland as well as on the faculty of Medicine at the University of Milan. He is a temporary advisor at the WHO’s International Agency for Research on Cancer (IARC) and at the World Health Organisation in Geneva.
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The development of sweet taste

The sense of taste is of immense importance in the lives of animals and humans as it determines food choices and influences the amounts consumed. Taste, in conjunction with other senses, plays a crucial role in decisions about whether a potential food will be accepted or rejected, while ensuring the intake of sufficient nutrients. In humans, taste has the additional value of contributing to the overall pleasure and enjoyment of a food or drink.

There are five basic tastes (Figure 1): Sweet taste permits the identification of energy-rich nutrients; Umami taste allows the recognition of savoury amino acids (protein-rich foods); Salty taste ensures proper dietary electrolyte (mineral) balance; Sour or Bitter tastes warn against the intake of potentially noxious and/or poisonous substances.

The taste system is complete at birth. Anatomically complete taste buds and olfactory neurons can be identified in the human foetus by the 15th week of gestation, while olfactory neurons are apparently functional by about the 25th week. Although it is difficult to establish at what stage in gestation the foetus actually begins to experience taste and flavour, it has been known for more than 150 years that premature infants respond to some tastes.

The sensation of taste results from the chemical stimulation of specialised cells called Taste Receptor Cells (TRCs), which are grouped in small clusters called taste buds. Taste buds can be found throughout the oral cavity, but are mainly located on the human tongue. Taste buds are composed of 50-150 TRCs (depending on species) and are distributed across different papillae (Figure 2).

Circumvallate papillae are found at the very back of the tongue and can contain hundreds (for mice) to thousands (for humans) of taste buds. Folate papillae are present at the posterior lateral edge of the tongue and contain a dozen to hundreds of taste buds. Fungiform papillae contain one or a few taste buds and are found in the anterior two-thirds of the tongue (Figure 2).

It was thought for a long time that different parts of the tongue were sensitive to different tastes. Recent molecular and functional data have revealed that, contrary to popular belief, there is no ‘tongue map’. Responsiveness to the five basic tastes – sweet, salty, sour, bitter and umami – is present in all areas of the tongue.
From birth, taste and familiarity influence behaviour toward food. Scientists have proven that a liking for sweetness and a dislike for bitterness at birth are innate human traits. Taste preferences and food aversions develop later in life, through experiences that are influenced by our attitudes, beliefs and expectations.

**Responsiveness to sweetness**

Responsiveness to sweetness is a primitive response, observable even in the simplest of organisms. In humans, sweet taste exerts a profound influence on behaviour. The innate pleasure response to sweet taste, observable at birth, serves to orientate the feeding response and provides a motivation for continued feeding.

Scientists believe that our preference for sweetness may be an evolutionary survival mechanism, ensuring the acceptance of breast milk, with its slightly sweet taste from the milk sugar lactose, the primary carbohydrate found in human milk.

The most convincing scientific evidence for this in humans comes from studies of premature and newborn infants that demonstrate conclusively, using a variety of different research techniques, that infants are sensitive to and prefer sweetness on their very first tasting.

Investigations of newborns’ taste responses have uniformly indicated that they respond even to dilute sweet tastes, are able to differentiate varying degrees of sweetness and will consume more of a sweet-tasting sucrose solution when compared to water.

Several findings support the conclusion that positive facial expressions elicited by sweet tasting substances are reflex-like. Firstly, single response components, such as tongue movements, can be reliably elicited in newborns by sweet tastes in a concentration dependent manner.

Secondly, infants born with severe developmental malformations of the central nervous system react to the sweet taste like a normal term-born infant. Consistent with the above observations for intake and facial expression, infants can express that they perceive and respond positively to sweet stimuli via a variety of other behaviours.

Studies performed in the 1990s using sweet taste solutions of sugar and the low calorie sweetener aspartame showed that they encourage mouthing and sucking movements and hand-to-mouth contacts, both of which are feeding-related behaviours.

**Figure 3: Infant facial expressions**

Facial expressions, elicited from a 3-day-old infant, which suggest contentment and liking or discomfort and rejection, have been used to assess the newborn’s responsiveness to taste stimuli in some of the earliest investigations on human taste development.
The preference for sweetness beyond infancy

Sweetness makes foods more appealing and is often used to introduce new foods into children’s diets\(^1\),\(^6\). Children learn to accept new tastes and flavours when they are associated with sweetness or with energy-dense nutrients, such as starch\(^2\).

It is important to highlight that preference for very sweet solutions continues during childhood and adolescence\(^2\) and gradually decreases by early adulthood, as shown by both cross-sectional and longitudinal studies\(^2\).

The evidence to date suggests that, on average, children and adolescents prefer higher sucrose (sugar) concentrations in water than adults\(^2\),\(^4\),\(^6\),\(^7\),\(^8\).

In 1999 De Graaf & Zandstra\(^9\) showed that 9 to 10-year-old children preferred higher concentration levels of sugar in water and lemonade than adolescents (14 to 16-years-old) who, in turn, liked sweetness more than young adults (20 to 25-years-old).

One plausible biological explanation is that children require more energy than adults and that sweetness preferences are a marker for biological growth. In general, sweetness preferences decline once growth is completed\(^9\).

Ageing

Sweet taste preferences persist from birth until old age, although the liking of intensely sweet taste declines substantially between early childhood and adult life. Fewer studies have been conducted on sweet taste preferences in the elderly, although anecdotal reports suggest that the liking for sweetness increases in older people. The evidence suggests that adults have a reduced preference for sweet substances, preferring less intensely sweet foods and drinks compared to children\(^2\),\(^8\)-\(^9\).

The ability to detect and perceive sweet taste shows remarkable stability across the lifespan. On average, an individual’s ability to detect the sweet taste of sugars and to judge taste intensity is little changed between 20 and 80 years of age\(^9\). By contrast, sensitivity to smell sharply declines with age, robbing some elderly persons of the ability to enjoy the flavour and aroma of foods. Since aroma is a major component of taste, the decline in olfactory ability means that older adults may rely more on sweetness for their enjoyment of foods. However, preferences for sweetness never disappear. Even in old age, sweetness provides the motivation to eat \(^9\).

The pleasure reaction to sweet taste is observed across individuals of all ages, races, and cultures

In 1999 De Graaf & Zandstra\(^9\) showed that 9 to 10-year-old children preferred higher concentration levels of sugar in water and lemonade than adolescents (14 to 16-years-old) who, in turn, liked sweetness more than young adults (20 to 25-years-old).

One plausible biological explanation is that children require more energy than adults and that sweetness preferences are a marker for biological growth. In general, sweetness preferences decline once growth is completed\(^9\).
Sweet taste and today’s lifestyles

Our diets and food production have changed significantly since early humans hunted for and gathered food. Today’s rigorous food safety standards mean that we no longer need to rely on sweet taste to act as a screening mechanism to identify foods that are safe to eat. In fact, many food and drink delicacies are bitter, salty and sour due to evolving consumer preferences. However, throughout time, one thing has remained constant – our preference for sweetness.

Energy density of the diet is often perceived through the sensation of taste. Sweetness, the traditional sensory indicator of both nutrients and calories, adds to the sensory appeal of a given food. Indeed, the concepts of palatability and good taste have long been synonymous with the sweetness of foods.

Given a choice, young children prefer energy-dense foods over those that deliver fewer calories per unit weight. The main reason is that the presence of sweetness also signals the presence of calories, a major physiological reinforcement for the active, growing child. Energy density and palatability are therefore linked. In general, foods are palatable because their energy density is high. For low energy density foods to be liked, they need to be sweet. In general, sweet fruits are preferred over low energy density foods that are not sweet (e.g. spinach). Sweetened yogurt is preferred over plain yogurt.

The human preference for sweetness should be managed with care. All foods and drinks can fit into a healthy, active lifestyle that includes a sensible, balanced diet and regular physical activity. However, high obesity rates in children and adults highlight the need for most individuals to balance calories consumed with calories burned during daily activities. While reducing the energy density of foods might seem a logical approach to the prevention of obesity, the fact is that a bulky low-energy diet has low sensory appeal for the consumer.

As the energy density of foods decreases, so does palatability.

The modern food industry has managed to provide highly palatable foods and drinks with lower energy density. One of the best examples is the development of food ingredients called low calorie sweeteners. These sweeteners are added to many foods and drinks, but their impact on drinks is potentially the most significant – they can reduce the energy content to zero, while maintaining both palatability and sweetness. Foods and drinks sweetened with low calorie sweeteners represent a growing segment of the food supply.

Low calorie sweeteners and sweet taste

Sweetness clearly increases the palatability of numerous foods and beverages, and stimulates intake. Over the years, it has been suggested that low calorie sweeteners might enhance the natural appetite for sweetness, exacerbate the liking for and intake of sweet products of all kinds, and prevent consumers from managing their response to sweetness. These early results have been tested by numerous later works and failed to show any stimulatory effect of low calorie sweeteners on appetite, the liking for sweet taste and/or energy intake.

The vast literature in this field has been reviewed periodically over the last 30 years and systematic reviews and meta-analysis concluded that the use of low calorie sweeteners induced a decrease in daily energy intake and appetite for sweet foods or drinks and facilitated weight loss in dieters.

Observational Studies

In 1998, data from the large-scale epidemiological SuViMax study, which included 12,000 French adults, indicated that users of low calorie sweeteners had a lower daily energy intake (about 4%) than non-users, and ingested significantly less sugar. According to food choices reported in dietary questionnaires, it appeared that users of low calorie sweeteners ingested less potatoes and less sweets. The results provided no evidence suggesting enhanced appetite in general, or higher appetite for sugar and sweet products in users of low calorie sweeteners.

In the USA, the National Weight Control Registry was established in 1993 to study formerly obese individuals who successfully lost a significant proportion of their body weight (10% or more) and maintained a normal body weight periodically over the last 30 years and systematic reviews and meta-analysis concluded that the use of low calorie sweeteners induced a decrease in daily energy intake and appetite for sweet foods or drinks and facilitated weight loss in dieters.

Why do we like sweet tasting foods and drinks?

