

BIOACCUMULATION OF TOXIC SUBSTANCES

XENOBIOTICS released to the environment can be taken up by organisms

BIOACCUMULATION produces higher concentration of a chemical in an organism than in its immediate environment, including food

BIOCONCENTRATION

A process which leads to higher concentration of Xenobiotics in the organisms than in the environment

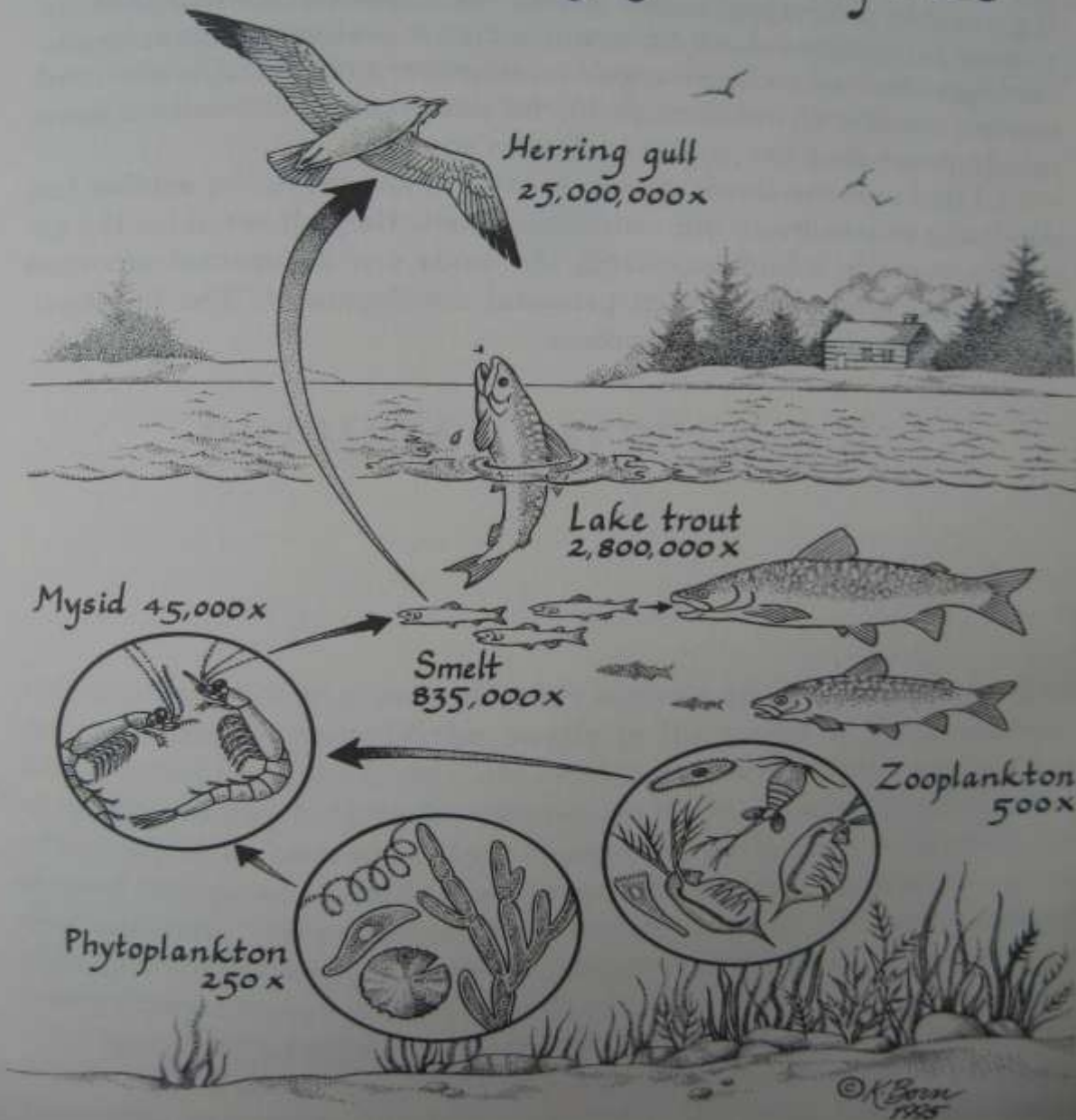
BIOMAGNIFICATION

A successive increase of bioaccumulation along the food chain - occurs when food is the major source of bioaccumulation

BIOACCUMULATION

- different routes (*via* air, water, solid & sediment)
- depend on environmental & physiological factors

