

AN EARLY HUMAN OVUM (THOMSON) *IN SITU*

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IN March 1913 the late Prof. Arthur Thomson received the uterus, which contained this ovum, with its adnexa from Prof. H. M. Turnbull of the London Hospital. He briefly demonstrated it to the Anatomical Section of the XVIIth International Congress of Medicine, which was held in London in the summer of that same year, but he never published any detailed description of it. Some five years ago he suggested to me that it was well worth further examination, but it was not until after his death that the slides of this specimen with his serial microphotographs and notes about it were brought to my notice. I shall call it in this paper after his name, and I hope that, if it is ever referred to in the future, it may be known as the Thomson ovum.

Prof. Turnbull discovered it in the decidua of the posterior wall of the uterus of an unmarried woman, aged 20, who died 5 hours after severe burns on 4 March 1913. The right ovary contained a corpus luteum, which measured 24.5×17.5 mm. The woman's menstrual history was as follows. It first occurred at 15 years of age and had always been regular. Her last period commenced on 3 February, exactly 29 days before her death. No dates of coitus were available. The post-mortem examination took place 26 hours after death. The specimen was fixed in 10 per cent. formalin and cut into sections 10μ thick.

From the menstrual history the maximal age of this ovum must be about 16 days. The inside measurements of the blastocyst are $2.1 \times 1.51 \times 0.7$ mm.; the embryonic disc measures 0.26×0.31 (?) $\times 0.16$ mm. (?)

In the following list it is placed among its contemporaries. Dr S. Zuckerman has kindly computed for me from the inside measurements of the blastocyst cavity a rough estimate of its cubic capacity in the several ova, the cavities being regarded for this purpose as perfect spheres, of which the diameters were taken as the mean of the dimensions given. These figures are added in a separate column.

	Inside measurements in mm. of blastocyst cavity	Capacity in c.mm. of blastocyst cavity
(1) Linzenmeier (1914)	$0.61 \times 0.52 \times 0.75$	0.13
(2) Stieve (1931)	$0.44 \times 1.08 \times 0.42$	0.14
(3) Peters (1899)	$1.6 \times 0.9 \times 0.8$	0.70
(4) v. Möllendorff Ei OP (1921)	$1.5 \times 1.15 \times 1$	0.94
(5) Petzer (1910)	1.6×0.9	1.02
(6) Falkiner (1932)	No measurements: estimated age 15 days	
(7) Tennant & Ramsey (1934)	$2.17 \times 1.4 \times 0.6$	1.41
(8) Thomson (1913)	$2.1 \times 1.51 \times 0.7$	1.55
(9) Herzog (1909)	$2.32 \times 1.2 \times 0.8$	1.56
(10) Heine & Hofbauer (1911)	$2.38 \times 0.98 \times 1.2$	1.84
(11) Schlagenhauer & Verocay (1916)	$2 \times 1.6 \times 1$	1.89
(12) Jung (1907)	$2.5 \times 2.2 \times 1.1$	3.78

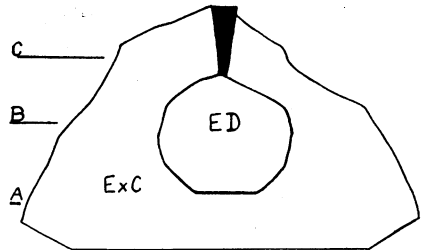
As is seen in the general view (Pl. I, fig. 1), the blastocyst cavity and the trophoblast are well preserved, but the embryonic disc, particularly the ectoderm, has suffered a good deal of disorganization, the amniotic cavity being partly obscured by cellular detritus. While this forbids any detailed description of the disc itself, I wish to draw attention here to:

- (1) the formation of the extra-embryonic coelom, and
- (2) the wall of the yolk sac.

I. THE EXTRA-EMBRYONIC COELOM

On either side of the amnio-embryonic vesicle a space is seen (Pl. I, fig. 2), the one on the left being triangular in section, while the right one appears more quadrangular in shape.

These are marked off from the rest of the blastocyst cavity by a regular layer of mesothelial cells, which vary in different sections in their number and their shape, most of them being round, while a minority are flatter and spindle-shaped. They are partially filled by coagulum similar to that of the magma. If the almost symmetrical channels thus formed are traced in one direction, they



Text-fig. 1.

are seen to coalesce and surround this end of the embryonic disc (Pl. II, fig. 1), the common channel extending for thirteen sections beyond the disc: traced in the opposite direction they are observed to narrow and finally disappear, the right before the left, sixteen sections from the disc, but here they are always separated from each other by a strand of mesenchyme and never join (Pl. II, fig. 4).

