

# The Alkaline Diet and the Warburg Effect

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## Abstract

The changing diets accompanying our modern life style have increased the content of foods that form acidic metabolic waste residues in the body. Wastes from these metabolic processes are released into the interstitial fluids and the blood, slightly changing their pH temporarily. This link may in turn have an impact on the incidence of non-communicable diseases (NCDs).

According to the Warburg Effect, an acidic cellular and circulatory environment may cause various specific health problems such as hypoxia and cancer, whereas an oxygen-rich optimum-alkaline environment could retain healthy cells. However, the mechanisms by which the diet may be influential on blood pH-related parameters and on health have remained largely unknown.

This paper begins with a detailed presentation of the concepts, issues and the existing evidence regarding alkaline and acid forming diets, and summarizes the three main mechanisms by which the diet influences the acid-base balance in the body. It then presents the findings of a small exploratory study in which one author (HB) followed diets traditionally thought to produce alkaline or acidic residues. After each diet period of one month (with a two-month wash out period), pH, oxygen saturation, and carbon dioxide partial pressure were measured for arterial and for venous blood.

The resulting data indicated that the diets followed changed blood pH-related parameters in the expected directions according to the acid-base theory of health. Increased intake of acid-forming foods resulted in a slightly lower pH level, but a significant reduction of oxygen saturation in the blood, whereas increased consumption of alkaline forming foods maintained the high oxygen saturation in the blood that, according to the Warburg Effect, may, if maintained, reduce the incidence of NCDs. Further cross-over research of this kind is needed, utilizing large samples and testing various dietary modifications.

## Introduction

### The impact of diet on the acid base balance in the body

Diet is partially responsible for serious non-communicable diseases (NCDs) [1, 2, 3, 4, 5, 6], but the mechanisms for this epidemiological link are not well known. This paper explores the potential effect of diet on temporary pH imbalances at cellular and circulatory levels as a possible partial explanation for this link. In particular, this may be important in the development of cancer.

There are three major mechanisms by which the diet has an impact on the maintenance of the acid-base balance at various levels: (1) the varying impacts of diet on digestion, resulting in a temporary “acid tide” or “alkaline tide” in parts of the circulatory system, (2) the acidity or alkalinity of food and drink as ingested, and (3) temporary acid-base imbalances at cellular level caused by the pH of the residues produced in processes of respiration and other metabolic functions, which may be influenced by the overall diet, as well as exercise. The most powerful of the three is likely the third, which is also the most poorly researched and most often misunderstood. In the remainder of this introduction, we will summarize the major issues in acid-base balance and then describe these three mechanisms of pH regulation in detail.

### Basic acid-base issues

The pH refers to the concentration of hydrogen ions in a solution, measured from 0 to 14 on a logarithmic scale, 0-7 being acidic and 7-14 being alkaline or base [7]. This may vary considerably from one area of the body to another. For example, the fluids in the stomach are very acidic (pH~1.5-3.5), but are neutralized when its contents enter the intestine, which is slightly alkaline [8].

The blood pH is normally maintained in the range of 7.35-7.45 [8]. The arterial blood is more alkaline and has a higher oxygen content than venous blood. If blood pH goes below 7.35, it is said to be acidic (acidemia), and if it goes above 7.45, it is alkalotic (alkalemia). A pH distant from this normal range means there is a severe, sometimes life-threatening acid-base disorder [9, 10].

Blood pH-related parameters are controlled by various buffering mechanisms in order to maintain the pH level within strict limits [11, 12, 13, 14, 15, 16]. These include regulating concentrations of bicarbonates, alkali salts and acids, and adjusting saturations for oxygen and carbon dioxide [17].

The primary method of pH regulation is via an increased rate of respiration [16]. When carbon dioxide levels are excessive, faster breathing increases oxygen saturation and decreases carbon dioxide levels. This mechanism is rapid and partially overcomes the problem, but cannot safely be complemented via a conscious effort to breathe faster.

The secondary mechanism that the body uses to deal with a longer term excessively acidic state is to excrete acids via the kidneys (in urine), via saliva, and through the skin (in sweat) [11, 18] and to reabsorb bicarbonate. If excess alkalinity is present, the kidneys excrete more bicarbonate into the urine [16, 17, 19, 20].

The tertiary mechanism of controlling blood pH over an even longer term is to use bone minerals such as calcium compounds in order to neutralize acids [21, 22]. However, this may result in reduction of bone density, and osteoporosis. There is also a risk of formation of kidney stones when excessive acid-forming foods and drinks are routinely consumed [23, 24].