Lake Ontario Biomagnification of PCBs



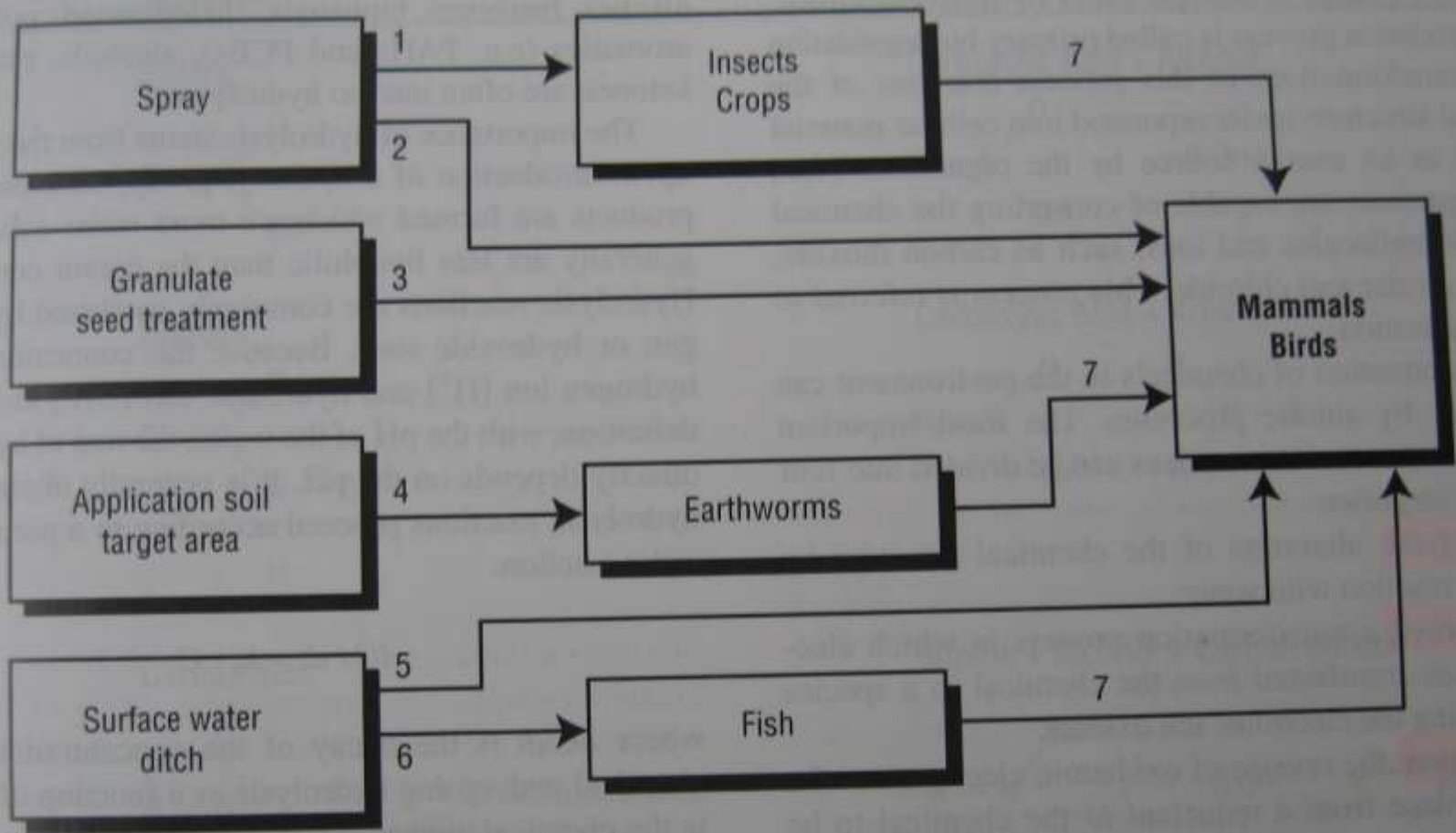


Figure 3.28. Food as a major source of contaminants for mammals and birds in a simplified food web. 1 = Application of spray, 2 = Drinking from leaves/crops, 3 = Ingestion of granules/treated seeds, 4 = Bioconcentration soil-worm, 5 = Drinking from surface water, 6 = Bioconcentration water-fish, 7 = Consumption. From USES [44]. With permission.



FUGACITY OF A CHEMICAL

The ratio of concentration to storage capacity

WATER

- Low concentration
- small storage capacity (solubility)

—————→ HIGH FUGACITY

ORGANISM

- Low ~> high concentration
- high storage capacity (per unit of volume)

—————→ LOW FUGACITY

HIGH FUGACITY

**CHEMICAL
FLUX**



LOW FUGACITY

ABSORPTION, DISTRIBUTION, AND EXCRETION OF TOXICANTS

Karl K. Rozman and Curtis D. Klaassen

INTRODUCTION

CELL MEMBRANES

Passive Transport

Simple Diffusion

Filtration

Special Transport

Active Transport

Facilitated Diffusion

Additional Transport Processes

ABSORPTION

Absorption of Toxicants by the
Gastrointestinal Tract

Absorption of Toxicants by the Lungs

Gases and Vapors

Aerosols and Particles

Absorption of Toxicants through the Skin

Absorption of Toxicants after Special Routes
of Administration

DISTRIBUTION

Volume of Distribution

Storage of Toxicants in Tissues

Plasma Proteins as Storage Depot

Liver and Kidney as Storage Depot

Fat as Storage Depot

Bone as Storage Depot

Blood-Brain Barrier

Passage of Toxicants across the Placenta

Redistribution of Toxicants

EXCRETION

Urinary Excretion

Fecal Excretion

Nonabsorbed Ingesta

Biliary Excretion

Intestinal Excretion

Intestinal Wall and Flora

Exhalation

Other Routes of Elimination

Cerebrospinal Fluid

Milk

Sweat and Saliva

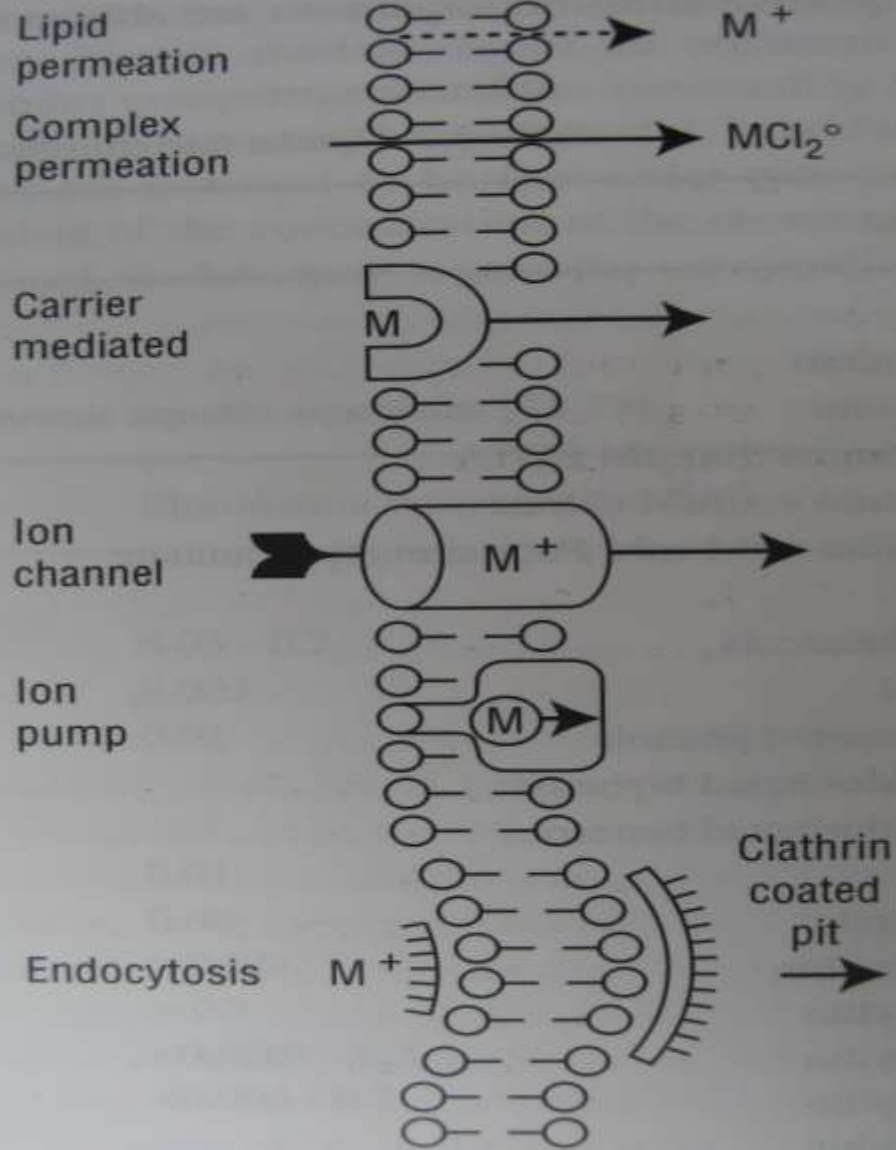
CONCLUSION



UPTAKE PROCESSES

- Several processes involving the passage of compounds across a biological membranes, mediated by a carrier or as a single solute
- Passive diffusion
- Fugacity
- Complex permeation
 - by carrier mediated processes
 - by ion channel
 - by ATPase





