

## **EXPOSURE TO NEPHROTOXINS**

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A nephrotoxin is any substance that causes injury to the kidneys. The mammalian kidney is an extremely complex organ, both anatomically and functionally. In addition to the excretion of wastes, the kidney plays a significant role in the regulation of total body homeostasis; it is the predominant organ involved in regulation of extracellular fluid (ECF) volume and electrolyte composition. The kidney also is a major site of formation of hormones that influence systemic metabolic functions, including erythropoietin, 1,25-dihydroxy-vitamin D<sub>3</sub>, rennin, and several vasoactive prostanoids and kinins. A toxicological insult to the kidney could affect any or all of these functions. However, the effects usually reported following toxic insult reflect decreased elimination of wastes, that is, an increase in blood urea nitrogen (BUN) or an increase in plasma creatinine. This does not necessarily mean that excretory functions are primarily affected by nephrotoxins; rather, these are renal functions that are measured rapidly and reliably. Thus, the use of BUN and plasma creatinine as clinical indices of nephrotoxicity reflects the state of technology, not necessarily the primary sites of nephrotoxicity.

### **Specific Nephrotoxins**

As indicated above, not all compounds that influence renal function affect the kidney directly. Renal function may be altered secondarily to changes in blood pressure, blood volume, or neural or hormonal influences and to a variety of destructive systemic effects. A focus will be on compounds that produce specific effects on the kidney. Many of these agents affect the kidney directly. In other cases, the ultimate toxicant might be a metabolic product formed within the kidney or produced in an external organ and transported to the kidney where it acts directly or requires further metabolism to a nephrotoxic product. (See Table 1.1) Some of these types of compounds are listed below and are not limited to:

**Heavy Metals** - mercury, cadmium, chromium, arsenic, gold, lead, iron, antimony, uranium, lead, zinc, nickel, and thallium.

**Halogenated Hydrocarbons** – chloroform, bromobenzene, hexachlorobutadiene, petroleum hydrocarbons.

**Therapeutic Agents** – microbicides, nanocomposites, analgesics, anesthetics, antibiotics/antivirals, antifungals, cephalosporins, tetracyclines, demeclocycline, Amphotericin B, antineoplastic agents (cisplatin)

**Immunosuppressants** – cyclosporine and others

**Environmental Contaminants** - herbicides, pesticides, hazardous wastes, substances, Chem Trails residues, fungicides, biological and chemical agents, advanced nano materials and biomaterials (microbicides), botanicals, mycotoxins, organic solvents, experimental tools/miscellaneous polymers.

## General Information on Kidney Disease (Failure)

Kidneys remove waste products from the body, keep body chemicals in balance, and help maintain the body's water balance. There are a number of different renal (kidney) problems that may occur. The kidneys may be damaged by exposure to certain drugs or toxins, including heavy metals, solvents, chemotherapy agents, snake or insect venom, poisonous mushrooms, and pesticides. Renal failure can also accompany or result from many other disorders, such as congestive heart failure, diabetes, chronic hypertension, liver disease, lupus, and sickle-cell anemia.

*Bright's disease* is a kidney disease marked by the presence of blood protein in the urine, along with hypertension and edema (retention of water in the tissues), Glomerulonephritis is an inflammation of tiny blood vessels within the kidney that filter out wastes from the blood. This may occur as a result of an immunological response to infection (severe septicemia), which may be triggered by exposure to microorganisms (pathogens), microbivores (urine bubbles), hazardous materials, venoms, mycotoxins, endotoxins, exotoxins, and many other toxic substances. In some cases of microbivores (bacteriophage/nanorobotics) the immune system does not respond at all. If a true infectious material, such as Streptococcus throat infection, it can have an affect upon the kidney. Pyeloneophritis is a kidney infection that may be caused by a birth defect. Both glomerulonephritis and pyelonephritis can be chronic or acute, and can be serious. *Hydronephrosis* is a condition in which the kidney and the renal pelvis (the structure into which urine is discharged from the kidney) become filled with urine due to an obstruction of urinary flow. *Polycystic kidney disease* (PKD) is stated as being an inherited disease in which cysts grow on the kidneys as balloons. This ballooning effect may be the direct cause of environmental pollutants being stuck within the nephrons (filters) of the kidney and ballooning in the area. Cysts are formed from the death of the white blood cells responding to a particular infectious material or hazardous material (such as heavy metal). The cysts grow on the kidneys, rendering them incapable of functioning. Kidney stones are mineral accumulations (primarily calcium) in the kidneys. In renal tubular acidosis, the kidneys fail to reabsorb bicarbonate normally, causing impaired ammonia production and acid excretion. Severe acidosis, potassium depletion, and bone disorders may result. Nephrotic syndrome is not a disease itself, but can be a sign of kidney disease. It is marked by edema and excess protein in the urine. It can be caused by lesions of glomeruli (small structures in the kidney made of capillaries) that become inflamed, or by chronic diseases such as diabetes or lupus.