Text-fig. 1 is a diagram drawn roughly to scale to show the disposition of these channels (*ExC*) around the embryonic disc (*ED*) viewed from the dorsal aspect. The letters *A*, *B*, *C* correspond to the levels of the microphotographs (Pl. II, fig. 1; Pl. I, fig. 2; and Pl. II, fig. 4). While, as I have said, these channels are well marked out from the rest of the blastocyst cavity for the most part, there are in the case of either of them gaps in their parietes where they become continuous with the adjacent magma spaces. When traced in serial sections, such spaces are seen to be increasingly outlined by mesothelial cells so that eventually they appear as little bays in the lateral wall of the channel which is thereby correspondingly enlarged.

The only other specimen which shows anything exactly like this one is the Peters ovum, but that of Tennant & Ramsey presents a condition very comparable to it. The former of these was re-examined by Prof. O. Grosser in 1908, and he has been good enough to send me his microphotograph of this ovum and very kindly permits me to reproduce it here for comparison (Pl. II, fig. 2).

The only differences in the appearance presented by these two sections are

that in the Peters ovum (1) the lateral wall of the right channel has been torn away, (2) the containing wall of the intact left channel appears to be more cellular, and (3) it extends on to the ventral wall of the yolk sac, while in the Thomson ovum, although they are wider, these channels are confined to the neighbourhood of the amnio-embryonic vesicle. Grosser described the channel in the Peters ovum as horseshoe-shaped; it surrounds the cranial end of the disc as a small fissure. This is rather different from the wide communication presented by the present specimen.

Similarly, Tennant & Ramsey described "a space around the embryonic mass which is limited by a single layer of flattened mesodermal cells having the appearance of an endothelium (Pl. II, fig. 3). This layer not only surrounds the space but likewise completely encloses those surfaces of the embryonic mass not directly attached to the mesenchyme." In their specimen, too, the space is horseshoe-shaped, as must be inferred from the fact that they found that "the left end of the embryonic disc stops bluntly and is entirely free of mesothelial attachment while the right is completely encased in mesenchyme".

The three ova, Thomson, Peters and Tennant & Ramsey, seem to represent three stages in the development of these channels. In the first they border the amnio-embryonic vesicle only, in the second they have extended on to the lateral side of the yolk sac, while in the last they practically enclose the yolk sac on all sides. Here, although the authors describe a space, I think there is a fine strand in the microphotograph they reproduce between the mesothelial layer covering the ventral wall of the yolk sac and the parietes of the space to the right of the midline which may indicate its bilateral origin (Pl. II, fig. 3), while in their reconstruction they show the ventral surface of the embryo as almost entirely free of mesenchyme, "with the exception of a small band of attachment extending about half way along this surface from the encapsulated end".

In 1908 Keibel & Elze and Grosser described these channels as the commencement of the extra-embryonic coelom. Grosser (1924) suggested that they gradually grow ventrally round the yolk sac, their opposed walls possibly forming, when they meet, the "distaler Nabelblasenfaden", the mesodermal strand which may connect the ventral pole of the yolk sac with the chorion. Further, he thought that these exocoelomic channels extend lateralwards by taking up the magma spaces nearest to them so that eventually the whole cavity of the blastocyst becomes the extra-embryonic coelom. He wondered if the loss of the lateral wall in the left channel in the Peters embryo might not be due to its having already broken down so that on this side the exocoelom was continuous with the intercellular spaces of the magma. I have mentioned above that in the Thomson ovum there is evidence of the commencement of the same process, while Tennant & Ramsey wrote that in the specimen they described "on the surface towards the main chorionic cavity the continuity of this membrane-like structure is less well maintained and irregularly distributed openings in it offer probable avenues of communication between the larger and the smaller spaces".

