WHITE PAPER



Leveraging the Antioxidant Properties of Fruits and Vegetables

Companion animals form an integral part of family life and have much more in common with their pet parents than shared social activities and mutual affection. As they get older, pets can experience some of the same types of age-related health problems as humans. And these health issues can be related to oxidative stress.

Oxidative stress is triggered when the animal's body doesn't have enough antioxidants to neutralize free radicals. This excess of free radicals can lead to cell and/or tissue damage and is related to chronic conditions such as cancer, diabetes and heart disease to name a few.

The body produces free radicals during its normal metabolic processes. However, several factors can exacerbate oxidative stress by stimulating excess production of free radicals. These factors can include diet, lifestyle, physical injury or environmental conditions like pollution, noise and crowds.

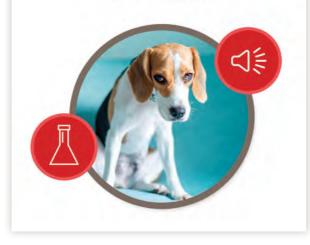


Urban Woes

Pollution, noise and crowds are found to a greater degree in urban areas than rural, pointing to one primary stressor, which is the environment. Evidence suggests that urban pets (and people) are exposed to conditions more likely to cause oxidative stress than those that inhabit rural areas. One study¹ links five urban factors with triggering an inflammatory response and subsequent oxidative stress. These include: chemical, noise and light pollution; infectious disease; and diet quality.

A recent study examined oxidative stress levels in the house sparrow. The house sparrow was selected because it is a globally distributed species and is well adapted to urban life. Despite this adaptation however, when blood samples were drawn and compared among birds that inhabit three different environments common to the house sparrow—urban, suburban and rural—the urban sparrows displayed higher levels of oxidative damage and lower antioxidant capacity compared to rural birds. This finding demonstrates some of the urban stressors common to not only birds but other living creatures, such as pets.²

> Causes of oxidative stress for urban pets: chemical, noise and light pollution; infectious disease; and diet quality.



Simple Aging

Current estimates suggest there are more than 52 million senior or geriatric dogs in the U.S. today, or dogs above the age of seven years. Improved nutrition and veterinary care have extended the average lifespan of companion animals. Just as scientists have had to study the effects of aging on humans, researchers are gathering information about senior animals as well. One study conducted with 80 Labrador retrievers from adulthood to the end of life found that aging dogs undergo changes similar to humans in terms of inflammatory response and an increase in oxidative damage. Senior dogs can struggle with pathologies or health issues, such as arthritis, diabetes, dementia, obesity, blindness and cognitive decline.³

Antioxidants: The First Line of Defense

Clearly part of the solution for combatting oxidative stress is to find its antidote. In the case of oxidation, it is the antioxidant, sourced from certain fruits and vegetables, that can be added to the diet and help neutralize free radicals.

Antioxidants supply multiple health benefits, including but not limited to anti-inflammatory effects, positive impact on immunity, retinal responses and physiological changes in the brain.

Immunity

Inflammation or an inflammatory response is the first line of defense to combat harmful stimulus. The types of stimulation that can trigger an inflammatory response can range from physical wounds like cuts and bruises to infections, exposure to pollution or even noise. When the inflammatory response is triggered, this enhances the production of free radicals which in turn causes oxidative stress, particularly when it happens on a repeated basis. Persistent or chronic inflammation has been linked to multiple major diseases, such as arthritis, cancer, heart disease and Alzheimer's to name a few.

Anti-inflammatory Foods

Many fruits and vegetables contain a diverse array of phytochemicals known as phenolics. This includes flavonoids, such as anthocyanins. Some anthocyanins, like quercetin for example, can cross the blood-brain barrier. Phytochemicals have a complex molecular structure allowing them to bind free radicals, so they can not attack metabolic systems within the body.

Some of the fruits experts recommend for their antiinflammatory properties can include pineapples, apples, mangos, cantaloupes and berries, particularly blueberries. Vegetables like pumpkin, squash, sweet potatoes and carrots can also be very helpful for inflammation due to the beta-carotene content.

Visual Response

Antioxidants are now documented as influencing retinal response, particularly in dogs. A specific study published in the *Journal of Nutritional Science*⁴ found, "Antioxidant supplementation...may be beneficial and effective in the long-term preservation and improvement of various functions of the canine eye." The study looked at beagles given a feeding regimen with or without antioxidant supplementation. Compared with the control group, the antioxidant-supplemented group showed improvement to varying degrees for retinal function and "significantly less decline in refractive error." Dogs do experience vision decline as they age. Retinal degeneration and a cloudy lens are "common forms of eye problems that results in a decline of visual function in dogs."



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Heart Disease



Heart disease is another health issue related to oxidative stress. A study published in *Veterinary Medicine International* ⁵ said that heart failure was one of the main causes of death for domestic carnivores. Well adapted nutrition it said, "constitutes a major asset to improve the well-being and life expectancy" of these companion animals. A study found that in dogs with heart failure, their cellular metabolism isn't working properly, leading to production of a large number of free radicals. However, a proper diet supplemented with the right blend of vitamins and polyphenols can increase the concentration of antioxidants to help neutralize free radicals and improve the canine's cellular metabolism.

