Placenta – a silent witness: clinical and forensic importance of placental examination

Marina Kos
Clinical Department of Pathology “Ljudevit Jurak” Clinical Hospital Center “Sestre milosrdnice”, Zagreb, Croatia
University of Zagreb Medical School, Zagreb, Croatia

Introduction

There is no doubt that obstetrics carries high medical liability risk. In many countries, gynecologists-obstetricians who attend childbirths and perform complex obstetric procedures are faced with increasing malpractice insurance premiums and litigation risk. The American College of Obstetricians and Gynecologists (ACOG) publishes its Survey of Professional Liability since 1983, with the objective to analyze the effect that malpractice litigation has had on the practice of obstetrics and gynecology in the United States. According to the 2003 ACOG survey, 76.3% of the members who answered the questionnaire have been involved in a lawsuit at least once in their professional career; gynecologists/obstetricians have been sued a total of 2.64 times per individual over the course of their careers. In the 2006 ACOG Survey, 89% of respondents indicated that they had been sued during their careers. The average number of claims per obstetrician was 2.6 (3). The ACOG’s 2009 Survey on Professional Liability showed that nearly 91% of gynecologists/obstetricians had experienced at least one liability claim filed against them during their professional careers, with an average of 2.69 claims per physician. In 2009, 62% percent of the total reported claims were for obstetric care as opposed to gynecology, the same as in the 2006 Survey. In the 2003 ACOG survey, fetal monitoring, neurologically impaired children, neonatal death, shoulder dystocia, uterine rupture, and “decision-to-incision” time were identified as clinical factors frequently present in obstetric malpractice cases. In both 2006 and 2009 Surveys the reasons for claim were neurologically impaired infant (in 31% of cases in both surveys), stillbirth/neonatal death (with 16 % of cases in both surveys), and delay or failure in diagnosis (in 11% of cases in 2009 vs. 14% in 2006).

In all the Surveys, neurological impair is the leading cause of the reasons for liability claim, with the cerebral palsy being the most serious damage. The possible etiologies have been discussed for years, and although the damage to neural tissue is undebatable, there is still no agreement upon the timing of the damage. Some authors think that 90% of the cases of cerebral palsy are not due to intrapartum events, while in the opinion of others most of the devastating events occurred in the perinatal period. It is still impossible to firmly determine in each single case whether the hypoxic insult has developed during delivery, in the first few hours after birth, or was already present before the labor began, as a consequence of long lasting hypoxia during pregnancy.

Careful gross and histopathological examination of the placenta in chosen cases can elucidate the events that occurred some time before labor, and help to connect and reconstruct the course of disease.

Importance of pathological examination of the placenta

Today, it is undisputable that all the samples of diagnostic value removed from the human body should be histologically examined, with only a few exceptions, one of them being the healthy human placenta. The placenta forms a functional unit between the mother and the fetus and any pathological event that concerns one of the two will influence the normal function of the placenta, resulting sometimes in morphological gross and/or histological change(s). It is the most important fetal organ because it is responsible for exchange of all nutrients, oxygen, and fluid from mother to fetus and removal of fetal waste products. It has also been called the “diary of gestational life”. The placenta provides important information’s on the timing and
etiology of many adverse events, including neurologic injury, fetal distress, infections, growth restriction, demise and many other fetal conditions. It also reflects the intrauterine environment, helps in identification of unsuspected maternal disorders, such as lupus or maternal vascular disease, and primary placental disorders, such as maternal floor infarction or chronic villitis. Furthermore, the placenta, being a fetal organ, expresses the fetal genotype and thus may provide diagnostic information on various genetic, chromosomal, congenital metabolic, or hematologic disorders.10

Severe abnormalities of the placenta may lead to adverse fetal outcome. Fortunately, the vast majority of pregnancies and newborns are normal, so only a subset of placentas requires submission to the pathology department for gross and histological examination. However, the clinical indications for placental examination have no gold standards. Some organizations have offered more or less similar guidelines, but the choice whether the placenta will be sent for histopathological examination is still left to the attending obstetrician.

