A REPORT OF FOUR CASES OF MEMBRANOUS DYSENORRHEA.

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The term "membranous dysmenorrhea" is applied to cases in which considerable portions of the endometrium are thrown off repeatedly at the menstrual period, usually with pains simulating those of labor. The cases vary in severity from the typical form in which a complete cast of the uterine cavity is discharged at each period with great pain, to the milder ones in which small fragments of the endometrium are passed with only slight symptoms at intervals of several months. Painless cases have occasionally been reported. The passage of a single membrane from the uterus does not constitute a case of membranous dysmenorrhea.

The affection is not a disease sui generis; but a condition which develops under varying circumstances, complicates different pathological processes, and presents a variety of microscopic appearances. Some writers have, therefore, suggested that the purely clinical term "membranous dysmenorrhea" be abandoned, and "exfoliative endometritis" (Wyder) (9), or "exfoliation of the menstrual mucosa" (Löhein) (17) be substituted.

The etiology and pathogenesis of the condition are obscure, and the treatment has been attended with little success. This is due partly to the fact that the affection is in reality rare, and the specimens available for study still more infrequent. The milder cases are often not brought to the physician's attention, while, on the other hand, the diagnosis is frequently made on insufficient clinical data, and without a microscopic examination of the membrane. The latter point is illustrated by the fact that, of eleven specimens sent to the gynecological laboratory of the Johns Hopkins Hospital with the diagnosis of "membranous dysmenorrhea," only three proved, after investigation of the patients' history and examination of the membrane, to be genuine cases. The other specimens showed decidua in four instances, vaginal epithelium in two, and one example respectively of uterine polyp and blood-clot.

The disease was first recognized by Morgagni (1), who reported a case and gave an excellent description of its clinical course. Denman in 1791, also described membranes, but thought them identical with decidual casts, previously described by Hunter. The first microscopic study of menstrual membranes was made in 1842 by Ernst Heinrich Weber, and the term "membranous dysmenorrhea" was applied to the condition by Olsham and Simpson in 1846 (3). A certain resemblance to decidual tissue impressed observers more and more strongly, so that a prolonged discussion was carried on, especially in Germany, as to whether all cases of membranous dysmenorrhea were not in reality merely repeated early abortions. Only within the last thirty years have the two conditions been clearly differentiated. The first adequate histological study in the modern literature on the subject is that by Wyder (1878) (9). Von Franqué, in 1899 (28), reported five cases and made an elaborate study of the pathological anatomy. Since his article numerous isolated cases have been reported, but little new information has been added to the subject.

The most important etiological factor in the condition is a preceding endometritis, arising after child-birth, an abortion or a gonorrheal infection. In other instances there is a retroflexion or some abnormality of the appendages. A considerable number of cases, however, occur in young unmarried women, with no history of infection and with pelvic organs apparently normal on examination. There is in this group, of course, the possibility of an overlooked vaginitis in childhood, or an endometritis accompanying the exanthema.

The clinical history is usually somewhat as follows: Menstruation was regular and normal until after a labor or an abortion followed by fever, or after a gonorrheal infection. Dysmenorrhea then developed, followed in a few months by the appearance of membranes. These may be discharged only at intervals of three or four months. Usually, however, one is passed at each period; in exceptional cases two or more. Menstruation is accompanied by pain, and the expulsion of the membrane, which usually occurs on the third or fourth day, is preceded by intermittent uterine contractions, similar to those of labor. These cease after the discharge of the membrane, which is often followed by copious hemorrhage.

The prognosis as to recovery, either with or without treatment, is not good, as the condition usually persists until the menopause. Sterility is the rule, although a few patients recover and become pregnant.

The treatment most often adopted in cases where the underlying condition is obscure, is curettage a few days before menstruation, followed by the application of tincture of iodine or carbolic acid to the uterine cavity. This procedure frequently gives temporary relief, but the patients usually relapse within a few months. Any associated lesions of the uterus or appendages should of course receive appropriate treatment.

The menstrual membrane, when discharged entire, forms a triangular sac, having the shape of the uterine cavity, and sometimes showing rounded holes at the sites of the tubal openings. The outer surface (i. e., towards the uterine wall) is ragged; the inner is smooth. The thickness of the membrane varies from that of tissue paper to two or three millimeters. Membranes of greater thickness suggest decidua. Complete casts of the uterine cavity are rarer in membranous dysmenorrhea than in pregnancy, the tissue in the former condition being usually passed in fragments.