Although these pH regulation mechanisms try to balance acidity levels, there are limitations to their functioning [24]. The response of the body to a metabolic acid load may not be immediate [25, 26, 27] and during the time that the blood remains slightly acidic, a number of cells may be damaged, die or mutate [11]. In addition, the relatively high concentration of calcium ions released in response may precipitate in the kidneys or in blood vessels, partially blocking them and reducing the flexibility of their interior surfaces, resulting in increased blood pressure and cardiac health issues [28]. If the kidneys and lungs cannot keep the body pH in balance due to acidosis, this can cause inflammation of the veins or even coma [29].

Within the small changes of blood pH in the range 7.2-7.5, the concentration of oxygen and other blood gases may still vary, with potential long-term consequences for health [30]. While it is normal for relatively high concentrations of acids to form and pH to slightly drop around muscle cells during intense physical activity [25], here we are concerned with whether a long-term impact may result from acid-forming dietary habits and whether alkaline-forming diets can reduce these acidity levels and benefit health.

## The impact of diet on acid-base balance

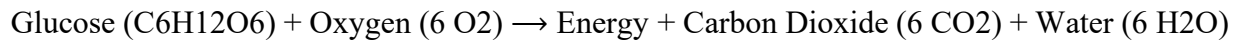
### Waste products of respiration at the cellular level

The cells use nutrient molecules such as sugars, fatty acids and excess amino acids from the diet to produce energy [31]. The preferred source is glucose [32]. Other types of sugars we ingest include fructose and sucrose (found in sugarcane, sugar beets, as well as in fruits), maltose (found in barley), galactose and lactose (found in milk and dairy products), mannose (found in fruits and vegetables) [33], and trace amounts of N-Glycolylneuraminic acid (neu5Gc) from eating animals that metabolize it (humans do not)--which may be associated with cancer and other diseases [34, 35].

These nutrient molecules are converted into energy via cellular respiration processes that take place as aerobic where sufficient oxygen is available (healthier), but as anaerobic in the absence of adequate oxygen--in particular in muscles during high intensity exercise--producing lactic acid residues [36, 37]. The energy production from nutrient molecules also forms cell metabolic wastes as by-products [14, 15], which may include water, carbon dioxide, carbonic acid, lactic

acid, bicarbonates, hydrochloric acid, sulphuric acid, phosphates and nitrogen compounds such as nitrates [38, 39, 40, 41, 42].

In aerobic respiration, the cellular fuels may be glucose, fatty acids or amino acids [13]. When cells use glucose as fuel, six molecules of oxygen are required for efficient combustion of each sugar molecule, as per the following equation [31, 32]:



However, where oxygen is not sufficient or it cannot reach the cells quickly enough to keep up with demand, conversion of glucose into energy inside the cells utilizes anaerobic respiration and fermentation processes. The following equation shows how energy is commonly produced from glucose in these processes [31, 32]:



When other oxidants like nitrates are in excess, anaerobic respiration will not only form lactic acid as a metabolic waste [18], but also nitrogen wastes such as ammonia [39, 40]. In addition, the lactic acid from anaerobic respiration is more acidic than the carbonic acid from aerobic respiration [40, 43, 44].

As by-products of cellular respiration, some foods may form acidic cell metabolic wastes (so-called acid-forming foods) and some may form alkaline cell metabolic wastes (so-called alkaline-forming foods), both temporarily affecting pH in some cellular environments and in blood compartments [18, 41]. The characteristics of these cell metabolic wastes depend in large part on the chemical structure, digestive processes, and metabolic pathways related to each nutrient molecule [32, 36]. The alkaline components of foods may include alkali salts such as potassium bicarbonate and potassium citrate, of which fresh vegetables and on-tree ripened natural fruits are good sources [41]. Acid-forming components of foods include protein, phosphate, and sulfur [18].

Amino acid metabolism releases hydrogen ions, causing a significant lowering of blood pH. The quantity and type of acid released will depend on which amino acids are present: lysine, arginine, and histidine generate hydrochloric acid; cysteine and methionine, which contain sulfur, generate small amounts of sulfuric acid residues [41, 42].

In traditional systems of medicine, foods that tend to form acidic metabolic waste, such as animal protein, are thought to cause certain health problems when predominant in the diet [36, 40, 42]. Foods that form predominantly alkaline metabolic waste are believed to improve health and are even used in treatments for some diseases [41, 42]. But modern research is also beginning to confirm that these dietary differences do indeed influence the acid-base balance of several body compartments [25].

