# THE TRANSMISSION OF MEPERIDINE ACROSS THE HUMAN PLACENTA\* |

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QUANTITATIVE data on the transmission of certain anesthetic and analgesic drugs from the maternal to the fetal circulation have been cited.<sup>1-3</sup> Such information is of value to the obstetrician and anesthesiologist for a better understanding of the effects upon the newborn baby of drugs administered to the mother during labor.

Hitherto, such knowledge was unavailable with regard to meperidine (Demerol). This drug has been used with increasing frequency for the relief of labor pain because it is said to produce little or no effect upon the respiration of the newborn baby when employed in this manner. Because of the important place meperidine has in the management of labor, the current studies of the pattern of its placental transfer from mother to fetus were undertaken.

#### Methods

Nine healthy pregnant women at term were used for this study. Two patients were delivered by cesarean section, the remainder spontaneously, with episiotomy. Meperidine in doses of 50 to 100 mg. was administered intramuscularly or intravenously during labor. The average total quantity of drug per patient during labor was 175 mg. and ranged from 50 to 300 mg. The last dose of meperidine varied from 40 minutes to 3 hours and 38 minutes prior to delivery. The first dose varied from 1 hour and 40 minutes to 15 hours before delivery. The method of drug administration was adjusted to the need of the patient for satisfactory analgesia.

Five patients were anesthetized for delivery with nitrous oxide-oxygen (a minimum concentration of 25 per cent oxygen was given); two with cyclo-

propane, and one each with low spinal anesthesia and caudal block.

At delivery, samples of blood were drawn simultaneously from the mother and the umbilical cord. The latter was considered representative of the fetal blood. Meperidine in plasma was determined according to a method to be published elsewhere. This method has been shown to be specific in that it included none of the metabolites of meperidine in the measurement.

#### Results

Plasma levels of meperidine indicate clearly that the drug crossed the placenta in significant concentrations (Table I). These varied from 45 to 106

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TABLE I

			_	TIME PRIOR				MATERNAL	CORD
			ROUTE OF	TO DELIVERY	ANALGESIC	DEPRESSANT		PLASMA	BLOOD
	TYPE OF	MEPERIDINE	ADMINIS-	(IN HR. AND	EFFECT ON	EFFECT ON		LEVEL	LEVEL
PATIENT	DELIVERY	(IN MG.)	TRATION	MIN.)	MOTHER	INFANT	ANESTHESIA	(IN MG./L.)	(IN MG./I
He	Vaginal	100	IM	2:00	Good	0	Cyclopropane	0.40	0.22
$Z_i$	Cesarean	50	$\mathbf{IM}$	1:43	Good	0	Cyclopropane	0.38	0.24
Po	Vaginal	50	IM	2:40	Good	0	Nitrous oxide	0.29	0.13
	Ŭ	50	IM	1:15					
$\mathbf{Fr}$	Vaginal	100	IM	1:40	$\mathbf{Good}$	Moderate depression	Nitrous oxide	0.45	0.38
Ni	Vaginal	75	IM	4:08	Poor	0	Nitrous oxide	0.50	0.39
	v agmai	75	ĬM	2:00	1 001	U	Tittous oxide	0.00	0.03
		50	īv	0:43					
St	Vaginal	100	ĨM	7:00	Good	0	Nitrous oxide	0.76	0.55
	· aginar	100	ĬM	4:00	dood	U	Title Ods Oxide	0.70	0.00
		100	īv	2:45					
Pa	Cesarean	100	ĬM	15:00	Good	0*	Low spinal	0.73	0.76
	00000	100	IM	9:00	Good	· ·	130W Spillar	0.70	0.10
		50	īv	0:40					
$\mathbf{B}\mathbf{u}$	Vaginal	100	ÎM	4:52	$\mathbf{Good}$	0†	Nitrous oxide	0.23	0.26
	,	100	IM	3:38	aooa	VI.	Titious onide	0.20	0.20
Al	Vaginal	100	ĬM	15:00	Good	0	Caudal	1.02	0.72
	. agmui	50	IM	11:00	Good	V	Outuui	******	0.12
		50	IM	6:40					
		50	ÎM	3:15					

\*Spontaneous breathing early. Aspirated meconium. No sequelae.
†Difficult to evaluate. Delivery was complicated by impaction of the shoulders of the fetus in the birth canal. A difficult delivery was followed by fetal anoxia requiring 20 minutes of intratracheal oxygen insufflation. Fetal recovery was satisfactory.

per cent (average 77 per cent) of the levels measured simultaneously in the maternal blood.

Satisfactory analgesia for the mother was achieved in all but one patient. In only one instance was there a clearly demonstrable depression of fetal respiration, presumably due to meperidine. Even so, the fetal plasma level of 0.38 mg. per liter was not the highest encountered. There were two other complications in the newborn. One was attributable to early intrauterine respiration and aspiration of meconium; the other to a difficult delivery caused by shoulder impaction of the fetus. Neither complication was harmful to the baby.

### Comment

With the aid of a sensitive method for its estimation, it was demonstrated that meperidine enters the fetal circulation relatively freely after parenteral administration to the mother. It also appears that doses of the drug which can produce satisfactory analgesia in the mother do not necessarily cause significant depression of respiration in the newborn baby. The rate of transplacental movement of the drug is unknown.

## Summary

- 1. Meperidine crosses the human placenta in significant concentrations.
- 2. In therapeutic doses administered to the mother, meperidine appears to cause little or no depression of respiration of the newborn child.

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