

International Journal of Advanced Research in Science, Communication and Technology (IJARSCT)

International Open-Access, Double-Blind, Peer-Reviewed, Refereed, Multidisciplinary Online Journal

Volume 3, Issue 2, November 2023

Impact on Food and a Healthy Lifespans, Research on Nutrition's

Priya Srivastava¹ and Dr. Savita Sangwan²

Research Scholar, Department of Home Science¹ Research Guide, Department of Home Science² Shri JJT University, Jhunjhunu, Rajasthan, India

Abstract: One of the best and least expensive methods to reduce the burden of numerous diseases and the risk factors that go along with them, such as obesity, is through proper diet. Nutrition research has the potential to significantly impact economies and global health since it holds the key to deepening our understanding of the causes of obesity and its associated comorbidities. Following a survey of seventy-five influential individuals, the American Society for Nutrition (ASN) organised a Working Group to determine which areas of nutrition research need to be advanced in order to have the biggest anticipated influence on the health and welfare of future generations worldwide. The high priority categories of ASN's Nutrition Research Needs are as follows: 1) individual variations in food and diet responses; 2) normal growth, development, and reproduction; 3) health maintenance; 4) medical care; 5) nutrition-related behaviours; and 6) food supply/environment. With the tremendous potential for translation and impact on public health, ASN expects that the Nutrition Research Needs will spur collaboration among scientists from various disciplines to pursue this ambitious research agenda. ASN also expects that the results of the Nutrition Research Needs would encourage the creation and acceptance of fresh, cutting-edge approaches to the diagnosis, treatment, and prevention of diseases linked to nutrition. Because nutrition research is multidisciplinary, different areas of expertise of stakeholders must work together to produce evidence-based nutrition guidelines and policies that will improve the health of the world's population through a variety of multidimensional techniques. ASN also recognised five instruments that are essential to the advancement of the Nutrition Research Needs in addition to the specified research needs: Databases, bioinformatics, omics, biomarkers, cost-effectiveness analysis, and 3) databases round out the list. Adv. Nutr. 4 (2013): 579–584.

Keywords: Nutrition

I. INTRODUCTION

Getting enough food is reliant on and involves the whole food chain. Foods derived from plants and animals, along with their numerous components, are the main sources of nutrition for humans. In addition to being essential for animal and human growth and development, nutrition also plays a critical role in illness prevention and treatment. Another essential component of maintaining functionality and excellent health is nutrition. The foundations for healthy populations and strong economies are represented by basic and applied research on the relationships between nutrition and the absence of infectious diseases, nutrient composition, and nutrition monitoring. As a result, cutting-edge nutrition research and instruction serve as the cornerstone for solutions to more significant health-related problems, enabling people to lead longer, healthier lives.

Only recently has the significance of nutrition become fully understood as a crucial component of the solutions to many of the societal, environmental, and economic problems that the world faces. The "grand" issues affecting nutrition research and science in the twenty-first century have been recognised by the American Society for Nutrition (ASN) and are referred to as "Nutrition Research Needs." The results of these Nutrition Research Needs will clarify approaches to the treatment and prevention of infectious and non-communicable diseases, such as diabetes, cancer, and cardiovascular disease. With the potential to significantly impact global economies, nutrition research holds the key to deepening our understanding of the underlying causes of obesity and its associated comorbidities. By direct and intentional agricultural practises, knowledge about adequate nutrition also plays a significant role in mitigating or eliminating food

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insecurity on a national and international level. Unquestionably, as the world's population grows, there will be a greater need for a food supply that is secure, readily available, sustainable, and reasonably priced, while also requiring adequate nutrition.

The Public Policy Committee of ASN first conceived of the ASN Nutrition Research Needs project to determine the global needs for nutrition research. The purpose of this endeavour is to inform and convey to decision-makers and interested parties the importance of funding nutrition research at a higher level in order to address societal demands. In September 2011, the Public Policy Committee of ASN contacted approximately 75 influential individuals to create a preliminary inventory of nutrition research requirements.

ASN assembled a Working Group of nutrition scientists and researchers in February 2012, drawing from a diverse membership of the Society, to identify the areas in need of nutrition research that will have the biggest effects on the health and welfare of the world's citizens. The Acknowledgments contains a list of the members of the Working Group. Starting with the draft list, the Working Group narrowed down and pulled together 6 nutrition research needs for which advancement would have the greatest projected impact on future health and well-being.