Anthocyanin: A Powerful Polyphenol

Among antioxidants, one particular group of compounds called anthocyanins has excited the medical and scientific community as specific benefits associated with them are identified and studied. Anthocyanins are found abundantly in brightly colored fruits and vegetables and responsible for bright red-orange and blue-violet pigmentation. The concentration of anthocyanin within any given food type will vary. Links have been found between anthocyanins and cardiovascular disease, cancer and cognitive function.

Flavonoids like anthocyanins can aid with memory and help stem the tide of age-related cognitive decline. Several factors are at work here, including their ability to inhibit neuroinflammation and improving blood flow to the brain. One more important factor is that certain anthocyanins can cross the blood brain barrier, making them more effective in a direct fashion in terms of beneficial cognitive effects.

One animal study of aged rats, which were fed blueberries,⁶ revealed a relationship between cognitive performance tested in a maze, and the total number of anthocyanin compounds found in the cortex. The anthocyanins or polyphenolic compounds in the blueberries were able to cross the blood brain barrier and localize in various brain regions important for learning and memory.

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A series of studies looked at the memory response or brain aging in older dogs and found that an antioxidantenriched diet enabled the dogs to learn complex tasks with greater success than the dogs fed a control diet without antioxidants.⁷ And yet another study found that older dogs fed an antioxidant-enriched diet displayed cognitive improvement within two weeks of starting the enriched diet, and when combined with mental stimulation, the effects were "even greater."

Dogs fed a diet high in antioxidants in another study, could recognize family members and other animals more easily than the control group. In addition, the dogs displayed a greater degree of agility with the anti-inflammatory effects of antioxidants in play. Antioxidants even have been shown to help dogs and cats that suffer from allergy or coat and skin problems. Experts recommend that owners may want to start antioxidant supplementation while dogs are still young, rather than waiting until they're older, to lower the chance of the dog developing cognitive dysfunction syndrome (CDS) later in life.⁸

One study in a veterinary journal about oxidative stress in canines emphasized "the most important aspect of this work is the discovery that cognitive performance can be improved by dietary manipulation, and that "antioxidants may potentially act, therefore, to prevent the development of ...age-associated behaviors and possibly even neuropathology by counteracting oxidative stress."

Role of Antioxidants and Natural Products in Inflammation. Oxidative Medicine and Cellular Longevity 2016.⁹

Antioxidant Sources

Clearly, the addition of antioxidant-containing ingredients to kibble or in pet treats can be an effective means of supplementing the companion animal diet to effectively counter oxidative stress. Fruit in pet treats, for example, can add fiber, aid with texture and humectancy, enhance color and flavor, and of course, supply antioxidant components that can combat health concerns related to oxidative stress. Adding to this list of benefits, fruit and vegetable ingredients contain no fat and are typically lower in calories than many other ingredients.

It is possible to add a greater percentage of fruit to a mix-in or a treat to maximize the potential phytonutrient content than if it were added within the formulation of standard kibble. Different formulations can tailor the fruit content and type to design treats or mix-ins that target different animal demographics, such as senior pets.

A listing of some of the potential fruits and vegetables suitable for canine consumption are listed in the sidebar. The various phytonutrient benefits associated with the individual fruits and vegetables are also listed, although a more thorough explanation is available through personal consultation with one of Tree Top Fruit Ingredients Specialists.

Mix-ins or Treats

One option for adding functional benefits to a pet's diet is with a superfood mix-in or topper made with simple and easily recognizable ingredients, such as fruits mixed with vegetables and/or whole grains. Mix-ins supply consumers with an easy method of adding to standard kibble, a scientifically balanced blend of simple, wholesome ingredients to help impact weight management or improve certain health aspects.

Typically, these mix-ins or toppers contain antioxidantrich ingredients that support overall health and wellbeing, while the free-from formulations appeal to picky pet parents. These toppers can be formulated free from soy, flavor enhancers, added sugar, artificial colors, and preservatives, and are both gluten- and GMO-free.

The second method of adding antioxidant-rich ingredients to a pet's diet is via a chew or treat. A chew can also combine more than a single fruit ingredient and be designed for a specific health issue. For

BENEFICIAL FRUITS/VEGETABLES:

