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**ABOUT COVER**

Editorial Board Member of *World Journal of Clinical Cases*, Luca Morelli, FACS, FASCRS, MD, Associate Professor, Division of General Surgery, Department of Translational Research and of New Surgical and Medical Technologies, University of Pisa, Pisa 56124, Italy. [luca.morelli@unipi.it](mailto:luca.morelli@unipi.it)

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## Prenatal ultrasonographic findings in Klippel-Trenaunay syndrome: A case report

Hou-Qing Pang, Qian-Qian Gao

**ORCID number:** Hou-Qing Pang  
0000-0003-1707-9600; Qian-Qian Gao  
0000-0002-7464-6635.

**Author contributions:** Pang HQ carried out the studies, collected data and drafted the manuscript; Gao QQ and Pang HQ participated in acquisition, analysis, or interpretation of data and revised the manuscript; all authors read and approved the final manuscript.

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**Hou-Qing Pang, Qian-Qian Gao**, Department of Ultrasound, West China Second University Hospital, Sichuan University, Chengdu 610041, Sichuan Province, China

**Hou-Qing Pang, Qian-Qian Gao**, Key Laboratory of Obstetric & Gynecological and Pediatric Diseases and Birth Defects of Ministry of Education, Sichuan University, Chengdu 610041, Sichuan Province, China

**Corresponding author:** Qian-Qian Gao, MD, Associate Professor, Department of Ultrasound, West China Second University Hospital, Sichuan University, No. 20 Third Section, South Renmin Road, Chengdu 610041, Sichuan Province, China. [gym471689@126.com](mailto:gym471689@126.com)

## Abstract

### BACKGROUND

Klippel-Trenaunay syndrome (KTS) is a rare congenital disorder. A detailed prenatal ultrasound examination plays an important role in the diagnosis of KTS and the subsequent counseling and follow-up of the patient.

### CASE SUMMARY

A 25-year-old woman attended our department for a regular examination. The whole of the right lower extremity and right buttock were observed to be markedly thicker compared to the left one at 18 wk of gestation. However, the lengths of the right femur, tibia and fibula were in the normal range. No marked edema and fluid/cystic spaces were detected in the lower limbs. There were no other organ abnormalities. The vasculature in the right limb was visibly dilated, with much higher intensive blood flow signals. No congenital embryonic veins were visible in both limbs. The right lower limb exhibited much more hypertrophy compared to the left limb two weeks later. Amniocentesis and genetic tests showed normal results with 46 XX. Despite the normal karyotype, the family opted to terminate the pregnancy. The post-mortem examination confirmed asymmetric hypertrophy of the right limb in the fetus and revealed a large area of marked dark-purple superficial capillary malformations occupying the skin of the right lower extremity. The enlargement of veins and soft tissue hypertrophy were also seen on postnatal X-ray and Magnetic Resonance Imaging. Autopsy revealed severe congestion in the right lower limb. A final diagnosis of KTS was made.

### CONCLUSION

KTS may be diagnosed prenatally based on the typical features observed during ultrasound examination.

Grade A (Excellent): A  
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 Grade D (Fair): 0  
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**Core Tip:** Klippel-Trenaunay syndrome (KTS) is a rare congenital disorder. The prenatal ultrasound features include hypertrophy of one extremity, a difference between the length of the bones of the extremities, multiple cystic lesions of the extremities or internal organs, increasing blood flow signals and dilated veins or persistence of the embryonic veins. KTS may be diagnosed based on these typical characteristics *in utero*.

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## INTRODUCTION

Klippel-Trenaunay syndrome (KTS) is a rare congenital disorder characterized by a classical triad of port-wine stains, varicosities, bone and soft tissue hypertrophy, and other complications such as lymphedema and lymphangiomas, which may involve any region of the body[1]. The etiology of this disease remains unclear. Most cases can be diagnosed through a complete history and clinical examination after birth[2]. The detailed prenatal images and follow-up are crucial for the diagnosis of KTS and the subsequent counseling and management of the patient. Ultrasonographic detection is an essential method for the diagnosis of KTS *in utero*[3]. The present study documents the prenatal ultrasound findings of a case of KTS.

## CASE PRESENTATION

### Chief complaints

A 25-year-old woman attended our department for a regular examination.

### History of present illness

The patient was 18 wk of gestation, with regular menstrual period and was gravida 1 parity 0. There was no history of other diseases during pregnancy.

### History of past illness

The patient had no significant medical history.

### Personal and family history

The personal and family history did not reveal any problems, including a history of vascular disorders.