A variety of microscopic appearances have been described in this condition. The membranous discharges from the uterus may be divided, however, into two groups (excluding decidua for the present) : exfoliated mucoa and fibrinous casts.

In membranes composed of altered mucosa, two pictures
may be differentiated in a general way. The first is that of
an interstitial endometritis. The stroma cells are of normal
size and appearance, and there is an infiltration of lympho-
cytes. Hemorrhage, exudate, and fibrin are usually present
in addition. In the second type the stroma cells show a
strong resemblance to decidua. They are enlarged, oval or
polygonal in shape, with a large vesicular nucleus and abun-
dant protoplasm. All gradations may be traced between these
decidua-like forms and the normal stroma cells. The entire
membrane may be composed of these altered cells, or glands
may also be present. Occasionally a compact and a spongy
layer can be distinguished. The two pictures—interstitial
endometritis and decidua-like areas—are often seen in the
same membrane. It is these enlarged stroma cells which
have given rise to so much confusion and to a prolonged dis-
cussion as to whether there is any criterion by which they
may be differentiated from true decidual cells. They are
usually interpreted in menstrual membranes, as the result of
chronic hyperemia and irritation. They are not, however,
peculiar to this condition, being found also in glandular
hypertrophy and edema of the endometrium, where they are
accounted for by circulatory changes.

Most membranes show signs of degeneration, varying from
a hydropic appearance of the cells and inability to take ordi-
nary stains, to coagulation necrosis. A “budding” of the
interl glandular tissue has been described by Von Franché (23)
and also by Hegar and Maier; i.e., compact foci of stroma
cells in which growth is more active, and which penetrate
independently in different directions through the loose stroma.
These authors found them also in the hyperplasia of the
decidua associated with hydrometra gravidarium. They con-
sider it a change which may be found in any plastic inflam-
mation.

Amyloid change in the vessel walls has been found in mem-
branes from a case of prolonged and severe pelvic inflamma-
tion. Large numbers of eosinophiles and also mitoses in the
stroma cells have been reported by several observers.

Fibrinous casts are composed of a network of fibrin con-
taining in its meshes red corpuscles, leucocytes, and remnants
of the cells of the mucosa. There is some doubt as to whether
these casts should be classified with true cases of membranous
dysmenorrhea. Some authors exclude them entirely. They
develop, nevertheless, in connection with endometriotic
processes, and are passed with the same symptoms as the organ-
ized membranes. In fact, cases have been reported, in which
the same patient passed at one time a fibrinous cast, and at
another a membrane of altered mucosa. It is impossible, also,
to separate the two varieties anatomically, as many transitio-
nal forms are found between the simple fibrinous cast and the
well-preserved, exfoliated endometrium. Schönheimer
(23) considers the process to be an acute fibrinous inflamma-
tion occurring under the influence of menstrual congestion,
after a terminated endometritis, and subsiding at the end of
the period.

The mechanism of separation of the membrane is obscure.
The most generally accepted theory is that the hyperplasia
of the stroma cells causes an obstruction to the escape of blood
into the superficial layers; hence it spreads out in the deeper
portions of the mucosa. The tissue, friable because of chronic
hyperemia and its young connective-tissue cells, yields in its
weakest part, and the membrane is dissected off by hemor-
rhage. The free bleeding which frequently follows the expul-
sion of the cast is thought to confirm this theory. An ab-
normal density in the superficial layers of the endometrium
would also hinder the escape of the blood, and lead to the
same result. Speaking against this mechanical theory of
separation is the fact that the blood is often distributed
equally through all parts of the membrane.

The degenerative changes found in the greater number of
membranes must also be an important factor in causing separa-
tion,—possibly quite as important as they are in the separa-
tion of the decidua. Regressive processes in the decidua—
chiefly coagulation necrosis with the appearance of fibrin—
appear in the second half of pregnancy and are marked at
term. The membranes from our first case show widespread
coagulation necrosis in the stroma, and in the second case the
process is beginning. In both cases there are pronounced
changes in the blood-vessels,—many are occluded with fibrin
and their walls degenerated.