- Apples (acetylcholine) Supply a source of fiber, phytonutrients, flavonoids, vitamin C, and potassium.
- Blackberries America's home-grown "super" fruit, blackberries contain high levels of antioxidant phytonutrients (particularly anthocyanins). Blackberries supply a good source of manganese, vitamin C, folate acid and magnesium, and as one of the brightly colored berries, a good source of anthocyanins.
- Blueberries High in vitamins and minerals, including vitamins C and K, and manganese, as well as a good source of dietary fiber.
- Tart Cherries Supply a source of fiber, manganese, vitamin A, vitamin C, and are high in antioxidants. The anthocyanins in tart cherries may reduce inflammation, according to some studies.
- Cranberries A rich source of several vitamins and minerals, especially vitamin C, manganese, vitamin E, vitamin K1, and a trace element of copper, as well as a source of both soluble and insoluble fiber.
- Pears Contain a sweet natural flavor and are a good source of potassium and dietary fiber, while also supplying vitamin C and vitamin A.
- Pumpkin A good source of fiber that helps support healthy digestion.
- Spinach High in vitamins A, B, C and K. Spinach also contains iron, antioxidants, beta-carotene and roughage, which stimulates the digestive tract.
- Strawberries Packed with vitamins, fiber and high levels of antioxidants; among the top 20 for fruits in antioxidant capacity. Good source of vitamin C, manganese, folate and potassium.
- Sweet Potatoes Low in sodium and very low in saturated fat and cholesterol. Sweet potatoes provide a good source of dietary fiber, vitamin B6 and potassium and a very good source of vitamin A, vitamin C and manganese.



example, a combination of berries with a high anthocyanin content could be designed for the urban dog to combat the daily stressors that can cause an oxidative/free radical imbalance. In one sample, human grade ingredients blend beef broth, with a variety of fruits selected for their phytonutrient content, mixed with peanut butter, some puffed cereal of brown rice and select spices.

Leveraging the antioxidant properties of fruit and vegetable ingredients for companion animal formulations can help improve quality of life, help combat diseases, and help mitigate the memory loss for a happier, healthier canine. These naturally tasty ingredients offer functional benefits, clean label attributes and an impressive portfolio of phytonutrients that will please pet parents as they see improvements in their dog's health and well-being.

Tree Top Forms Available

Single Strength Fruit Purées Concentrated Fruit Purées Drum Dried Fruit Flakes Regular/Evaporated Moisture Apples Low Moisture Apples Low Moisture Air-Dried Powders Low Moisture Puffed Apples Apple Sauce

Custom Pouch Solutions for Your Contract Co-packaging Needs

Fruit & Veggie Blends Pouch Sizes: 3.2 oz, 4.0 oz, 6.5 oz Minimum run size 84,000 pouches



Bio

Doug Webster, Director of Research and Development

Doug Webster has been with Tree Top, Inc. since 2000 and is currently Director of Product Development. In this role, he leads a research and development team, creating custom fruit solutions for customers of Tree Top. Doug earned a bachelor's degree in biology at Western Washington University and a Master of Science in food science from Washington State University. After completing his masters, he moved to the Yakima Valley, focusing his endeavors in the hop, wine and fruit industries. Doug is a winemaker, and until recently, owned his own winery. He now only makes wine for family and friends.

References

- 1. Isaksson, C. (2015), "Urbanization, oxidative stress and inflammation: a question of evolving, acclimatizing or coping with urban environmental stress." Funct Ecol, 29: 913-923. doi:10.1111/1365-2435.12477
- Herrera-Dueñas Amparo, Pineda-Pampliega Javier, Antonio-García María T., Aguirre José I. (2017), "The Influence of Urban Environments on Oxidative Stress Balance: A Case Study on the House Sparrow in the Iberian Peninsula." Front Ecol Evol, 5. doi.org/10.3389/fevo.2017.00106
- Alexander, J. E., Colyer, A., Haydock, R. M., Hayek, M. G., & Park, J. (2017). "Understanding how dogs age: Longitudinal analysis of markers of inflammation, immune function, and oxidative stress." The Journals of Gerontology: Series A Biological Sciences, 73 (6): 720-728 https://doi.org/10.1093/gerona/glx182
- 4. Wang, Wei et al. (2016) "Antioxidant supplementation increases retinal responses and decreases refractive error changes in dogs." Journal of Nutritional Science vol. 5 e18. 10, doi:10.1017/jns.2016.5

 Sagols, Emmanuelle, and Nathalie Priymenko. (2011) "Oxidative stress in dog with heart failure: the role of dietary fatty acids and antioxidants." Veterinary Medicine International vol. 2011 180206. doi:10.4061/2011/180206

- 6. Andres-Lacueva, Cristina et al. (2005) "Anthocyanins in aged blueberry-fed rats are found centrally and may enhance memory." Nutritional Neuroscience 8(2):111-120, doi.org/10.1080/10284150500078117
- 7. Dowling, Amy L S, and Elizabeth Head. (2012) "Antioxidants in the canine model of human aging." Biochimica et biophysica acta vol. 1822,5: 685-9. doi:10.1016/j.bbadis.2011.09.020
- 8. Cotman CW, Head E, Muggenburg BA, et al. (2002) "Brain aging in the canine: a diet enriched in antioxidants reduces cognitive dysfunction." Neurobiol Aging 2002;23:809–818.
- 9. Arulselvan, Palanisamy et al. (2016) "Role of Antioxidants and Natural Products in Inflammation." Oxidative Medicine and Cellular Longevity, doi:10.1155/2016/5276130

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Combating Oxidative Stress in Companion Animals: Leveraging the Antioxidant Properties of Fruits and Vegetables

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