## The impact of diet on digestion-caused acid or alkali “tides” in the blood

Via energy-expensive processes, the stomach converts salt, water and carbon dioxide into hydrochloric acid (pH ~1), which creates a corresponding concentration of bicarbonate ions ( $\text{HCO}_3^-$  with an alkaline pH) that are secreted into the blood stream. As result, for a few hours, the venous blood around the stomach is more alkaline, referred to as the “alkaline tide” [26]. The pancreas also secretes additional bicarbonate into the duodenum [45] and a corresponding quantity of acid into the blood stream, causing a temporary ‘acid tide’ [27]. These  $\text{H}^+$  ions eventually neutralize the earlier alkaline tide in the blood that was caused by the stomach [26]. These tides are also more gradually regulated by other pH regulation mechanisms as needed to maintain blood pH within normal parameters [26, 27].

For easy to digest meals (such as fruits and vegetables), the alkaline and acid tides are less extreme, because fewer hydrogen ions are needed for the stomach and fewer bicarbonate ions from the pancreas. But for heavy meals such as animal protein that require more acids to digest (or in the case of vomiting due to sickness or bulimia), these tides would be more pronounced, because the stomach has to produce greater amounts of acid [26]. Hence, frequent consumption of heavy meals may exacerbate acid-base disorders.

## The impact of the pH of foods ingested

Alkaline supplements such as bicarbonate salts cannot be effectively used to alkalize the blood stream, as they are immediately neutralized by stomach acid. For the same reason, the body’s cells do not receive any alkalinity directly from “alkaline-water” (pH~8.5-9.0) that is sometimes advertised as a body alkalizing drink [46]. Indeed, as result of their neutralizing stomach acid, it is forced to produce extra acid, forming a temporary alkaline tide in the blood stream around the stomach as a by-product. If this increase in blood pH takes place at the same time as sustained high-intensity exercise, it may reduce the muscle pains associated with a high blood lactate response [47]. However, neutralizing the stomach acid may in the long term increase the risk of food- or water-borne illnesses by allowing bacteria to build up [48]. If the bicarbonate source is sodium, this may also contribute to blood pressure issues and even damage the stomach lining, increasing the risk of stomach cancer [49]. Bicarbonate salts may also cause mucosal damage to the gastrointestinal tract when used long-term, and even be life-threatening in patients with abnormal renal function or chronic kidney disease [50]. Of crucial importance in this context, the use of alkaline substances like bicarbonate cannot be an effective solution for diseases related to metabolic acidosis, as the alkaline tide that results from consuming them is short-term [27], but the body needs the sustained presence of normal alkalinity. In contrast to short-term approaches, high enough intake of alkaline foods or drinks over the long term, in particular those naturally rich in potassium bicarbonate, can help maintain this optimum alkaline environment.

Consuming too much acidic foods or drinks over a long period of time may also negatively affect stomach conditions as well as altering the body pH. For instance, drinking too many acidic carbonated beverages may worsen acid reflux. Furthermore, they may increase body acidity,

leading to reduced bone mineral density and increased risk of fracture [51]. This is probably because the pancreas has to produce more bicarbonates to neutralize the extra acids from the acid-rich carbonated drinks after they leave the stomach. Hence, a more significant acid-tide is experienced in venous circulation, which is neutralized by mobilizing minerals from bone. The same issue has been reported in case of chronic consumption 1000 mg vitamin C supplements, which may eventually increase the risk of kidney stone formation [52].

Regardless of the original pH of the diet, acidic or alkaline metabolic wastes are formed at cellular level, and the pH of these residues (discussed in the next section) is independent of the total pH of the actual food or drink as consumed. Hence, while the pH of foods as ingested has impacts discussed above, the pH in body compartments at any one time is mainly dictated by the processes related to food digestion (acid and base “tides”) and the chemical reactions and products related to cellular respiration, rather than to the pH of their parent foods.

### Alkaline-Forming Diets

The most poorly understood of the body’s mechanisms for regulating acidity relates to the formation at cellular level of acidic or basic waste products of respiration or other metabolic functions, which we propose can be influenced by the composition of the diet. Some natural healers believe that acid-forming foods may play a large role in this, while alkaline-forming foods may be preventative [41, 53]. This may be especially important in older people, because they gradually face loss of some renal acid-base regulatory function [22, 25].

Specific fruits and vegetables are commonly cited as healthy in the alternative medical literature [41, 46, 53, 54]. For example, apples, oranges and green beans have a substantial content of mannose, a natural sugar that may slow down growth of tumors [33]. The alkalizing characteristics of fruits may depend on their content and composition and can even be affected by various production and harvest-related factors. The effect is different for climacteric fruits that can still ripen after being picked, versus non-climacteric fruits that can ripen little if at all once removed from the plant [55, 56]. Early picking in this latter case may lead to less alkalizing characteristics and greater acidity [54]. On-tree ripened fruits may have higher levels of some minerals and vitamins [66], because some elements may further be translocated into the fruits from the leaves and possibly other parts of the plant [67]. In addition, fruit ripened under light has a higher sugar content [68] and lower acid content [69, 70].