These three ova are peculiar then in being the only ones so far described which exhibit definite symmetrical coelomic channels.¹ The Fetzner (Grosser, 1924) and Herzog (Lewis, 1912) ova and the much more advanced embryo described by Waterston (Mall, 1916) have all been considered by these different observers to show something similar, but I would suggest that at all events the illustrations, which accompany the descriptions of these specimens, are far from convincing from this point of view. In all other specimens the whole of the blastocyst cavity seems to represent the extra-embryonic coelom (Bryce, 1924-5). An accumulation of magma fluid appears according to this author's account near its centre close to the embryonic disc, which gradually leads to the stretching and absorption of the mesodermal bands traversing it and to the isolation of the embryo, so that ultimately all the magma spaces run together and the whole of the magma cavity becomes the exocoelom. Grosser (1913) and v. Möllendorff (1921) both suggested that there are thus two alternative ways in which this coelom may be formed.

It is, however, perhaps pertinent to observe that in the Peters ovum the trophoblast is already lined with "morula" mesoderm (Stieve, 1926), which also covers the ventral wall of the yolk sac beyond the limits of the exocoelomic channel. The Thomson ovum presents the same appearance, although the mesodermal cells covering the yolk sac are few. In either of these cases, apparently, a gradual liquefaction of the magma would result in a perfect coelom lined with mesoderm without the help of the mesothelium of the channels at all.

With regard to the later fate of these exocoelomic channels there is only Grosser's surmise to guide us, and one cannot help thinking that their formation as seen in the three specimens in which they occur serves no useful purpose. In any case their appearance in the Tennant-Ramsey and Thomson ova prove that the Peters ovum is no longer quite exceptional.

II. THE YOLK SAC

As seen in Pl. I, fig. 2, this appears as a space beneath the amnio-embryonic disc, outlined from the rest of the magma cavity by a thin membranous wall. It is flattened dorsoventrally. On the left side, its ventral wall is connected by an irregular strand of mesenchyme cells to the chorion, but this is the only evidence of a commencing "Nabelblasenfaden" so obvious in the Schlagenhauser-Verocay specimen. While the dorsal wall of the yolk sac beneath the embryonic disc has a lining of cubical entodermal epithelium its lateral and ventral walls are formed by a sparse layer of flattened spindle-shaped cells mesothelial in character. These latter appear to be similar to many of the cells lining the extra-embryonic coelomic channels, and the coagulum within the yolk sac seems to be exactly like that filling these

¹ Since this was written K. Hiramatsu (*Fol. Anat. Japon.*, 1936, Bd. XIV, p. 15) has published an account of the ovum, Ei-Ando. In this he describes two small cavities on either side of the dorsal portion of the yolk sac, which he thinks correspond to these exocoelomic channels.

channels and the magma spaces. Clothing the lateral walls and extending on to the ventral angles of the sac are a few round mesodermal cells. It is interesting to compare the condition found here with that noted or illustrated in the other ova in the list given above.

Linzenmeier. There is no mention of this point in the text, but the accompanying illustration shows flattened cells on the ventral side of the yolk sac with a cubical entoderm under the embryonic disc.

Stieve. "The yolk sac is outlined by spindle-shaped mesoderm cells, while the entoderm is only represented by a single layer beneath the ectoderm of the disc."

Peters. Here the whole sac is lined by a cubical epithelium of the same type all round (Pl. II, fig. 2). In Peters' original drawing the cells on the ventral wall are more irregularly spaced than are those beneath the disc.

v. Möllendorff, Ei OP. "In its whole circumference the yolk sac is lined with markedly flattened cells with lens-shaped nuclei." His figure shows a condition very much like that I have described in the Thomson ovum.

Fetzer. The sac is lined except beneath the disc with a "thin low entoderm". Sporadically the cells are larger, cubical or short cylindrical in type (Fetzer & Florian).

Falkiner. He gives no description of this point, but his illustration shows the lining of the yolk sac very much like that in the Thomson ovum.

Tennant & Ramsey. "The yolk sac is lined with entodermal cells, which are cuboidal on the dorsal surface and flattened on the ventral surface" (Pl. II, fig. 3).

Herzog. Lewis (1912) described the sac in this ovum thus: "Over the greater part of it the entoderm forms a very thin layer resembling endothelium."

Heine & Hofbauer. The yolk sac is lined by a single layer of flattened cells except under the disc.

Schlagenhauser & Verocay. The yolk sac is lined throughout with entoderm.

Jung. The yolk sac shows a simple layer of flat entoderm cells.