The diagnosis of membranous dysmenorrhea, although it
may be very probable from the clinical history, should not be
made without a microscopic examination, as there are two
other discharges from the genital tract, which may simulate
menstrual membranes macroscopically. These are vaginal
casts, or fragments of vaginal epithelium, and decidual casts.
The former are thrown off, either as the result of an exfolia-
tive vaginitis, or treatment of the vagina with strong chemi-
cals, such as silver nitrate. The tissue in exfoliative vaginitis
may be passed either during menstruation or independently
of it. If the pieces are passed with pain during the period,
the case may be considered one of membranous dysmenorrhea.
This was the case in two specimens sent to the laboratory,
with the diagnosis of “membranous dysmenorrhe,” and
proving, on microscopic examination to be of vaginal origin.
An exfoliative vaginitis may accompany a true membranous
dysmenorrhea. Leopold (11) reports a case of this kind, and
considers the cause of the two processes the same, i.e., a
superficial hemorrhage, arising from extreme hyperemia,
and extending through the cervix into the vagina. Hoggan (6)
describes a membrane, the upper part of which was composed
of uterine mucosa, the lower of vaginal epithelium. If small
portions of vaginal epithelium are passed together with endo-
metrium, the former may be overlooked, in case only the
larger pieces in the specimen are examined.

As a rule vaginal casts and pieces of vaginal tissue are
thinner, tougher, and more parchment-like than membranes
from the uterus, and no glandular openings are seen on the
surface. The diagnosis can be made immediately with the
microscope.

Decidual casts are expelled in abortion, extra-uterine preg-
nancy, and in the rare cases of pregnancy in one horn of a
double uterus. The typical decidual cast is larger, thicker,
and more vascular than the dysmenorrheic membrane. If chorionic villi are found on microscopic examination, the diagnosis of intra-uterine pregnancy is of course clear. If decidua alone is present, the only diagnosis that can be made from consideration merely of the cast, is that of the existence of pregnancy,—either in a normal uterus, a rudimentary horn, or a tube.

The greatest difficulty, however, has arisen over the differential diagnosis of an early abortion from a menstrual membrane containing large decidua-like cells. The question arises chiefly in cases of early pregnancy, before the decidua has reached its full development and typical form. Cells, which in size, form, nucleus, and staining properties closely resemble true decidual cells are found apart from pregnancy, not only in menstrual membranes, but also in edema of the endometrium, glandular hypertrophy, and inflammatory conditions of the mucosa. There may be no difference in size between these enlarged forms and decidual cells, as was proved by Von Fracqué (23) in a series of cases; but the former do not show the epithelioid appearance found so often in the mature decidual cell, which has more abundant protoplasm and a much more sharply-defined outline. The protoplasm of the decidual cell also loses its fibrillated appearance and takes a deeper eosin stain. Although the differential diagnosis of a menstrual membrane from typical decidua is usually clear on microscopic examination, there are many confusing cases, in which the question of early pregnancy cannot be excluded without the aid of the clinical history.

There are specimens from three cases of genuine membranous dysmenorrhea in the collection of the gynecological laboratory of the Johns Hopkins Hospital, and also one specimen of a fibrinous cast, which, from the patient’s history, should be included in this group.

The cases are as follows:

Case I.—L. R., age 20. Gynecological history No. 12,427. Sent by Dr. G. K. Vandervpke, Phoebeus, Va. Was admitted to the Johns Hopkins Hospital, October 11, 1905, and discharged October 20, 1905.

Complaint.—Pain in left side, painful menstruation and backache.

The family history was unimportant.

Past history and present illness.—Patient has always been well, aside from the present trouble. There is no history of severe infectious diseases.

Had a tumor removed from the right breast in 1904. Menstruation began at twelve years, is regular every four weeks, lasting seven days, and has been painful and profuse from the beginning. Ever since the onset, so far as patient can remember, she has passed with each period “pieces of flesh.” Dysmenorrhea is severe on the first day, less on the second, and on the third day the membrane is passed during a paroxysm of pain, lasting about one hour. This pain, which is always located a little to the left of the mid line, is sharp, and, as patient expresses it, “feels like raking over a raw surface.” It is relieved immediately by expulsion of the membrane, which is sometimes followed by considerable hemorrhage; at other times none.