Another factor traditionally thought to have an impact on the alkalizing effect of certain foods is the use of chemical fertilizers [62, 63]. High nitrate intake may shift the body toward increased anaerobic respiration, increasing acid environments at cellular level, and fertilizer residues may add to this. To the extent that nitrates are converted into nitrites, they directly reduce the oxygen-carrying capacity of the blood [64, 65].

A diet rich in alkaline-forming foods may also improve calcium balance and reduce risk for osteoporosis and stone formation in the kidneys by reducing bone resorption [42, 67, 68, 69]. In contrast, diet-induced metabolic acidosis is associated with increased risks of insulin resistance

and type 2 diabetes [66]. Even some types of cancers have been linked to metabolic acidosis and acid–base imbalance [36, 37].

### Determining whether foods are acid or alkaline forming

Understanding the acid and alkaline forming character of foods (independent from the original pH of the food or drink) requires measuring which foods have which kind of corresponding effect. One method would be to measure the pH of urine or saliva following consumption of each type of food. However, this is an indirect measurement and not very accurate because some of the metabolic wastes such as carbon dioxide leave the body via the lungs and blood acidity may partly be neutralized by bone calcium compounds; acidity of the saliva is usually neutralized by the teeth; and some acids are excreted via sweating [11, 16, 13]. Hence, the pH of neither urine nor saliva may be reliable indicators for blood pH [19, 70].

Another approach is to estimate the acid/alkaline forming character of foods based on their composition [19, 53, 71]. Methods used include calculation of values for DAL (dietary acid load) or PRAL (potential renal acid load), NAE (net acid excretion) and EAA (excess acid/alkaline ash) for each foodstuff [19]. These methods are based on the composition of the food regarding magnesium, potassium, fiber, saturated fat, simple sugars, sodium, chloride, and protein/potassium ratios. For instance, the equation for calculating PRAL is  $(\text{mEq}/100 \text{ g}) = 0.49 \times \text{protein (g}/100 \text{ g}) + 0.037 \times \text{phosphorous (mg}/100 \text{ g}) - 0.021 \times \text{potassium (mg}/100 \text{ g}) - 0.026 \times \text{magnesium (mg}/100 \text{ g}) - 0.013 \times \text{calcium (mg}/100 \text{ g})$  [25]. The PRAL for vegetables is negative (indicating alkalinity), and for cheese and red meat, positive (indicating acidity) [41]. These values are in general agreement with the observations of urine pH when vegetable-rich or protein-rich diets are followed. EAA calculations are useful but not always predictive of urine pH [19]. None of these methods is comprehensive, each having their own limitations and uncertainties.

Another method used by natural health practitioners is Live (Cell) Blood Analysis [53]. However, this type of analysis is not accepted as a standard laboratory practice because its validity has not been established [72].

A more accurate technique than the ones discussed above is to directly measure venous blood pH-related parameters following consumption of a specific food or diet [25]. Since the cell metabolic wastes are all excreted into the venous blood, taking a blood sample from the veins can more accurately indicate acidity of the metabolic wastes and provide more accurate and realistic results. To determine venous blood gas (VBG) parameters in a blood sample, pH, oxygen saturation, and partial pressure of oxygen and carbon dioxide need to be measured [9, 10], and thus this may currently be considered to be too time-consuming and expensive to use in determining whether a large number of foods are acid- or alkali-forming (It would nevertheless be cost effective if future research suggests that alkaline-forming diets have a substantial impact in cancer prevention). In addition, the release of bone calcium or excretion of acids into the urine may eventually regulate the blood pH to a normal level, making the reliability of detecting acidosis based only on blood pH potentially inaccurate. However, in the short term, as mentioned



above, the pH of cellular and circulatory environments will be influenced by the waste products of respiration and other metabolic processes and these in turn will be influenced by the food eaten within the past day or less. These temporary effects are what need to be measured; more research on humans is needed to determine the optimal time to take such measurements after meal consumption [12, 73, 74].

Table 1 provides a categorized list of foods and drinks based on the effect they are thought to have on body pH (based on likely impact of metabolic residues). In each case, we have chosen the most reliable available data in the literature and performed a qualitative comparison based on agreements between the different references [19, 20, 41, 42, 46, 52, 53, 71]. However, most of these data came from sources which used measurement methods that have not been verified.

## The Warburg Effect on Acid-Base Disorders

Cellular respiration requires appropriate conditions in terms of pH, temperature, oxygen saturation, and concentration of essential minerals, vitamins, enzymes and co-enzymes in order to be effective and controlled [31, 37]. If cellular respiration processes are disturbed or damaged, some cells may die. This is not a major problem, as long as not too many of them die, because they can be replaced. The potentially dangerous cells are the ones that mutate to survive such damage [75], developing into cancerous cells as a result of impaired respiration [76, 77].