With the Peters and Schagenhauser ova standing out as exceptions this analysis confirms Bryce's (1924-5) statement that in the first instance the yolk sac is always lined with flattened endothelial-like cells. In older embryos, as Grosser (1924) pointed out, there is a complete change; in these (e.g. Rossenbeck, Grosser and Graf Spee ova) while the peripheral part of the sac is lined with cuboidal cells the portion of it beneath the disc has the flatter type of epithelium. Similarly Florian & Hill (1934-5) described the yolk sac wall in their specimen—"while the endoderm of its cranial and lateral walls is formed of flattened cells, that of its ventral wall is distinctly higher."

In attempting to explain these differences Stieve interpreted the appearance presented by the ovum (Werner) he described as follows. The future yolk sac is here represented by an enlarged magma space lying beneath the embryonic disc and outlined by mesoderm cells. Later the entoderm of the disc must gradually grow ventrally and surround this space on its inner aspect and so form a definite yolk sac. He followed Corning (1925), who described the entoderm in the human ovum as beginning as an entodermal plate—the lowest

layer of cells of the embryonic disc—rather than as an entodermal vesicle. In favour of such a conception are, I think, the following points:

(1) It brings the formation of the human yolk sac in line with that which has been found to occur in most other mammals (Stieve). The only observations in comparative embryology, as far as I am aware, which suggest the development of the yolk sac by the dehiscence or vacuolization of a primitively solid entodermal mass are those of Hubrecht (1889–90) in *Erinaceus* and Hill (1932) in *Nasalis larvatus*.

(2) The entodermal vesicle of the Teacher-Bryce ovum (TB. I.) lying in the magma and divorced from the amnio-embryonic vesicle might very well be described as a magma space (Stieve). Bryce (1924–5) himself described it as being lined with flattened cells precisely like those forming the wall of the sac in Linzenmeier's embryo.

(3) In the Tennant-Ramsey ovum the cubical entoderm appears to be surrounding the yolk sac and to have left only a portion of its ventral wall lined by flattened cells (Pl. II, fig. 3).

(4) In nearly all the early ova the yolk sac, lined with flat mesothelial-like cells, is filled with a loose coagulum which seems to be indistinguishable from that of the magma spaces (Triepel, 1916). On the other hand Schlagenhauser & Verocay found in their ovum that the contents of the yolk sac stained more deeply with eosin than did that of the magma. Tennant & Ramsey described the yolk sac in their specimen as "partially filled by pink-staining finely granular material, while in other sections large yolk granules are present in the cavity". Peters figured some drops of (?) yolk-like material in his original drawing of the yolk sac in his ovum. In all these three instances, as has been noted already, the sac is lined more or less completely with a cubical entoderm. One might, therefore, suggest that there is some correlation between the character of the contents of the yolk sac and of its cellular lining.

The appearance of the yolk sac in the Thomson ovum is very much like that shown in Stieve's illustration. On Prof. J. P. Hill's advice I ventured to send the serial microphotographs of this ovum to Prof. Grosser, and he was kind enough to give me his opinion on this point after examining them. His criticism of both the Thomson and the Stieve specimen is the same. In both, according to him, the entodermal lining of the yolk sac has disappeared except under the disc. I confess I think this explanation is a difficult one. Allowing that 26 hours elapsed between the woman's death and the fixation of the Thomson ovum, the entoderm could scarcely have vanished in this time without leaving a trace. If the cells had been shed, they or their debris should surely still be found in the yolk sac, unless its walls were broken. But the yolk sac appears to be intact, filled with coagulum, but containing only an occasional detached cell. If there had been a dehiscence in its wall, the fluid content would certainly have escaped before its cellular lining. Further, as I have shown above, in most of the early ova the cells lining the yolk sac are described as being of this same character, and, while in different specimens they vary

somewhat both quantitatively and qualitatively, most embryologists would follow Bryce in describing them as flattened and endothelial-like. But, if one accepts Grosser's explanation for the Stieve and Thomson ova, is one to condemn others of about the same age which show a similar condition? One is left apparently with two alternatives. There is, on the one hand, the view generally held that these mesothelial-like cells lining the early yolk sac, so similar to many of those outlining the exocoelomic channels and to those scattered elsewhere throughout the blastocyst cavity, grow eventually into the cubical or low columnar entoderm of the older sacs. On the other hand, there is the conception of Corning and Stieve of a yolk sac originally preformed as a magma space and outlined in mesoderm, which later acquires an epithelial lining from the growing margins of an entodermal plate. This latter idea apparently has much to recommend it and I should prefer myself to interpret this specimen as rather corroborating it.

