



Sildenafil induced liver injury – a case report

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Drug induced liver injury (DILI) can result from medications, dietary supplements, or various herbal products. When the injury is dose-dependent, it is referred to as intrinsic liver injury. Idiosyncratic liver injury is not dose-dependent and occurs in a small subset of individuals, with a variable latency period (1–2).

A 59-year-old male with a history of arterial hypertension was admitted to our department due to a two-week history of abdominal pain, jaundice, and elevated liver enzymes.

In the months preceding the onset of symptoms, he had modified his lifestyle by adopting a new diet, increasing physical activity, and initiating the use of a dietary supplement aimed at lowering cholesterol. He reported occasional use of sildenafil. His usual intake of alcohol was at least one unit daily.

Imaging and laboratory diagnostics excluded diseases of the gallbladder and biliary tree, hepatic vascular disorders, infectious causes, inherited metabolic disorders, and immune-mediated liver diseases. Histological examination of the liver biopsy was consistent with DILI. We confirmed hepatocellular toxic liver injury, most likely due to sildenafil, although a possibility of a combined effect of multiple agents cannot be excluded. At the most recent follow-up, liver enzyme levels had returned to normal. Table 1 and 2 present investigations and their values at the initial and last outpatient assessment.

A broad spectrum of substances has the potential to cause liver injury. Obtaining a comprehensive medical history and excluding alternative etiologies are essential steps in establishing an accurate diagnosis. The primary therapeutic interventions include discontinuation of the suspected agent and close patient monitoring. Despite an extensive diagnostic

Table 1: Investigations and Their Values at the Patient's Initial Assessment in Our Institution

| Investigation | Result | Ref. values |
|--|--------|--------------|
| Blood count | | |
| Leukocytes (10 ⁹ /L) | 6.35 | 4.00 – 10.00 |
| Erythrocytes (10 ¹² /L) | 5.53 | 4.50 – 5.50 |
| Haemoglobin (g/L) | 162 | 130 – 170 |
| MCV (fL) | 86.1 | 83.0 – 101.0 |
| Thrombocytes (10 ⁹ /L) | 279 | 150 – 410 |
| Biochemical investigations (Blood serum) | | |
| Glucose (mmol/L) | 5.7 | 3.6 – 6.1 |
| Urea (mmol/L) | 3.2 | 2.8 – 7.5 |
| Creatinine (µmol/L) | 67 | 64 – 104 |
| oGF (mL/min/1.73m ²) | >90 | 80 – 120 |
| CRP (mg/L) | 4 | <5 |
| Total bilirubin (µmol/L) | 185 | <17 |
| Direct bilirubin (µmol/L) | 143 | <5 |
| AST (µkat/L) | 15.19 | <0.58 |
| ALT (µkat/L) | 39.63 | <0.74 |
| γ-GT (µkat/L) | 7.3 | <0.92 |
| Alkaline phosphatase (µkat/L) | 2.48 | 0.50 – 2.00 |
| Lipase (µkat/L) | 1.55 | <1.25 |
| Sodium (mmol/L) | 135 | 135 – 145 |
| Potassium (mmol/L) | 3.83 | 3.5 – 5.3 |
| Chloride (mmol/L) | 105 | 97 – 110 |
| Magnesium (mmol/L) | 0.86 | 0.6 – 1.1 |
| Lactate dehydrogenase (µkat/L) | 4.72 | <4.13 |
| Albumin (g/L) | 36 | 32 – 55 |
| Ammoniac (µmol/L) | 42 | <50 |
| Iron (Fe) (µmol/L) | 4.19 | <4.13 |
| Feritin (µg/L) | 3082 | 30 – 300 |
| Transferrin (g/L) | 2.41 | 2.00 – 3.60 |
| Transferrin saturation (%) | 82.1 | 16 - 45 |
| Ceruloplasmin (g/L) | 0.30 | 0.22 – 0.61 |
| Copper (Cu) (µmol/L) | 20.6 | 11.0 – 22.0 |
| CDT (%) | 2.01 | 1.19 – 2.47 |
| Tumor markers | | |
| Total PSA (µg/L) | 0.72 | <4 |
| Free PSA (µg/L) | 0.258 | <0.934 |
| AFP (kU/L) | 2.8 | <11.1 |
| CEA (µg/L) | <2 | <5 |
| CA 19-9 (kU/L) | 5 | <37 |
| CA 72-4 (kU/L) | 2.20 | <8.82 |
| β2-microglobuline (mg/L) | 2.61 | <8.2 |