### Physical examination

On admission, her temperature was 36.7°C, respiratory rate was 23 breaths/min, and blood pressure was 120/80 mmHg. The obstetric examination revealed that the fundal height and abdominal circumference of the patient were consistent with her gestational age.

### Laboratory examinations

Her nuchal translucency measurement and triple test results were normal.

### Imaging examinations

The whole of the right lower extremity and right buttock were observed to be

markedly thicker compared to the left one at 18 wk of gestation. However, the lengths of the right femur, tibia, and fibula were in the normal range. No marked edema and fluid/cystic spaces were detected in the lower limbs (Figure 1A and B). There were no other organ abnormalities. On the basis of these characteristics on ultrasound images, a diagnosis of KTS was suspected. Therefore, a detailed examination of the lower-limb vascular system was performed. The external iliac veins, great saphenous veins, femoral/popliteal veins, and the lateral thigh area in both lower limbs were carefully examined. The vasculature in the right limb was visibly dilated, with much higher intensive blood flow signals. No congenital embryonic veins were visible in both limbs (Figure 1C and D). Examination of the vascular system also supported the diagnosis of KTS. In the examination conducted two weeks later, the right lower limb exhibited much more hypertrophy compared to the left limb. Amniocentesis and genetic tests showed normal results with 46 XX.

The post-mortem examination confirmed asymmetric hypertrophy of the right limb and revealed a large area of marked dark-purple superficial capillary malformations occupying the skin of the right lower extremity (Figure 1E). Enlargement of the veins and soft tissue hypertrophy were also observed in the postnatal X-ray and magnetic resonance imaging (MRI), although embryonic veins were absent (Figure 1F and G). Autopsy revealed severe congestion in the right lower limb (Figure 1H).

## FINAL DIAGNOSIS

Following consultation with the geneticists and the professor of a vascular malformation clinic, a final diagnosis of KTS was made.

## TREATMENT

Despite the normal karyotype, the family opted to terminate the pregnancy after a counseling session with the obstetric/pediatric team, considering a range of possible outcomes, including significant disability.

## OUTCOME AND FOLLOW-UP

The patient underwent termination of the pregnancy. Two years later, the patient became pregnant and delivered a healthy female infant.

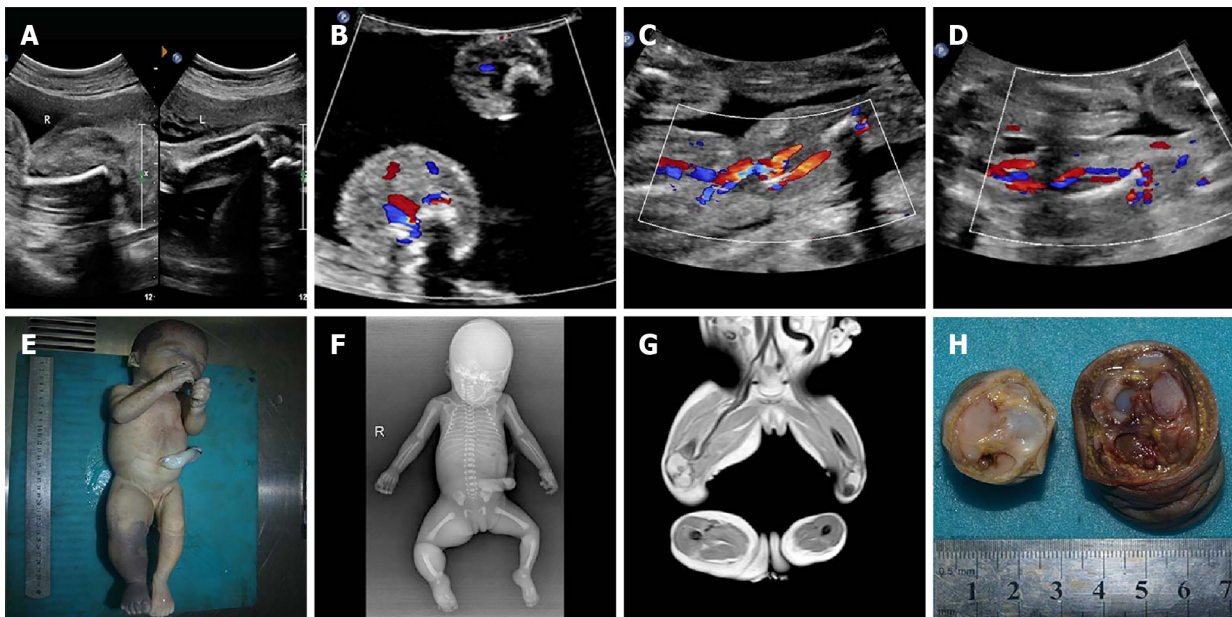
## DISCUSSION

KTS is a rare and sporadic congenital disorder, which presents at birth, early infancy or childhood. The proposed pathogenesis of the disease mainly includes deep vein atresia, chronic venous hypertension, persistence of the embryological vascular system and mesodermal anomaly[1].