The ASN then informed its membership of the 6 priority research needs and sought further member input. A workshop was held during ASN's 2012 Scientific Sessions and Annual Meeting in San Diego, CA, with nearly 250 attendees. The research needs were also shared via ASN's member newsletter, which reaches the entire membership base of nearly 5000 individuals, to inform and seek input from members who did not attend the annual meeting or the workshop. Member feedback on the Nutrition Research Needs was incorporated during development of the final document.

II. THE TOP NUTRITION RESEARCH NEEDS

The top 6 nutrition research needs cut across the entire re- search spectrum from basic science to health policy, from discovery to application. Specific research areas are listed under each research need. These 6 nutrition research needs are highlighted in the hope that they will prompt scientists from all disciplines to collaborate to advance these challenging research needs that have high potential for translation and public health impact. Although the topics presented focus principally on human nutrition research, the Working Group recognized that nutrition research using animal models is an essential foundation for making new discover- ies that can be translated to advances in human nutrition. Further, the importance of animal nutrition research is emphasized within these research needs in particular: "Understanding the role of nutrition in health maintenance" and "Understanding the food supply/environment." The research community will benefit from clearly articulated nutrition research priorities that will lead to science-based information, help to shape policy and enhance future funding for nutrition research, and thereby further promote the field of nutrition science.

Understanding variability in individual responses to diet and foods

A top priority for future nutrition research is the need to better understand variability in metabolic responses to diet and food. Enormous variability exists in individual responses to diet and food components that affect overall health. Discoveries underpinning this variability will lead to advances in personalized nutrition interventions and will better inform health and food policies, including Dietary Reference Intakes (DRIs) for nutrient needs and, ideally, future recommendations for known bioactive food components. Research in the following areas is necessary to deter- mine the origins and architecture of variability and to explain similar or dissimilar responses to diet and food components by subpopulations, as influenced by genetic, epigenetic, and ethnic and/or racial differences.

Omics. Omics research, such as nutrigenomics and nutrigenomics (e.g., epigenetics, transcriptomics, proteomics, and metabolomics), will help to determine how specific nutrients interact with genes, proteins, and metabolites to predict an individual's health. Omics provide information on individualized nutrient requirements, including how nutrients are digested, absorbed, and metabolized, and their functions in the body. Omics will help to determine and reflect an individual's nutritional status and will aid in the creation of new nutritional and disease biomarkers.

Microbiome. Diverse microbes, such as bacteria and vi- ruses, live in and on the body and contribute to the microbiome, which is estimated to have 10 times as many cells as the body itself (1). Microbes can vary in type and quantity, making each organism's microbiome unique although subpopulations may have similar microbiome characteristics.

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The microbiota needs to be better defined, and change due to diet, age, physiologic state, and disease need to be determined. Research is needed to determine the micro- biome's role in varying biological responses to diet and food components and its importance in disease prevention and progression. Conversely, research is also needed to determine how the microbiome is influenced by diet and other environmental factors.

Biological networks. Basic research is needed to provide a better understanding of biological networks, such as an individuals' genome (DNA/RNA protein profiles), and how these networks affect metabolic responses to diet and food. Environmental interactions, including nutrients and other dietary components, bacteria, viruses, and chemical contaminants, all may affect the responsiveness of biological networks to specific foods and the entire diet.

Tissue specificity and temporality. Research is needed to describe the mechanisms by which dietary factors affect variability in development and functioning, including which tissues are most influenced by dietary factors and when during the most critical stages in life this influence occurs.

Understanding the impact of nutrition on healthy growth, development, and reproduction *Epigenetics/imprinting*. Epigenetics and imprinting re- search examines how exposures to dietary components during critical periods of development may "program" long- term health and well-being. Research is needed to determine how early nutritional events contribute to disease later in life and alter normal developmental progression.

Early nutrition. Research is necessary to better understand the role of diet and individual food components on normal growth and development. This includes the role of parent's preconception diets, the maternal diet during pregnancy, and early nutritional events. Studies indicate that the timing of an infant's introduction to solid foods may increase the likelihood of becoming obese later in life (2). These findings are important given that the number of overweight children in the United States has increased dramatically in recent years (3). Research is now needed to determine the best approaches to influence these factors during early life. The important role of nutrition throughout early life on growth and development, as well as on health and well-being, needs to be continually assessed.

