

UPTAKE AND DISTRIBUTION OF INHALATIONAL ANAESTHETICS

THESIS

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Master Degree of Anaesthesia*

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To the memory of
my father



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Introduction

I N T R O D U C T I O N

Inhaled anaesthetics produce anaesthesia when their tension in the brain reaches a certain level.

The level of anaesthesia produced is governed by the height of anaesthetic tension in the brain, which is under control of the attendant anaesthetist.

The tension gradient between the inspired and alveolar gas is built up by three factors (Ventilation, uptake and concentration). The effect of ventilation in increasing the alveolar concentration of the anaesthetic agent is countered by the effect of the blood removing it from the alveoli.

In this thesis, we will discuss the subject under three main headings, anaesthetic uptake, ventilation and the concentration effect, with special consideration of the effect of anaesthetics on air and gases in closed body cavities, and the effect of abnormalities of cardiopulmonary function, with added notes to placental transfer of inhaled anaesthetics.

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UPTAKE AND DISTRIBUTION OF VOLATILE ANAESTHETIC AGENTS

Anaesthesia results from the development of an appropriate brain anaesthetic partial pressure. To achieve this brain partial pressure we administer a higher pressure from the anaesthetic machine. A difference exists between machine and anaesthetic circuit, between circuit and lungs, between lungs and arterial blood and finally between blood and body tissues.

These can be considered in a logical sequence of four interrelated steps:

- 1) The production and delivery of a suitable concentration of anaesthetic drug for inhalation.
- 2) The factors influencing distribution of the agent to the lungs.
- 3) Uptake from lungs.
- 4) Delivery of anaesthetic agent from circulation to brain.(Fig.1)
(Wylie and Churchill-Davidson, 1979)

UPTAKE OF ANAESTHETICS FROM THE LUNGS

Uptake of anaesthetics from the lungs is product of three factors:

- 1) Solubility of the anaesthetic in blood.
- 2) Cardiac output.
- 3) The difference between the partial pressure of the agent in venous blood and its partial pressure in alveolar gas.