Dr. Otto Warburg a German scientist, hypothesized that a repeatedly oxygen-poor, acidic cellular environment may result in acidosis in the micro-environment of tissues, damage the surrounding normal tissues, and cause cell mutations, leading towards cancer [36, 77, 78]. This so-called Warburg Effect is a form of modified cellular metabolism found in cancer cells, which tends to favor a specialized fermentation over the aerobic respiration pathway that most other cells of the body prefer.

Anaerobic respiration processes are faster but less efficient, and also consume greater amounts of glucose than aerobic respiration to produce similar levels of energy (ATP) [32]. Anaerobic respiration is found in virtually all tumors, which means cancer cells use larger amounts of sugars. Cancer cells may also use excessive quantities of other nutrients as well, including fats and some amino acids such as glutamine [79].

Cancer cells need large quantities of sugar to grow quickly and multiply at a fast rate, which takes a lot of energy. Hence, most simple tools for staging and diagnosis of cancer work by revealing the places in the body where cells are consuming extra sugar [79]. Thus, maintaining the blood sugar levels within the lower end of the healthy range (i.e., a fasting blood sugar of 72-79 mg/dl) may weaken cancer cells, while normal cells can manage to survive such conditions [76, 80, 81, 82, 83]. In contrast, having too much sugar in the blood for long periods of time may favor cancer cells, as well as lead to other serious health problems such as heart disease, type 2 diabetes, and kidney disease [76, 80, 84].

The Warburg Effect suggests that one of the conditions which favors growth of cancer cells is



Table 1: Acid/Alkaline forming characteristics for certain foods [19, 20, 41, 42, 46, 52, 53, 71]

Category	Strong Acid (least healthy)	Medium Acid	Weak Acid	Weak Alkaline	Medium Alkaline	Strong Alkaline (healthiest)
<b>Fruits</b>	Prunes <sup>1,2</sup> Canned fruits Factory juices <sup>1,2</sup>	Sour cherries <sup>1,2</sup> Sour plums <sup>1,2,3,5</sup>	Sweet Plums <sup>1,2</sup>	Oranges <sup>1,2,3</sup> Bananas <sup>1,2,3</sup> Cherries <sup>1,2,3</sup> Peaches <sup>1,2</sup> Pomegranates <sup>2</sup>	Grapes <sup>1,2</sup> Apples <sup>1,2</sup> Pears <sup>1,2</sup> Melons <sup>1,2</sup> Raisins <sup>1,2</sup>	Lemons <sup>1,2</sup> Limes <sup>1</sup> Dates <sup>1,2,5</sup> Figs <sup>1,2</sup> Mangoes <sup>1,2</sup>
<b>Vegetable, Beans, Legumes</b>	Fried potatoes and chips <sup>1,2</sup>	Peas <sup>1</sup> Lima beans <sup>1</sup> Lentils <sup>1,2</sup> Boiled potatoes <sup>1</sup>	Kidney beans <sup>1</sup> Cooked vegetables <sup>1</sup>	Cucumber <sup>1,3</sup> , Carrots <sup>1,2</sup> , Tomatoes <sup>1,2</sup> , Mushrooms <sup>1</sup> Cabbage <sup>1</sup> , Green peas <sup>1,2</sup>	Olives <sup>1</sup> Green Beans <sup>1</sup> Okra <sup>1</sup> Turnip <sup>3</sup> Celery <sup>1,2,3</sup> Pumpkin <sup>2</sup>	Garlic <sup>1</sup> Onions <sup>1</sup> Spinach <sup>1,2,3</sup>
<b>Nuts and Seeds</b>	Peanuts <sup>1,2,3</sup> Cashews	Walnuts Pistachios <sup>1,2,3</sup>	Seeds: Pumpkin <sup>1,2</sup> Sunflower <sup>1</sup> Sesame <sup>1</sup>	-	-	Raw almonds <sup>1,2,3,5</sup>
<b>Meats</b>	Pork meat <sup>1,3</sup> Shellfish <sup>1</sup> Rabbit <sup>1,3</sup>	Beef <sup>1,2,3</sup> Lamb <sup>1</sup> Turkey <sup>1</sup> Veal <sup>1</sup>	Fish <sup>1,2</sup> Chicken <sup>1,2</sup>	-	-	-
<b>Eggs and Dairy</b>	Cream <sup>1,2</sup> Ice Cream <sup>1,2</sup>	Homogenized Cow's Milk <sup>1,2</sup> Cheese <sup>1,2,3</sup> Store eggs <sup>1,2</sup>	Yogurt <sup>1</sup> Butter <sup>1</sup> Raw Milk <sup>1</sup> Farm Eggs <sup>1</sup>	Goat milk <sup>1,2</sup> Goat cheese <sup>1,2</sup>	-	-
<b>Grains and Cereals</b>	Pastries <sup>1</sup> Pasta <sup>1</sup> Cereals (corn etc.) <sup>1</sup>	White rice <sup>1,2</sup> White flour <sup>1,2</sup> Oats <sup>1,2</sup> Bread <sup>1,2</sup>	Whole wheat <sup>1</sup> Brown rice <sup>1,2</sup>	-	-	-
<b>Oils</b>	Frying oil <sup>1,2</sup>	Sunflower oil Sesame oil <sup>1,2</sup>	-	-	-	Olive oil <sup>1</sup>
<b>Drinks</b>	Liquor <sup>1,2</sup> Beer <sup>1,2</sup> Soft Drinks <sup>1,2</sup> Vitamin C Supplements (1000 mg) <sup>4</sup>	Black Tea <sup>1,2</sup> Coffee <sup>1,2</sup> Wine <sup>1,2</sup> Alkaline water <sup>1</sup>	Bottled mineral water <sup>1</sup>	Spring mineral water <sup>1</sup> Ginger tea <sup>1</sup>	Lemon juice drink <sup>1,2</sup>	-
<b>Other foods</b>	Chocolate <sup>1,2</sup> Ketchup <sup>1,2</sup> Mayonnaise <sup>1,2</sup>	Jam <sup>1,2</sup> Sugar <sup>1,2</sup> Vinegar <sup>1,2</sup>	Processed honey <sup>1</sup>	Ginger <sup>1</sup> Natural honey <sup>1</sup>	-	-