Dr Adam Drewnowski: The liking of sweet taste is a universal human trait. Infants, children, and teenagers of all races and cultures find sweet foods and drinks highly appealing. By associating pleasant taste sensations with nourishment, nature made sweetness a powerful driver of human eating behaviour. The taste response to sweetness was essential to survival. By contrast, bitter taste, associated with toxic compounds, was nature’s signal for dietary danger and led to instant food rejection by the child.

Whereas young children like very sweet tastes, adults do not. Sweet taste preferences typically decline once the child is grown. Adolescents and adults prefer less intensely sweet foods, experiment with other flavours, and even learn to tolerate bitter taste. However, preferences for sweetness never disappear. Even in old age, sweetness provides the motivation to eat.
Low calorie sweeteners do not increase our liking for sweetness, evidence suggesting that they actually lead to a decrease in appetite for sweet products.

A 2015 study by Antenucci and Hayes involving 401 participants found that low-calorie sweeteners, when used in amounts typical for preparing foods and beverages, do not enhance a person’s sweet taste buds and do not lead to sweet cravings and can be a helpful part of a healthy lifestyle. Participants perceived the sweetness of sugar substitutes at lower concentrations than real sugar, and the intensity of these sensations was not sweeter than sugar.

Randomised Controlled Trials
In the CHOICE study, a weight loss 6-month intervention, participants (n=104) were asked to replace daily intake of sugar-sweetened beverages with low calorie sweetened beverages, while another group (n=106) was asked to replace sugar-sweetened beverages with water. The six-month intervention did not include any other dietary prescription (no caloric restriction). The hypothesis tested was that consumption of low calorie sweeteners in beverages would enhance the consumption of sweet-tasting foods and beverages.

The dietary changes recorded in the intervention group did not support the hypothesis. The participants exposed to a high level of intake of low calorie sweetened beverages for six months significantly reduced their sugar intake (i.e. dessert items) during the intervention. Notably, intake of sugar-containing beverages, including tea and coffee, dropped significantly as it did in the water group. In the low calorie sweetened beverage group, however, intake of desserts and caloric sweeteners decreased more than it did in the water group. In this study, spontaneous intake data brings no support to the notion that low calorie sweeteners in beverages exert an enhancing effect on appetite in general, and on appetite for sweet-tasting products in particular. Actually the results suggest a broader suppression of appetite for sweetness in participants with a high daily intake of low calorie sweetened drinks rather than promoting it.

The DRINK study in 2012 was a randomised double blind trial carried out in school children. During an 18-month intervention 641 mostly normal weight children were randomised into two groups. One group received and drank a sugar-containing beverage (104 kcal) every day, whereas the control group received and drank a placebo drink (sweetened with low calorie sweeteners). Over the intervention phase, no compensation appeared for the absence of energy from the low calorie sweeteners containing drink, and experienced satiety was the same in both groups. Again this randomized controlled trial in children shows no support for the hypothesis that low calorie sweeteners might exacerbate liking or wanting of sweet tasting products.

A newly published trial by Peters et al. (2014 & 2016) of a 12-wk randomized behavioural intervention for weight loss and a 40-wk weight maintenance indicates a higher weight loss and maintenance in a group of obese participants required to drink low calorie sweetened beverages every day, compared to participants who were requested to drink equivalent amounts of water.

The greater weight loss in the low calorie sweeteners group was accompanied by a larger reduction in subjective feelings of hunger, contrary to what was observed in the water group. It is suggested by the authors that limiting access to sweetness (in beverages) in the water group may have promoted a desire to seek sweetness from other aspects of the diet, thereby leading them to consume more sweet foods resulting in greater energy intake and less weight loss.

Overall, the existing studies, using widely differing methodologies in various types of consumers (men, women, children, lean, obese, and formerly obese), reach largely convergent conclusions:

- The use of low calorie sweeteners shows no consistent association with a heightened appetite for sugar or sweet products.
- In many instances, the use of low calorie sweeteners is associated with a lower intake of sweet tasting foods or drinks.
- Recent intervention studies in children and adults confirm that low calorie sweeteners use tends to reduce rather than increase the intake of sugar-containing foods, and to facilitate, rather than impair, weight loss.
Low calorie sweeteners: the development of sweet taste

References

The use and role of low calorie sweeteners

Low calorie sweetener is the term used to describe compounds that taste sweet and provide few or no calories, or compounds that have such an intensely sweet taste that they can be used in food products at concentrations low enough not to contribute significantly to caloric content.

All low calorie sweeteners have a much higher sweetening power than sugar. This offers one major advantage to food and drink manufacturers and ultimately consumers – sweet taste whilst eliminating or substantially reducing the calories in a food or drink.

Low calorie sweeteners are used in a variety of food and drink products including soft drinks, chewing gum, confectionery, frozen desserts, yoghurts, dessert mixes and puddings. They are also widely used in healthcare, making many medicines more palatable. Low calorie sweeteners are clearly labelled on the packaging of food, healthcare and drink products that contain them.

Low calorie sweeteners have been safely used and enjoyed by consumers all over the world for more than a century. The first commonly used low calorie sweetener, saccharin, was discovered in 1879. Since then, a number of other low calorie sweeteners, including acesulfame K (ace-K), aspartame, cyclamate and sucralose, have been discovered and are now in widespread use worldwide.

In the European Union (EU), the most frequently used low calorie sweeteners are acesulfame-K, aspartame, cyclamate, saccharin, sucralose and steviol glycosides. A detailed look at the key characteristics of these low calorie sweeteners can be found on pages 30 and 31.

Each low calorie sweetener used in food and drink production has its own unique taste profile, technical characteristics and benefits. Low calorie sweeteners can be used alone or in combination with each other as a blend. By combining two or more low calorie sweeteners, it is possible for food and drink manufacturers to tailor the taste and characteristics of sweetness to the demands of a product and consumers’ tastes, whilst taking into account factors such as stability and cost.

When low calorie sweeteners are combined with each other they continue to be safe. Health authorities around the world have concluded that there is no scientific basis to expect any physiological effects to arise from the use of blends of approved sweeteners.

All of the low calorie sweeteners used in food production in Europe today have been subjected to rigorous safety testing. This is discussed in more detail in the next section. The regulatory process for low calorie sweeteners is thorough and gaining approval of a new low calorie sweetener is time and cost intensive, with the very stringent approval process often taking from 10 to 20 years.

Today’s more sedentary lifestyles and an increased interest in weight management mean that low calorie sweeteners can play an important part in achieving an active, healthy lifestyle. By replacing the equivalent sweetness of sugar without the equivalent calories and increasing palatability of healthy and low calorie foods, low calorie sweeteners, if used consistently to reduce calories, can act as an aid to weight reduction, weight maintenance and oral health. Furthermore, as low calorie sweeteners do not affect insulin levels, they may be used to provide sweet-tasting foods and drinks for people who must carefully monitor carbohydrate intake, such as people with diabetes. These benefits are covered in more detail on pages 17-23.
Low calorie sweeteners: their use and role

Q&A

What are low calorie sweeteners?

Prof. Andrew Renwick: Low calorie sweeteners are substances added to foods and drinks to provide sweet taste without calories, or with very few calories. Most low calorie sweeteners are several hundred times sweeter than table sugar, meaning that only small quantities need to be added to achieve a sweetening effect.

Though they are generally referred to as low calorie sweeteners by the scientific community, they are sometimes also referred to as sweeteners, artificial sweeteners, intense sweeteners and non-nutritive sweeteners.

Which foods and drinks contain low calorie sweeteners?

A wide variety of products contain low calorie sweeteners, including soft drinks, dairy products such as yoghurt and ice cream, desserts, chewing gums, condiments such as salad dressing, mustards and sauces and many other products including chewable multivitamins, mouthwashes and cough syrups.

Why do some foods and drinks contain blends of low calorie sweeteners?

Prof. Andrew Renwick: Each low calorie sweetener has its own unique taste profile, characteristics and benefit. By using different blends of low calorie sweeteners, subtle taste options are also available. Food and drink manufacturers choose which low calorie sweetener to use, either on its own or as a blend, based on taste considerations, stability and cost. Blending low calorie sweeteners is safe and the use of a blend reduces the amount of each individual sweetener that would be needed to sweeten foods and drinks.

How do you know if a low calorie sweetener has been added to foods or drinks?

As with any other food ingredient, they are shown on the label and the ingredients list with their full name and other times together with their E number. European legislation requires that foodstuffs and drinks containing a low calorie sweetener must bear the labelling ‘with sweetener(s)’. For the benefit of individuals with Phenylketonuria (PKU), where the low calorie sweetener aspartame is present in food and drinks, the label states that the product ‘contains a source of phenylalanine’. Individuals with PKU are diagnosed at birth and have to control the intake of phenylalanine in their diet.

In the EU, the most frequently used low calorie sweeteners in foods and drinks are ace-K, aspartame, cyclamate, saccharin, sucralose and steviol glycosides.

References

The safety and approval of low calorie sweeteners

EU legislation on sweeteners
Sweeteners were first regulated at European level in the 1990’s with the entry into force of Directive 94/35/EC of the European Parliament and the Council on sweeteners in foodstuffs, also known as the “Sweeteners Directive”.1

More recently, the European Parliament and the Council adopted a framework regulation, Regulation 1333/2008, to consolidate all current authorisations for sweeteners and food additives into one legal text.2 Annex II of this legislation, established by Commission Regulation 1129/2011, provides a Community list of sweeteners approved for use in foods, beverages and table-top sweeteners and their conditions of use. Where appropriate, maximum use levels are specified.3 Sweeteners must also meet EU purity criteria specifications.4

Within the EU, the 11 low calorie sweeteners currently authorised for use are acesulfame-K (E950), aspartame (E951), aspartame-acesulfame salt (E962), cyclamate (E952), neohesperidine DC (E959), saccharin (E954), sucralose (E955), thaumatin (E957), neotame (E961), steviol glycosides (E960) and advantame (E969).

At the request of the European Commission, the European Food Safety Authority is currently carrying out an ambitious re-evaluation of the safety of all food additives which were approved on the EU market before 20th January 2009. Aspartame is the first sweetener to have undergone this re-evaluation process. Opinions on the other sweeteners can be expected to be published by EFSA in the next few years.

