Calcifying nested stromal epithelial tumor of the liver in a patient with Klinefelter syndrome: a case report and review of the literature

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Abstract

Background: Calcifying nested stromal epithelial tumor (CNSET) is a primary neoplasm of the liver, characterized by well-demarcated nests consisting of spindle and epithelioid cells with calcification and bone formation. An association of Cushing syndrome with CNSET has drawn attention, but the origin of CNSET has not been clarified.

Case presentation: We report here the case of a 20-year-old male with Klinefelter syndrome who underwent liver resection for an increasing liver tumor that was pathologically diagnosed with CNSET. He was postoperatively followed up and received several examinations, and recurrences and extrahepatic lymph node metastases were detected on the 64th day after surgery. Chemoembolization and chemotherapy were not effective, leading to tumor progression with development of progressive liver failure, and the patient finally died 164 days after hepatectomy.

Conclusions: This case suggests that an imbalance of hormones affects the genesis and progression of CNSET, and indicates the importance of closely following patients with CNSET by imaging with attention to hepatic recurrence and extrahepatic metastases.

Keywords: Calcifying nested stromal epithelial tumor (CNSET), Hormone imbalance, Klinefelter syndrome, Liver, Neoplasm

Background

Calcifying nested stromal-epithelial tumor (CNSET) is an uncommon primary hepatic tumor that is characterized by a nested morphologic growth pattern composed of spindled and epithelioid cells with various shape of calcification or ossification. Most liver cancer is hepatocellular carcinoma (HCC), followed by intrahepatic bile duct cancer. In 2001, Ishak et al. first described a non-hepatic and non-biliary tumor resembling CNSET [1]. This tumor is known by several other names, including ossifying stromal-epithelial tumor, desmoplastic nested spindle cell tumor of the liver (DNSTL), nested stromal epithelial tumor (NSET), and ossifying malignant mixed epithelial and stromal tumor [2–23]. As far as we are aware, 38 cases have been reported in the literature. These tumors have similar morphology, immunohistochemistry, and molecular profiles, and Misra et al. suggested that they may be related, but with a spectrum of morphologic features [24]. The reported tumors have been found predominantly in females and commonly in children, and most arose from the right hepatic lobe. In a number of cases, an association between these tumors and Cushing syndrome has also been described. Here, we report a case of postoperatively recurrent CNSET with aggressive clinical behavior and extrahepatic lymph node metastasis in a patient with Klinefelter syndrome. To our knowledge, this is the first case of a patient with CNSET concurrent with Klinefelter syndrome. CNSET is generally described as a tumor with low malignant potential, but the severe and progressive clinical course in our case indicates that...
the pathogenesis of CNSET may be related to hormone imbalance.

Case presentation
The patient was a 20-year-old male who had been a low-birth-weight infant, and had a history of Klinefelter syndrome and pulmonary valve stenosis. He was introduced to our hospital for further examination of a liver tumor that was increasing in size. The tumor had been found incidentally after laboratory findings in a health checkup showed impairment of liver function. The patient had declined treatment due to his employment situation, and had instead been followed up for 1 year.

At the first visit, he was completely asymptomatic with normal vital signs. A physical examination revealed a palpable right upper mass without tenderness. No symptom related to Cushing syndrome was observed. In blood tests, hepatitis B virus surface antigen and hepatitis C virus antibody were negative. Liver function tests indicated mild dysfunction. Regarding tumor markers, serum alpha-fetoprotein (AFP) and carcinoembryonic antigen (CEA) were normal; however, neuron-specific enolase (NSE) was elevated.