The above table has been integrated from several different sources, which include:

- 1- pH food chart published by Corriher [46]
- 2- pH food chart published by Bridgeford [53]
- 3- pH food chart published by Remer [71]
- 4- Thomas et al. [52]
- 5- Venous blood examination by the author (HB) to partially verify the table

hypoxia. In contrast, an oxygen-rich, optimum-alkaline environment can maintain cells in a healthy state [36, 78, 85]. For this to be taken advantage of, appropriate conditions need to be provided internally around the cells by enhancing cellular respiration efficiency (i.e., aerobic instead of anaerobic) [36, 77, 79]. However, practical means of attaining this have been elusive. If both oxygen and carbon dioxide are provided, the blood tends to become more acidic, as the natural solubility of carbon dioxide in the blood is much higher than that of oxygen [74]. Yet long-term increases of oxygen intake via the lungs may result in oxygen toxicity or lung damage [86]. In addition, red blood cells' ability to carry oxygen from the lungs is limited. Using pressurized oxygen chambers may temporarily result in a small increase of arterial oxygen saturation [76], but is not a sustainable solution, and cannot help with a chronic acidic venous circulation.

Enthused by this still poorly understood Warburg Effect, some doctors have used it incorrectly, assuming mistakenly (as we explained above) that consumption of an alkaline substance would balance pH at cellular level. This has caused the death of some patients. In one case in 2012, for example, a 27-year-old British woman paid thousands of dollars to a doctor for alkaline treatment using baking soda (sodium bicarbonate), but after the treatment, her condition worsened due to severe alkalosis and soon afterwards she died [87].

Treatments of this kind were based on the incorrect belief that blood pH in cancer patients could be in the range of 4-5, hence, an extreme alkalinity can kill cancer cells [88]. But in fact, any blood pH less than 7.00 or above 7.65 is life-threatening and may result in death [16, 21, 89].

In order to be effective, the alkalinity required by the Warburg Effect would have to utilize a precisely increased oxygen saturation in the blood around the affected body cells. This turn would require measuring and monitoring of blood pH-related parameters to effectively overcome the acidity and provide a healthy body pH in the range of 7.40-7.45 [16, 82, 88].

In order to properly address the acid-base disorder based on blood pH-related parameters, multiple blood parameters need to be measured. For instance, in normal people with a healthy blood pH, the arterial oxygen saturation is usually high. But in patients diagnosed with hypoxia whose kidneys and lungs cannot keep the acid-base balance, there will usually be a drop of blood pH and a significant reduction in oxygen saturation (acidosis leading to hypoxia) [11, 12]. In some cases, blood pH may appear to be normal due to increased bicarbonate concentration by the kidneys or minerals released from bones into the blood, while oxygen saturation may still remain low (i.e., the acidosis is corrected, while hypoxia remains a major health issue) [13]. Oxygen saturation is also influenced by the quality of the blood cells, meaning that if many are damaged for any reason, the overall ability of the red blood cells to carry oxygen is reduced [14]. Acidosis due to acid forming dietary habits, even if mild, may worsen hypoxia [15, 4, 5].