One important symptom of kidney problems is edema. Edema results when the kidneys produce less urine because they are unable to properly excrete salt and other wastes, and fluid builds up in the body. Ankles and hands may swell, and the person becomes short of breath. Toxic wastes may accumulate in the blood stream due to kidney malfunction, a condition known as uremia. Symptoms of kidney problems included abdominal pain, appetite loss, back pain, chills, fever, fluid retention (bloating), nausea, urinary urgency, and vomiting. The urine may be cloudy, bloody, or bubbles (microbivores). Back pain may be sudden and intense, occurring just above the waist and running down the groin.

## Nutrients

The following supplements aid in controlling urinary tract infection, exposure to hazardous materials and other related toxins to help maintain proper kidney function. Unless otherwise specified, the dosages recommended here are for adults. For a child between the ages of twelve and seventeen, reduce the dose to three-quarters of the recommended amount. For a child between six and twelve, use one-half of the recommended dose, and for a child under the age of six, use one-quarter of the recommended amount.

| Supplement                          | Suggested Dosage  | Comments   |
|-------------------------------------|---|--|
| Acidophilus                         | As directed on label, 3 times daily. Take on an empty stomach.  | Especially important if antibiotics.   |
| Coenzyme A                          | As directed on label.   | Acts as an antioxidant & removes harmful substances from the body.             |
| Restorex Complex                    | As directed on the label, 3 times daily.  | Aids in building cellular compounds.   |
| Vitamin B <sub>6</sub>              | Pain, 500 mg/day<br>50 mg, 4 times/day  | To reduce fluid retention  |
| Choline                             | 50 mg daily   |  |
| Inositol (IP <sub>6</sub> )         | 100 mg daily  |  |
| Vitamin C with Bioflaonoids         | 2,000 – 4,000 mg daily  | Acidifies the urine, boosts immune function, and aids in cell membrane healing |
| Calcium & Magnesium                 | As directed on bottle, "Calm"<br>1,500 mg daily (Ca)<br>750 mg (magnesium) daily  | For proper mineral balance; calcium and magnesium should be in a 2 to 1 ratio  |
| L-Arginine<br>And<br>L-methionine   | 500 mg, 4 times day<br><br>As directed on label, on an empty Stomach. Take with water or juice. Do Not take with milk. Take with 50 mg Vitamin B <sub>6</sub> and 100 mg Vitamin C for better absorption. | For kidney disease<br><br>For improved kidney circulation                      |
| Lecithin granules<br>or<br>Capsules | 1 tbsp. 3 times daily<br><br>2,400 mg or IU's, 3 times a day before meals   | Needed for nephritis.  |

| SUPPLEMENT   | DIRECTIONS   | COMMENTS  |
|--|--|---|
| Digestive-Ease   | As directed on bottle  | Necessary for digestion   |
| Apple cider vinegar but prefer Pineapple Vinegar 1 teaspoon in a glass of water ½ hour before meals. |  |   |
| Multi Vitamin or VIBE  | As directed on bottle  | Necessary for cellular support.   |
| VermiiZyme   | As directed on bottle  | Necessary for mitochondria Support.   |
| Griminex   | As directed on bottle  | Hormonal support.   |
| Nettles, herb leaves and roots   | As directed on bottle  | Kidney & bladder cellular repair  |
| Vitamin A with<br>With<br>Mixed carotenoids  | 100,000 IU daily for 3 days<br>then 50,000 IU daily for 5<br>days, then reduce to 25,000 IU<br>Daily. If you are pregnant, do not<br>Exceed 10,000 IU daily. | Important in healing of urinary<br>tract lining and in immune<br>function. Use emulsion form<br>for easier assimilation and<br>greater safety at high doses.<br>DO not take this amount in pill<br>form. (soft gels – emulsion) |
| Vitamin B Complex  | 100 mg of each major<br>B vitamin daily (amounts of<br>Individual vitamins   | B vitamins work best when<br>taken together. Use a high<br>potency formula.   |
| Plus extra<br>Vitamin B <sub>2</sub>   | in a complex will vary)<br>25 mg 3 times daily.  | Needed for nephritis.   |
| 1 Tablespoon Olive Oil<br>or Vitamin E emulsion<br>or capsules                                       | 800 IU daily.<br>start with 200 IU daily and gradually<br>and increase to 1,000 IU daily   | Promotes immune function.<br>An important free radical.<br>destroyer. Use d-alpha-<br>tocopherol form.  |
| Zinc   | 50 – 80 mg daily. Do not<br>Exceed 100 mg daily<br>From all supplements  | An immunostimulant necessary<br>for healing and an important<br>inhibitor of crystallization and<br>Crystal growth. Use zinc<br>gluconate lozengers or<br>Coldezze, Opti Zinc for best<br>Absorption                            |
| Plus Copper  | for SOD-1 utilization<br>3 mg daily.   |   |
| Zinc & Copper may be substituted with LipoSOD  | As directed on bottle  | SOD Supplement & Potent<br>Anti-Oxidant Nutrient Repair<br>And support formula.   |

|                                      |                       |                                      |
|--------------------------------------|-----------------------|--------------------------------------|
| Phase 2/L-Carnitine with SX-Fraction | As directed on Bottle | Phase II Liver Detox                 |
| Super Tremella                       | As directed on Bottle | Nourishes the lung, stomach & Kidney |

