1	Left retrocaval ureter around the ipsilateral limb of a double
2	caudal vena cava in a cat
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19 Summary

Necropsy of an adult, neutered male cat, which was euthanized due to its FIV-positive status, demonstrated the presence of a left retrocaval ureter that was entrapped around the left limb of a double caudal vena cava. These associated anomalies originate from the complex embryofoetal development of the caudal vena cava. Since no clinical signs had been reported and no gross lesions related to this anomaly were observed, this manifestation should be considered as a mere anatomical variation.

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27 Keywords: Double caudal vena cava; retrocaval ureter; cat

28 An adult, neutered male cat (*Felis catus*) was euthanized after being tested positive for 29 FIV (feline immunodeficiency virus). The cadaver was presented at the Department of 30 Veterinary Sciences of the University of Antwerp for educational purposes. No gross macroscopic lesions potentially associated with FIV positivity were found. However, the cat 31 32 presented a double caudal vena cava. More specifically, the left and right caudal vena cava 33 merged only at the level of the first lumbar vertebra to form a single vessel. The left and right 34 caudal vena cava individually drained the ipsilateral common iliac veins. No anastomoses 35 were found between the left and right counterparts at the level of the iliac veins, nor between 36 any other segment of the double caudal vena cava. Just prior to its confluence with its 37 contralateral part, the left caudal vena cava crossed the abdominal aorta ventrally (Fig. 1).

38 The complex embryofoetal development of the caudal vena cava lies at the basis of the 39 observed vascular anomaly. For years, the supracardinal model that was published (for the 40 cat) in 1920 by Huntington and McClure was considered the gold standard. This model that is applied in both human and veterinary embryology states that the right-sided counterparts of 41 42 three pairs of embryonic veins (i.e. supracardinal, subcardinal and caudal cardinal veins) form 43 the abdominal part of the caudal vena cava by means of fusion, anastomosis and asymmetric 44 vascular degeneration during prenatal life (Huntington and McClure, 1920; Butler, 1927; 45 Cornillie and Simoens, 2005; Cornillie et al., 2006; Hyttel, 2010; Sadler, 2012).

However, a recent study revisiting the development of the inferior vena cava and vena azygos by means of three-dimensional reconstructions of human embryos (Hikspoors et al., 2014) has revealed various inaccuracies in the supracardinal system of Huntington and McClure (1920). All dorsal veins, formerly indicated as the supracardinal veins, have to be considered as actual segments of the caudal cardinal system that gradually adopts a different topography due to allometric growth of surrounding organs and tissues. What is important for the present case is that, according to these recent insights, two paired longitudinal channels can be found in the region of the kidneys, namely both caudal cardinal veins, which course at
the dorsomedial side of the kidneys, and the caudal extensions of the subcardinal veins, which
also run medially but ventral to the kidneys.

56 At the pelvic inlet, just caudal to the bifurcation of the aorta into the umbilical arteries, 57 these four veins are united in a huge anastomosis. At a certain point during development the 58 caudal cardinal venous segments draining into the subcardinal anastomosis at the 59 craniomedial poles of the kidneys, the infrarenal segments of the subcardinal veins collecting 60 and interconnecting the left and right common iliac veins, and tributaries of both venous 61 systems form a vascular ring around the ipsilateral kidney and ureter (Cornillie and Simoens, 2005; Hikspoors et al., 2014). This complex venous configuration evolves from a bilateral 62 63 symmetric system to a single, right-sided vein, in a process during which the left-sided 64 counterparts of all aforementioned veins gradually disappear (Cornillie and Simoens, 2005) 65 (Fig. 2). The presence of a double caudal vena cava is therefore considered to result from the persistence of the left-sided venous segments, in combination with the absence of the 66 67 aforementioned iliac anastomosis (Bass et al., 2000; Cornillie, 2008).

Moreover, careful dissection during necropsy revealed another curiosity. The left ureter was entrapped by the left (limb of the) caudal vena cava, a condition known as retrocaval or circumcaval ureter (Gramegna et al., 2003). It slightly deviated towards the midline to cross the left caudal vena cava dorsally, at the level of the fifth lumbar vertebra. The left ureter resumed its more lateral course towards the urinary bladder after emerging between the vein and the left psoas musculature. The right ureter presented a normal course along the right psoas muscles (Fig. 3).

A rare case of retrocaval ureter in an intact male cat was reported in 2006 by Cornillie and co-workers. This cat, however, presented a right retrocaval ureter associated with a single caudal vena cava. The recently gained insights into the development of the infrarenal part of

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78 the caudal vena cava allow formulation of the following hypothesis on the origin of a 79 retrocaval ureter. The basis is that the infrarenal segment of the subcardinal vein, which forms 80 the ventral branch of the circumureteric vascular ring, persists instead of the dorsally located 81 caudal cardinal vein. When the subcardinal vein contributes in the formation of the caudal 82 vena cava, it will cross the ureter ventrally and sweep it along its migratory path towards the 83 midline. This results in the medial deviation and entrapment of the ureter between the caudal 84 vena cava and psoas musculature. In our case with occurrence of a left retrocaval ureter 85 associated with a double caudal vena cava, it is hypothesized that this occurred on the left 86 side. More specifically, the infrarenal part of the left limb of the caudal vena cava was formed 87 by the persisting infrarenal segment of the subcardinal vein that runs ventral to the left ureter, 88 while the right limb was formed by the caudal cardinal vein that runs dorsal to the right ureter 89 (Fig. 4).