**Indications for pathological examination of the placenta**

According to the guidelines issued by College of American Pathologists at their XIX Conference dedicated exclusively to the examination of the placenta and some other considerations, the main indications for placental examination are shown in Table 1.11,12

<table>
<thead>
<tr>
<th>Maternal conditions</th>
<th>Fetal conditions</th>
<th>Placental conditions</th>
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<tbody>
<tr>
<td>Systemic disorders (e.g. diabetes mellitus, hypertensive disorders, collagen vascular disease)</td>
<td>Stillbirth/neonatal death</td>
<td>Abnormalities of the placental shape</td>
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<td>Unexplained third-trimester bleeding</td>
<td>Multiple pregnancy</td>
<td>Retroplacental hematoma</td>
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<td>Severe oligohydramnios</td>
<td>Congenital malformations, dysmorphic phenotype or abnormal karyotype</td>
<td>Small or large placental size or weight for gestational age</td>
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<td>Peripartum fever and/or infection</td>
<td>Intrauterine growth restriction</td>
<td>Abnormalities of the umbilical cord (length, appearance)</td>
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<tr>
<td>Premature delivery</td>
<td>Prematurity</td>
<td>Abnormalities of the fetal membranes (e.g. amnion nodosum, meconium staining, etc)</td>
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<td>Drug addiction/alcohol abuse</td>
<td>Hydrops</td>
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<td>Invasive procedures with suspected placental injury</td>
<td>Meconium</td>
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<td>Placental abruption</td>
<td>Admittance to neonatal intensive care unit</td>
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<td></td>
<td>Apgar &lt;=3 in 5. minute</td>
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<tr>
<td></td>
<td>Neurological problems (seizures)</td>
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<td></td>
<td>Suspected infection</td>
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**Table 1.** The main indications for placental examination (modified from ref. 11, 12)
Others recommend that placentas from pregnancy complicated with cholestasis, hepatitis B, human immunodeficiency infection, other maternal diseases with normal pregnancy outcome, placenta previa and postpartum hemorrhage should not be sent for pathological examination in spite of moderate cost of this examination. These and other guidelines are only recommendations, and each institutions may accept all or only some of them, to meet the needs of the population they serve.

Even with valid indications the human placenta is one of the most under examined specimens. However, all placentas should be examined grossly, immediately after delivery, by the attending physician or a nurse. To conclude whether the placenta is normal, or should be sent for pathological examination, the person examining the placenta should have at least a basic knowledge of placental anatomy and pathology. The description of the placental shape, the color of the fetal membranes, the insertion and length of the umbilical cord and placental weight must be recorded in the clinical history. Even though the placenta should be left suspended on the umbilical cord for at least an hour after delivery, this practice is never followed, so the placental weight measured immediately after delivery is about 10 to 20% greater than the real weight because of the blood it contains. No matter the indications that are followed, the chosen placentas are sent to the pathology department.

Placentas submitted to pathological examination should be accompanied by a specimen requisition form containing clinical information. The importance of providing the clinical information cannot be overemphasized, because the absence of clinical informations prevents the appropriate evaluation and hampers the conclusions that are drawn from it. The informations that must be included are gravidity and parity, obstetric history, obstetric estimate of gestational age, route of delivery, fetal birth weight, gender, Apgar scores, maternal and fetal complications of pregnancy, labor, delivery, and total umbilical cord length. Some institutions even have a dedicated specimen requisition form for the placenta that facilitates the provision of these informations to the pathologist. The clinician should also state the indication(s) for which the placenta is being submitted.

The obstetricians should bear in mind that in case bacterial or/and viral cultures, cytogenetic and metabolic studies are needed the samples must be taken from the fresh placental tissue and in sterile conditions, immediately after delivery, in the delivery room. Depending on a pathologist that is going to examine the placenta, it can be submitted fresh, without fixative, or in the appropriate amount (10 times placental volume) of fixative, usually 10% buffered formalin. If there is no possibility of the pathological examination of the placenta in the near future, or a delay is anticipated between delivery and receipt at the pathology department the placenta can be stored in a refrigerator at 4°C for a week. It would be ideal, but is practically impossible in most institutions, for all placentas not requiring pathological examination to be stored in a refrigerator for a week in case the clinical status of the newborn changes, so that the placenta can still be pathologically examined. The placenta should never be deep frozen, because the ice crystals distort the villi, and the fine pathological changes are impossible to appreciate on histological examination.