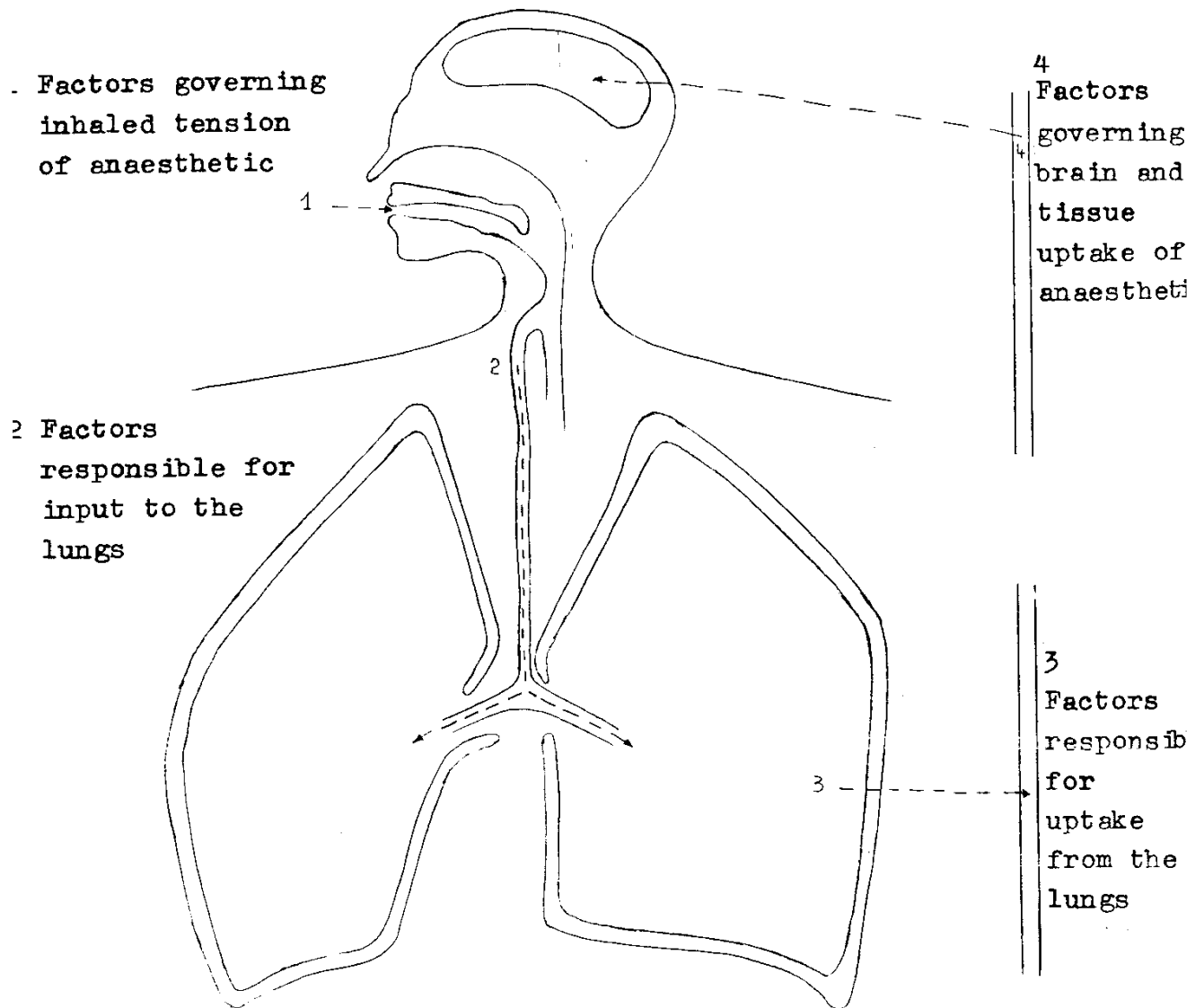


Fig. 1. Diagrammatic illustration of the four factors influencing the brain tension of volatile anaesthetic agents and gases.
(Reproduced from Wylie and Churchill-Davidson, 1979)

SOLUBILITY

Solubility is defined as the ratio of anaesthetic concentration existing in two phases when the two phases are in equilibrium. (Eger and Larson, 1964). The greater the blood / gas partition coefficient, the greater the uptake of anaesthetic and the slower the rate of increase in the alveolar concentration so a higher coefficients result in slow inductions. (Table 1).

Anaesthetic gas	Blood/Gas	Brain/Blood
Cyclopropane	0.4 - 0.6	0.76
Nitrous Oxide	0.47	1.1
Isoflurane	1.4	2.6
Enflurane	1.8	1.4
Halothane	2.3	2.3
Chloroform	8.4	1.4
Trichloroethylene	9	1.7
Diethyl ether	12	1.0
Methoxyflurane	12	1.7

Table 1. Partition coefficients of some anaesthetic gases at 37°C.
(Adapted from Eger, 1974)

Since solubility varies with temperature, the temperature at which the determination of solubility was made must also be stated. (Gregory and Eger, 1968).

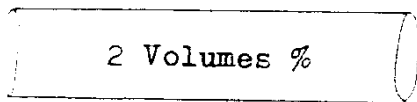
The Ostwald solubility coefficient, refers to a special partition coefficient. When one hundred per cent of a gas is equilibrated with a liquid or tissue, the volume of gas dissolved per unit volume of liquid or tissue is called the Ostwald solubility coefficient.

The Bunsen coefficient equals the Ostwald solubility coefficient times 273 divided by absolute temperature at which the Ostwald coefficient was measured. (Eger, 1974).

The importance of solubility in clinical practice is illustrated in the following example: If an anaesthetic is totally insoluble in blood (Blood/gas partition coefficient = 0) then none of it will be taken into the circulation consequently the alveolar concentration will rise. On the other hand if an anaesthetic has a low blood solubility, then only small quantities can be carried by the blood and both alveolar concentration and tension will rise rapidly. If the gas or vapour has a high solubility then large amounts can be absorbed by the blood stream, so that it is difficult for the alveolar concentration to rise rapidly. (Fig. 2).