UPTAKE PROCESSES

Example:

Cadmium (Cd^{2+}) may be taken up by Ca^{2+} -ATPases

or as

Cadmium Xanthate complex (in fish)

FACTORS AFFECTING BIOCONCENTRATION

1. MOLECULAR SIZE

2. MOLECULAR CHARGE

3. Chemical SPECIATION: Cadmium [Cd Cl_2 , Cd NO_3]

4. SURFACE/VOLUME RATIO

~> smaller organisms take up and eliminate chemicals faster than large organisms

5. MORPHOLOGY, PHYSIOLOGY (e.g. filter feeder)

6. BIOTRANSFORMATION

~> degrades the chemical to more polar products catalyzed by enzymes



Chemical SPECIATION: Cadmium [Cd Cl_2 , Cd NO_3]

Cr^{3+} less toxic

Cr^{6+} highly toxic

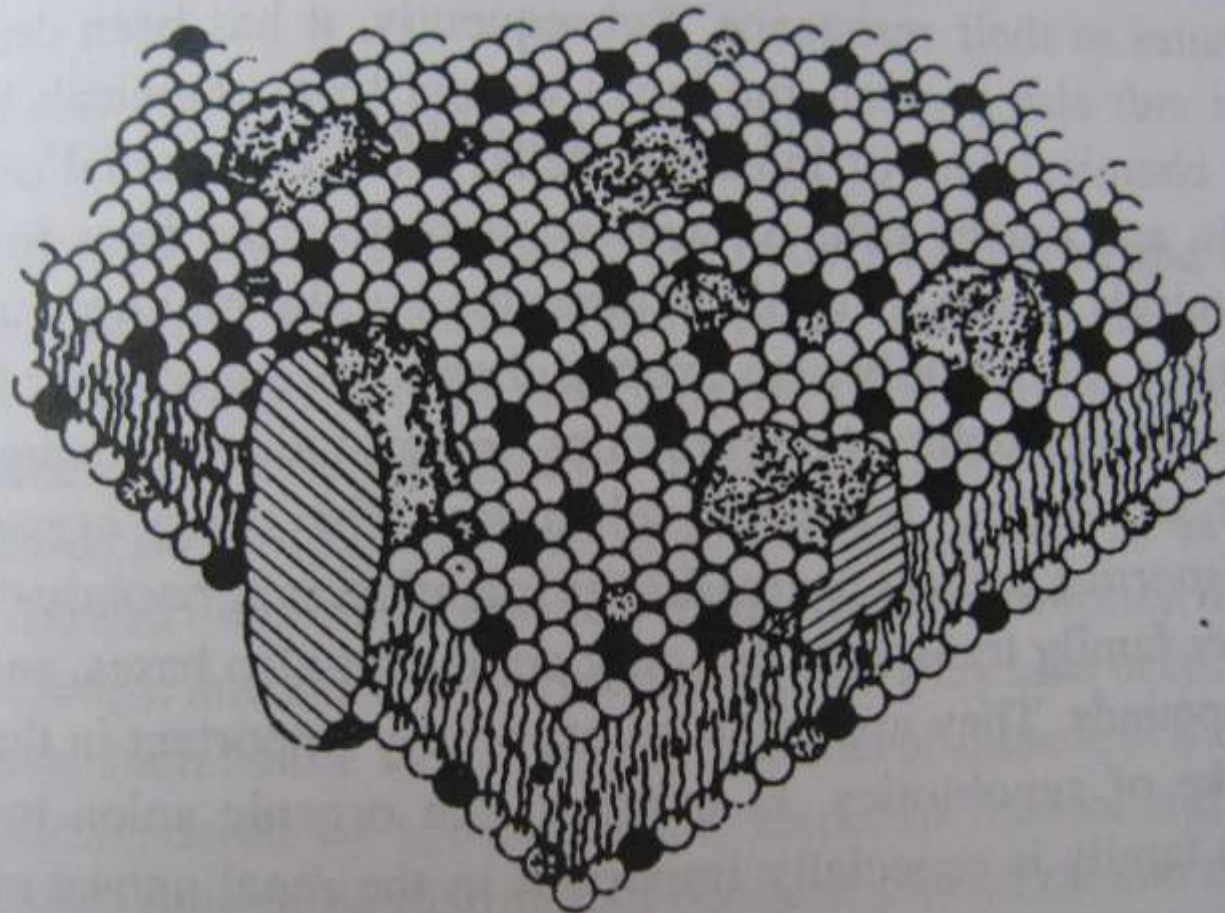


Figure 5-2. Schematic model of a biological membrane.



