

Severe fetal cardiomegaly caused by multiple non-giant placental chorioangiomas

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21 6 **Running head**
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24 7 Cardiomegaly by non-giant placental chorioangiomas
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For Peer Review

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24 **Abstract**

25 A pregnant woman was referred to our hospital due to fetal cardiomegaly. We
26 recognized a dilated umbilical vein, which raised a suspicion of placental chorioangioma.
27 A male neonate was delivered at 37 weeks of gestation. The cardiomegaly was gradually
28 improved. Pathological examination identified five non-giant placental chorioangiomas.
29 Multiple non-giant chorioangiomas may cause fetal complications despite the difficulty
30 of prenatal diagnosis.

31 Main text**32 Introduction**

33 Causes of fetal cardiomegaly include not only fetal cardiac disease, but also
34 extra-cardiac shunt disease.¹ Among these, abnormal fetal circulations such as placental
35 chorioangioma are greatly different from other diseases in terms of the requisite clinical
36 management, because delivery may reduce the cardiac burden. We report a case in which
37 fetal cardiomegaly due to multiple non-giant placental chorioangiomas was unable to be
38 diagnosed prenatally, but was strongly suspected from the dilated umbilical vein.

40 Case

41 A 39-year-old pregnant woman with no past or family history was referred to
42 our hospital due to fetal cardiomegaly at 31 weeks of gestation. On presentation, fetal
43 echocardiography showed a cardiothoracic area ratio of 0.55 (Fig.1a), but neither
44 structural abnormality nor prominent valve regurgitation were detected, and contractility
45 of both ventricles was maintained. Neither fetal echography nor magnetic resonance
46 imaging revealed no extra-cardiac shunt disease. We recognized a thickly expanded
47 umbilical cord and dilated umbilical vein and artery (Fig.1b, c), which raised a suspicion
48 of shunt diseases in the placenta such as placental chorioangioma. However, we were

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6 49 unable to reach a definitive diagnosis. The flow pattern of umbilical artery was normal
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9 50 (Fig.1d). However, the flow **pattern** of umbilical vein was pulsatile (Fig.1e), and that of
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12 51 inferior vena cava had retrograde component (Fig.1f), so it has been suggested cardiac
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15 52 function declines. We continued careful monitoring for the occurrence of hydrops or fetal
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18 53 heart failure under hospitalization. Up to the time of delivery, findings from fetal
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21 54 echography did not show any marked changes. No fetal arrhythmia was seen, and anti-
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24 55 SS-A and anti-SS-B antibodies were negative in maternal blood testing. Non-invasive
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27 56 prenatal testing showed no abnormalities.

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30 57 A male neonate was delivered by selective caesarean section at 37+1 weeks of
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33 58 gestation with a birth weight of 3,114 g without asphyxia. No external malformations
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36 59 were identified, but the umbilical cord was thick with a diameter of about 25 mm (Fig.2a).
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39 60 The cardiothoracic ratio on chest X-ray was 0.63. Echocardiography showed no structural
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42 61 abnormality, left ventricular ejection fraction was 50%, and left ventricular diameter at
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45 62 end diastole was 20 mm. The value of N-terminal pro-B-type natriuretic peptide was
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48 63 2,156 pg/mL. Hemoglobin level and platelet count were normal. We started continuous
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51 64 intravenous infusion of milrinone after birth, considering the possibility of increased
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54 65 afterload due to withdrawal from the fetal circulation. A few days later, when we tried to
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57 66 reduce milrinone, his cardiac function apparently deteriorated, so it gradually declined
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7 67 over two weeks. Although the value of N-terminal pro-B-type natriuretic peptide
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10 68 decreased to 143 pg/mL in two weeks, it rose to 236 pg/mL after stopping milrinone, so
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13 69 he was started to take pimobendan and discharged. Cardiothoracic ratio improved to 0.50.