Patient has also marked paroxysms of pain in the intermenstrual interval. These attacks last frequently but a few minutes, and may occur several times a day, or at intervals of several days. The pain is located in the same spot as during menstruation. Patient thinks that the pieces of membrane passed at present are smaller than formerly, and that she is gradually improving.

About a year before admission patient noticed that the uterus was situated low down, and since then she has been troubled with backache and leucorrhea.

Physical examination on entrance showed a well-nourished girl. The heart, lungs, and abdomen were normal.

Pelvic examination under ether gave the following: Outlet, vaginal; no signs of infection; cervix low, 1½ in. from the outlet; fundus in anteposition; freely movable; both ovaries easily felt and apparently normal.

A dilatation and curetting were performed, and a very small amount of endometrium, normal in appearance, was obtained (nineteen days after the last period). The specimen unfortunately was not saved.

Patient made an uneventful recovery, and was discharged ten days after admission. She sent to the laboratory some material passed during the period in November, which on microscopic examination showed nothing but blood clot.

The laboratory possesses three specimens from this patient, the first passed when she was fifteen years old, the second in August, 1902, at the age of seventeen, and the last in March, 1905. There is a certain similarity in the membranes, all of which show marked vascular, inflammatory and degenerative changes, and the first two also glandular hypertrophy.

The first membrane (gyn. path. No. 8315), has been imper-
stroma. The blood-vessels are numerous and comparatively large. Some are filled with well preserved red corpuscles; others with fibrin. Shadows of red cells are found throughout the tissue; also numerous leukocytes.

The membrane passed in August, 1902 (gyn. path. No. 8399).

FIG. 2.

FIG. 3.

FIG. 2.—Case I. Gyn. Path. 8399. Section through entire thickness of membrane, showing glandular dilatation, changes in blood vessels and degeneration in stroma.

FIG. 3.—Case I. Gyn. Path. 8400. Section through entire thickness of membrane, showing extensive necrosis and changes in blood vessels.

consists of a piece of tissue about 3 mm. thick, and approximately the size and shape of the uterine cavity. Microscopically it presents a striking picture, on account of the marked glandular hypertrophy and the extensive degenerative changes.

The surface epithelium is mostly lacking, being represented only by a few groups of cuboidal cells. The stroma immediately beneath the surface is thickened, and contains compressed and atrophied glands, which run parallel to the surface. In the deeper portions of the membrane are groups of tortuous and greatly dilated glands. They are lined with a single layer of low-cylindrical epithelium, and their lumina are filled with fibrin, red cells and leukocytes. In the neighborhood of these glands are numerous very large thin-walled vessels distended with blood. The stroma cells are somewhat enlarged. In a few areas near the large blood-vessels they stain clearly. For the most part, however, the cell outline and nucleus are indistinct and stain faintly and diffusely. Many of the cells are swollen. Some nuclei are shrunken and take an intense stain; others are undergoing fragmentation. The stroma cells are widely separated by fresh hemorrhage and exudate, and there is everywhere an abundant network of fibrin and a thick infiltration with polymorphonuclear leukocytes. The blood-vessels are numerous and large. Many are filled with fibrin and the walls of some of the smaller stain homogeneously with eosin, as if undergoing hyaline change.

The membrane expelled in March, 1905 (gyn. path. No. 8400), shows more advanced degenerative changes than the preceding. It is composed of thickened stroma, showing areas of coagulation necrosis, especially in the region of the blood-vessels. Clinging to the edges of the membrane are fragments of well preserved glands and stroma. The cells of the membrane are somewhat larger than normal. They all stain diffusely, are indistinct in outline, and show shrinking or fragmentation of their nuclei. The blood-vessels, which are numerous in the deeper portions of the tissue, show marked degeneration, their walls stain homogeneously with eosin, and their lumina are filled with fibrin. The entire membrane is traversed with a network of fibrin and thickly infiltrated with polymorphonuclear leukocytes.

CASE II.—The membrane from this case was sent to the laboratory February 7, 1905, by Dr. J. B. Beeson, of Livingston, Montana, with the following history: The patient, age 30, is a well-formed, healthy-appearing woman, married five years. Her menstrual history was normal until the appearance of the membranes. She had one or two abortions (probably artificially induced), during her early married life; for the past two or three years, she has not been pregnant. She began to pass membranes eight or nine months ago, and since then has discharged one at each period. Severe dysmenorrhea, confining her to bed for two or three days, developed coincidently with the appearance of the membranes. No pelvic examination was made and the patient has been lost sight of.