Ultrasonography showed a large low-echoic solid tumor with a vertical diameter of >80 mm with partial calcification implied by an acoustic shadow in an anterior lesion of the liver. A computed tomography (CT) scan of the chest, abdomen, and pelvis revealed an 81 × 76 × 72 mm large, heterogeneously enhanced mass in the right lobe of the liver with dense partial calcification (Fig. 1a). Subsequent positron emission tomography (PET)/CT showed a large hepatic mass in the right lobe with a maximum standardized uptake value (SUV) of 22.4 and no extrahepatic metastasis. In magnetic resonance imaging (MRI), most of the tumor was weakly enhanced in T1-weighted images and strongly enhanced in T2-weighted images. Part of the tumor had early enhancement and washout in enhanced MRI. These findings suggested HCC, and especially fibrolamellar HCC, but without evidence of distant metastasis.

Right hepatic lobectomy and cholecystectomy were performed 11 months after the initial detection of the tumor. The patient received no adjuvant chemotherapy or radiotherapy. The postoperative course was characterized by respiratory failure that required reintubation on postoperative day (POD) 2. X-ray and bronchofiberscopy showed pneumonias due to pulmonary atelectasis and pulmonary edema. The subsequent hospital course was uneventful. On POD 7, a CT scan of the abdomen was interpreted as negative for hemoperitoneum and tumor recurrence, and the patient was discharged on POD 12.

The patient was followed up as an outpatient and received several examinations. On POD 62, a CT scan showed multiple, obscure, and circumscribed recurrent lesions in the remnant liver with contrast enhancement. The largest of these lesions had a diameter of 42 mm in segment 1 (S1) (Fig. 1b). In addition, a hypermetabolic para-aortic lymph node with possible metastasis was identified. On PODs 70 and 73, the patient underwent transcatheter arterial chemoembolization (TACE), but a second CT scan in the outpatient department on POD 84 revealed enlargement of recurrent tumors and the para-aortic lymph node. Chemotherapy (protocol for HCC) was started, but was unsuccessful because of side effects. At this time, there were no further surgical options and no other chemotherapy that was likely to be effective. Therefore, the patient received palliative care. The patient died 164 days after hepatectomy from tumor progression with development of progressive liver failure.

Grossly, the tumor was confined to the right liver lobe. The resected specimen weighed 1180 g. The lesion had a maximum diameter of 100 mm, and was a well-circumscribed solitary mass with multiple small calcifications that were sharply demarcated from surrounding uninvolved liver parenchyma (Fig. 2). The surgical margin was tumor-free. Microscopically, the tumor was characterized by an organoid arrangement of cellular nests of epithelioid cells and areas of sheet-like cell overgrowth (Fig. 3a). These cells had oval-like nuclei with no clear nucleolus and eosinophilic cytoplasm. Transition zones between epithelioid and spindle cells were observed, and a framework of spindle cells surrounded nests of epithelioid cells (Fig. 3b, c). Bile ducts were not intermingled.

Fig. 1 CT findings. a A large, heterogeneously enhanced mass with focal calcification was present in the right lobe of the liver. b Multiple recurrent lesions in the remnant liver and an enlarged para-aortic lymph node on postoperative day 62.
with the tumor region. There were extensive regions of necrosis and calcification (or ossification) in the center of the tumor (Fig. 3d).

In immunohistochemical staining, epithelioid cells were positive for CD56, cytokeratin AE1/AE3 (focal), WT-1 (diffuse or dot-like in cytoplasm), β-catenin (diffuse in nucleus), vimentin, NCAM, and NSE (Fig. 4a, b). Spindle cells in mesenchymal components such as the septum were diffusely stained with α-smooth muscle actin (α-SMA) (Fig. 4c). The AFP level was within the normal range. Staining for glypican-3 was negative. The proliferation index on MIB-1 (Ki-67) immunostaining was < 5%. Staining was negative for hepatocyte paraffin-1, CK7, adrenocorticotropic hormone (ACTH), estrogen receptor (ER), and progesterone receptor (PR). The morphological and immunohistochemical features led to diagnosis of CNSET.

**Discussion and conclusions**

CNSET of the liver is an uncommon tumor that was first described by Ishak et al. in 2001. Predicting the clinical behavior of this tumor is difficult because only a few cases have been reported under a variety of names.