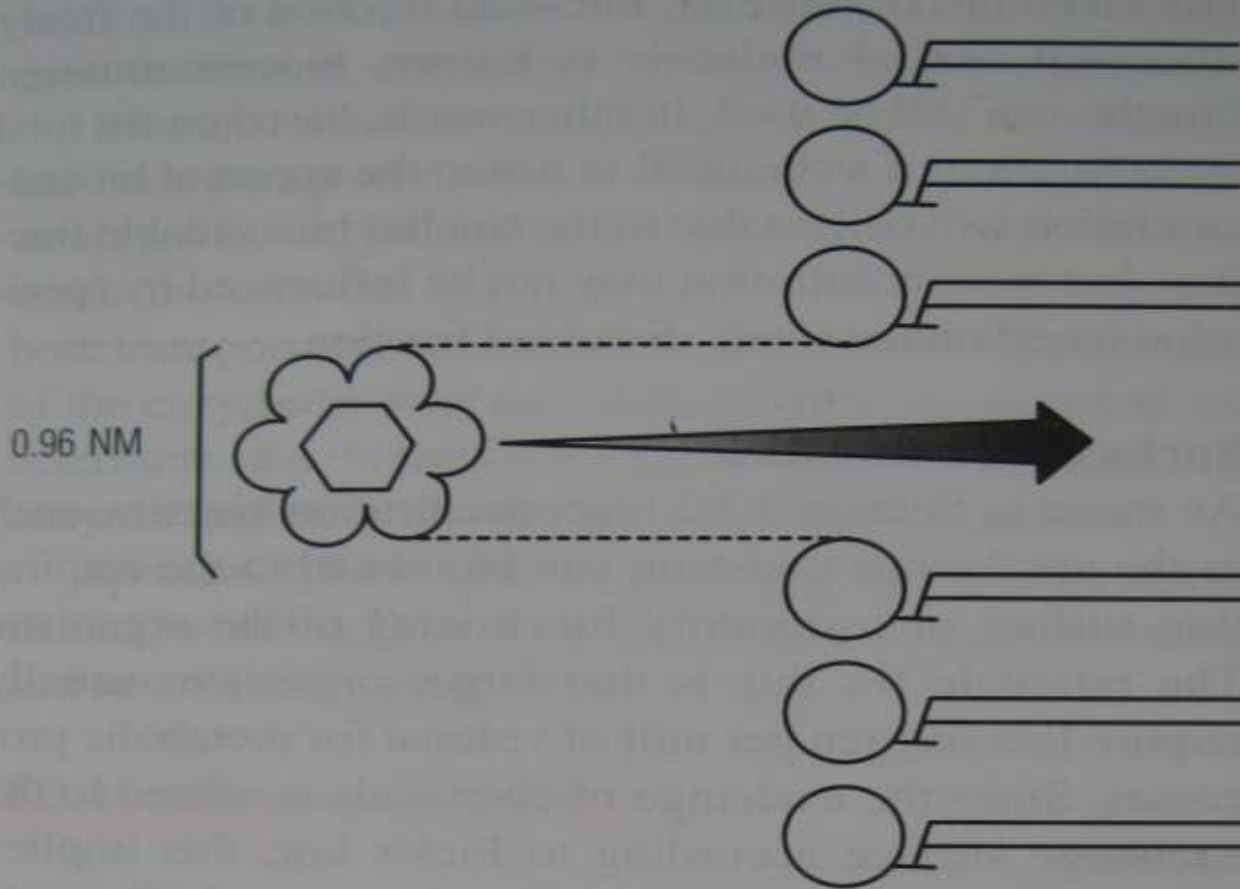


Figure 3.20. Diagram showing the transfer of a hydrophobic molecule across the polar heads of a bilipid membrane in relation to the effective cross-section of the membrane's cavity for neutral organic chemicals. Reprinted from [25]. With kind permission from Elsevier Science Ltd.



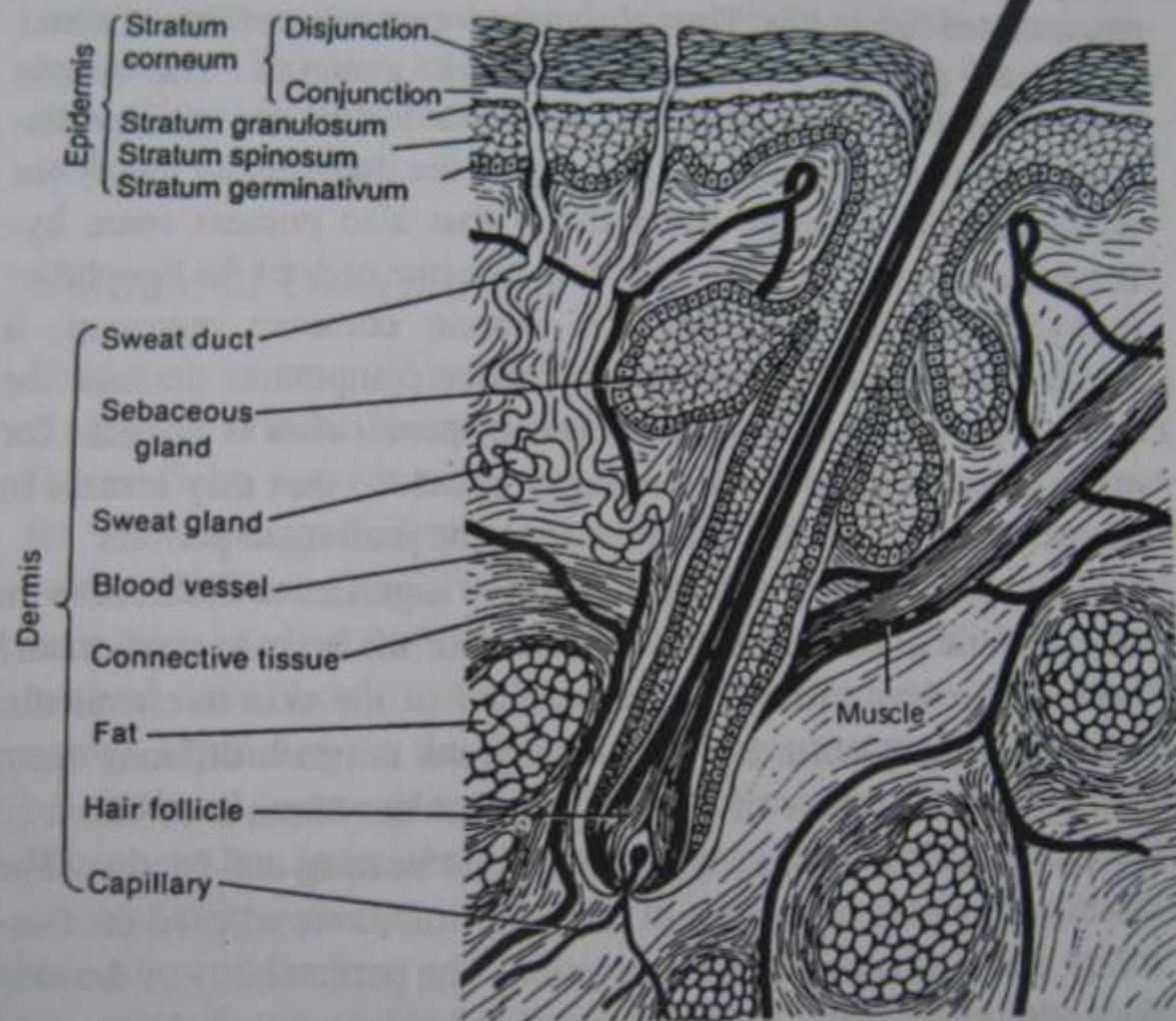
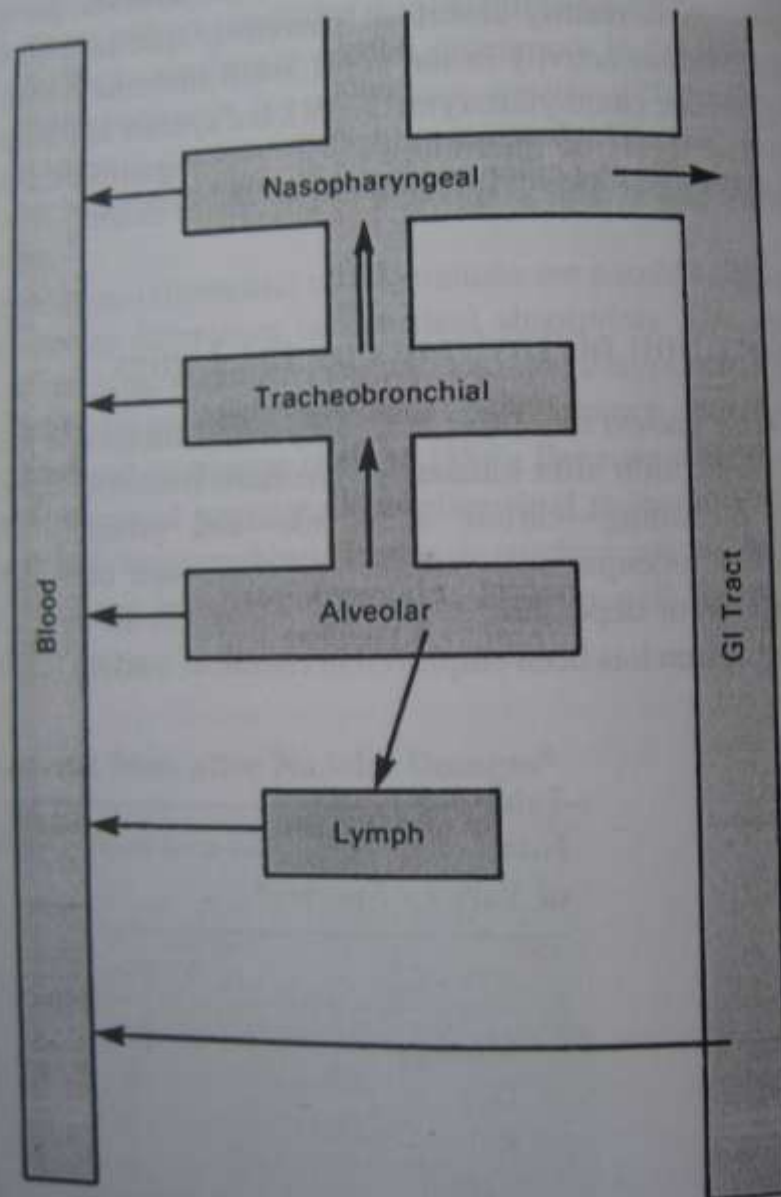