EFSA opinion on aspartame
In December 2013, as part of the re-evaluation process and following one of the most comprehensive scientific risk assessments undertaken on a food additive, EFSA published its opinion on aspartame, re-confirming that aspartame is safe for consumers at levels currently permitted.5 Highlighting the publication of the opinion on its website, EFSA pointed out, ”Experts of ANS Panel have considered all available information and, following a detailed analysis, have concluded that the current Acceptable Daily Intake (ADI) of 40mg/kg bw/day is protective for the general population”. EFSA also highlighted that the breakdown products of aspartame (phenylalanine, methanol and aspartic acid) are also naturally present in other foods (for instance, methanol is found in fruit and vegetables).

The regulatory bodies involved
Regulatory approval of low calorie sweeteners in the EU is prepared by the European Commission on the basis of the scientific advice of the European Food Safety Authority (EFSA). The EFSA panel dealing with the safety of sweeteners is the ANS Panel (Food Additives and Nutrient Sources added to food), an independent panel comprised of members appointed on the basis of proven scientific excellence. The process is explained in more detail on page 14. Previously, the EU relied on the Scientific Committee on Food (SCF). The SCF was the scientific guarantor for the safety of food additives in use within the EU from 1974 to March 2003. Since April 2003, this has been the responsibility of EFSA. At a worldwide level this responsibility rests with the Joint Expert Scientific Committee on Food Additives of the United Nations Food & Agriculture Organisation and the World Health Organisation (JECFA). More details on these organisations can be found in the glossary on page 32.

Figure 1: Chronology of EU food additive law
How a low calorie sweetener is approved for use in foods and drinks in the EU

The authorisation and conditions of use of a low calorie sweetener, like any other food additive, is harmonised at EU level.

EFSA is responsible for the provision of scientific advice and scientific technical support for European Union legislation and policies in all fields that have a direct or indirect impact on food and food safety. Applicants (e.g. ingredient manufacturers) can only apply for approval of a low calorie sweetener after extensive safety tests have been completed and evidence provided of the product’s safety and utility. The petition provides technical details about the product and comprehensive data obtained from safety studies.

At a minimum, data is needed to answer the following:

- How the product will be consumed and how much will be consumed?
- Who, including vulnerable groups such as children or pregnant women, will consume the ingredient and how much will each group consume?
- Is the ingredient suitable for food processing use?
- What does the ingredient do, as an additive to food?
- Is there evidence to show that the ingredient does not cause adverse effects at the relevant use levels?

In the approval process, an Acceptable Daily Intake (ADI) is set for each low calorie sweetener by EFSA. The ADI is a guideline quantity that represents the amount of low calorie sweetener that can be safely consumed on a daily basis throughout a person’s lifetime without any health effects.

The safety data are then examined by EFSA. At any time, questions raised by EFSA must be answered by the applicant. Sometimes this may require additional studies. Completing and analysing the safety studies may take up to 10 years. Following the publication of a scientific opinion by EFSA, the European Commission drafts a proposal for authorisation of use of the low calorie sweetener in foods and drinks available in European Union countries.

After following the required procedure and only if the regulators are fully satisfied that the product is safe, will approval be given. This means that all of the low calorie sweeteners available in the EU market are safe for human consumption.

How acceptable daily intake (ADI) is established

The EU evaluation process establishes the ADI of low calorie sweeteners. The ADI is a measure of the amount of an approved additive that can be consumed daily in the diet, over a lifetime, without any health problems.

ADIs are expressed in milligrams (mg) per kilogram (kg) of body weight (bw) per day.

The ADI is usually based on the daily maximum intake that can be given to test animals throughout life without producing any adverse effects, known as the No Observed Adverse Effect Level (NOAEL); the ADI is calculated as the safe intake divided by a 100-fold safety factor to cover species differences and sensitive groups of the population such as children and the elderly. The use of the ADI principle for toxicological evaluation and safety assessment of food additives is accepted by all regulatory bodies worldwide.

Usage levels are set and use is monitored so that consumption does not reach ADI levels. As the ADI relates to lifetime use, it provides a safety margin large enough for scientists not to be concerned if an individual’s short-term intake exceeds the ADI, as long as the average intake over long periods of time does not exceed it. The ADI is the most important practical tool for scientists in ensuring the appropriate and safe use of low calorie sweeteners.

Consumption of low calorie sweeteners in Europe

Studies published in this area show that the average intakes of all low calorie sweeteners are well below the ADI values.

Recent studies have focused on children because of their higher intakes of foods and drinks on a bodyweight basis, and on both children and adults with diabetes, because of their higher potential intakes of low calorie sweeteners.
Section 3

Low calorie sweeteners: safety and approval

The NOAEL is divided by 100

The ADI is obtained

Our average consumption is more than 10 times lower than the ADI

NOAEL
No Observed Adverse Daily Effect Level
4000mg/kg/day

ADI
Acceptable Daily Intake
40 mg /kg/day

Average aspartame consumption in EU
4mg /kg/day

Figure 2: Aspartame Consumption Compared with ADI

In Belgium, a study by WIV (the government’s Scientific Institute for Public Health), concluded that low calorie sweeteners do not pose a health risk for adults, or for people with diabetes or for heavy consumers of light/diet products. The study showed that adults who regularly consume products containing low calorie sweeteners only reach a maximum of 25% of the ADI for cyclamate, 17% for acesulfame-K, 5% for aspartame, 11% for saccharin and 7% for sucralose.

In order to reach the established ADI for aspartame (40 milligrams (mg) per kilogram (kg) bodyweight (bw) per day), a woman weighing 60kg would have to consume every day during her lifetime 280 tabletop sweetener tablets or 20 cans of low calorie sweetened soft drinks. See Figure 2.

What is the meaning of an E number?

Prof. Andrew Renwick: If a food additive has an E number this is assurance that it has passed stringent safety tests and is approved for use throughout the EU. This approval is monitored, reviewed and amended in the light of new scientific data. In order to be authorised it must be demonstrated that an additive is both harmless and useful.

The ‘E’ reference for each additive refers to Europe and shows that the additive is authorised and regarded as safe in Europe. In effect, the E is a guarantee of safety. Food additives must be included either by name or by an E number in the ingredient list.

Why is there some speculation and concern over the general safety of low calorie sweeteners?

Dr Carlo La Vecchia: Over the past decades, various reports have claimed that low calorie sweeteners are associated with a range of adverse health effects. However, the evidence for these claims has been reviewed by international agencies, such as EFSA, and they have concluded that such claims are without substance. Much of the potentially frightening misinformation about low calorie sweeteners seems to be based on misunderstandings, data dredging or selective use of information, rather than a balanced view of all the information. The claimed adverse effects have not been found in subsequent studies. Nonetheless, unsubstantiated anecdotal reports have been widely covered in the media and online, leaving some consumers unsure as to whether low calorie sweeteners are safe.

Regulatory agencies, such as EFSA, continue to advise the European Commission that the use of low calorie sweeteners in foods and drinks, consumed within acceptable daily intake allowances, pose no threat to human health.

Is it true that low calorie sweeteners can cause neurological and mood problems?

Prof. Andrew Renwick: Anecdotal reports have suggested that low calorie sweeteners such as aspartame are associated with a range of behavioural and neurological problems such as headaches and epilepsy seizures. But research into aspartame and brain function does not support this. Controlled clinical studies have found no evidence of any neurologic or behavioural effects of aspartame in healthy adults or children, no effect of aspartame on cognition or behaviour in children with attention deficit disorder and no association between aspartame and seizures in individuals with seizure disorders.

Organisations representing the interests of people living with epilepsy, multiple sclerosis, Parkinson’s disease and Alzheimer’s disease have reviewed these anecdotal claims and concluded that there is no scientific basis to advocate the exclusion of aspartame and other low calorie sweeteners from the diet. This position is also supported by expert scientific committees of European agencies such as EFSA and national food safety agencies including AFSSA, the French Food Safety Agency, which in 2002 concluded that there was no evidence to link aspartame with the occurrence of epileptic seizures.
Can the consumption of low calorie sweeteners increase the risk of developing certain cancers?

**Dr Carlo La Vecchia:** No, there is no scientific evidence that links the consumption of low calorie sweeteners to cancer. Silvano Gallus and co-workers from the Institute of Pharmacological Research Mario Negri, published a study that further supports the claim that there is no indication that low calorie sweeteners cause cancer.

They studied the intakes of low calorie sweeteners in patients with a range of different cancers. Data were collected over a 13-year period on over 11,000 cases after taking into account various confounding factors (such as smoking), and it was determined that consumers of low calorie sweeteners were not at an increased risk of any of the cancers. Furthermore, when they divided low calorie sweetener use into saccharin, aspartame and other low calorie sweeteners, none of the results suggested a significant increase in any of the cancer forms. A subsequent report in 2009 found no association between low calorie sweeteners and gastric, pancreatic and endometrial cancer.

**Saccharin**

Saccharin safety was questioned after studies conducted in the early 1970s showed that high doses (equivalent to hundreds of cans of diet soft drinks a day for a lifetime) increased the incidence of bladder cancer in male rats. Subsequent laboratory studies demonstrated that this was specific to male rats; investigations in humans determined that there was a species-specific mode of action of saccharin on the male rat bladder. Epidemiology studies have shown there is no association between saccharin consumption and urinary bladder cancer, even in high intake consumers. Thus, research conducted over the past 25 years convincingly demonstrates that saccharin does not cause cancer in humans.

**Aspartame**

A paper published in 2005, by the European Ramazzini Foundation (ERF) claimed that there was a link between aspartame and cancer in rats and mice. However, the data showed no consistent dose-risk relationship had been established, nor was an adequate survival analysis completed. Furthermore, the claimed effects were not found in previous studies conducted according to regulatory guidelines. In response, EFSA’s panel on Food Additives and Nutrient Sources (ANS) conducted a detailed review of the ERF data and determined that there were significant flaws in the study, such that it did not provide evidence that aspartame causes cancer. There was therefore, no reason to revise the previously established ADI for aspartame of 40mg/kg bw/day, so aspartame safety was reaffirmed. The ANS panel has recently evaluated a subsequent study performed at the European Ramazzini Foundation (ERF) on mice and concluded that the data does not give reason to reconsider the previous evaluations of aspartame.

