

AIN SHAMS UNIVERSITY
FACULTY OF MEDICINE
1995

INFLUENCE OF ANAESTHETIC DRUGS ON THE FETUS

An Essay Submitted in Partial Fulfilment For
Master Degree in Anaesthesia

BY
Ahmed Aly Fawaz
(M.B.B.CH.)

A.A.

Supervised by

Prof.Dr Anissa Khamis Azmy
Prof.of Anaesthesia & I.C.U.
Ain Shams University

Assist.Prof.Dr. Mohamed Aly Ahmed Zaghloul
Assit.Prof. of Anaesthesia & I.C.U.
Ain Shams University

Dr. Alaa Eldin Abdel-Wahab Korraa
Lecturer of Anaesthesia & I.C.U.
Ain Shams University



AIN SHAMS UNIVERSITY
FACULTY OF MEDICINE
1995

INFLUENCE OF ANAESTHETIC DRUGS ON THE FETUS

An Essay Submitted in Partial Fulfilment For
Master Degree in Anaesthesia

BY
Ahmed Aly Fawaz
(M.B.B.CH.)

Supervised by

Prof.Dr Anissa Khamis Azmy
Prof.of Anaesthesia & I.C.U.
Ain Shams University

Assist.Prof.Dr. Mohamed Aly Ahmed Zaghloul
Assit.Prof. of Anaesthesia & I.C.U.
Ain Shams University

Dr. Alaa Eldin Abdel-Wahab Korraa
Lecturer of Anaesthesia & I.C.U.
Ain Shams University

بِسْمِ اللَّهِ الرَّحْمَنِ الرَّحِيمِ

TO
MY PARENTS

CONTENTS

	PAGE
1 - Acknowledgment	
2 - Introduction	1 - 2
3 - Maternal -Placental -Fetal Unit	3 - 17
4 - Placental transfer of Drugs .	18 - 28
5 - Effect o f Anaesthetic Drugs on The Fetus	29 - 62
6 - <i>Effect of Non-Anaesthetic Drugs on the Fetus</i>	63 - 70
7 - Fetus & neonatal Assessment & Resuscitation .	71 - 94
8 - Conclusion .	95 -102
9 - Summary .	
10 - References .	
11 - Arabic Summary.	

List of Figures

		Page
1	Fetal circulation A	9
	B	10
2	Effect of local anaesthetics on uteroplacental blood flow	53
3, 4, 5, 6	Fetal heart rate pattern	78
7	Scanlon Early Neonatal Neurobehavioral Assessment Scale	86
8	New neurological and adaptive capacity scoring system	87
9	Overview of steps taken in neonatal resuscitation	90

List of Tables

		Page
1	Molecular weight of several drugs used in anaesthesia	21
2	Protein binding capacity of amide-type local anaesthetics	50
3	Scoring system for evaluation the nonstress test	72
4	Biophysical profile scoring	75
5	Fetal heart rate pattern	81
6	Fetal scalp blood values	83
7	The APGAR scoring system	85
8	Normal term neonate blood gas data	88
9	Reported drug effects on the human fetus	96- 99

Acknowledgment

First , thanks are all to **God** for blessing me this work until it reached its end , as a little part of his generous help throughout life .

I would like to express my deepest gratitude as I am deeply indebted to my professor and supervisor Prof.Dr. **Anissa Khamis Azmy** , professor of anaesthesia , Ain Shams University , for her kind moral support , enthusiastic encouragement and help to pursue this effort .

Also , I am particularly very grateful to Assist.Prof.Dr. **Mohamed Aly Ahmed Zaghloul** , Assistant Professor of anaesthesia , Ain Shams University , for his great support , really, it is a great honour to work under his guidance and supervision .

I would also like to express my very sincere thanks to Dr. **Alaa Eldin Abdel-Wahab Korraa** , lecturer of anaesthesia , Ain Shams University , for his great support , patience and the tremendous effort he has done in the meticulous revision of the whole work .

Finally , I sincerely hope this work achieve something in our work in anaesthesia .

Ahamed Aly Fawaz

INTRODUCTION

INTRODUCTION

It is estimated that between 0.5 % and 2.0 % of pregnant women will at some time during pregnancy undergo anaesthetic or surgical procedures for incidental medical illness (*Brodsky et al , 1980*) .

There are many practical difficulties in studying anaesthetic drugs and techniques to determine what may be best for both the mother and the fetus . Physiological changes of pregnancy may alter the pharmacokinetics and pharmacodynamics of anaesthetics and the fetal disposition of drugs is largely unknown . The normal fetus appears able to withstand a variety of anaesthetic techniques , but there is little information regarding the compromised fetus or premature neonate (*Gin , 1993*) .

The past few years have seen the development of major new diagnostic technology for examination of the fetus in utero . Ultrasonography and fetoscopy in particular can now provide detailed information about the fetal condition . However , investigation of the effects of anaesthesia on the fetus is difficult for a number of reasons . The fetus is relatively inaccessible in utero and invasive studies in human are difficult to justify on ethical grounds (*Nandi et al , 1990*) .

The period of gestation is a time of multiple physiological changes , many of which may be disturbed by anaesthetic and surgical techniques (*Delaney , 1983*) . Maternally administered drugs may have drastic and far -reaching effects on the developing fetus . The fetus may be subject not only to the pharmacological effects of drug after it is transferred across the

placenta , but also to alterations in its inutero environment because of the physiological effects of the drug on the mother (*Peter et al. , 1986*) .

It is prudent to choose an anaesthetic technique which minimizes fetal exposure to the drugs and use agents which can be eliminated quickly by the neonate (*Gin , 1993*) .

MATERNAL-PLACENTAL -FETAL UNIT

MATERNAL -PLACENTAL- FETAL UNIT

Anatomy of the Maternal -Placental-Fetal Unit

The maternal-placental -fetal unit can be divided into three components :

1 - The maternal component .

2 - The placental component .

3 - The fetal component .

constant interaction between these components is required for fetal well-being .

The Maternal Component

Every maternal biologic system is dramatically altered during pregnancy
The most important for perinatal pharmacology are the alterations in the maternal cardiovascular system and uterine anatomy .

Cardiovascular and hemodynamic changes :

1 - Increase in total blood volume (25 - 40 %) due to increased plasma volume (40 - 50 %) and increased red cell mass (20 %) .

- 2 - Increase in cardiac output (30 - 50 %) due to increased heart rate (12 to 15 beats/ min.) , increased stroke volume (30 %) and decreased systemic vascular resistance (15 %) .
- 3 - Increase in uterine blood flow and a redistribution of uterine perfusion (*Osthiemer , 1992*) .

In the nonpregnant uterus , blood flow averages 50 ml / min. , increasing with pregnancy to 500 - 700 ml / min. . This change represents at least a 10 - fold increase in blood flow . At term , 10 to 20 % of maternal cardiac output is allotted to utero-placental circulation . The placenta receives more than 80 % of uterine perfusion . Whereas the myometrium receives only 20 % .

The cardiovascular and hemodynamic changes of pregnancy influence the pharmacologic characteristics of maternally administered drugs (*Osthiemer , 1992*) .

Changes in uterine anatomy :

The gravid uterus is an enlarging abdominal mass requiring an enlarging vascular supply . Arterial supply to the gravid uterus is provided by the uterine arteries , which arise from the hypogastric and ovarian arteries . The vessels in turn arise from the abdominal aorta . Anastomoses occur between uterine and ovarian arteries along the lateral border of the gravid uterus in the broad ligament . Uterine vessels then penetrate the muscular layer of the uterus , known as myometrium (*Osthiemer , 1992*) .