14
15 70 Histopathological examination showed intimal thickening of vessels in the
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18 71 umbilical cord. Five chorioangiomas were identified in the placenta, with a maximum
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21 72 diameter of 3 cm (Fig.2b). We determined that these were the cause of fetal cardiomegaly.
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27 74 **Discussion**

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30 75 Placental chorioangioma is the most common placental tumor, found in 1% of
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33 76 all pregnancies. This disease is reported to cause fetal complications such as growth
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36 77 restriction, anemia, thrombocytopenia and hydrops, especially in cases with
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39 78 chorioangioma >4 cm in diameter, known as giant chorioangioma. However, most
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42 79 chorioangiomas are small and can be found in postpartum placental pathology
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45 80 examination without any complications ever presenting.^{2,3}

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48 81 What was noteworthy in our case was that multiple non-giant chorioangiomas
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51 82 may cause fetal complications despite the difficulty of prenatal diagnosis. While color
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54 83 Doppler echography is considered useful for diagnosing chorioangioma, prenatal
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57 84 diagnosis is sometimes difficult for small lesions.⁴ When fetal cardiomegaly and a dilated
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85 umbilical vein are identified, this disease should be suspected even if a definitive
86 diagnosis cannot be reached from echography.

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94 or not-for-profit sectors.

95
96 **Conflicts of Interest**

97 None.

98
99 **Ethical Standards**

100 The authors assert that all work reported complies with the ethical standards of
101 the Helsinki convention, and institutional and national research committee.

102
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113 **Figures**

114 **Figure 1.** Fetal echography findings

115 **a)** Cardiothoracic area ratio is 0.55 in fetal echocardiography.

116 **b, c)** The umbilical cord is thick, with an umbilical vein diameter of 12 mm (white arrow)
117 and an umbilical artery diameter of 5 mm (black arrow).

118 **d-f)** The flow pattern. (d): umbilical artery, (e): umbilical vein, (f): inferior vena cava.

119 Note the pulsatile pattern in the umbilical vein and the retrograde flow in the inferior vena
120 cava, which suggests the decreased cardiac function of the fetus.

121

122 **Figure 2.** Findings after birth

123 **a)** Diameter of the umbilical cord at birth is about 25 mm.

124 **b)** The largest placental chorioangioma is about 3 cm along the major axis.

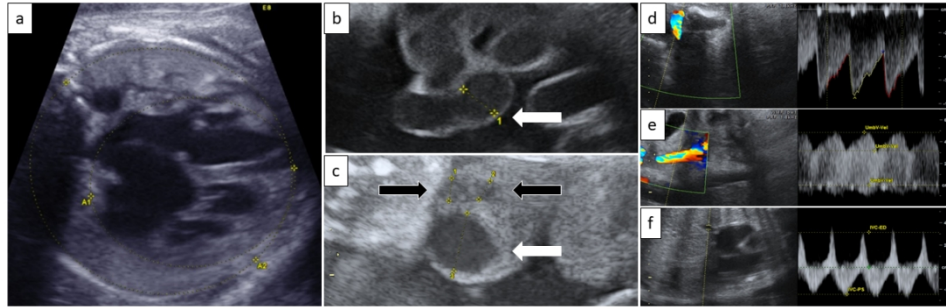


Figure 1

Fetal echography findings

a) Cardiothoracic area ratio is 0.55 in fetal echocardiography.

b, c) The umbilical cord is thick, with an umbilical vein diameter of 12 mm (white arrow) and an umbilical artery diameter of 5 mm (black arrow).

d-f) The flow pattern. (d): umbilical artery, (e): umbilical vein, (f): inferior vena cava. Note the pulsatile pattern in the umbilical vein and the retrograde flow in the inferior vena cava, which suggests the decreased cardiac function of the fetus.

338x190mm (96 x 96 DPI)

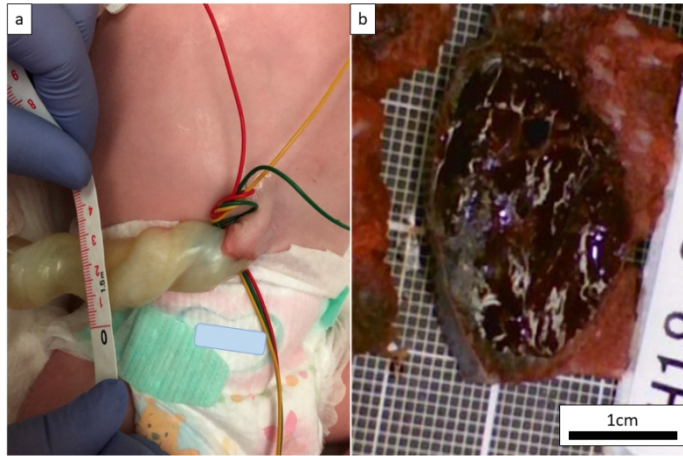


Figure 2

Findings after birth

a) Diameter of the umbilical cord at birth is about 25 mm.

b) The largest placental chorioangioma is about 3 cm along the major axis.

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