Perhaps an alkaline-producing diet could help achieve a balanced blood pH, but first we must find out whether such a diet appears to have the expected effect. So far, there is little experimental evidence backing the claim made by some practitioners of traditional medicine that certain foods are acidic and others are basic with respect to the impact of their metabolic

residues on biochemical parameters and thereby on indicators of pH in the blood. Hence, one of us (HB) experimented with such diets on himself, in hopes that this might help motivate other researchers to examine the issue in large-scale crossover trials.

## **The Exploratory Study's methods**

The experimental subject (author HB) was a male, 37 years old, non-smoker, with a body mass index of 22-23, free of illness and in good physical condition according to blood examinations and a general health check-up performed prior to the experiment. In addition, he had not taken drugs or supplements of any kind for almost a year.

For the purpose of examining the effects on blood pH of alkaline forming and acid forming foods, the diets tested in the experiment were based in part on the foods labeled as alkaline or acid in Table 1. Substantial amounts of foods thought to be alkaline and acid forming, respectively, were added to his normal diet, with no attempt made to change other aspects such as total calorie intake. These diets were each consumed for one month and separated by a two-month wash out period. Larger quantities of alkaline/acidic foods were consumed in the final week of each month.

The alkaline diet in first period consisted of increased quantities of those foods identified as such in Table 1 including on-tree ripened fruits, fresh raw leaf vegetables and natural plant-based foods in usual diets (boiled or steam-cooked), plus daily consumption of 5-10 raw almonds (produced without any fertilizers, as almonds are particularly susceptible to an increase in HCN content in the presence of excess nitrogen). The acid-forming diet in the second period consisted of increased quantities of those foods identified as such in Table 1, plus 4-6 sour plums per day (considered to be a strong acid-forming food).

Immediately after each of the month-long test periods, measurements were performed for blood pH and blood gas saturations and partial pressures. These measurements were performed for the arterial blood gases (ABG) and venous blood gases (VGB) separately. All the blood samples related to this study were taken and analyzed by Massoud Clinical Laboratory in Tehran, which provides comprehensive clinical services.

To check and confirm the reliability of the earlier tests for effect of alkaline diets, the experiment was repeated one year later on the same person. This time, only the alkaline diet was consumed for one month and only VGB was checked. This diet included daily consumption of approximately 10-15 dates (the Khasoei variety), 3-5 apples and 3-5 oranges (on-tree ripened), and the intake of alkaline foods in meals was increased as much as possible; acid-forming foods were minimized and any food with artificial sweeteners such as factory-juices, biscuits, soft drinks, cakes and chocolates were completely avoided.

## Results

The following measurements were taken and are reported here: venous and arterial blood samples following the acid forming diet, venous and arterial blood samples following the alkaline forming diet, and an additional venous blood sample following another experiment of alkaline diet alone; and pH of venous and arterial blood measurements taken after each diet. In addition, at the end of the second experiment, the fasting blood sugar concentration was 74 mg/dl, which is at the lower end of the healthy range, 72-99 mg/dl.

Table 2 summarizes blood pH, blood oxygen saturation, partial pressure of oxygen and partial pressure of carbon dioxide, and Figure 2 illustrates the relationships among the parameters. The relative difference in blood pH between the two measurement times was also determined and reported in Table 2.

These results can be summarized as follows:

- The oxygen saturation of venous blood is always significantly lower than arterial blood, as should be the case.
- The acid-forming diet reduced the venous blood pH down to 7.30, compared to 7.42 after the alkaline-forming diet (typical venous blood pH on average is 7.35-7.40 according to the literature [90, 91, 92, 93, 94, 95]), suggesting that the diet type could meaningfully change blood acidity level in the veins.
- Oxygen saturation of venous blood appears to be significantly lowered by an acid-forming diet down to 33%, and increased following an alkaline-forming diet up to 81%. Presumably, this is due to diet-related metabolic wastes from the cells excreted into the veins.
- The oxygen saturation of arterial blood from acid-forming diet to alkaline-forming diet, increased from 94% to 98%. The change in oxygen saturation in arteries due to change of diet type is expected to be minor and in any case is already close to the theoretical ceiling of 100%.
- The alkaline-forming diet appeared to increase partial pressure of oxygen and also reduce partial pressure of carbon dioxide in both arterial and venous blood.

These data were all from measurements taken at a laboratory located in Tehran, around 1500m above the sea level. At higher altitudes, carbon dioxide loss is slightly greater, and entry of oxygen from the air into the blood is slightly reduced [96, 97, 98, 99, 100]. For instance, based on the literature, the commonly reported range for partial pressure of carbon dioxide at sea level is around 40-45 mmHg and at 1500m above sea level is about 35-40 mmHg. Hence, the above values should be looked at with this expected effect of altitude on overall typical ranges.