![Fig. 2](image1.png) Gross findings for the lesion. An inspection showed a well-circumscribed solitary mass with multiple small calcifications that were sharply demarcated from surrounding liver parenchyma.

![Fig. 3](image2.png) Histopathological findings for the tumor. a Nests of epithelial cells surrounded by a desmoplastic stroma with focal calcifications (×40). b Epithelioid nests (×200). c Desmoplastic stroma of spindle cells (×200). d Necrosis and regions of calcification (×100).
Klinefelter syndrome occurs in males with at least one Y chromosome and at least two X chromosomes. It is the most common sex chromosomal disorder in males and affects 1 in 660 men [25]. This syndrome was first suggested that CNSET begins as a gradually enlarging hepatic calcification since childhood. Makahlouf et al. reported a liver adenoma in a young patient with Klinefelter syndrome [28].

Four of the patients with CNSET (Table 1) had a history of oral contraceptives, which is of interest because high serum estradiol and low serum testosterone occur in Klinefelter syndrome and in oral contraceptive use. An association between hepatic adenoma and oral contraceptives has been proposed, based on suspected carcinogenic effects of estrogen and enzyme induction of progesterone [29]. Wang et al. also proposed a relationship between oral contraceptive use and development of CNSET in adult women [13]. There are no reports of a relationship of CNSET with sex hormones, but we suggest that CNSET may be related to imbalance of sex hormones. A relationship between oral contraceptive use and breast cancer risk has been shown, and Beaber found that long-term use of oral contraceptives may be more strongly related to a risk of ER− (× 3.5) and triple-negative (× 3.7) cancer compared to ER+ cancer; although the differences were not significant [30]. Our case was negative for ER in immunostaining. An imbalance of sex hormones, such as a high level of estradiol, might initiate occurrence and development of CNSET via a non-sex hormone receptor pathway, and it is possible that the constitutive imbalance of sex hormones affected the aggressive clinical behavior in our case.

In six of the cases of CNSET, patients had a history of hepatic calcification since childhood. Makahlouf et al. suggested that CNSET begins as a gradually enlarging small calcified lesion [7]. In our case, calcification of the liver was not observed in childhood. In blood tests, serum levels of AFP and CEA were in the normal range in all investigated cases. Imaging shows a typically large and well-circumscribed lesion with a macrolobulated margin, as in our case; with a distinctive large mass with heterogeneous enhancement and dense calcification on CT; and similar features with predominant T1 hypointensity and T2 hyperintensity on MRI. Dynamic postcontrast MRI may help to distinguish CNSET from other diseases with similar enhancement patterns. Radiologic differential diagnoses include hepatic vascular formation, fibrolamellar HCC, and hepatoblastoma [20]. Fibrolamellar HCC, which we first suspected in our case, is often detected at a similar age and has similar imaging findings with a central scar on CT and MRI. Calcifications are seen in 35–68% of cases of fibrolamellar HCC, but these tend to be small and fewer than three in number [31].