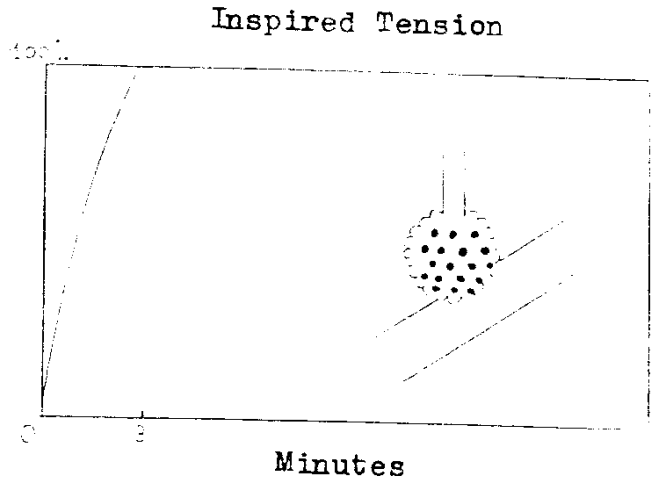
(Wylie and Churchill-Davidson, 1979)

1 Volume%

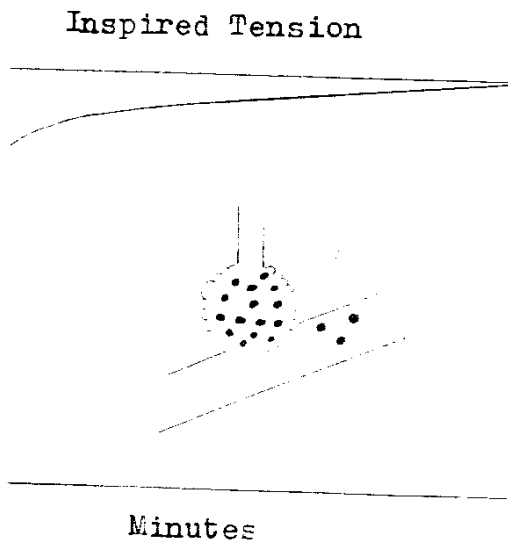


Blood	=	2
Gas		1

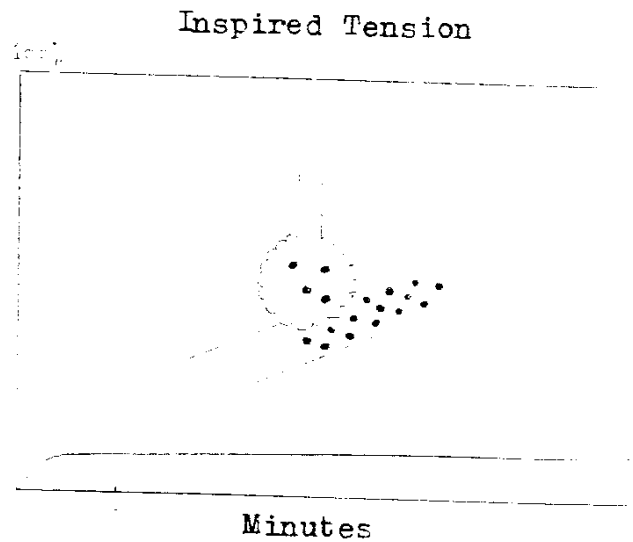
PARTITION COEFFICIENT = 2



Rate of increase in alveolar tension of a gas which is totally insoluble in blood.



Rate of increase in alveolar tension of gas which is poorly soluble in blood. (e.g. N_2O).



Rate of increase in alveolar tension of gas which is highly soluble in blood. (e.g. Ether).

Fig. 2.
Adapted from Wylie and Churchill, 1979)

CARDIAC OUTPUT

Because blood carries anaesthetic away from the lungs, the greater the cardiac output, the greater the uptake, and consequently the slower the rate of rise of alveolar level. (Yamamura, 1968). Thus induction of anaesthesia in a patient with a high cardiac output, as for example in extremely nervous or thyrotoxic patient, will take longer than usual when a soluble agent such as ether is used. In shock the reverse occurs and the unusually rapid rise in alveolar tension that results may lead to anaesthetics overdose, specially with the more soluble agents.

VENOUS ANAESTHETIC LEVELS

The last factor which determines uptake is the anaesthetic partial pressure difference between alveolus and returning venous blood. The larger this difference, the greater the uptake.

When all tissues are equilibrated with the alveolar anaesthetic partial pressure, there is no difference between anaesthetic partial pressure in the alveoli and in the returning venous blood, and uptake is zero regardless of the solubility or cardiac output. Complete equilibration occurs only after an infinitely long anaesthetic time. (Gray, Nunn and Utting, 1979)