| Investigation | Result | Ref. values |
|---|---|--------------|
| Hormones | | |
| B12 (pmol/L) | 34 | 6.1 – 32.6 |
| Folates (nmol/L) | 1342 | 132 – 857 |
| TSH (mU/L) | 1.13 | 0.27 – 4.2 |
| 17-beta oestradiol (nmol/L) | 0.11 | |
| Progesterone (nmol/L) | <1.6 | |
| Prolactin (µg/L) | 5.9 | 1.6 – 17.7 |
| Testosterone (nmol/L) | 35.76 | |
| SHBG (nmol/L) | 102.8 | 13.5 – 71.4 |
| DHEAS (nmol/L) | 1.96 | |
| β-HCG (IU/L) | 0 | <15 |
| FSH (U/L) | 5.1 | |
| LH (U/L) | 5.0 | |
| Immunology | | |
| ANA, ANCA, ASMA, LKM, liver profile (M2, gp210, sp100, LKM1, LC1, SLA, F-actin) | Negative | |
| AMA | Suspected | |
| IgG (g/L) | 12.10 | 7.67 – 15.90 |
| IgA (g/L) | 4.09 | 0.61 – 3.56 |
| IgM (g/L) | 0.35 | 0.37 – 2.86 |
| Toxicology | | |
| Opiates, Amphetamines, Cocaine, Cannabis, Methadone, Benzodiazepines | Negative | |
| Coagulation profile | | |
| Prothrombin time (PT) (E) | 0.82 | 0.7 – 1.2 |
| INR | 1.11 | |
| Thrombin time (TT) (s) | 17 | 15 – 21 |
| Activated Partial Thromboplastin Time (aPTT) (s) | 33 | 26 – 36 |
| Fibrinogen (g/L) | 2.58 | 2.2 – 4.2 |
| Microbiology | | |
| Hepatitis E | Negative | |
| Hepatitis A | Post-Infectious/ Post-Vaccination Status | |
| Hepatitis B | Non-reactive | |
| Hepatitis C | Non-reactive | |
| HIV | Non-reactive | |
| HSV-1 | Negative | |
| HSV-2 | Negative | |
| CMV | Negative | |
| EBV | IgG – positive, IgM – negative | |
| Leptospira | Negative | |
| Hantavirus | Negative | |
| Haemocultures | Negative | |

Table 2: Investigations and Their Values at the Patient's Last Outpatient Assessment in Our Institution

| Investigation | Result | Ref. values |
|--|-------------|--------------|
| Blood count | | |
| Leukocytes (10 ⁹ /L) | 6.35 | 4.00 – 10.00 |
| Erythrocytes (10 ¹² /L) | 5.53 | 4.50 – 5.50 |
| Haemoglobin (g/L) | 162 | 130 – 170 |
| MCV (fL) | 86.1 | 83.0 – 101.0 |
| Thrombocytes (10 ⁹ /L) | 279 | 150 – 410 |
| Biochemical investigations (Blood serum) | | |
| Glucose (mmol/L) | 6.3 | 3.6 – 6.1 |
| Urea (mmol/L) | 3.9 | 2.8 – 7.5 |
| Creatinine (µmol/L) | 77 | 64 – 104 |
| eGFR (mL/min/1.73m ²) | >90 | 80 – 120 |
| CRP (mg/L) | <3 | <5 |
| Total bilirubin (µmol/L) | 11 | <17 |
| Direct bilirubin (µmol/L) | 3 | <5 |
| AST (µkat/L) | 0.79 | <0.58 |
| ALT (µkat/L) | 1.21 | <0.74 |
| γ-GT (µkat/L) | 0.95 | <0.92 |
| Alkaline phosphatase (µkat/L) | 1.09 | 0.50 – 2.00 |
| Sodium (mmol/L) | 136 | 135 – 145 |
| Potassium (mmol/L) | 4.51 | 3.5 – 5.3 |
| Chloride (mmol/L) | 104 | 97 – 110 |
| CHE (µkat/L) | 193 | 117 – 317 |
| Cholesterol (mmol/L) | 5.23 | 4.0 – 5.7 |
| Triglycerides (mmol/L) | 0.96 | 0.6 – 1.7 |
| Albumin (g/L) | 38 | 32 – 55 |
| Iron (Fe) (µmol/L) | 14 | <4.13 |
| Feritin (µg/L) | 151 | 30 – 300 |
| AFP (kU/L) | 2.4 | < 11.1 |
| Coagulation profile | | |
| Prothrombin time (PT) (E) | 0.86 | 0.7 – 1.2 |
| INR | 1.09 | |

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