While I realize to the full the weight that rightly should be attached to Prof. Grosser's opinion, I would suggest that the appearance of the cellular lining of the yolk sac in this ovum is not so remarkable.

SUMMARY

1. The interior measurements of the blastocyst cavity in the Thomson ovum are $2.1 \times 1.51 \times 0.7$ mm.; its maximal age is about 16 days.

2. The extra-embryonic coelom is here seen to be commencing as fairly symmetrical channels on either side of the amnio-embryonic vesicle. A comparable appearance to this has been observed previously only in the Peters and Tennant-Ramsey ova.

3. The yolk sac has a lining of cubical entoderm beneath the disc; elsewhere it is merely outlined by a sparse layer of spindle-shaped mesothelial-like cells. This condition is shown not to be exceptional in the earlier ova and is held to corroborate Stieve's conception of an entodermal plate spreading ventrally to form the proper entodermal wall of the yolk sac as seen in later embryos.

I am very much indebted to Prof. J. P. Hill for his kindness in examining the slides of this specimen, and to Prof. O. Grosser for giving me his opinion on the serial microphotographs of it, and for his kind permission to reproduce his own photograph of the Peters ovum. I must also thank the publishers of *Surgery, Gynecology and Obstetrics* for allowing me to reproduce the illustration, Pl. II, fig. 3, which appeared originally in that journal. The other microphotographs are the work of Mr W. Chesterman of this Department.

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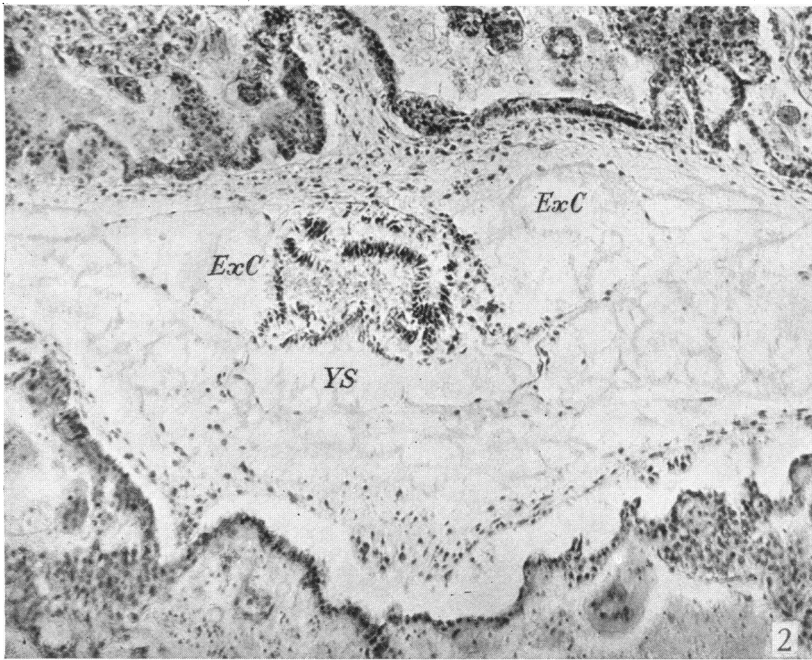
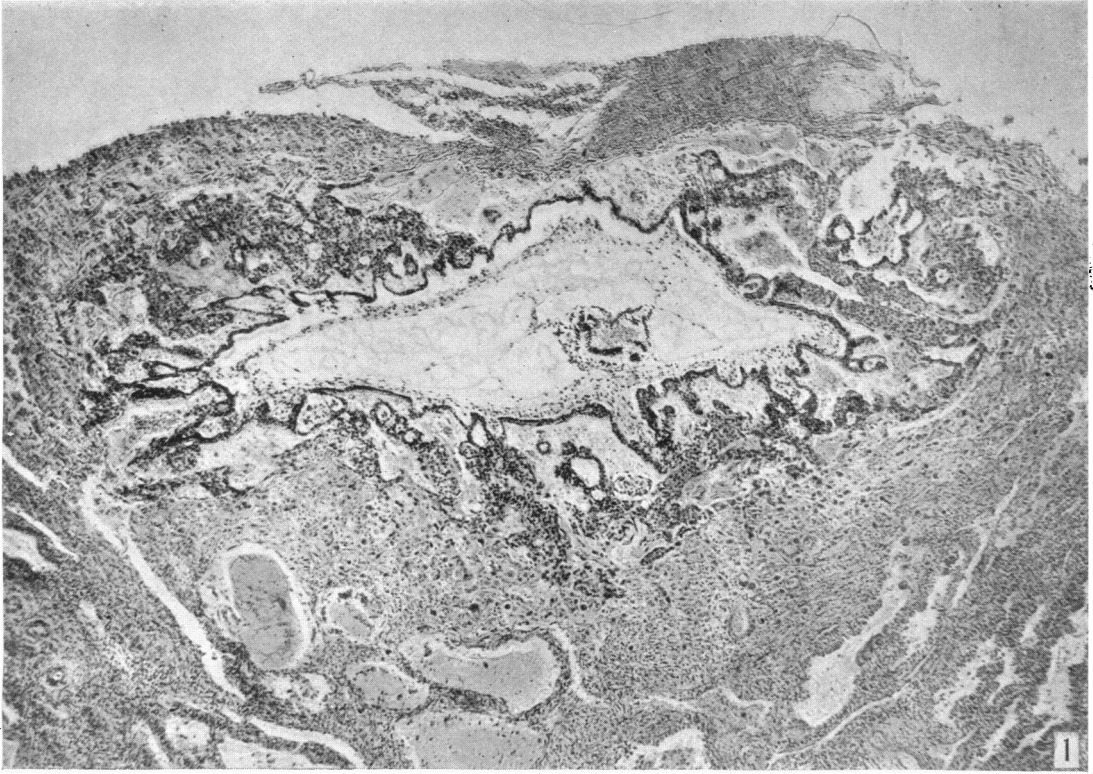
EXPLANATION OF PLATES