Pathological evaluation and reporting

The fact of life is that in most pathological departments, other biopsies have precedence over placentas, so the obstetrician often receives the report on a placenta several weeks after submission. The report usually contains only placental measures and gross description of the placenta and the umbilical cord (frequently not even that) and the description of the histological appearance is „immature placenta“ or „mature placenta“, or „placenta without pathological changes“. After reading such reports, many obstetricians that are not convinced in the usefulness of the placental pathological examination in the first place, are discouraged to continue seeking pathological consultation. Placental lesions associated with adverse perinatal outcome can be roughly divided into those with abnormal blood flow in the maternal circulation, abnormal blood flow in the fetal circulation, inflammatory processes, and primary placental lesions. Each of these categories is associated with identifiable pathologic lesions. Intrauterine demise or neurologic injury can occur by sudden and possibly devastating events, by chronic processes that lead to decreased placental and fetal reserves, or by a combination of both. It is helpful to distinguish between pathologic placental lesions that result in acute and chronic intrauterine compromise (Table 2.)
Acute
Normal placental weight or weight appropriate for fetal weight
Acute villous edema
Intravillous hemorrhage
Acute retroplacental hemorrhage
Acute meconium staining

Chronic
Abnormal placental weight in relation to fetal weight
Chorangiosis
Fetal normoblastemia
Chronic meconium staining
Meconium associated myonecrosis of cord vessel(s)
Acute or necrotising funisitis
Significant chronic villitis
Amnion nodosum
Significant placental ischemia or infarction
Decidual vasculopathy
Maternal floor infarction/massive perivillous fibrinoid deposition
Fetal thrombotic vasculopathy

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<th>Table 2. Placental findings indicating acute and chronic in utero compromise</th>
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The quality of reports on the investigation of the placenta also varies greatly from institution to institution. One study showed that general surgical pathologists have a higher rate of under diagnosis of placental lesions compared to pediatric pathologists. These findings underline the fact that the reporting pathologist should be adequately trained and experienced in placental pathology. Templates and checklists for the reporting of placentas might help to improve the completeness and uniformity of reporting. The pathological report on placenta is usually sent only to the attending obstetrician, while the neonatologists or pediatricians caring for a newborn are not aware of the findings. In the opinion of the author, the copy of the placental pathology report should be filed in both the mother’s and newborn’s clinical history.

Discussion

Of the ACOG Survey respondents who reported making changes to their obstetric practice as a result of the risk or fear of professional liability claims or litigation, 30% decreased the number of high-risk obstetric patients that they accepted. Performing more cesarean sections was reported by 29% of respondents that changed their obstetric practices, and 25.9% stopped offering/performing vaginal births after cesarean (VBACs). An additional 13.9% decreased the number of total deliveries. About 8% of survey respondents reported that they had stopped practicing obstetrics altogether.

Other studies addressing this problem showed that malpractice claims led to a small reduction in physician delivery volume, but they did not have a significant impact on cesarean section rates. Litigation risk
and high malpractice premiums certainly affect negatively the gynecologist-obstetricians’ career satisfac-
tion. The result of a relatively recent study revealed that 43.7% of gynecologist-obstetricians had become le-
ss satisfied over the last 5 years and 34.0% would not recommend obstetrics/gynecology to students seeking
career advice\textsuperscript{20}.

Gynecologists_obstetricians are not the only doctors involved in delivery that are sued very frequently.
Anesthesiologists, and pediatricians/neonatologists are also often accused of malpractice in cases of unfavor-
able outcome of pregnancy. Before 1990 maternal death and newborn death/brain damage were the most
common complications in obstetric anesthesia malpractice claims. Newborn death/brain damage has decrea-
sed, yet it still remains also a leading cause of obstetric anesthesia malpractice claims\textsuperscript{21}.