The methodology and conclusions of the ERF’s recent and previous studies on aspartame have been dismissed by EFSA and other independent scientific bodies. A statement released in February 2011, by the ANSES (French Food Safety Authority) concluded, “The methodology used (exposure to very high doses over a lifetime) is not used by any other group of researchers and, importantly, does not follow accepted reference methodology (OECD)”.

The 2007 review undertaken by Silvano Gallus and co-workers concluded that there is no new evidence that requires a revision of the existing opinions indicating a lack of genotoxic/mutagenic potential of aspartame.

There is extensive evidence to support the safety of aspartame. An epidemiological study from the National Cancer Institute in 2006 concluded that there is no link between aspartame consumption and leukaemias, lymphomas and brain tumours. The study evaluated over 500,000 men and women between the ages of 50 and 69 over a five-year period. The researchers found that there was no evidence of an increased risk of leukaemias, lymphomas and brain tumours among those who used aspartame.

Furthermore, a review published in 2007 concluded that suggestions of adverse effects had ‘no credible scientific basis’. The review was conducted by a panel of eight eminent experts over an 11-month period. It considered more than 500 studies, articles and reports conducted over the last 25 years, including work that was not published but that was submitted to government bodies as part of the regulatory approval process.

In April 2010, EFSA and its Advisory Forum in cooperation with a group of distinguished European Experts (18 experts from 10 EU countries), reviewed all papers published on aspartame since 2002 (SCF Review 2002) to address any remaining questions raised about the safety of aspartame for use in food. The National Experts concluded that no new evidence had been identified which required a recommendation to EFSA that the previous Opinions of EFSA and the SCF need to be reassessed.

Lastly, in December 2013, as part of the re-evaluation process and following one of the most comprehensive scientific risk assessments undertaken on a food additive, EFSA published its opinion on aspartame, re-confirming that aspartame is safe for consumers at levels currently permitted.

**Cyclamate**

Fewer allegations have been raised about cyclamate, and consequently less epidemiological data are available than for saccharin or aspartame. There is no evidence that cyclamate is associated to elevated risk at any cancer site, and no regulatory agency has raised concern on carcinogenic risk of cyclamate use. There is no evidence of toxicity of cyclamate nor of its carcinogenicity in animal experiments nor of cancer risk in humans. Independent scientists of the Joint FAO/WHO Expert Committee on Food Additives (JECFA) have consistently affirmed the safety of cyclamate for use as a sweetener in foods and drinks.
References

34. Aspartame : point d’étape sur les travaux de l’Anses : http://www.anses.fr/ (Translation EN)

Section 3

What is the meaning of ADI?

Prof. Andrew Renwick: ADI stands for Acceptable Daily Intake. The ADI is an estimate of the amount of an approved additive that can be consumed daily in the diet, over a lifetime, without any health problems. ADIs are expressed in milligrams (mg) per kilogram (kg) of body weight (bw) per day. The ADI is usually based on the daily intake that can be given to test animals throughout life without producing any adverse effects. The ADI is calculated as the safe intake divided by a 100-fold safety factor to cover species differences and sensitive groups of the population such as children and the elderly. The use of the ADI principle for toxicological evaluation and safety assessment of food additives is accepted worldwide by all regulatory bodies.

Who sets the ADI?

Prof. Andrew Renwick: JECFA introduced the concept of the ADI for the safety regulation of all food and drink additives in 1961. Different international scientific authorities such as JECFA, EFSA, and AFSSA (now ANSES) use the same method of deriving the ADI independently, guaranteeing consistency of food safety worldwide.

What if someone exceeds the ADI on any given day?

Prof. Andrew Renwick: The ADI for a food additive is not intended to define the maximum dose that would be safe to consume on any given day – it is a guideline quantity level. It is implicit that a person may occasionally consume an additive in quantities in excess of the ADI without adverse effects on the health.

In that case, why does the ADI matter?

Prof. Andrew Renwick: The ADI provides reassurance that a food additive, in this case a low calorie sweetener, is safe to consume. Setting ADI values and assessing consumption of low calorie sweeteners and other food additives according to their ADIs is a way of ensuring that the actual intake of food additives is well below a level that may be considered harmful to health. The presence of an ADI should not cause alarm, it is actually a guarantee of safety – only when a low calorie sweetener has been found safe by an independent scientific body and approved by the EU it is given an ADI.
Benefits of low calorie sweeteners for diet and health

Low calorie sweeteners, palatability and calorie intake

It is clear that to lose weight or to maintain a healthy weight, individuals need to pay careful attention to energy balance. It is important to balance the calories you consume with the calories you burn by maintaining a sensible, balanced diet combined with regular physical activity. In order to lose body weight you need to either burn more calories or consume fewer calories, or both. Energy density (kcal/g) of foods is an important determinant of energy intake in a meal1, 2 or over the course of the day3. By substituting low calorie sweeteners for sugar (kcal/g), it is possible to lower the energy density of foods and drinks. As a result, low calorie sweeteners can eliminate or substantially reduce the calories in some foods and drinks, offering an easy method of reducing calories while maintaining the palatability of the diet.

Where sugar is the main source of energy, as in soft drinks, low calorie sweeteners help to bring the energy density of drinks close to a negligible caloric content4. By contrast, sugar is only one of the ingredients of medium or higher energy density foods such as yoghurt or ice cream, with the remaining calories derived from protein or fat5. In high energy density foods such as chocolate, replacing sugar with a low calorie sweetener leads to relatively minor reductions in energy6.

Effect of low calorie sweeteners on appetite, satiety and food intake

The influence of low calorie sweeteners on hunger, satiety and food intake has been addressed in many laboratory studies and reviews6-13. While the use of low calorie sweeteners does not, in itself, result in a rapid weight loss, it may promote long term dietary compliance by improving the diversity, variety and the overall palatability of a reduced energy diet6.

In the 1980s, it was suggested that low calorie sweeteners may stimulate appetite, thereby increasing food intake and promoting weight gain. The appetite stimulation theory first drew attention when in 1986, Blundell and Hill15 reported that individuals who were consuming highly sweetened solutions perceived themselves to be hungrier than when they were consuming water alone16.

The study, however, relied only on subjects’ hunger ratings and did not measure their actual food intake, which is considered essential by psychologists and obesity experts. Blundell and colleagues conducted a subsequent study using solutions sweetened with several low calorie sweeteners and no increases in actual food intakes were observed17.

More than 30 years of research and several studies have examined the acute effects of low calorie sweeteners on hunger and food intake. They concluded that replacing sucrose (sugar) with low calorie sweeteners in foods or drinks does not increase food intake or hunger in children18-19 nor has it been shown to increase food intake in normal-weight20-24 or overweight men and women24-25.

Table 1: Difference in Calorie Content of Foods and Drinks with Sugar or Substituted with Low Calorie Sweeteners

<table>
<thead>
<tr>
<th>TYPE OF FOOD OR DRINK</th>
<th>CALORIE CONTENT: FOOD OR DRINK WITH SUGAR</th>
<th>CALORIE CONTENT: FOOD OR DRINK WITH LOW CALORIE SWEETENERS</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cola Soft Drink (330ml)</td>
<td>139 kcal</td>
<td>0.7 kcal</td>
</tr>
<tr>
<td>Fruit Drink (250ml)</td>
<td>184 kcal</td>
<td>27 kcal</td>
</tr>
<tr>
<td>Drinking Yogurt (250ml)</td>
<td>180 kcal</td>
<td>105 kcal</td>
</tr>
<tr>
<td>Strawberry Yogurt (125g pot)</td>
<td>118 kcal</td>
<td>84 kcal</td>
</tr>
<tr>
<td>Rasberry Jelly (100g)</td>
<td>80 kcal</td>
<td>5 kcal</td>
</tr>
<tr>
<td>Orange Squash (250ml)</td>
<td>110 kcal</td>
<td>5 kcal</td>
</tr>
<tr>
<td>Tea/Coffee with Sugar or Tabletop Sweetener</td>
<td>16 kcal (1 Teaspoon)</td>
<td>1 kcal (1 Tablet)</td>
</tr>
</tbody>
</table>
The primary evidence appeared in a number of studies with adults in the early '90s which had shown that familiar low calorie drinks sweetened with aspartame do not affect short-term appetite or food intake when they are consumed before lunch or with meals, compared with the effects of water\textsuperscript{20,25}. All of these studies reported either unchanged or reduced motivation to eat regardless of whether the low calorie sweetener was delivered in a solid or liquid medium. For more details, refer to Table 2 on page 21.

Studies throughout the '00s continued to affirm that low calorie sweeteners neither promote nor suppress appetite\textsuperscript{1,26-27}. Because of their volume, low calorie beverages may suppress appetite for about an hour, but do not appear to affect food intake at the next meal. By contrast, a caloric beverage will suppress appetite and may or may not reduce energy intake at the next meal. Satiety (fullness after consumption) was similar for children given low calorie sweeteners (LCS) containing beverages or sugar-sweetened beverages (SSB) in an 18-month intervention trial\textsuperscript{28} and similar effects have been found among adults\textsuperscript{27}. For example, in a crossover study by Anton \textit{et al} (2010) adults given foods sweetened with LCS (290 kcal) or sucrose (490 kcal) before lunch and dinner reported similar hunger and satiety ratings among participants who ate similar amounts of food at the next meal\textsuperscript{30}.

Another meal test study by Maersk \textit{et al} (2012)\textsuperscript{31} showed that after drinking 500 ml of a sugar sweetened soft drink (SSB) total energy intake (from the drink and the following \textit{ad libitum} meal) was higher compared with a diet soft drink or water. It was concluded that the energy provided by SSB was not fully compensated by decreased energy intake at the following meal, emphasizing the risk of generating a positive energy balance by frequent consumption of energy-containing beverages. Furthermore, there were no indications that the low calorie sweetener used in this study (aspartame) increased appetite or energy intake, compared with water.