On gross examination, a CNSET is a lobulated mass with variable calcification and is generally well-circumscribed within the liver. Some previous reports presented no evidence of calcification. The size of the tumors has
| No | Authors | Age | Sex | Symptoms                                      | Surgical treatment | Chemotherapy | Outcome/follow-up | Associated history or disease                                      |
|----|---------|-----|-----|-----------------------------------------------|--------------------|--------------|-------------------|------------------------------------------------------------------|
| 1  | Heywood et al. [2] 2002 | 28  | F   | Incidental                                    | Trisegmentectomy IV, V, and VI wedge resection VII | Recurrence/72 months | Persistent fetal hemoglobin, OCP                                |
| 2  | Hill et al. [4] 2005    | 2   | M   | Abdominal mass                                 | Partial hepatectomy | Post          | Alive/84 months    |                                                                  |
| 3  | 6       | F   | Incidental                                | Gross excision     |               | Alive/58 months  |                                                                  |
| 4  | 6       | F   | Incidental                                | Gross excision     |               | Alive/8 months   |                                                                  |
| 5  | Heerema-McKenney et al. [3] 2005 | 14  | F   | Abdominal mass                                 | Left lobectomy     | Post          | Recurrence/11 years | Nephroblastomatosis and Wilms tumor                               |
| 6  | 3       | F   | Incidental                                | Gross excision     | Pre/post       | Alive/8 months  |                                                                  |
| 7  | 4       | M   | Incidental                                | Enucleation        |               | Alive/36 months  |                                                                  |
| 8  | 11      | F   | Cushingoid features, abdominal mass       | Left lobectomy     |               | Alive/24 months  | Cushing syndrome                                             |
| 9  | 12      | F   | Cushingoid feature, abdominal mass        | Right hepatectomy  |               | Alive/168 months | Cushing syndrome                                             |
| 10 | 14      | F   | Ileus                                      | Gross excision     | Post          | Recurrence/12 months | BWS, hypoplastic kidney, omphalocele                           |
| 11 | Brodsky et al. [5] 2008 | 17.5| F   | Cushingoid feature, abdominal pain            | Left lobectomy + partial hepatectomy | Recurrence/12 months | Cushing syndrome                                             |
| 12 | Wirojanan et al. [6] 2008 | 2   | M   | ND                                            | Resection          | Post          | Alive/84 months    | Fragile X syndrome                                             |
| 13 | Meir et al. [8] 2009    | 2.5 | F   | Incidental                                  | Right lobectomy    |               | Alive/8 months    | Asymptomatic hydronephrosis                                     |
| 14 | 2       | F   | Incidental                                | Partial hepatectomy |               | Alive/6 months  |                                                                  |
| 15 | Maikhlof et al. [7] 2009 | 14  | F   | Incidental                                | Partial hepatectomy |               | Alive/151 months   |                                                                  |
| 16 | 15      | F   | Incidental                                | Partial hepatectomy | Post          | Alive/264 months |                                                                  |
| 17 | 16      | M   | Cushingoid feature                         | Partial hepatectomy |               | Alive/56 months  | Cushing syndrome                                             |
| 18 | 18      | F   | Incidental                                | Transplant         |               | Death/40 months (no recurrence)                              |                                                                  |
| 19 | 19      | M   | Incidental                                | No (needle biopsy) |               | Lost to follow-up |                                                                  |
| 20 | 32      | F   | Incidental                                | Partial hepatectomy |               | Alive/13 months  |                                                                  |
| 21 | 33      | F   | Incidental                                | Partial hepatectomy |               | Alive/14 months  |                                                                  |
| 22 | Rod et al. [9] 2009     | 17  | F   | Cushingoid feature, palpable right upper abdominal mass | Resection          |               | Alive/30 months    | Cushing syndrome                                             |
| 23 | Grazi et al. [10] 2010  | 25  | F   | Diarrhea and recurrent abdominal pain         | Right lobectomy extending the caudate lobe segment IV | Alive/6 months | OCP                                                          |
| 24 | Ramirez et al. [11] 2010 | 33  | M   | Unspecific abdominal pain and dyspepsia       | Left lobectomy     |               | Alive/15 months    | HBV(+)                                                         |
| 25 | Wang et al. [13] 2011    | 34  | F   | Incidental                                | Left lobectomy     |               | Alive/42 months    | OCP                                                          |
| 26 | Hommann et al. [12] 2011 | 14  | F   | Incidental                                | Resection, transplantation | Post          | Recurrence/28 months | Moderate hypoxic brain injury, omphalocele                     |
| No | Authors                  | Age | Sex | Symptoms                                      | Surgical treatment      | Chemotherapy | Outcome/follow-up     | Associated history or disease          |
|----|-------------------------|-----|-----|-----------------------------------------------|-------------------------|--------------|----------------------|-----------------------------------------|
| 28 | Assmann et al. [14] 2012 | 16  | M   | Palpable abdominal mass                       | Transplantation          | Pre          | Alive/24 months      | Cushing-like habitus                    |
| 29 |                        | 3   | F   | Unclear obstipation                           | Partial hepatectomy      | Post         | Alive/5 years        |                                         |
| 30 | Geramizadeh et al. [15] 2012 | 8   | M   | Cushingoid feature                            | Right extended hepatectomy |              | Death 10 days (no recurrence) | Cushing syndrome                        |
| 31 | Ghodke et al. [16] 2012 | 9   | M   | Abdominal pain, fever, jaundice, weight loss, anorexia | Segmental hepatectomy  | ND           | Alive/12 months      |                                         |
| 32 | Malowany et al. [17] 2013 | 2   | F   | Incidental                                   | Resection               | ND           | No recurrence         | BWS                                     |
| 33 | Procopio et al. [18] 2014 | 23  | F   | Abdominal distension and dyspepsia            | Extended left hepatectomy|              | Alive/21 months      | OCP                                     |
| 34 | Samarghandi et al. [19] 2015 | 11  | F   | Weight gain, increased appetite, abdominal pain | Segmental hepatectomy  | ND           | ND                   |                                         |
| 35 | Schaffer et al. [20] 2016 | 14  | F   | Abdominal distention and swelling of cheeks   | Transplant              |              | Alive/10 months      | BWS, Cushing syndrome                   |
| 36 | Weeda et al. [21] 2016  | 16  | M   | Cushingoid feature, weight gain, distended abdomen | Trisegmentectomy         |              | Alive/13 years       | Cushing syndrome                        |
| 37 | Khoshnam et al. [22] 2017 | 14  | F   | Cushingoid feature, abdominal swelling and pain | Transplantation          | Pre          | Alive/ND             | BWS, Cushing syndrome                   |
| 38 | Tehseen et al. [23] 2017 | 13  | F   | Abdominal pain and distention, Cushingoid features | Transplantation          | Pre          | Alive/28 months      | Developmental delay, Cushing syndrome    |
| 39 | Our case 2017           | 20  | M   | Incidental                                   | Right hepatectomy        | Post         | Death recurrence/2 months | Klinefelter syndrome                    |