Figure 5-5. Diagram of a cross section of human skin.





Schematic Diagram of the absorption and translocation of chemicals by lungs



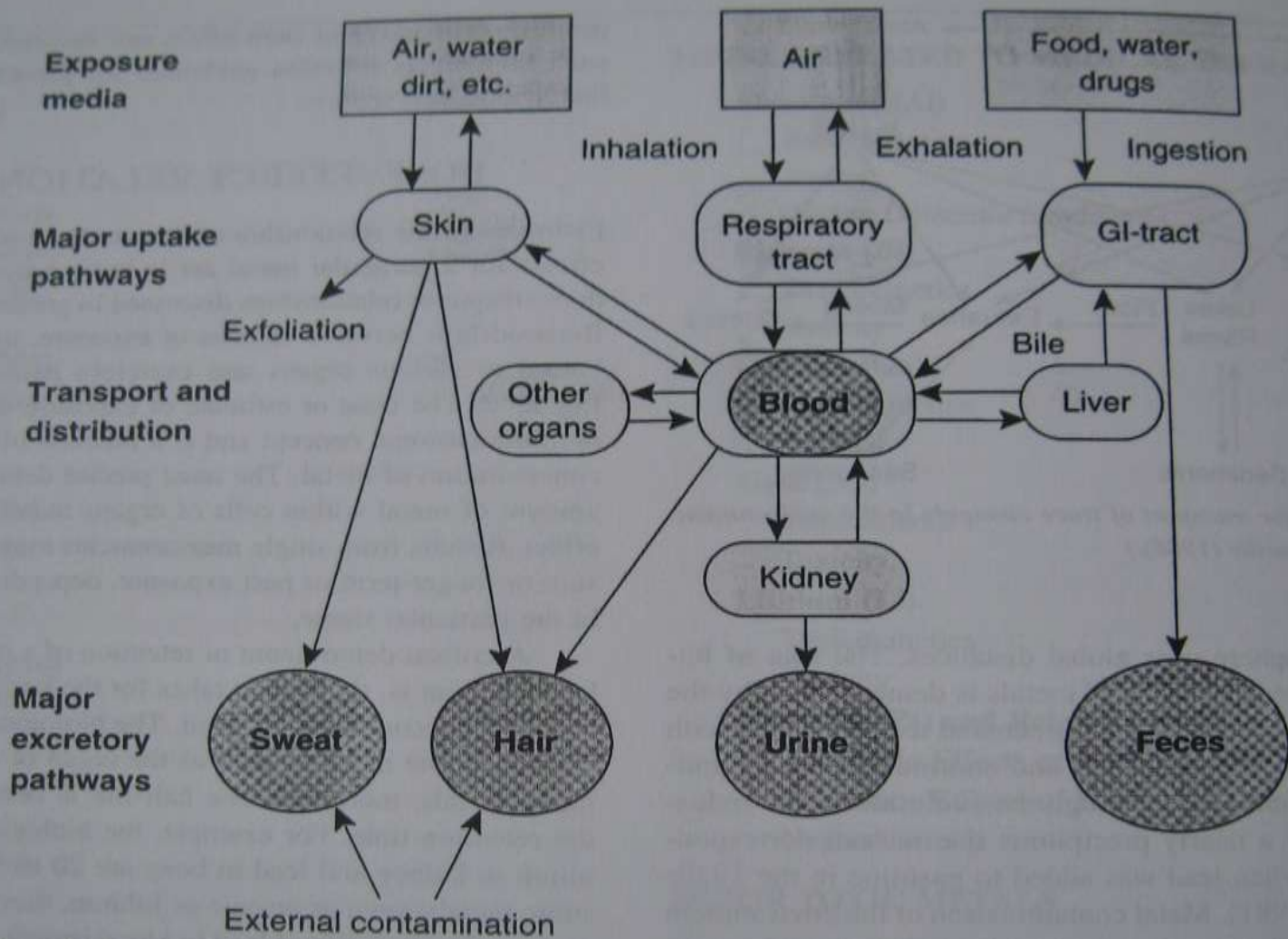


Figure 23-2. Metabolism after exposure to metals via skin absorption, inhalation and ingestion.

The arrows indicate how the metals are transported and distributed. Tissues that are particularly useful for biological monitoring are identified in shaded areas. [From Elinder et al. (1994), with permission.]