The basis of litigation claims against obstetricians, anesthesiologists and neonatologists is the notion that
fetal death or neurological disabilities are the result of failure or delay in intervention or inappropriate ma-
agement of injuries believed to have occurred during the process of delivery. The intense fetal monitoring
and changes in methods of delivery have decreased the incidence of cerebral palsy, but not substantially\textsuperscript{22-24}.\textsuperscript{25} One of the reasons for this is that most of the fetal brain injuries occur before hospital admission and the be-
ginning of labor, many of them being the result of intrauterine infection and inflammation, or reduced or in-
terrupted placental vascular perfusion\textsuperscript{25}). The majority of cases of cerebral palsy, particularly in term infants,
are now considered to be due to ante partum events\textsuperscript{26,27}.

In the context of stillbirth or neurological impairment, examination of the placenta may be helpful in se-
veral ways. Firstly, the placenta itself may be abnormal and contribute directly to the adverse outcome, (e.g.
when there is a tight umbilical cord knob, maternal floor infarction or a large chorangioma). The category of
primary placental lesions also contains massive perivillous fibrinoid deposition, decidual vasculopathy lead-
ing to placental ischemia and/or infarction. Placenta itself may sometimes function normally, but the patho-
logic findings reflect abnormal intrauterine environment (e.g. intervillus thrombosis in a pale and hydropic
placenta whose villous capillaries contain nucleated fetal red blood cells that reflects fetomaternal hemorrha-
ge). Some findings (such as chorangiosis) reflect an adaptation to adverse intrauterine conditions (hypoxia).
The adverse outcome may be due to pathologic processes that are not placental in origin but that lead to ab-
normal placental function such as maternal under perfusion and fetal thrombotic vasculopathy. Many placen-
tal lesions develop during the prenatal period, long before labor and delivery. They cannot be prevented even
by the best of obstetrical care, but they can be identified and documented by pathological examination of the
placenta\textsuperscript{28}. In case of adverse pregnancy outcome, the normal placental findings on pathological examina-
tion are also very important, because in such a case certain conditions may be ruled out and the attention sho-
uld be directed elsewhere to look for the cause of injury.

During legal proceedings, the pathologist functioning as an expert witness may be asked to specify a time
frame for a placental lesion. It is often impossible to provide answers accurate to days or hours, but the rou-
gh distinction and framework of placental pathologies that result in acute or chronic compromise can be used
for determining the timing of the fetal insult. Pathologic examination provides more information on chronic
than on acute events. However, many acute events can also be diagnosed or confirmed on placental examina-
tion. Acute lesions may be associated with sudden catastrophic events whereas chronic lesions develop over a
period of time leading to decreased placental reserves at a minimum. Markedly depleted reserves will render
the infant susceptible to stresses of labor and to more acute events and therefore may also be associated with
significant injury or death. In stillborn or neurologically impaired infants, multiple placental lesions are often
present. By the timing of all lesions found, a sequence of events can be reconstructed in the development of
an adverse intrauterine environment. The decrease of placental reserves and function is usually the result of
synergistic action of multiple lesions of different etiologies that involve different aspects of placental functi-
on (fetal or maternal blood circulation). Acute events also frequently occur in combination with chronic pro-
cesses\textsuperscript{29,31}. Multiple placental lesions greatly increase the susceptibility of the fetus to neurologic damage\textsuperscript{29-31}.

The type and significance of placental lesions determine the extent to which placental pathology can be
helpful in understanding adverse antenatal and perinatal events. Interpretation of these lesions is complex and
requires experience and insight into clinicopathologic correlation with outcome. However, the most impor-
tant part of placental examination is ensuring that it is performed, because the slides and paraffin blocks of
placental tissue remain in the archives and can be retrieved even after a couple of years, should the need for revision or expertise arise. A photographic record of gross pathologic findings is also very useful. The gross and histological pathologic report of the placenta may not completely explain the etiology and timing but is an important and essential witness in understanding adverse pregnancy outcome. One also has to bear in mind that placental lesions are not necessarily the cause of unfavorable pregnancy outcome, and some structural changes may be the consequences of poor fetal condition. The placenta is an easily available specimen and the costs of a routine pathological examination are moderate, so in all doubtful cases, the clinicians should not hesitate to ask for a pathological analysis and opinion.

References

18. Gimm GW. The impact of malpractice liability claims on obstetrical practice patterns. Health Serv Res 2010; 45:195-211.