Table 2: Blood gases and pH for the candidate from a single blood draw after one month on each diet

Blood type	Diet type	Blood pH	$\Delta(\text{pH}) = \text{pH}-7.4$	P CO <sub>2</sub> mmHg	P O <sub>2</sub> mmHg	HCO <sub>3</sub>	Oxygen Saturation%
Arterial	Alkaline-forming	7.47	+ 0.07	31	111	25	98
	Acid-forming	7.44	+ 0.04	37	67	25	94
Venous	Alkaline-forming	7.42	+ 0.02	35	44	24	81
	Acid-forming	7.30	- 0.1	53	22	23	33 <sup>a</sup>
	Alkaline-forming (2 <sup>nd</sup> experiment)	7.40	0	34	43	21	79
Typical common ranges for VBG parameters <sup>b</sup>		7.35-7.41	-	35-45	35-45	21-29	60-85

<sup>a</sup> While this value appears to be far below the commonly measured ones, we checked and several people with largely acid-forming diets had similar values.

<sup>b</sup> These ranges come from combining various sources of published data from different laboratories, as each has reported the normal range differently [90, 91, 92, 93, 94, 95].

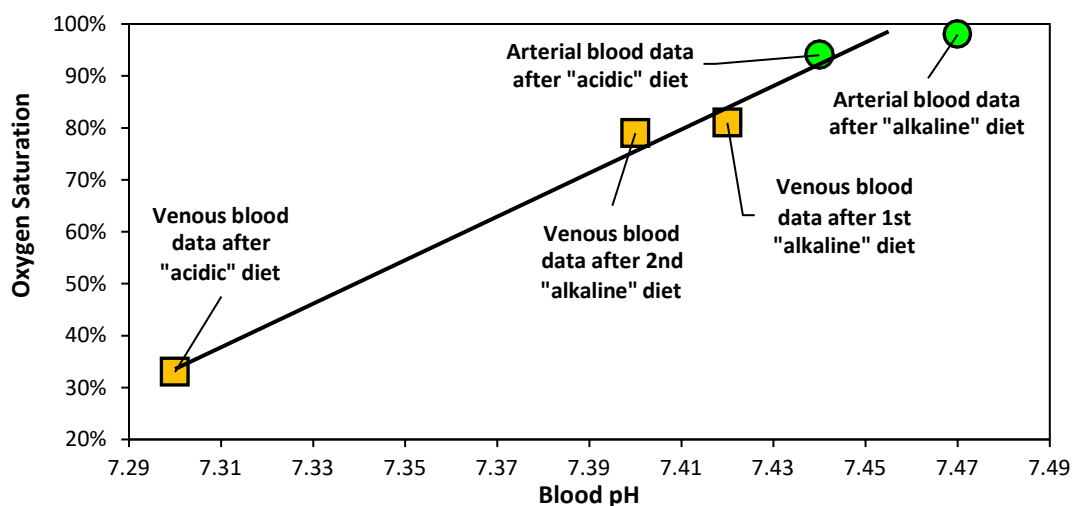
## Discussion and conclusions

Of course, this study was extremely limited in scope, with an  $n$  of only one and measurements only taken after each diet, not before the diets. This exploratory study found the hypothesized relationship between diet type with pH and oxygen saturation of venous blood, suggesting that these parameters are significantly affected by diet type. It also suggests that the type of diet chosen was effective and the data consulted from acid/alkaline forming food tables are plausible.

The present study was intended to provide data on a first-time basis to determine whether a full-blown study might be justifiable of the potential of using an alkaline diet to achieve improved

health and prevention of disease, as predicted by the Warburg Effect. We hope this will inspire future researchers to conduct cross-over trials on an adequate sample size.

Figure 1: Relationship between blood pH and oxygen saturation for the venous and arterial blood samples



## Acknowledgments

We would like to acknowledge and show our gratitude to International Agency for Research on Cancer (IARC), Cancer Research UK, Dr. Rachel Thompson (World Cancer Research Fund's), Dr. Gordon Edwards (Canadian Coalition for Nuclear Responsibility), Thomas Corriher (Health-Wyze USA), Dr. Ali Karami (National Foundation of Healthy Lifestyle), Dr. Roghayeh Javanmard and Mazdafar Momeni for sharing valuable information that were used in the study.

We are also thankful to Hanieh Habibfar, Kian Bahrami, Nikan Bahrami, Radin Shouri, Chantele Siavoshi, Lucas Waters, Gemma Byfield, Sahar Mosaffa, Soraya Mohamadzadeh and Maryam Alibeigi for their great help and support in the research work.

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