ORGANISMS (BIOACCUMULATION)

- METALS bind to protein (metallothionein)
- ORGANIC CHEMICALS are stored in lipids
- ORGANOMETALS are stored in either lipids or protein

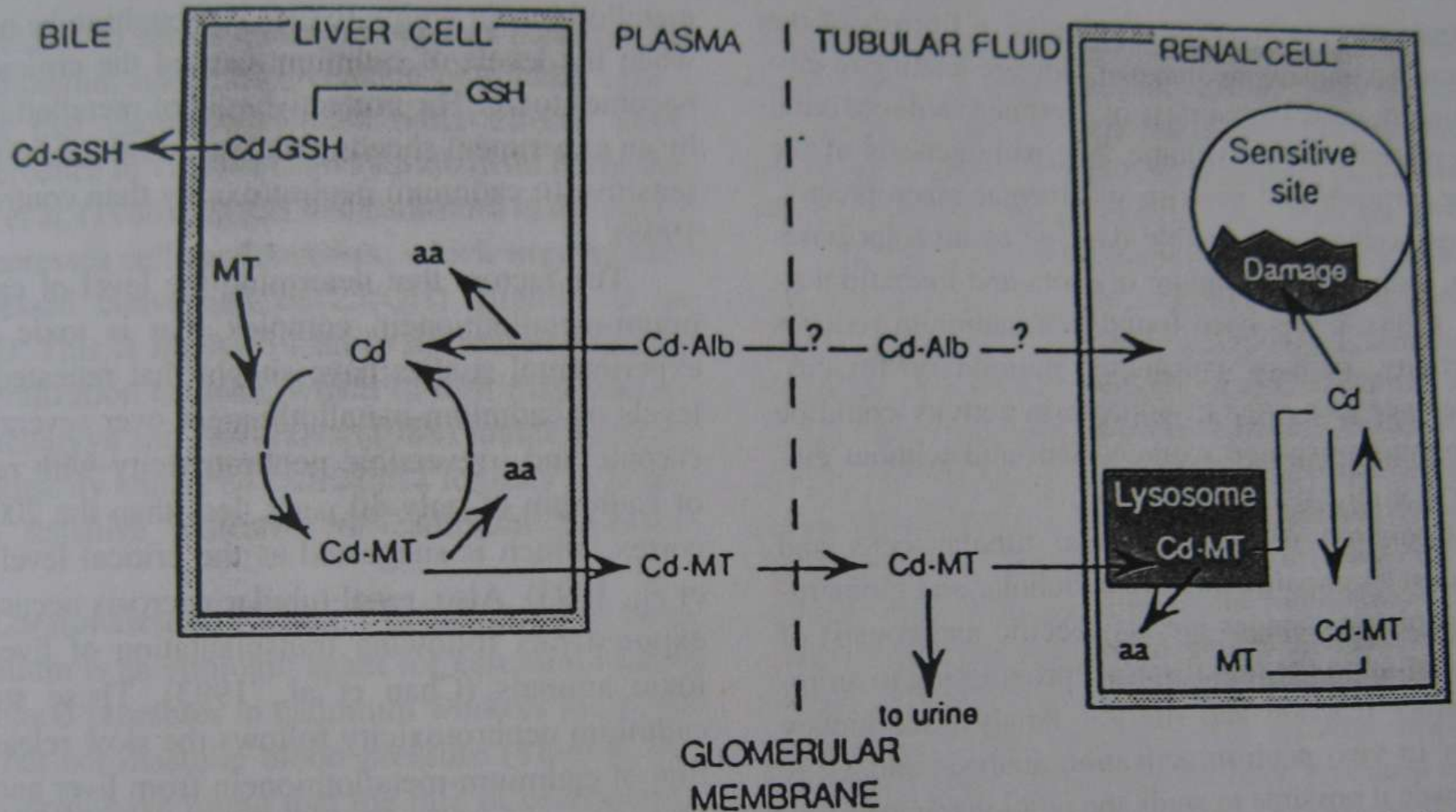


Figure 23-3. A schematic representation of the role of metallothionein in the disposition of cadmium in the liver and kidney. [Adapted from Jarup et al. (1998).]



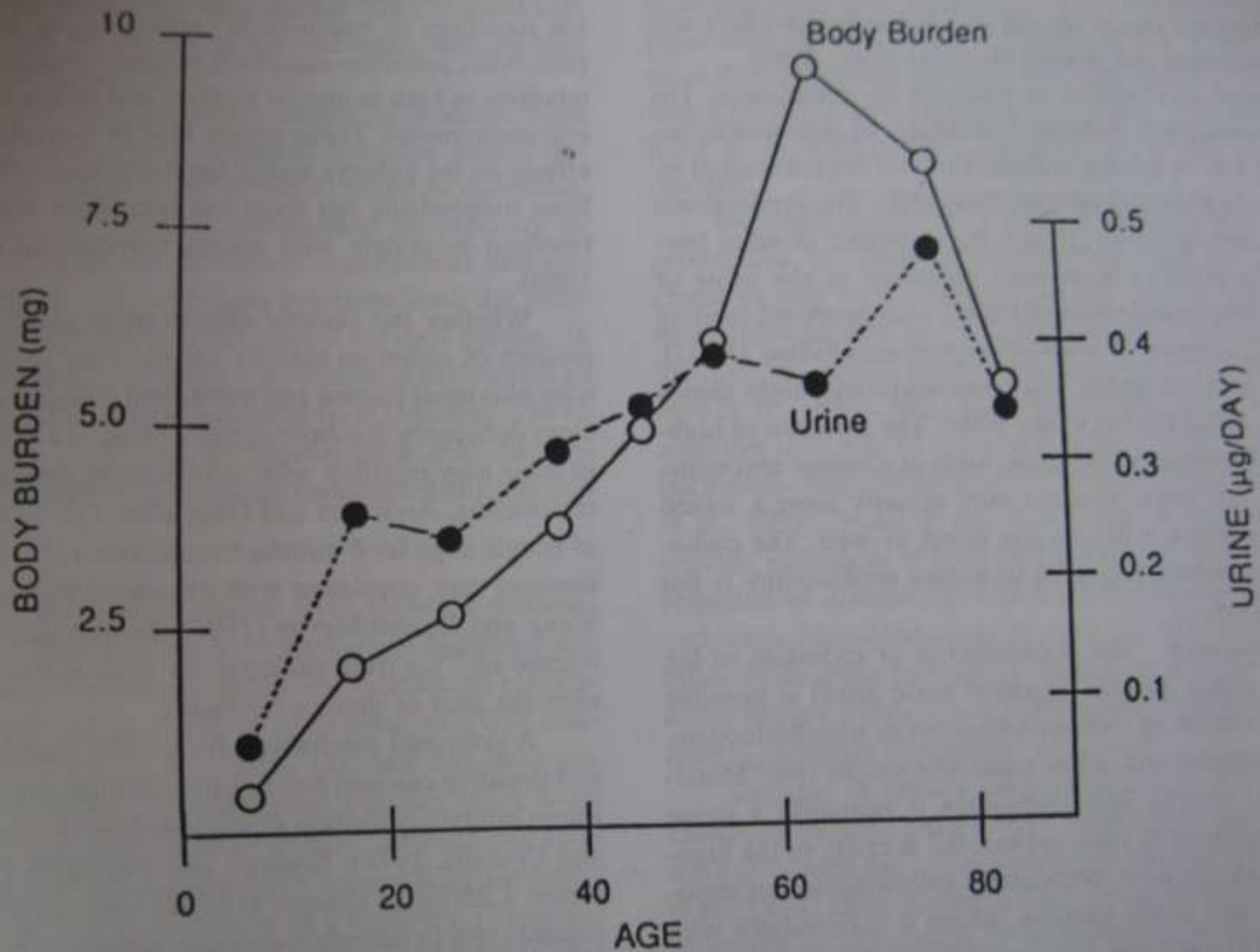


Figure 23-4. Illustration of the accumulation of cadmium with age.