When it comes to energy intake studies, they suggest that food and beverages sweetened with low calorie sweeteners may help reduce energy intake if used in place of more energy-dense food and drinks\textsuperscript{63}. Hence, the benefit of low calorie sweeteners will tend to be greater for drinks (where sugar is the main or only energy source) than for foods (where other macronutrients may be needed to replace sugar and provide bulk).

Reviews of randomised controlled trials in which low calorie sweeteners were used in place of sugar over several days or weeks conclude that \textit{ad libitum} energy intakes are lower with low calorie sweeteners because people only partly compensate for the missing calories\textsuperscript{27,32-33}. The degree of compensation of the food or drink is likely to vary depending on the physical form, composition, amount, timescale and individual factors. Variation in design may explain why some studies find a larger energy deficit than others\textsuperscript{27}.

Although more data from longer-term interventions are needed, it appears that low calorie sweeteners, especially in beverages, can be a useful aid to maintain reduced energy intake\textsuperscript{27,34-35}.

### Low calorie sweeteners in weight management and obesity

Strategies to reverse the upward trend in obesity rates need to focus on both reducing energy intake and increasing energy expenditure. The provision of low or reduced calorie foods is one way of helping people to reduce their energy intake and so assist weight maintenance or weight loss. Most studies investigating the role of low calorie sweeteners in weight control have shown that replacing foods and drinks in the diet with light or diet versions as part of a calorie controlled programme can result in an overall reduced caloric intake.

Pioneering work by Porikos and colleagues and by other researchers confirmed the positive effect of low calorie sweeteners on reducing caloric intake. In 1977 to 1984, Porikos \textit{et al}, showed that lean and obese participants living in a metabolic ward consumed fewer calories overall when all the available sources of sugar in their \textit{ad libitum} diet were replaced with aspartame\textsuperscript{36-37}. Research by Tordoff and Alleva\textsuperscript{38} found a similar effect and reported that consuming aspartame-sweetened soft drinks significantly reduced energy intake in both males and females eating their normal diet, and significantly decreased the body weight of males over a 3-week period when compared with a no-drink condition.

Further research had revealed that the use of low calorie sweeteners may help increase compliance with longer term weight management control programmes. Kanders \textit{et al}, measured weight loss, perceived feelings of energy and wellbeing, among 59 free-living obese men and women who were knowingly on a weight-control programme for 12 weeks\textsuperscript{39}. 

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\textsuperscript{1} References 1, 26-27. 
\textsuperscript{2} References 20, 25. 
\textsuperscript{3} References 28, 30. 
\textsuperscript{4} References 31. 
\textsuperscript{5} References 63. 
\textsuperscript{6} References 27, 32-33. 
\textsuperscript{7} References 27, 34-35. 

Low calorie sweeteners: benefits in diet and health
The experimental group was encouraged to use low calorie sweeteners, whereas the control group were encouraged to avoid all products with low calorie sweeteners. At the one-year follow-up, sustained weight loss was associated with increased low calorie sweetener consumption, a decreased desire for sweets and increased physical activity levels.

In addition, Blackburn et al., conducted the first large, randomised, controlled, prospective outpatient clinical trial investigating whether the addition of low calorie sweeteners to a multidisciplinary weight control programme would improve weight loss and long-term control of body weight in 163 obese women. The women were randomly assigned to groups that either consumed or abstained from foods sweetened with aspartame.

The results indicated that although both groups lost an average of 10% of their initial body weight (~10kg), those who consumed low calorie sweeteners were more successful in keeping the weight off in the long term. After 3 years, the group that consumed food sweetened with aspartame had kept off about half of the lost weight, maintaining a medically significant average weight loss of 5 percent of their initial bodyweight while the group that abstained from foods sweetened with aspartame had on average regained almost all the weight. This finding has important clinical implications given the poor long-term success rates of dietary treatments of obesity.

A review paper by Mattes and Popkin published in the American Journal of Clinical Nutrition in 2009 analyzed findings from 224 studies on the effects of low calorie sweeteners on appetite, food intake and weight. They found that longer-term trials consistently indicate that the use of low calorie sweeteners results in slightly lower energy intakes and that if low calorie sweeteners are used as substitutes for higher energy-yielding sweeteners, they have the potential to aid in weight management.

A review by Bellisle and Drewnowski published in 2007 examined whether reducing the energy density of sweet drinks and foods through the introduction of low calorie sweeteners can be a useful aid for weight control. Their review of clinical and epidemiological studies concluded that, although they are not a ‘silver bullet’, low calorie sweeteners can help people reduce their calorie intakes. A review of studies by De la Hunty et al, 2006 demonstrated that, “using foods and drinks sweetened with intense sweeteners instead of sucrose results in a significant reduction in both energy intakes and body weight”.

The Academy of Nutrition and Dietetics (formerly the American Dietetic Association) updated its position paper on nutritive and non-nutritive sweeteners in 2012 and concluded that “there is good evidence to support the use of aspartame and aspartame-sweetened products as part of a comprehensive weight loss or maintenance program by individuals may be associated with greater weight loss and may assist individuals with weight maintenance over time”.

In 2015, the International Journal of Obesity published one of the most thorough systematic reviews of the past decade, on low calorie sweeteners consumption and the reduction of energy intake and body weight. Rogers et al conducted a systematic review including meta-analyses of animal, human observational and human intervention studies providing information on low calorie sweeteners consumption and energy intake and/or body weight.

A considerable weight of evidence was found in favour of the consumption of low calorie sweeteners as helpful in reducing relative energy intake and body weight. On the other hand, no evidence was found from the many acute and sustained intervention studies in humans that the consumption of low calorie sweeteners increases energy intake. Most importantly, the effects of low calorie sweetened beverages on body weight also appear neutral relative to water or even superior in some contexts.

The review by Rogers et al confirms the outcomes of another systematic review and meta-analysis published one year before, in 2014 by Miller and Perez, which concluded that data from RCTs, which provide the highest quality of evidence for examining the potentially causal effects of LCS intake, indicate that substituting LCS options for their regular-calorie versions results in a modest weight loss and may be a useful dietary tool to improve compliance with weight loss or weight maintenance plans.
In terms of the long-term effectiveness of low calorie sweeteners, a new randomised clinical study by Peters et al., published in the peer-reviewed journal ‘Obesity’ in early 2016, provides strong evidence that low calorie sweetened beverages can help people to successfully maintain their body weight in the long-term. The study evaluated the effects of water versus low calorie sweetened beverages (LCS beverages) on body weight in a sample of 303 overweight and obese adults during a year-long behavioural weight loss and maintenance programme. The participants were randomly assigned to one of two groups: those who were allowed to consume LCS beverages (710ml/24 ounces daily) and those who were in a control group drinking only water. Results from the one-year trial, which was completed by 222 subjects (no difference between groups), showed that the LCS group had greater maintenance of weight loss and higher reduction in waist circumference (on average -8.67 cm vs -4.17 cm). In particular, participants drinking LCS beverages had better outcomes at 52 weeks (6.21±7.65 kg), compared to the group drinking water alone (2.45±5.59 kg; P < 0.001). Forty-four percent of the subjects in the LCS beverages group lost at least 5% of their body weight from baseline to year one compared to only 25% in the water group.

This new publication by Peters et al builds on the results from the first phase of the study, which found that subjects who consumed diet drinks lost 45 percent more weight than those in the water group (on average 5.95 kg vs. 4.09 kg, P < 0.001) during a 12-weeks weight loss programme. Forty-four percent of the subjects in the LCS beverages group lost at least 5% of their body weight from baseline to year one compared to only 25% in the water group.

Low calorie sweeteners also help people with diabetes or those on calorie restricted diets to feel less alienated because of their dietary requirements. Low calorie sweeteners and diabetes

Diabetes is recognised as a group of heterogeneous disorders with the common elements of hyperglycaemia and glucose intolerance, due to insulin deficiency, impaired effectiveness of insulin action, or both.

Diabetes is a chronic disease that occurs when a person’s body 1) can no longer make insulin, or 2) can no longer make enough insulin, or 3) cannot use insulin properly. Insulin is produced by the pancreas.

The importance of low calorie sweeteners in the diet of people with diabetes is undisputed. People with diabetes may, however, consume moderate amounts of sugar. If the sugar contained in food products stays for a longer time in the stomach and intestines (as is the case, for example, with chocolate), then people with diabetes may also enjoy some of these products.

Equally, when consumed with glucose, low calorie sweeteners do not appear to modulate the glycaemic response. Indeed, EFSA (European Food Safety Authority) has approved the health claim that low calorie sweeteners help reduce post-prandial glycaemic response.

Low calorie sweeteners offer people with diabetes broader food choices by providing the pleasure of the sweet taste without raising blood glucose. As low calorie sweeteners have no impact on insulin and blood sugar levels and do not provide calories, they can also have a role in weight loss and weight control for people with type II diabetes.