Nutrition and reproductive health. The impact of nutrition on reproductive health, including before and after conception, requires further research. Nutrition has a direct impact on both maternal and paternal fertility and the ability to conceive and also plays a key role in preventing diseases related to reproductive organs, including prostate and ovarian cancers. Although numerous studies have investigated how fruit and vegetable consumption may affect risk of breast, prostate, and other cancers, there is no clear consensus in the scientific literature. Thus, well-designed controlled intervention studies are needed to determine whether effects are limited to subpopulations, what factors influence a response and what mechanisms may account for changes in health.

Understanding the role of nutrition in health maintenance

Health maintenance includes noncommunicable disease prevention and treatment as well as weight management. The role that food components, particularly novel ingredients, contribute to health maintenance requires continuing research. Researchers and the public rely on dietary guidance, including the DRIs, to guide nutrition recommendations and health policy. Research is needed to better define the nutrient needs that best support health maintenance in all populations and their subgroups, from infancy through- out life. Nutrition across life is a fundamental issue that requires investigation so that recommendations will "match" with true biological needs.

Optimal bodily function. Research is needed to determine the roles that nutrition and fitness, both singularly and together, have in maintaining bodily functions, including cognitive, immune, skeletal, muscular, and other functions. Evolving research areas include prevention of disease-related processes, such as inflammation, and definition of mechanisms that have an important role in health maintenance, such as immunocompetence. Animal models are used to understand the requirements for optimal health in humans and production animals.

Energy balance. Research is also needed to examine the use of a systems approach to achieve energy balance including and integrating environmental, biological, psychosocial, and food system factors. A systems approach is preferable because the standard experimental approach of varying one factor at a time has accomplished little to address the population wide problem of energy imbalance. A solution- oriented approach that is comprehensive in nature and takes into account the complexities of achieving energy balance must be created. Although far more research is needed to identify system wide changes that maximize energy balance, intriguing examples exist. "Shape Up Somerville, MA," effectively reduced weight gain in high-risk children through a multifaceted community-based environmental change

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campaign (4). Shape Up Somerville increased the community's physical activity and healthful eating through physical infra- structure improvements and citywide policy and programming changes.

Understanding the role of nutrition in medical management

The rapid translation of nutrition research advances into evidence-based practice and policy is a priority for ensuring optimal patient care and effective disease management. Nutrition researchers have a key role in bridging the gap between disease prevention and disease treatment by fostering clinical research, providing innovative education for caregivers and patients, and delineating best practices for medical nutrition in primary care settings.

Disease progression. To improve the medical management of disease, research is needed to determine how nutritional factors influence both disease initiation and progression, as well as how nutrition affects a patient's response to therapy. Genetic and epigenetic variations among individuals can result in both positive and negative responses to diets, to specific foods, and to novel food components. The issue of individual variability is of considerable importance in refining medical management, including nutrition support, and requires continuing research.

Expanded research will allow us to better understand and minimize unfavorable impacts of both reduced and elevated nutrient intakes on disease progression and overall health. Disease/mortality response curves are U-shaped for many nutrients (that is, there is an increased risk of adverse out- comes if the nutrient is ingested in either too low or too high amounts). The importance of achieving a proper nutrient balance is seen in the example of chronic inflammation. Chronic inflammation contributes too many noncommunicable diseases and can result from high intakes of proinflammatory omega-6 fatty acids in the face of low intakes of anti- inflammatory omega-3 fatty acids (5). Research will help to determine the desired intake for essential and nonessential nutrients alone and when combined with other nutrients in the diet.

Nutrition support for special subgroups. Nutrition re- search is needed to establish the required nutritional needs that best support survival, growth, and development in sub- populations, such as in chronically diseased patients, in children, and in aging adults. With the success of medical advances, as have been seen with in vitro fertilization and neonatal care, caring for preterm infants presents a new challenge in early nutritional management. Preterm infants have special nutrition needs that will greatly affect their future growth and development, as well as their eventual health status as adults.