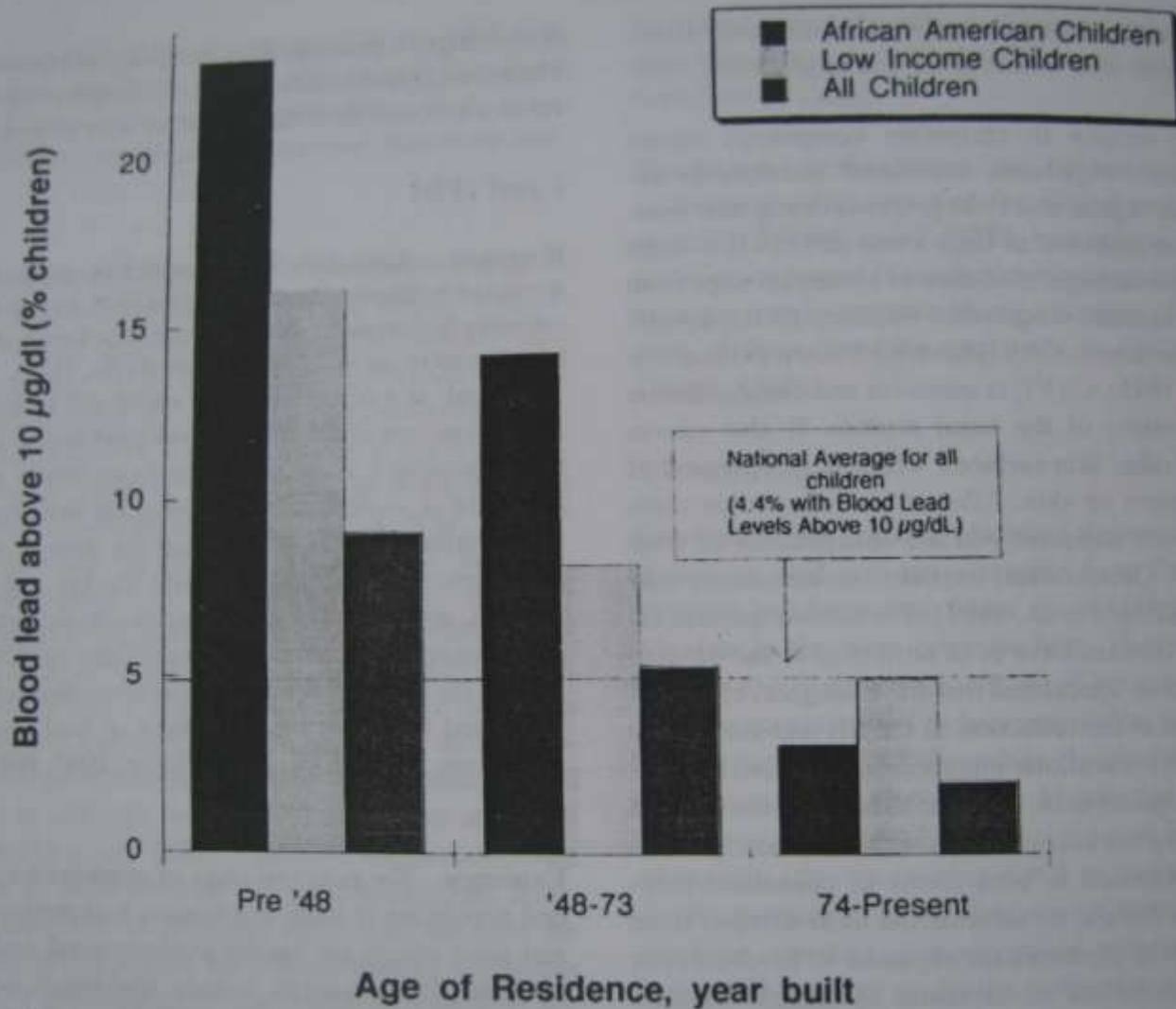


Figure 23-5. National blood lead levels. National Health and Nutrition Examination Survey, Phase 2, 1991-1994. [Adapted from CDC (2000).]



ELIMINATION

- Passive & active mechanisms
- Hydrophobic chemicals: passive diffusion (water, faeces)
- Growth
- Biotransformation