Furthermore, in 2012 the American Heart Association (AHA) and the American Diabetes Association (ADA) issued a joint scientific statement on low calorie sweeteners and their potential usefulness in helping people achieve and maintain a healthy body weight and help people with diabetes to control their glucose level. More details on diabetes can be found on page 23.
Table 2: Studies showing an impact on hunger, food intake or weight with the consumption of low calorie sweeteners

<table>
<thead>
<tr>
<th>AUTHOR</th>
<th>STUDY</th>
<th>CONCLUSION</th>
</tr>
</thead>
<tbody>
<tr>
<td>Rolls et. al (1989)</td>
<td>Healthy body weight; jelly or pudding with aspartame, 2hrs before meal; half aware others not</td>
<td>Reduction of hunger; aspartame helped reduced total kcal intake</td>
</tr>
<tr>
<td>Rolls et. al. (1990)</td>
<td>Healthy body weight; 8-16 oz of lemonade (sucrose or aspartame) or water</td>
<td>No difference in hunger or intake</td>
</tr>
<tr>
<td>Mattes (1990)</td>
<td>Healthy body weight; cereal with unsweetened cereal, or cereal with aspartame or sucrose; some aware others not</td>
<td>No difference in hunger or food intake</td>
</tr>
<tr>
<td>Canty et. al. (1991)</td>
<td>Healthy body weight; 7 oz of water or soda with saccharine, aspartame or sucrose (after breakfast, 1 hr. before lunch)</td>
<td>No increase in hunger ratings; No difference in kcal intake</td>
</tr>
<tr>
<td>Porikos et al, (1984)</td>
<td>Obese; calories diluted with low calorie sweetener Healthy wt; calories diluted with low calorie sweeteners</td>
<td>kcs reduced by 16%</td>
</tr>
<tr>
<td>Kanders et al. (1988)</td>
<td>Obese men and women. 12 weeks LCS group were given, in addition to weight loss diet, low-calorie, aspartame sweetened puddings or milkshakes and encouraged to use diet drinks</td>
<td>LCS group sustained weight loss at 1 yr follow-up</td>
</tr>
<tr>
<td>Tordoff and Alleva (1990)</td>
<td>30 non-obese men and women 3 weeks. subjects were given 1150ml of soft drink (~4 cans) sweetened with either aspartame or high-fructose corn syrup (HFCS) or no soft drink in a crossover study</td>
<td>LCS group (men and women) overall kcals reduced. LCS male group achieved weight reduction over the 3 –weeks</td>
</tr>
<tr>
<td>Kanders, Blackburn et al (1993, 1994)*</td>
<td>163 Obese women -19 weeks, weight loss programme groups that either consumed or abstained from foods sweetened with aspartame.</td>
<td>Both groups had 10% (~10kg) weight reduction. LCS group more successful in keeping the weight off in the long term.</td>
</tr>
<tr>
<td>Blackburn et. al (1997)*</td>
<td>Obese women, +3 year study. Intervention group were given, in addition to weight loss diet, aspartame-sweetened puddings or milkshakes and encouraged to use other aspartame-sweetened products</td>
<td>LCS group had sign. better weight maintenance (5% of weight loss) at +3yr follow-up.</td>
</tr>
<tr>
<td>De Ruyter et al. (2012)</td>
<td>A 18-month trial (DRINK) involving 641 normal-weight children 5-12 years of age. Each child was given daily either 250 mL [100 calories (kcals)] non-carbonated sugar-sweetened beverage (SSB) or 250 mL non-carbonated sugar-free beverage sweetened with sucralose and acesulfame potassium (Low Calorie Sweeteners). The provided beverage was given to replace their usual serving of SSB per day.</td>
<td>Average body weight was lower by 1 kg (2.2 lbs.) in the LCS group. When adjusted for differences in height, body weight in the LCS group was lower by 0.82 kg (1.8 lbs.). Children in the LCS group also gained less body fat (0.55 – 0.57 kg). All of these results were statistically significant.</td>
</tr>
<tr>
<td>Piernas et al. (2013)</td>
<td>Data derived from the CHOICE study. A sub-analysis evaluating the effect of water vs. diet beverages (DB) in a 6-month (total length) weight loss intervention, with analysis at 3 and 6 months. [water groups: n = 106 (94% women); DB group: n = 104 (82% women)].</td>
<td>The DB group decreased energy from all beverages more than the water group did only at month 3. The DB group had a greater reduction in dessert intake than the water group did at month 6.</td>
</tr>
<tr>
<td>Peters et al. (2014 &amp; 2016)</td>
<td>A randomized clinical study evaluating the effects of water versus low calorie sweetened beverages (LCS beverages) on body weight in a sample of 303 overweight and obese adults during 1 yearlong behavioural weight loss (12-weeks) and maintenance (40 weeks) programme.</td>
<td>Participants drinking LCS beverages had lost more weight and reduced waist circumference (WC) at 52 weeks (6.21±7.65 kg), compared to the group drinking water alone (2.45±5.39 kg; P &lt; 0.001). Nearly 19% more subjects in the LCS beverages group lost at least 5% of their body weight from baseline to year one. LCS beverages group subjects reported no increase in subjective hunger compared to water group subject.</td>
</tr>
</tbody>
</table>

*Longest trial to date on low calorie sweeteners and weight management
Low calorie sweeteners: benefits in diet and health

How can low calorie sweeteners benefit people with diabetes?
Dr Adam Drewnowski: The dietary control of blood glucose involves avoiding or sharply limiting foods that contain quick-release carbohydrates or simple sugars. Diabetic patients are advised to seek out sugar-free foods and drinks, and to select low-glycaemic index foods wherever possible. However, a sugarless diet of fibre and whole grains can lack in sensory appeal and may be hard to follow for long periods of time. Low calorie sweeteners offer an ideal way to preserve eating pleasure and promote dietary compliance. Replacing sugar in drinks with low calorie sweeteners maintains sweet taste and glycaemic control. Low calorie sweeteners can improve the quality of life of the diabetic patient.

Do low calorie sweeteners have an impact on appetite and food intake?
Dr Adam Drewnowski: Human appetite is influenced by both calories and volume of the just-consumed food or drink. A high calorie drink will suppress appetite and may reduce the amount of food eaten at the next meal. A zero calorie drink will suppress appetite for about an hour but will not affect the amount of food eaten in the next meal. Suggestions that low calorie sweeteners promote appetite and can result in overeating are incorrect. Numerous laboratory studies have shown no difference in appetite or satiety between drinks and yogurts that were plain and those that were sweetened with low calorie sweeteners.

Can low calorie sweeteners help people reduce their calorie intake and lose weight?
Dr Adam Drewnowski: Substituting sugar calories in a sugar-sweetened drink with a low calorie sweetener leads to a saving of some 150 kcal and reduces the energy density of the drink from 0.4 kcal per gram down to zero. Zero calorie drinks, in turn, help to bring down the energy density of the diet. Low energy density diets have been linked with higher nutrient density and with better management of body weight. In principle, a calorie saving of 150 kcal/day should lead to a substantial weight loss. However, it must be noted that low calorie sweeteners are most helpful when used in the context of an active, healthy lifestyle that includes a sensible, balanced diet and regular physical activity. Using low calorie sweeteners does not give permission to eat.

Low calorie sweeteners provide dental health benefits
When sugar-sweetened foods and drinks are consumed, the bacteria present in the dental plaque convert the sugar to acid. As this is repeated over time, this acid can attack the tooth enamel, leading to tooth demineralisation, which may contribute to increased risk of caries. Low calorie sweeteners are not fermentable, and do not contribute to tooth decay.

Studies show that products such as sugar-free chewing gum, which contains low calorie sweeteners, substantially reduce the risk of tooth decay. In 2010 as part of EFSA’s evaluation of health claims, EFSA concluded that there is a cause and effect relationship between the consumption of sugar-free chewing gum and reduction of tooth demineralisation and a reduction in incidence of caries.

Furthermore, by improving palatability, low calorie sweeteners can also encourage the use of toothpastes, mouthwashes and fluoride supplements that assist dental hygiene.
Low calorie sweeteners: benefits in diet and health

References


33. EFSA NDA (EFSA Panel on Dietetic Products Nutrition and Allergies (2011) Scientific opinion on the substantiation of health claims related to intense sweeteners and contribution to the maintenance or achievement of a normal body weight (ID 139, 1444, 4290), reduction of post-prandial glycaemic responses (ID 4298), maintenance of normal blood glucose concentrations (ID 1221, 4298), and maintenance of tooth mineralisation by decreasing tooth demineralisation (ID 1134, 1167, 1285) pursuant to Article 13(1) of Regulation (EC) No 1924/2006. EFSA Journal 9: 2229.


Low calorie sweeteners and special health considerations

Though the general population may not have any specific health concerns relating to the consumption of low calorie sweeteners, they may decide to seek the advice of a healthcare professional on whether they should or could include low calorie sweeteners in their diet. This section discusses the use of low calorie sweeteners by those sectors of the population with special health considerations: children, pregnant women, people with diabetes, or those suffering from the rare, inherited condition phenylketonuria (PKU).

Low calorie sweeteners and children

Low calorie sweeteners are not approved for use in foods for infants and young children. These foods are generally known as ‘baby foods’ and include foods specially formulated for infants and young children who are in good health, and foods for those whose digestive processes or metabolism is disturbed. ‘Infants’ are defined as children under the age of 12 months and ‘young children’ as children aged between one and three years.

The consumption of low calorie sweeteners by children is sometimes questioned for two main reasons – because of their physical size and relatively high food and drink intakes compared with adults. However, studies undertaken in this area indicate that children can safely consume low calorie sweeteners.

In comparison to adults, children tend to have the highest intake of low calorie sweeteners as calculated as milligrams (mg) of intake per kilogram (kg) of bodyweight (bw) per day. Though there is evidence that children’s intake of low calorie sweeteners is indeed greater than adult consumption, particularly within children with diabetes, there is no risk of exceeding the ADI levels.

In December 2002, European regulators stated that in the numerous studies focusing on the intake of aspartame by children in Europe, all found that even the highest consumption is well below the ADI. In the past, questions have been raised about whether low calorie sweeteners and specifically aspartame have behavioural effects on children. Studies conducted in this area demonstrate that aspartame has not been found to show behaviour effects. Controlled studies found no evidence of any neurologic or behavioural effects of aspartame in healthy adults or children, no effect of aspartame on cognition or behaviour in children with attention deficit disorder, and no association between aspartame and seizures in individuals with seizure disorders.

Key facts about aspartame, a widely used low calorie sweetener

- Aspartame is made from two amino acids (building blocks of protein) aspartic acid and phenylalanine (as the methyl ester).
- The two amino acids in aspartame are found in mother’s milk and occur naturally in foods such as meats, milk, fruits and vegetables.
- When aspartame is digested, it is broken down to common dietary components, including a small amount of methanol. More methanol is released by the metabolism of pectin in fruit juices, than is formed from a similar volume of an aspartame sweetened drink.
- The body uses these components in exactly the same way whether they come from aspartame or common foods.
- The foods we consume every day (e.g. chicken, milk, grape juice, tomato juice) provide much greater amounts of these components than aspartame does.
- The safety of aspartame has been confirmed by regulatory authorities in more than 100 countries. Most recently, the safety of aspartame was re-confirmed by the European Food Safety Authority as part of a re-evaluation programme for all food additives in the EU. See Section 3 for more information.
Low calorie sweeteners and pregnancy

The consumption of approved low calorie sweeteners within the ADI is safe during pregnancy.