ND: not determined, OCP: oral contraceptive pill use, BWS: Beckwith-Wiedmann syndrome
ranged from 2.8 to 30 cm, and have tended to be large. On the cut surface, the tumor might appear granular, homogeneous white or tan, with foci of softening, and cyst formation. Histologic analysis shows typical characteristics of well-demarcated nests of spindle and epithelioid cells surrounded by a desmoplastic stroma. Within the nests, tumor cells with epithelioid shapes have bland clear features. The desmoplastic stroma has morphologic characteristics of myofibroblasts, and the surrounding liver parenchyma largely shows no remarkable finding. Individual cell psammomatous calcification and regions of ossification have frequently been described in previous case reports.

Immunohistochemistry can also help with diagnosis of CNSET. The tumors tend to be positive for vimentin, pan-cytokeratin, and CD57. Staining for WT-1 protein in tumor cells is varied, with weak to moderate nuclear staining, dot-like paranuclear staining, and diffuse cytoplasmic staining. Nest cells are focally positive for NSE, CD56, and sometimes S-100. Stromal components of CNSET are consistently immunoreactive for α-SMA. The histological differential diagnoses of tumors with both epithelial and mesenchymal components and variable calcification include hepatoblastoma, synovial sarcoma, teratoma, desmoplastic round cell tumor (DSRCT), inflammatory myofibroblastic tumor of the liver, biliary rhabdomyosarcoma, metastatic Wilms tumor, and spindle cell carcinoid tumor [24].