PLATE I

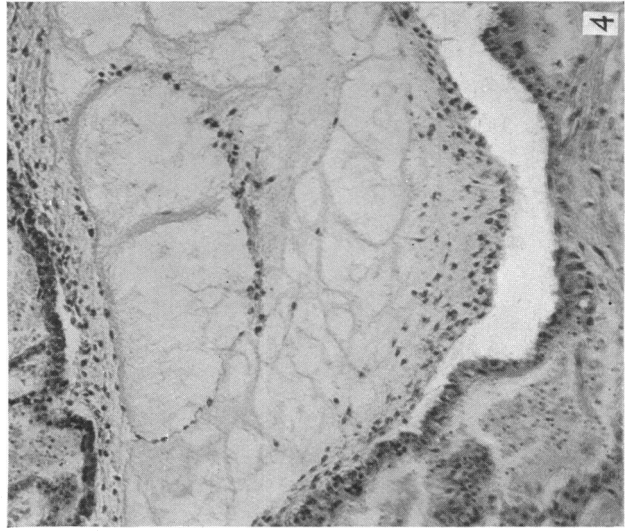
- Fig. 1. The section photographed is through the middle of the blastocyst cavity ($\times 44$).
- Fig. 2. The same section as fig. 1 ($\times 100$). This shows the exocoelomic channels, *ExC*, on either side of the embryonic disc and ventral to this the yolk sac, *YS*. Above the disc is the amniotic cavity obscured by cellular debris which partially fills it. This section corresponds to the plane lettered *B* in Text-fig. 1.

PLATE II

- Fig. 1. A section ($\times 100$) 20μ beyond one extremity of the embryonic disc. It corresponds to the plane *A* in Text-fig. 1 and shows that the two exocoelomic channels have joined round this end of the disc.
- Fig. 4. A section ($\times 100$) 20μ beyond the opposite extremity of the disc showing the two channels still separate from each other. This corresponds to the plane *C* in Text-fig. 1.
- Fig. 2. A photograph of a section ($\times 100$) of the Peters ovum, kindly sent me by Prof. O. Grosser. It shows the exocoelomic channel on the left intact, bordering the embryonic disc and the yolk sac, while that on the right has only its medial wall left.
- Fig. 3. A section ($\times 150$?) of the Tennant-Ramsey ovum (reproduced by courtesy of the Editors of *Surgery, Gynecology and Obstetrics*). It shows the amniotic cavity, the embryonic disc and the yolk sac surrounded on either side and ventrally by the extra-embryonic coelom.



ODGERS—AN EARLY HUMAN OVUM (THOMSON) *IN SITU*.



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