Scientific communities around the world, including in Europe, have conducted safety evaluations in this area, focusing on the possible effects of low calorie sweeteners on pregnant women as well as any effects on the developing foetus. No evidence has been found to indicate that there is any risk to the mother or the foetus.

The above is also reaffirmed by the position of the Academy of Nutrition and Dietetics published in 2012 which states: “Pregnancy is a time of special concern because the focus is on maternal and fetal health. All FDA-approved low calorie sweeteners are approved for use by the general public, which includes pregnant and lactating women. The position of the Academy is that use of low calorie sweeteners is acceptable during pregnancy. Any low calorie sweetener that was found to be unsafe at any stage of life would not be approved for use.”

In the case of aspartame, a sweetener made from two amino acids, further evaluation of safety in pregnancy has been conducted to evaluate foetal exposure to the components of aspartame: aspartic acid, phenylalanine or methanol. Unlike other approved low calorie sweeteners, aspartame is metabolised in the human body. Enzymes in the digestive tract break aspartame down into its components, each of which is then metabolised just as it would be if derived from other dietary sources. All three of the metabolites of aspartame are present in many foods consumed as part of a normal diet. As a result of these studies, it was concluded that, like other low calorie sweeteners, it is safe to consume aspartame within the ADI during pregnancy.

The allegation, made by a group of Danish researchers, that the intake of low calorie sweeteners increases the risk of preterm delivery is not consistent with the extensive body of scientific evidence that shows these products are safe.

The authors suggested that methanol formed from aspartame could be responsible for the association. However, the main human exposure to methanol arises from the digestion of pectin, which is present in high concentrations in fruit juices (300-600mg/day). There is more methanol released from the digestion of fruit juice than from an equal volume of a low calorie sweetened carbonated drink. In addition a recent statement on the effects of chronic dietary exposure to methanol, released in March 2011, by the UK –COT (Committee On Toxicity Of Chemicals In Food, Consumer Products And The Environment) concluded that the exposure to methanol at the levels found in the diet both naturally occurring and from currently permitted levels of aspartame would not be expected to result in any adverse effects.

The use of low calorie sweeteners has been very well studied both in humans and in animals. This research has shown no adverse effects on the mother or the developing baby related to the use of low calorie sweeteners.

There are many factors that increase the risk of premature births, including being overweight or obese, smoking, diabetes, poor nutrition, anaemia, stress, depression and many more. At a time when the consequences of obesity, including in pregnancy, pose a significant challenge to public health, unsettling potentially sensitive population groups about choices that help them control their weight is particularly irresponsible.

In February 2011, EFSA reviewed and dismissed the recent publications by Danish researchers (Halldorsson et al., 2010) alleging that low calorie sweeteners may be unsafe. The EFSA Panel concluded that “there is no evidence available to support a causal relationship between the consumption of low-calorie sweetened soft drinks and preterm delivery.”

More recently, EFSA re-confirmed the safety of aspartame for the general population, including for pregnant women, in its full re-evaluation of this sweetener which was published in December 2013. See Section 3 for more information.

Aspartame and people with phenylketonuria

Phenylketonuria (PKU) is a rare inherited condition affecting 1 in 10,000 people. Those who have it lack the enzyme that converts phenylalanine into the amino acid tyrosine. Phenylalanine is an essential amino acid and is a building block of protein. It is also a component of aspartame. For those with PKU, consuming protein-containing food leads to a build up of phenylalanine in the body. People with PKU must limit their intake of phenylalanine to prevent it reaching toxic levels in their blood and other tissues.

Figure 1: Example of a drink label carrying a statement indicating that the product contains phenylalanine

Ingredients: Water, Orange Fruit from Concentrate (11%), Citric Acid, Acidity Regulator (Sodium Citrate), Preservatives (Potassium Sorbate, Sodium Metabisulphite), Sweeteners (Aspartame, Saccharin), Flavourings, Stabiliser (Cellulose Gum), Colour (Beta-Carotene).

Contains a Source of Phenylalanine.
Management of PKU requires a low phenylalanine diet. This means that high protein foods such as meat, cheese, poultry, eggs, milk/dairy products and nuts are not permitted. Instead the diet is supplemented with artificial protein which contains a low level of phenylalanine.

For the benefit of individuals with PKU, foods, drinks and healthcare products that contain the low calorie sweetener aspartame must legally carry a label statement indicating that the product contains phenylalanine. (Figure 1).

Throughout most of Europe, PKU is screened for shortly after birth.

Low calorie sweeteners and people with diabetes

Based on the latest figures released by the International Diabetes Federation16, it is estimated that, in Europe in 2015, 59.8 million people between 20-79 years of age suffer from diabetes. A further 31.7 million people, 4.8% of adults aged 20-79, are estimated to be living with impaired glucose tolerance and are at increased risk of developing diabetes. Indeed, by 2040, it is predicted that there will be 71.1 million adults living with diabetes in the Europe Region16. (figure 1)

**Type 1 diabetes mellitus:**
Sometimes called insulin-dependent diabetes, type 1 diabetes is caused by destruction of the insulin producing cells of the pancreas, typically due to an auto-immune reaction, where these cells are attacked by the body’s defence system. The beta cells of the pancreas therefore produce little or no insulin, the hormone that allows glucose to enter body cells (Figure 2). The disease can affect people of any age, but usually occurs in children or young adults. People with type 1 diabetes need injections of insulin every day in order to control the levels of glucose in their blood. (figure 2)

**Type 2 diabetes mellitus:**
Characterised by insulin resistance and relative insulin deficiency, either of which may be present at the time that diabetes becomes clinically manifest. The diagnosis of type 2 diabetes usually occurs after the age of 40 years but could occur earlier, especially in populations with high diabetes prevalence and/or in individuals who are overweight or obese. There are increasing reports of children developing type 2 diabetes. Type 2 diabetes is often, but not always, associated with obesity, which itself can cause insulin resistance and lead to elevated blood glucose levels. (figure 2)

**Gestational diabetes:**
Appears during pregnancy and can lead to serious health risks to the mother and her infant and increases the risk for developing Type 2 diabetes later in life. (figure 2)

As low calorie sweeteners have no impact on insulin and blood glucose levels, their importance in the diets of people with diabetes is undisputed. Low calorie sweeteners offer people with diabetes the pleasure of sweet tasting foods and drinks without raising blood glucose. Food and drink products prepared with low calorie sweeteners are suitable for people with diabetes17. Scientists have also concluded that low calorie sweeteners help people with type 2 diabetes in the important and often challenging task of controlling their weight18.

In 2012, the American Heart Association (AHA) and the American Diabetes Association (ADA) issued a joint scientific statement on low calorie sweeteners and their potential usefulness in helping people achieve and maintain a healthy body weight and in helping people with diabetes to control their blood glucose levels 19.

The statement highlights that:

- Substituting low calorie sweeteners for sugars added to foods and beverages may help people reach and maintain a healthy body weight – as long as the substitution doesn’t lead to eating additional calories later as “compensation.”
- For people with diabetes, low calorie sweeteners used alone or in foods and beverages remain an option and when used appropriately can aid in glucose control.
Q&A

Should pregnant women avoid consuming products with low calorie sweeteners?

Dr Carlo La Vecchia: Women do not need to avoid or be concerned about consuming low calorie sweeteners whilst pregnant. Consumption of low calorie sweeteners, within the ADI set by EFSA, is safe during pregnancy, because all low calorie sweeteners have been subject to appropriate testing. The variety of foods and drinks sweetened with low calorie sweeteners can help satisfy a pregnant woman’s taste for sweetness while adding few or no calories. In any case, pregnant and breastfeeding women need to consume adequate calories to nourish the foetus or infant and should consult with a physician about their nutritional needs. It is important to remember that weight control remains a priority, particularly in pregnancy.

Should children be consuming products containing low calorie sweeteners?

Dr Carlo La Vecchia: Low calorie sweeteners are also safe for children, but it is important to keep in mind that children, particularly young children, need ample calories for rapid growth and development.

Are people with diabetes at risk of overconsumption of low calorie sweeteners in their diet?

Prof. Andrew Renwick: No, people with diabetes are not at risk of overconsumption of low calorie sweeteners. Overconsumption would require an individual to exceed the ADI. As low calorie sweeteners are very beneficial for people with diabetes and are regularly included in the diet, the consumption of low calorie sweeteners by people with diabetes has been carefully monitored by the scientific community. Studies have shown that the intakes of low calorie sweeteners across Europe are well below the ADI, including within people with diabetes. Even within children with diabetes, the group with the highest potential for intakes, various studies have shown that the intakes are still below the ADI.

In 2015, an estimated 59.8 million people in Europe between 20–79 years of age are suffering from diabetes. By 2040 it is estimated that this number will increase to 71.1 million.

References

Low calorie sweeteners and healthy lifestyle habits

Following a healthy eating pattern is the new trend in the area of nutrition and many experts agree it’s time to stop focusing on individual nutrients, and time to start talking about healthy eating patterns overall. A healthy eating pattern supports a healthy body weight and can help prevent or reduce the risk of chronic diseases by integrating healthy dietary behaviours and regular physical activity. A number of dietary behaviours have been linked to a better diet quality and a healthy eating pattern, including the use of low calorie sweeteners and foods and drinks containing them.1,2

Low calorie sweeteners and higher diet quality go hand-in-hand

In early 2016 Gibson et al.2 published evidence from 1590 participants of the UK National Diet and Nutrition Survey (NDNS). In the study they explored if people who consume low calorie sweetened beverages (LCBs) tend to follow healthier diets and have lower energy, saturated fats and sugars intake, not only compared to individuals who prefer sugar sweetened beverages (SSBs) but also versus people not consuming any kind of beverage at all (non-consumers) or those consuming both types of drinks.

The study2 found that diet drinks’ consumers had a better diet quality, which was similar to non-consumers, as both groups had higher fish, fruits and vegetables consumption, and lower meat and sugar intake, compared to SSBs consumers and/ or those individuals drinking both SSBs and LCBs. Most importantly, LCBs consumers had an identical mean total energy intake (1719 kcal/day) as non-consumers (1718 kcal/day) and a significantly lower energy intake compared to SSBs users (1958 kcal/day) and consumers of both type of beverages (1986 kcal/day).