90 According to an extended study performed by Bélanger and co-workers (2014), who examined the carcasses of 301 cats, a right retrocaval ureter is a rather common anatomic 91 92 variation. It was seen in 28% of the examined carcasses. Far more unique is the presence of a 93 left retrocaval ureter, having a prevalence of 1%. The presence of a double caudal vena cava 94 without entrapment of one or both ureters is seldom seen (0.7%). The combination of the 95 vascular and urinary variations is more typical as it has been observed in approximately 5% of 96 the examined carcasses. The combination of a left retrocaval ureter and a double caudal vena 97 cava was only observed in one female cat (Bélanger et al., 2014). So, this case report is the 98 first to describe these findings in a male cat.

99 The clinical relevance of the presented anomalies is unknown since no gross 100 macroscopic lesions were observed and no clinical signs had been reported. Nevertheless, it is 101 likely that cats with retrocaval ureters are more susceptible to ureteral obstruction and 102 consequently hydronephrosis. The former can be the result of the external pressure exerted on

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the ureter by the caudal vena cava that retracts the ureter to the midline (Cornillie et al., 2006). Moreover, humans that develop kidney stones may suffer from obstructing uroliths that cannot pass the ureter where it sweeps around the caudal vena cava to flow to the urinary bladder (Soundappan and Barker, 2004). In cats, the pathogenesis of urolithiasis is different from that in humans as cats are more prone to develop lower urinary tract disease. This is characterized, amongst others, by the formation of urinary bladder stones and the potential obstruction of the urethra (Lund et al., 2013).

110 In summary, retrocaval ureter(s) and/or a double caudal vena cava in cats are rarely 111 observed during necropsy. Since no clinical signs have yet been associated with these 112 anomalies, they should be considered as mere anatomical variations.

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152 Figure legends

Fig. 1. Ventral view of the opened abdominal cavity in a necropsied neutered male cat demonstrating the presence of a left caudal vena cava (l cvc) and a right caudal vena cava (r cvc). The left ureter (arrows) is entrapped by the former vessel. II = left lung, Ii = liver, st =stomach, sp = spleen, 1 rv = left renal vein, lk = left kidney, cd = colon descendens, j =jejunum, ub = urinary bladder, fh = femoral head.

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159 Fig. 2. Ventral schematic views of a human embryo (L = left side, R = right side) showing the 160 development of the caudal vena cava (after Hikspoors et al., 2014). First (left image), a 161 vascular ring around each kidney and ureter is formed by the caudal cardinal venous segments 162 (1: I and IV) that drain into the subcardinal anastomosis (a and b) at the craniomedial poles of 163 the kidneys (2), the infrarenal segments of the subcardinal veins (3: II and III) that collect (c 164 and d) and interconnect (e) the left and right common iliac veins, and tributaries of both 165 venous systems. The paired subcardinal vein anastomoses in the ventral midline (f). 166 Subsequently (right image), the infrarenal part of the caudal vena cava is formed by the 167 interconnection of the left and right common iliac veins (e). The renal part is made of the 168 right-sided caudal cardinal vein (I), whereas the suprarenal segment is formed by the right 169 side of the regressing anastomosis (f). The renal veins originate from the cranial anastomoses 170 between the caudal cardinal and subcardinal venous systems (a and b).

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Fig. 3. Ventral view of the abdominal cavity of the same cat as in Fig. 1. The structures of interest have been dissected to clearly show the retrocaval position of the left ureter. The kidneys have been removed and the urinary bladder was incised to investigate the potential presence of hydronephrosis and urolithiasis, respectively. cvc = caudal vena cava, l cvc = left 176 caudal vena cava, r cvc = right caudal vena cava, lu = left ureter, pm = psoas muscles, aa =
177 abdominal aorta, ru = right ureter, fh = femoral head, ub = urinary bladder.

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179 Fig. 4. Schematic representation of the formation of a left retrocaval ureter in the presence of 180 a double caudal vena cava. When the left limb of the caudal vena cava (l cvc) is formed by the 181 persisting subcardinal vein (III), it will cross the ureter ventrally and entrap it while making 182 the anastomosis with the caudal cardinal vein (d), which is located dorsal to the ureter, at the 183 pelvic inlet. The absence of a right retrocaval ureter is explained by the formation of the right 184 limb of the caudal vena cava by the right caudal cardinal vein (a, I and c) that runs dorsal to 185 the ureter. lk = left kidney, lu = left ureter, rk = right kidney, ru = right ureter, cvc = caudal186 vena cava, r cvc = right caudal vena cava.

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