Understanding nutrition-related behaviors

Drivers of food choice. Understanding the link between behavior and food choices can help tackle obesity and other nutrition-related issues that are a public health priority. Individual food choices can be influenced by a number of different drivers including the following:

Government policy Environmental cues Cultural differences

Communication tools, such as social networking and food marketing

Research is needed to identify the impact of these various drivers and understand how they work alone or together to influence nutrition-related behavior. Research will show how these drivers should be altered to have the highest positive influence on individual behavior and therefore public health. For example, the state of Mississippi recorded a 13% decline in obesity among elementary school students from 2005 to 2011 (6). Multiple changes in the environment occurred, such as the setting of standards for foods sold in school vending machines, setting a requirement for more school exercise time, mandating healthier environments in childcare settings, and establishing programs that encouraged fruit and vegetable consumption. The challenge now is to determine what effect these combined actions will have on obesity-related behaviors in the long run.

Nutrition and brain functioning. Further explorations of the biochemical and behavioral bases for food choices and intake over time are essential. Brain function as it relates to food desire and choice needs to be clarified through research, and the multiple hormones that affect eating require further study as well. Factors such as meal frequency and size, speed of meal consumption, and how these factors are influenced by social cues require objective data, which can only be provided by research. Understanding how the marketing of healthy behaviors could help consumers achieve

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dietary guidance goals should be a priority. As part of this approach, innovative and practical methods for accurately measuring and evaluating food purchases and eating occasions must be developed.

Imprinting. Because of the high propensity of obese children remaining obese as adults (7), additional research is needed to determine how eating and satiety behaviors are imprinted during critical periods of development and to show how food components affect neural biochemistry and brain functioning and therefore shape behavior. This research will provide us with a better understanding of how and why an individual makes particular food choices. Although scientists recently validated the concept that food availability during pregnancy has permanent effects on gene expression in children (8), human studies are needed to confirm or refute the hypothesis that fetal programming, resulting from maternal obesity, leads to excess weight in children and into adulthood.

Understanding the food supply/environment

Food environment and food choice. Simply knowing or understanding what constitutes a healthy diet is not enough to change an individual's diet or lifestyle. Understanding how the food environment affects dietary and lifestyle choices is necessary before effective policies can be instituted that will change a population's diet in a meaningful way.

III. FROM THE AMERICAN SOCIETY FOR NUTRITION

Examples of key questions that should be addressed include the following:

Is current dietary guidance an effective way of communicating dietary change?

Do food assistance programs promote positive dietary patterns or have negative dietary and health consequences?

What role does food advertising play in food decision-making among different age groups and educational levels?

How do farm-to-fork food systems, with an increased emphasis on local agricultural production and consumption, influence dietary patterns and behaviors?

How can farm-to-fork food systems ultimately be used to pro- mote healthy behaviors and improve public health?

How can we most effectively measure, monitor, and evaluate dietary change?

Food composition and novel foods and food ingredients. Having an affordable, available, sustainable, safe, and nutritious food supply is also an important underpinning for making significant changes to a population's diet and life-style. Examples of key research areas to address include the following:

Enhancing our knowledge of the nutrient and phytonutrient content and bioavailability of foods produced, processed, and consumed

Studying how to better align and foster collaboration between nutrition and agricultural production

Can shifting agricultural focus from principally agronomic to include quality factors (such as taste, flavor, and nutritional value) have positive effects on fruit and vegetable consumption?

Can we leverage technologies, such as biotechnology and nanotechnology, to develop novel foods and food ingredients that will improve health, both domestically and abroad, and provide credible, tangible functional health benefits?

Public/private partnerships. To tackle these enormous challenges requires the coordinated efforts of public and private partners. The development of public/private partnerships between food and agricultural industries, government, academia, and nongovernmental organizations has the potential to advance nutrition research, enabling meaningful changes to be made to American and global diets (e.g., increased fruit and vegetable consumption to match government recommendations). We need to examine successful examples of public/private partnerships that have resulted in improved nutritional status and food security in specific populations (9).

IV. CROSS-CUTTING TOOLS TO ADVANCE NUTRITION RESEARCH

Nutrition research is truly a cross-cutting discipline, and the Working Group identified several tools that are also necessary to advance the priority needs in nutrition research. Adequately powered intervention trials continue to be essential for validating research theories arising from experimental and epidemiologic studies. However, the development of new, impactful tools will help us to more effectively quantify dietary intake and food waste and to determine the effective- ness of nutrition standards, such as DRI values and the *Dietary Guidelines for Americans*. Although not a traditional tool, multidisciplinary partnerships among scientific societies government, industry,

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academia, and others are fundamental to advance the nutrition research agenda. ASN and its membership must be proactive not only in efforts to advance nutrition research (including initiating and leading partnerships) but also in developing the tools needed to enhance the field. ASN recognizes the need to facilitate effetive communication among academia, industry, government agencies, consumers, and other stakeholders to advance nutrition.