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During the first 0 – 30 days (one month) it may be difficult to adjust to the changes for your body. One must remember that the condition that has expressed itself as chronic effect from nephrotoxins may have taken years to create. Commitment to healing your kidney and body is necessary. Nothing is easy when you are making changes out of CHAOS within the cell. Cellular dysfunction, injury and death are the cellular global mechanisms to create disease within a living life system.

**REFERENCES:**

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**TABLE 1.1: EXAMPLES OF OCCUPATIONAL/ENVIRONMENTAL AND EXPERIMENTAL NEPHROTOXICANTS\***

(Taken from Chapter 11: Toxic Responses of the Kidney, page 376 by William R. Hewitt, Robin S. Goldstein, and Jerry B. Hook. *Casarett and Doull's Toxicology: The Basic Science of Poisons, 4<sup>th</sup> Edition*. Mary O. Amdur, Ph.D., John Doull, Ph.D, M.D., and Curtis D. Klaassen, Ph.D. Pergamon Press. New York © 1991) and relative referenced internet listings of research archives.

| TOXICANT                           | REFERENCE               |
|------------------------------------|-------------------------|
| Mycotoxins/kBotanicals             | Bermdt. 1987            |
| Aflatoxin B                        |                         |
| Citrinin                           |                         |
| Ochratoxin A                       |                         |
| Monocrotalin                       |                         |
| Pyroolizidine alkaloids            |                         |
| Rubratoxin B                       |                         |
| Halogenated Aliphatic Hydrocarbons | Lock, 1988; Kluwe, 1990 |
| Bromobenzene                       |                         |
| Bromodichloromethane               |                         |
| Carbon tetrachloride               |                         |
| Chloroform                         |                         |
| Chlorotrifluoroethylene            |                         |
| Dibromochloropropane               |                         |
| Dichloroethane                     |                         |
| Hexachlorobutadiene                |                         |
| Pentachloroethane                  |                         |
| Trichloroethylene                  |                         |
| Tetrachloroethylene                |                         |
| Tetrafluoroethylene                |                         |
| Tris (2,3-dibromopropyl)-phosphate |                         |
| Volatile Hydrocarbons              | Alden, et al, 1984      |
| Petroleum fuels                    | Swenberg, et al. 1989   |
| Metals                             |                         |
| Cadmium                            | Fowler, et al, 1987     |
| Gold                               | Wedeen, 1988            |
| Lead                               |                         |
| Mercury                            |                         |
| Nickel                             |                         |
| Chromium                           |                         |
| Uranium                            |                         |
| Strontium-90                       | Staninger, 2009         |
| Herbicides/Fungicides              | Ecker et al, 1975       |
| Paraquat                           | Berndt, 1982a           |
| Diquat                             | Rankin et al, 1989      |
| Sussinimides                       |                         |
| 1,4,5-Trichlorophenoxyacetate      |                         |

**Table 1.1 continued**

| <b>TOXICANT</b>                                | <b>REFERENCE</b>            |
|--|-----------------------------|
| Organic Solvents                               | Gosselin et al, 1984        |
| Ethylene glycol                                |                             |
| Diethylene glycol                              |                             |
| Toluene  |                             |
| Experimental Tools/Miscellaneous               |                             |
| p-Aminophenol                                  | Newton et al, 1982          |
| Adrimaycin                                     | Elema, et al, 1988          |
| Benzidine                                      | Zenser and Davis, in press  |
| Bromoethanamine hydrobromide                   | Sabatini, 1988              |
| d-Serine                                       | Carone and Ganote, 1975     |
| d-Lysine                                       | Racusen, et al, 1985        |
| Limonene                                       | Lehman-McKeeman et al, 1989 |
| Maleic acid                                    | Berliner et al, 1950        |
| Microbivores & Nanorobots                      | Staninger, 2008 in press    |
| Modified Food Starch (Waterborne polyurethane) | Staninger, 2009             |
| Nano tubes, gels, & composites                 | Staninger, 2006, 2007       |
| Phenylanthranillic acid                        | Bach and Gregg, 1988        |
| Puromycin aminonecleoside                      | Elema et al., 1988          |

\* Only agents that have a direct effect on the nephron are cited. Examples represent some, not all, commonly cited nephrotoxicants.