The etiology of KTS remains unknown. Most reported cases have not been associated with chromosome abnormalities. Whelan *et al*[4] reported a case of KTS associated with a single gene defect at 5q or 11p, thus, demonstrating that this genetic defect may result in increased angiogenesis leading to KTS[5]. In addition, a case with a terminal 2q37.3 deletion[6] and a case of loss of heterozygosity for 1q21.2 q44[7] were also reported.

Most of the reported cases had no genetic characteristics except a case with unproven autosomal dominant inheritance[8]. Most KTS cases were unilateral and affected the lower extremities. Both upper and lower limbs might be involved in 10% of patients. According to published reports, hemangioma lesions can also involve other organs and body parts, such as the liver, lung, spinal cord, cranial area, skull, intestinal tract, urinary tract, testis, adrenal glands, and peritoneal and retroperitoneal cavity[7,9-11].

Most patients have skin vascular nevi or wine stains to some extent and hypertrophy of the skeleton and soft tissue; thus, they were diagnosed according to these typical signs at birth[1]. A deep tissue cavernous hemangioma may gradually appear and aggravate in later childhood. The prenatal ultrasound features *in utero*



**Figure 1** Prenatal sonograms at 18 wk of gestation and images after induced-abortion. A and B: Marked thickening of the right thigh, no marked edema and fluid/cystic spaces in the lower limbs (A, sagittal section and B, transverse section); C and D: Enlarged vasculature and higher intensive blood flow signals in the right limb; E: The purple stains predominantly affecting the right lower limb and asymmetry of the lower limbs; F: X-ray image showing the right lower limb hypertrophy; G: Magnetic resonance imaging image showing enlargement of the vasculature and hypertrophy of soft tissue; H: Severe congestion in the right lower limb. R: Right lower limb; L: Left lower limb.

have been described in many studies[8,12-14]. They include marked asymmetrical hypertrophy of limbs, thickening of the subcutaneous soft tissue, multiple cystic lesions, increasing blood flow signals, dilated veins or persistence of embryonic veins. It is difficult to detect skin capillary malformations, we speculate that the fetoscope may be a useful tool to observe fetal skin vascular nevi or wine stains *in utero*.

One of the main features of KTS is anomaly of the limb vein system, which appears as the persistence of embryonic veins, and varicose enlargement hypoplasia of the limb venous system. Assimakopoulos *et al*[9] revealed a case associated with a hypertrophied great saphenous vein. We also found more blood flow signals and the dilation of deep veins, which was confirmed by MRI and autopsy after induced abortion in our case. These findings indicated that it is feasible to carry out a thorough prenatal ultrasound examination of the venous system in the lower extremities to obtain more information in order to diagnose KTS.

The clinical presentation varies from minimally symptomatic disease to life-threatening bleeding and embolism. The prognosis is correlated with the size of the masses and their growth. There were large and extensive anechoic areas or the involvement of internal organs in reviewed cases with poor prognosis[3]. Therefore, the ultrasound scan should be repeated and the area and progress of the cystic lesions monitored, which may increase the risk of intrauterine heart failure and bleeding in the postnatal stage.

It is worth noting that there are many more reported cases after birth than *in utero*. This indicates that many cases of fetal KTS are not diagnosed prenatally as the lesions are minimal or complex, and it is difficult to make an accurate diagnosis. KTS should be suspected when there is hypertrophy of one of the extremities, discrepancy in the bone length, surface masses on the limbs and/or trunk, and unexplained cystic lesions of the internal organs or limbs. Therefore, necessary examinations should be performed to determine the location and severity of the lesions and to provide a reliable basis for the selection of appropriate treatment methods. MRI is another useful method, which can offer more detail on the soft tissue and vasculature.

## CONCLUSION

KTS may be diagnosed prenatally based on the typical features on ultrasound examination. Timely prenatal diagnosis and follow-up are important for subsequent prenatal counseling and adjustment of medical care and choices according to each

case.

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