The specimen (gyn. path. No. 8239), consists of a few shreds of grayish-red tissue. Microscopically it is composed almost entirely of stroma. The surface epithelium is absent, and only an occasional remnant of a gland is seen. The stroma varies much in density; the cells in some areas being more closely set than normal; in others separated by exudate and their character changed. The rarified areas are, as a general rule, near the free surface of the membrane; the denser areas in the interior, where they occasionally surround the blood-vessels. The largest of the altered stroma cells have round, vesicular, faintly staining nuclei, two or three times the size of those of the usual stroma cells. The cell outline cannot be defined; but the protoplasm is small in amount and stains very faintly with eosin. Every gradation can be traced between them and the normal stroma cell. Numerous thin-walled vessels are present throughout the section, most of them occluded with fibrin. From them a delicate network of fibrin spreads out into the surrounding tissue. Indeed, the abundance of fibrin, both within and without the vessels is a striking feature. Scattered everywhere throughout the membrane are foci, in which the fibrinous network and the leukocyte infiltration are denser, and the stroma cells are indistinct and stain diffusely. In a few
places the process has advanced to actual necrosis. There is everywhere a marked infiltration with small round cells, especially about the thrombosed vessels.

Even in the absence of a pelvic examination, both the clinical history and microscopic appearance of the membrane point to an inflammatory origin in this case.

CASE III.—The following history is obtained from Dr. C. D. McLeod, of Chatfield, Ohio, who sent the specimen to the laboratory. The patient, age 38, was married at 16 years of age, and has four children. No history of puerperal infection. She has had marked dysmenorrhea for the past seven years, and during the last two or three years has passed shreds of membrane. She has also been subject to severe pain in the pelvis about ten days after menstruation. About twelve years ago she had some nervous trouble, diagnosed as "cerebro-spinal meningitis," which has left her with a partial paralysis of the left side.

gives the impression of having been separated in mass. The appearance is perhaps due partially to the fact that manual aid was given in the removal of the cast. The inner surface is smooth and the epithelium fairly well preserved. The surface towards the uterine wall is ragged and shows parts of glands, some of them abruptly torn across, projecting beyond the stroma. The glands are convoluted and dilated. Their epithelium shows no abnormality. The stroma is denser towards the surface; in the lower layers separated by exudate. The cells are considerably enlarged, oval or stellate in outline, and have a round, pale-staining nucleus and an increased amount of protoplasm. Many have a swollen and hydropic appearance. They are pushed apart by serous exudate and in some areas by a small amount of hemorrhage. Numerous lymphocytes are scattered everywhere throughout the membrane. The blood-vessels are abundant, and are filled with well-preserved red cells. The degenerative changes and fibrin, so conspicuous in the other membranes, are absent in this specimen.

CASE IV.—There is in the gynecological laboratory only one example of fibrinous cast of the uterus. This was sent by Dr. E. C. Soukert, of Chicago, with the following history: M. S., single, age 24. Born in Poland and came to America at the age of 18. Has worked in tailor-shops for the past five years.

Had diphtheria and scarlet fever at about eight years of age, with perfect recovery from both. She was very stout at the time of puberty. Menstruation began at thirteen and was irregular for several years. The flow has been slight, lasting for three days, and from the onset accompanied by dysmenorrhea. The pain has increased in severity, and on the third or fourth day of each period during the last three years she has passed stringy mucous and shreds of membrane. No leucorrhea.

Patient consulted the physician in January, 1905, for the increasing dysmenorrhea. She brought with her a complete fibrinous cast of the uterus,—the first one which had been expelled entire.

Physical examination at that time showed the patient to be well-nourished, and the chest and abdomen negative. The uterus was normal in size and position, and movable. Patient refused the treatment recommended, and it has been impossible to trace her subsequent history.

The specimen from the cast (gyn. path. No. 8470), shows mi-
crosscopically a heavy, loose-meshed network of fibrin, in which
remnants of cells are in places visible. There is a wide band of
fibrin on the surface; below that a layer of finer network; and
in the lowest stratum are numerous rounded openings, some of
which are artefacts, and others of which seem to have been
blood-vessels. The cells which are scattered throughout the net-
work are so degenerated that no idea of their nature can be
obtained.

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