Standard treatment for CNSET has not established, but all reported cases underwent gross total resection of the tumor, including wedge resection, partial hepatectomy, and hepatic lobectomy. In seven patients diagnosed with Cushing syndrome, cushingoid symptoms subsided after tumors producing ACTH were excised. Six cases with unresectable tumors received liver transplantation. A few cases received chemotherapy using a soft tissue sarcoma or hepatoblastoma protocol. However, the effect of using chemotherapy or radiotherapy has not been proved. The prognosis of CNSET is unclear, but the tumor is normally slow-growing and of low malignant potential. In contrast to our patient, most cases have long-term survival after resection. Five cases had local recurrence after excision of the primary tumor and two had metastasis. Brodsky et al. described a case with extrahepatic lymph node metastasis after resection of the primary liver tumor [5]. Hommann et al. described an unresectable tumor in a 16-year-old girl who underwent hepatic transplantation, but had lung metastasis at 28 months postoperatively and died due to lung metastasis 37 months after transplantation [12]. Makhlouf et al. described a patient with two local recurrences that were successfully treated by radiofrequency ablation [7]. Our case had local recurrence in the liver and extrahepatic lymph node metastasis immediately after resection and showed more aggressive clinical behavior than most cases of CNSET. Therefore, this case suggests that patients with CNSET should be carefully followed by imaging study with close attention to hepatic recurrence and extrahepatic metastases.

There is no conclusive evidence for the origin of CNSET, but several hypotheses have been proposed. In addition to the potential link between CNSET and oral contraceptives, it has been hypothesized that CNSETs are derivatives of hepatic mesenchymal precursor cells with possible differentiation along a bile duct lineage, based on CD56-positive staining of bile ducts and tumor nests [20]. It was also noted that WT-1 expressed in CNSETs might affect transformation of mesenchymal to epithelial cells [3]. Based on our case, we suggest that a continual imbalance of hormones influences the pathogenesis of CNSET and leads to aggressive behavior after resection. However, there is also no evidence for the histogenesis of CNSET, and further studies of this tumor are needed.

In conclusion, we have presented a case of calcifying nested stromal epithelial tumor of the liver, an uncommon tumor characterized by well-demarcated nests of epithelial and spindle cells surrounded by myofibroblastic stroma and various calcifications. To our knowledge, this is the first reported case of CNSET with Klinefelter syndrome. Since this tumor presented with a very aggressive clinical course with recurrences and metastasis, the genesis and progression of CNSET may be related to hormone imbalance. Additionally, this case indicates the importance of careful follow-up with imaging and close attention to recurrence and metastases in a patient with CNSET. More studies are needed to improve the diagnosis and treatment of CNSET.

### Abbreviations

- AFP: Alpha-fetoprotein; CEA: Carcinoembryonic antigen; CNSET: Calcifying nested stromal epithelial tumor; CT: Computed tomography; DNSTL: Desmoplastic nested spindle cell tumor of the liver; DSRCT: Desmoplastic round cell tumor; HCC: Hepatocellular carcinoma; MRI: Magnetic resonance imaging; NSE: Neuron-specific enolase; NSET: Nested stromal epithelial tumor; PET: Positron emission tomography; POD: Postoperative day; SUV: Standardized uptake value; TACE: Transcatheter arterial chemoembolization; α-SMA: α-Smooth muscle actin

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### Authors’ contributions

TS drafted the manuscript and examined the patient; KN assisted in the preparation of the manuscript and examined the patient; YT, SA, HN, and HK contributed to pathological diagnosis and assisted in the preparation of the manuscript; IK, KD, SK, ET, and HK critically reviewed the manuscript and examined the patient. All authors read and approved the final manuscript.
Ethics approval and consent to participate
Not applicable.

Consent for publication
Written informed consent for publication of their clinical details and clinical images was obtained from the patient’s family. A copy of the consent form is available for review by the Editor of this journal.

Competing interests
The authors declare that they have no competing interests.