Omics

Omics (especially genomics, proteomics, and metabolomics) will enable us to determine how specific nutrients interact with genes, proteins, and metabolites to predict the future health of an individual. A field of study that encompasses technological advances as well as omics-based research, it is sometimes referred to as personalized nutrition. Omics hold the keys to major nutrition breakthroughs in noncommunicable disease and obesity prevention. Omics provide information on how well nutrients are digested, absorbed, metabolized, and used by an individual. Moreover, omics will lead to new biomarkers that reveal a person's nutritional status and health status all at one time.

Bioinformatics

Bioinformatics is an interdisciplinary field that uses computer science and information technology to develop and enhance techniques to make it easier to acquire, store, organize, retrieve, and use biological data. Bioinformatics will enable nutrition researchers to manage, analyze, and under- stand nutrition data and to make connections between diet and health that were not previously possible. Databases are necessary to gain the full benefits of bioinformatics, because they make nutrition data easily accessible in a machine-read- able format.

Databases

Accurate, up-to-date food and nutrient databases are essential to track and observe trends related to the nutrition and health of individuals. Databases link food and supplement composition and intake data to health outcomes. Nutrient databases should be expanded to cover more foods and their bioactive components, including nonessential nutrients. Nutrition data must be incorporated into databases related to novel research areas, such as nutrigenomics and the microbiome, to adequately link these areas with nutrition. Data collection must also be improved with enhancements such as photographic food intake documentation, direct upload of food composition and sensory characteristics (if not proprietary) from food manufacturers, and biological sample collection.

Biomarkers

Intake, effect, and exposure biomarkers allow us to deter- mine and monitor the health and nutritional status of individuals and subpopulations, including ethnic and racial minorities. Biomarkers that are responsive to diet and nutrition will help assess disease progression and variability in response to treatment, while improving early diagnosis and prevention. Biomarkers must continue to be developed and validated to accurately track food and nutrient intake given our rapidly changing food supply.

Cost-effectiveness analysis

Cost-effectiveness analysis is a tool used to calculate and compare the relative costs and benefits of nutrition research interventions. Cost effectiveness analysis helps to determine the most cost-effective option that will have the greatest ben- efit to public health.

V. CONCLUSION

The multidisciplinary nature of nutrition research requires collaboration among research scientists with differing areas of expertise, many different stakeholders, and multifaceted approaches to develop the knowledge base required for establishing the evidence-based nutrition guidance and policies that will lead to better health and well-being of world populations. Proper nutrition offers one of the most effective and least costly ways to decrease the burden of chronic and noncommunicable diseases and their risk factors, including obesity. Although there is skepticism about the abil- ity to complete large, well-controlled dietary interventions at a reasonable cost in the United States, the success of the Lyon Diet Heart study in France (10,11) and the prevención con DIeta mediterránea (PREDIMED) study in Spain (12), both

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of which used variations of the Mediterranean diet, show this approach can be successful, even in the presence of drug treatment of cardiovascular risks in the latter study. Both of these studies showed significant reductions in cardiovascular disease (and cancer in the Lyon study) after relatively modest dietary changes.

Perhaps the greatest barrier to advancing the connections between food and health is the variability in individual responses to diet; it is also the origin of public skepticism to acceptance of dietary advice and the opportunity for entrepreneurship in the private sector. Imagine being able to identify, with certainty, those most likely to benefit from prescriptive nutrition advice through the various omic technologies and then providing these groups of people with customized nutrition advice based on their metabolic risk profiles. This is the new frontier of the nutritional science that offers the opportunity to predictably engineer our physiologic networks for health through diet. The confidence this approach would bring to the skeptical consumer would improve adherence to weight management and disease treatment techniques and improve the chances of success for disease prevention. To realize the full positive impact of achieving good nutrition on disease prevention and the health of populations, we must have the will to invest in and support the 6 key areas of nutrition research that have been outlined above.

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