Furthermore, in terms of macronutrients intake, in comparison to SSBs group and both beverages consumers, LCBs group had:

- significantly lower intakes of sugars (both as g/day and % of energy intake),
- significantly lower intakes of both fat and saturated fatty acids (on an absolute basis but not as % of energy intake),
- significantly higher protein intake (as % of energy intake).

Low calorie sweeteners use is associated with higher healthy eating index and more physical activity in further studies

More publications have found that the consumption of low calorie sweeteners and foods and drinks containing them is related to a healthier diet quality and overall lifestyle. For example, a study1 in Nutrients (2014) suggests that individuals who consume low calorie sweeteners tend to have better diet and exercise habits. Researchers from the University of Washington looked to gain insights into the health habits of low calorie sweeteners consumers. They used data from the National Health and Nutrition Examination Survey (NHANES) collected between 1999 and 2008. NHANES captures health and nutrition information from more than 22,000 people across the US. Using this large data source gave the researchers a thorough view of how Americans who consume low calorie sweeteners eat and exercise and of their overall characteristics. Low calorie sweeteners users (vs. no users) were found to be more likely females, older, educated, and have a higher socioeconomic status.

The researchers reviewed the participants’ diets using the Healthy Eating Index, a USDA tool to compare an individual’s diet to the Dietary Guidelines for Americans. Low calorie sweeteners consumers had much higher scores on the Index than those who did not consume LCS. This score means low calorie sweeteners users tend to be at the head of the class in having a healthy diet. LCS users also reported similar energy intakes but higher intakes of fruits, vegetables, calcium and magnesium, as well as lower intakes of fat, added sugars, and saturated fats, compared to non-users. So, overall, low calorie sweeteners users had a better diet quality (figure 1).

![Figure 1: Healthy Eating Index in consumers vs. non-consumers of low calorie sweeteners.](source: Center for Public Health Nutrition, University of Washington)
Further, analysis\(^1\) showed that individuals who consume LCS tended to be more physically active than those who don't. They were also less likely to be smokers. This study shows that low calorie sweetener consumption correlates with better and healthier lifestyle overall, therefore, suggesting that low calorie sweeteners consumption and healthful eating and exercise habits tend to go hand-in-hand (figure 2).

The above findings are supported by an earlier study by Sigman-Grant and Hsieh in 2005 that suggested people who regularly use low calorie sweeteners may choose healthier diets\(^5\).

In addition, in a 2014 online survey by Catenacci et al\(^6\), showed that consumption of low calorie sweeteners is high among successful “weight loss maintainers” (individuals who have not only lost weight, but also kept it off). Low calorie sweetener consumers try to manage their energy intake by choosing foods and drinks containing LCS instead of caloric sweeteners. They are also eating higher-quality, more balanced diets and getting more physical activity.

When combined with a healthy diet, being physically active and adopting a healthy lifestyle, using low calorie sweeteners to reduce calories is a winning strategy for improved weight management and better health.

Low calorie sweeteners use is related to the intent to lose and maintain body weight

A pioneering study published in 2016 by Drewnowski & Rehm\(^7\) showed that the intent to lose or maintain body weight was one likely predictor of current LCS use and also that low calorie sweeteners consumption was not unique to obese individuals but held at all levels of BMI. That suggests that LCS use was tied directly to dieting behaviors, regardless of whether the participants were overweight or obese.

The recently published analysis\(^7\) merged National Health and Nutrition Examination Survey (NHANES) dietary intake data with retrospective weight control histories, a rarely exploited resource within NHANES (data from five NHANES cycles in a representative US sample of 22,231 adults). The study findings confirm the main hypothesis that trying to lose or maintain body weight over a 12-month period was associated with higher LCS use, independent of body weight. Specifically, it was found that individuals who tried to lose weight during the past year were 64% more likely to consume any type of LCS product. Similar results were obtained with the ‘trying to not gain weight’ variable. Furthermore, LCS use was much more common among individuals who experienced significant weight change in the preceding 10 years as compared with those who did not. This new finding confirms what has been assumed for years, that people troubled by weight management turn to LCS as a strategy for weight control, rather than the other way around. As explained by the authors, this is a typical example of reverse causality.

Figure 2: Low calorie sweeteners consumers are less likely to smoke and more likely to engage in physical activity.

References

### Characteristics of low calorie sweeteners commonly used in Europe

<table>
<thead>
<tr>
<th></th>
<th>ACESULFAME POTASSIUM (Ace-K) E950</th>
<th>ASPARTAME E951</th>
<th>CYCLAMATE E952</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Composition</strong></td>
<td>A combination of an organic acid and potassium</td>
<td>Aspartame is made from two amino acids aspartic acid and phenylalanine. These are found naturally in the diet.</td>
<td>Cyclamic acid (an organic acid), sodium- or calcium salt</td>
</tr>
<tr>
<td><strong>Acceptable daily intake ADI</strong> (for children &amp; adults)</td>
<td>0-9 mg/kg</td>
<td>0-40 mg/kg</td>
<td>0-7 mg/kg</td>
</tr>
<tr>
<td><strong>Sweetening power compared to table sugar (sucrose)</strong></td>
<td>ca. 200 times sweeter</td>
<td>ca. 200 times sweeter</td>
<td>ca. 50 times sweeter</td>
</tr>
<tr>
<td><strong>Year discovered</strong></td>
<td>1967</td>
<td>1969</td>
<td>1937</td>
</tr>
<tr>
<td><strong>Metabolic and physiological properties</strong></td>
<td>Not metabolised by the human body and excreted unchanged</td>
<td>Digested like other proteins to its components all of which occur in the diet in greater quantities.</td>
<td>Generally not metabolised and excreted unchanged</td>
</tr>
<tr>
<td><strong>Caloric value</strong></td>
<td>Calorie free</td>
<td>4kcal/g (used in very small amounts)</td>
<td>Calorie free</td>
</tr>
<tr>
<td><strong>Stability</strong></td>
<td>Heat stable, suitable for cooking and baking</td>
<td>Loses sweetening properties when exposed to high temperature, therefore not recommended for baking</td>
<td>Good stability at high and low temperatures, can be used in cooking and baking</td>
</tr>
<tr>
<td></td>
<td>Readily soluble</td>
<td>Can be added to foods at the end of cooking cycle</td>
<td>Good solubility</td>
</tr>
<tr>
<td><strong>Uses</strong></td>
<td>Used in drinks, foods, table-top sweeteners, oral-care and pharmaceutical products</td>
<td>Widely used in beverages, dairy products, table-top sweeteners and confectionery incl. chewing gum, due to its sweet taste profile.</td>
<td>Used in table-top sweeteners, drinks, chewing gums, salad dressings and jams</td>
</tr>
</tbody>
</table>
| **SACCHARIN**
| **E954** |
| **Saccharin (an organic acid), sodium or calcium salt** |

| **SUCRALOSE**
| **E955** |
| Derived from sugar in a process that selectively substitutes three atoms of chlorine for three hydroxyl groups on the sugar molecule |

| **STEVIO GYLCOSIDES**
| **E960** |
| Steviol glycosides are natural sweet tasting constituents of *Stevia rebaudiana*, a plant native to South America. Steviol glycoside preparations usually contain as the major components the glycosides Stevioside and Rebaudioside A. |

| **0-5 mg/kg** |
| **0-15 mg/kg** |
| **0.4 mg/kg (expressed as Steviol)** |

| **ca. 500 times sweeter** |
| **ca. 600 times sweeter** |
| **ca. 200 to 300 times sweeter depending on the glycoside** |

| **1879** |
| **1976** |
| **1931 (first isolated)** |

| **Not metabolised and excreted unchanged** |
| **Not metabolised and excreted unchanged** |
| Steviol glycosides are broken down to steviol in the gut. Steviol is excreted in the urine as steviol glucuronide. |

| **Calorie free** |
| **Calorie free** |
| **Calorie free** |

| **Heat stable** |
| **Good stability in very high temperatures. Can be used in cooking and baking** |
| Steviol glycosides are heat stable |

| **Can be used in cooking and baking** |
| **Good solubility** |

| **Used in table-top sweeteners, drinks, desserts, confectionery and also in pharmaceutical products** |
| **Used in baked goods, desserts, table-top sweeteners, ice-cream and dairy products, breakfast cereals and confectionery** |
| **Used in foods, drinks and table-top sweeteners** |
Low calorie sweetener: The term used to describe compounds that taste sweet and provide no calories or compounds that have such an intensely sweet taste that they can be used in food products at concentrations low enough not to contribute significantly to calorific content.

SCF (The Scientific Committee on Food of the Commission of the European Union):
The SCF was established in 1974 and reformed in 1997. It advised the Commission of the European Union (EU) on issues relating to the protection of health and safety of persons arising from the consumption of food. It was responsible for risk assessment of food additives until establishment of the EFSA. [http://ec.europa.eu/food/fs/sc/scf/index_en.html](http://ec.europa.eu/food/fs/sc/scf/index_en.html)

Sugar: Sugars are the naturally-occurring nutrients that make food taste sweet. Sucrose (also referred to as table sugar) is a crystalline or powdered substance, white when pure, consisting of sucrose obtained mainly from sugar cane and sugar beets and used in many foods, drinks, and medicines to improve their taste. There are a number of other different sugars including glucose and fructose, found in fruit and vegetables, milk sugar is known as lactose, and maltose is found in malted drinks and beer. All types of sugar have the same nutritional value, 4 kilocalories per gram.

The European Food Safety Authority’s ANS Panel:
The panel on food additives and nutrient sources added to food (ANS) deals with questions of safety in the use of food additives, nutrient sources and other substances deliberately added to food, excluding flavourings and enzymes. EFSA’s panels are composed of independent experts appointed on the basis of proven scientific excellence.
The International Sweeteners Association is a non-profit making, industry funded organisation representing manufacturers and users of low calorie sweeteners. The ISA is recognised by the European Commission, national and international regulatory authorities, and the World Health Organisation, and has Non-Government Observer status with the Codex Alimentarius Commission which establishes international food standards. [www.sweeteners.org](http://www.sweeteners.org)