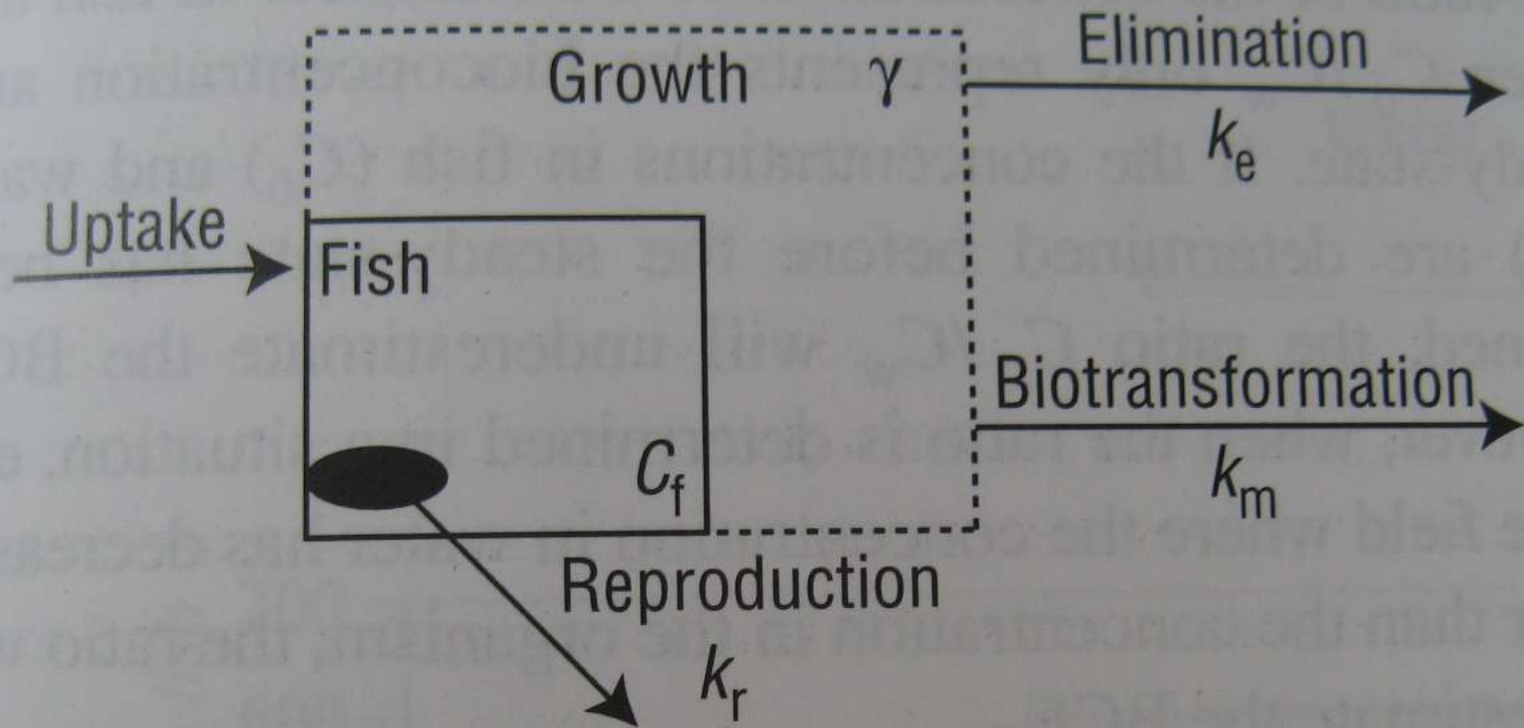
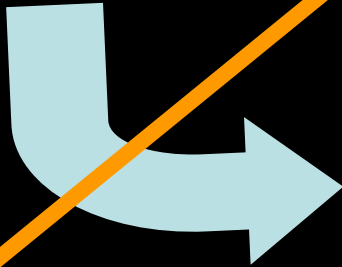


Figure 3.15. Different processes which reduce the concentration of xenobiotic contaminants in an organism (C_f): physico-chemical elimination (k_e), biotransformation (k_m), growth (γ) and reproduction (k_r). From [30]. With permission.

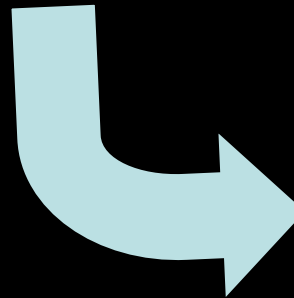


BAHAYA



RISIKO

**PAPARAN
(KONTAK)**



DAMPAK

**TAKARAN
(DOSIS)**



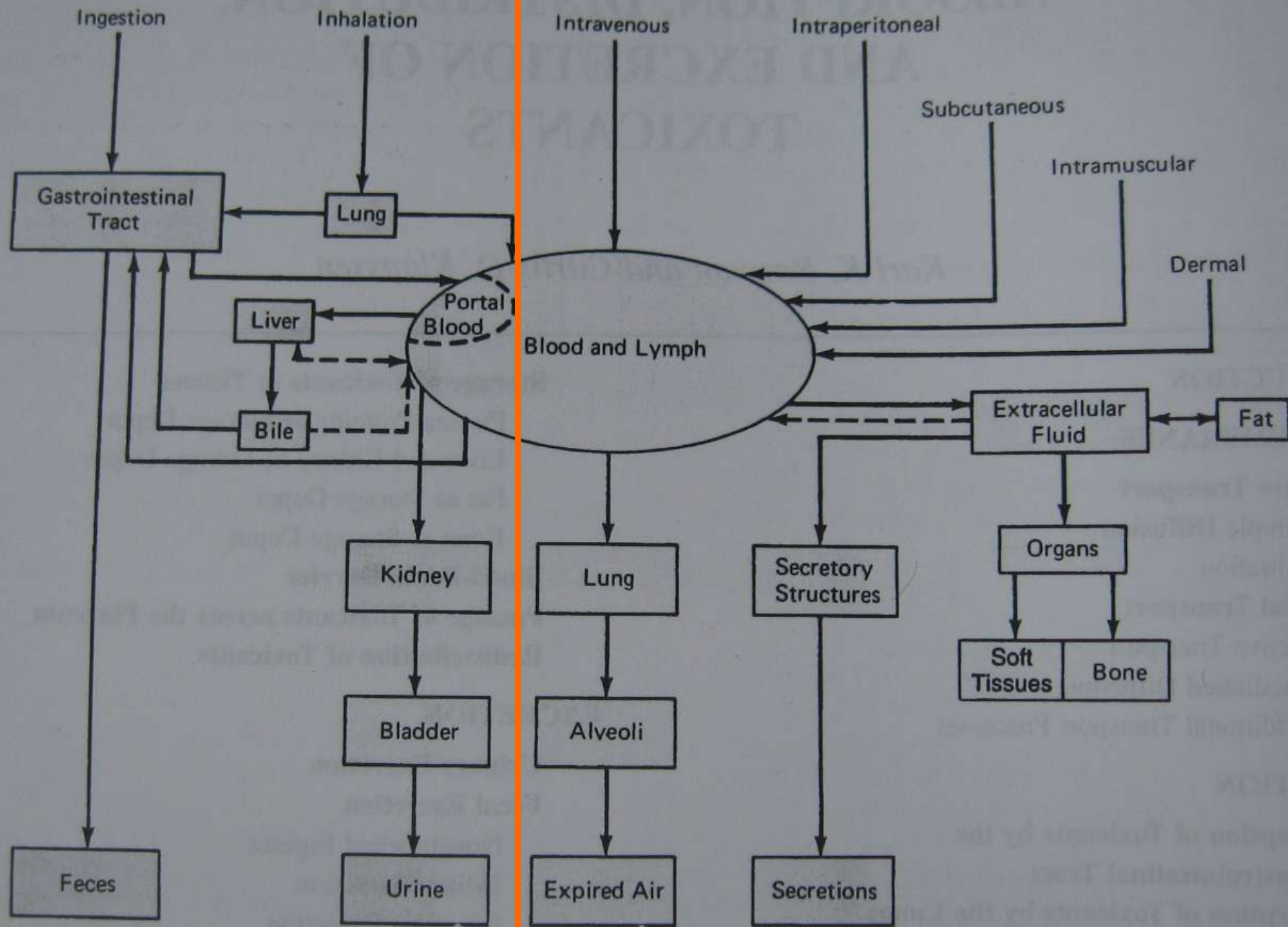


Figure 5-1. Routes of absorption, distribution, and excretion of toxicants in the body.