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Author details
1. Ishak K, Goodman Z, Stocker J. Tumors of the liver and intrahepatic bile ducts. Washington: Armed Forces Institute of Pathology; 2001. p. 276–8.
2. Heywood G, Burgart LJ, Nagorney DM. Ossifying malignant mixed epithelial tumor of the liver: a case report and review of literature. J Gastrointest Oncol. 2012;3:151–7. https://doi.org/10.4103/0975-1051.98928.
3. Makhlouf HR, Abdul-Al HM, Wang G, Goodman ZD. Calcifying nested stromal-epithelial tumor of the liver: a neoplasia with defective mesenchymal-epithelial transition. Hum Pathol. 2010;41:1831–6. https://doi.org/10.1016/j.humpath.2010.04.012.
4. Komatsu N, Robinson H, Clay MR, Schaffer LR, Gillespie SE, Shehata BM. Calcifying nested stromal-epithelial tumor (CNSET) of the liver in Beckwith-Wiedemann syndrome. Eur J Med Genet. 2017;60:136–42. https://doi.org/10.1016/j.ejmg.2016.12.001.
5. Theeume S, Rampin L, Schermankiewitz E, Maglloca JF, Romero R. Successful liver transplantation for non-resectable desmoplastic nested spindle cell tumor complicated by Cushing’s syndrome. Pediatr Transplant. 2017;21: Epub Jun 18. https://doi.org/10.1111/petr.12382.
6. Mista S, Bihari C. Desmoplastic nested spindle cell tumors and nested stromal epithelial tumors of the liver. Am J Med. 2012;130:245–51. https://doi.org/10.1016/j.amjmed.2011.12.001.
7. Groth KA, Skakkebaek BH, Bovier BH. Klinefelter syndrome and liver adenoma. J Clin Gastroenterol. 1991;13:214–6. https://doi.org/10.1097/00004836-199106000-00010.
8. Komatsu N, Robinson H, Clay MR, Schaffer LR, Gillespie SE, Shehata BM. Calcifying nested stromal-epithelial tumor (CNSET) of the liver in Beckwith-Wiedemann syndrome. Eur J Med Genet. 2017;60:136–9. https://doi.org/10.1016/j.ejmg.2015.12.003.
9. Theeume S, Rampin L, Schermankiewitz E, Maglloca JF, Romero R. Successful liver transplantation for non-resectable desmoplastic nested spindle cell tumor complicated by Cushing’s syndrome. Pediatr Transplant. 2017;21: Epub Jun 18. https://doi.org/10.1111/petr.12382.
10. Groth KA, Skakkebaek BH, Bovier BH. Klinefelter syndrome and liver adenoma. J Clin Gastroenterol. 1991;13:214–6. https://doi.org/10.1097/00004836-199106000-00010.
11. Theeume S, Rampin L, Schermankiewitz E, Maglloca JF, Romero R. Successful liver transplantation for non-resectable desmoplastic nested spindle cell tumor complicated by Cushing’s syndrome. Pediatr Transplant. 2017;21: Epub Jun 18. https://doi.org/10.1111/petr.12382.
12. Komatsu N, Robinson H, Clay MR, Schaffer LR, Gillespie SE, Shehata BM. Calcifying nested stromal-epithelial tumor (CNSET) of the liver in Beckwith-Wiedemann syndrome. Eur J Med Genet. 2017;60:136–9. https://doi.org/10.1016/j.ejmg.2015.12.003.
13. Theeume S, Rampin L, Schermankiewitz E, Maglloca JF, Romero R. Successful liver transplantation for non-resectable desmoplastic nested spindle cell tumor complicated by Cushing’s syndrome. Pediatr Transplant. 2017;21: Epub Jun 18. https://doi.org/10.1111/petr.12382.
14. Groth KA, Skakkebaek BH, Bovier BH. Klinefelter syndrome and liver adenoma. J Clin Gastroenterol. 1991;13:214–6. https://doi.org/10.1097/00004836-199106000-00010.