

1 Tuberculosis and adrenal insufficiency: the duo continues to reveal secrets!

2 Abstract

3

4 **Introduction:** Isolated adrenal tuberculosis is rare, particularly in countries where
5 tuberculosis remains endemic. The advent of modern imaging has made it possible to
6 identify atypical radiological forms that can mimic tumour lesions. This clinical case illustrates
7 the importance of a complete aetiological investigation in all cases of bilateral adrenal
8 involvement, particularly in countries where tuberculosis is endemic.

9 **Case study:** A 75-year-old patient with no history of tuberculosis presented with symptoms
10 of adrenal insufficiency that had been developing over a period of six months. Clinical
11 examination revealed an asthenic, hypotensive patient with no melanoderma. Laboratory
12 tests confirmed primary adrenal insufficiency with an 8-hour cortisol level of 52 µg/l (normal
13 range 62 to 194 µg/dL), adrenocorticotropic hormone (ACTH) of 148 pg/ml (normal range 9-
14 60 pg/ml), hyponatraemia of 130 mmol/l and hyperkalaemia of 7.8 mmol/l. Adrenal
15 computed tomography (CT) revealed atypical radiological features combining nodular,
16 calcified and pseudotumoral lesions. The diagnosis was confirmed by bacteriological testing.
17 Treatment with hydrocortisone and anti-tuberculosis drugs led to an improvement in
18 symptoms.

19 **Conclusion:** The combination of adrenal insufficiency and bilateral CT scan lesions, even if
20 atypical, should raise suspicion of tuberculosis, especially in endemic areas. An integrated
21 clinical, hormonal, radiological and microbiological approach is essential.

22 **Keywords:** Addison's disease; multifocal tuberculosis; diagnostic imaging; atypical imaging

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24 Introduction

25 Isolated adrenal tuberculosis is rare, particularly in countries where tuberculosis remains
26 endemic. Adrenal tuberculosis is the leading cause of Addison's disease, especially in
27 developing countries [1,2].

28 It is essential to obtain a rapid aetiological diagnosis of Addison's disease in order to prevent
29 patients from developing a potentially fatal adrenal crisis. Imaging methods are the
30 cornerstone of the initial diagnosis of adrenal tuberculosis. Computed tomography plays an
31 important role in diagnosis and follow-up [2,3].

32 Multifocal forms combining pulmonary, adrenal and sometimes lymph node lesions remain
33 rare, accounting for less than 5% of cases in some series [4].

34 We report a clinical case of an elderly patient with adrenal insufficiency revealing multifocal
35 tuberculosis with an atypical CT scan appearance.

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37 Observation

38 This is a 75-year-old patient with a history of hypertension and diabetes, no history of
39 tuberculosis or tuberculosis infection, a former smoker and alcoholic who quit 20 years ago.

40 The patient arrived at the emergency department with digestive symptoms including
41 abdominal pain, vomiting and diarrhoea, associated with severe asthenia. He was admitted
42 to the nephrology department for acute renal failure and underwent three haemodialysis
43 sessions, with good improvement in renal function.

44 Our opinion was sought due to the progressive worsening of generalised physical and mental
45 asthenia in the context of a deterioration in his general condition.

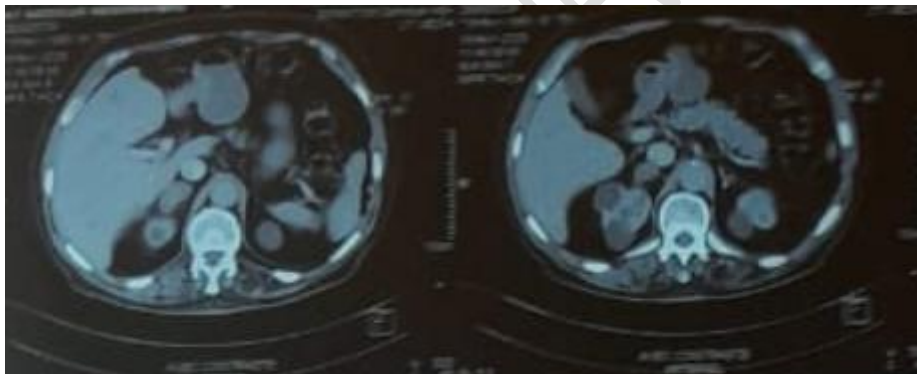
46 The interview revealed chronic asthenia for six months, hypoglycaemic and hypotensive
47 discomfort, polyuropolydipsic syndrome, constipation, disabling arthralgia of the lower limbs
48 and a negative infectious history.

49 The clinical examination revealed a conscious patient who was pale, asthenic, hypotensive,
50 with no signs of melanoderma or dehydration, hypoglycaemia at 0.60g/l and a body mass
51 index of 27kg/m².

52 Laboratory tests confirmed primary adrenal insufficiency with an 8-hour cortisol level of 52
53 µg/l (normal range 62 to 194 µg/dL), adrenocorticotrophic hormone (ACTH) at 148 pg/ml
54 (normal range 9-60 pg/mL), hyponatraemia at 130 mmol/l and hyperkalaemia at 7.8 mmol/l.
55 The rest of the assessment noted a CRP of 26 mg/l and a glomerular filtration rate of 51
56 ml/min.

57 Adrenal CT scan revealed a heterogeneous nodular mass measuring 36x22mm on the right
58 side, with a spontaneous density of 27 Hounsfield units (HU) and an absolute washout of
59 28.5%. On the left side, calcifications and an 11x12mm nodular lesion with a spontaneous
60 density of 25 HU and an absolute washout of 12.5% were observed, leading to a diagnosis of
61 suspicious nodular lesions (Figure 1).

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Figure 1: Adrenal CT scan showing:

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On the right, a heterogeneous nodular mass measuring 36x22mm, with a spontaneous density of 27 HU and
66 an absolute washout of 28.5%.

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On the left, calcifications and an 11x12mm nodular lesion with spontaneous density at 25 HU and absolute
68 washout of 12.5%.

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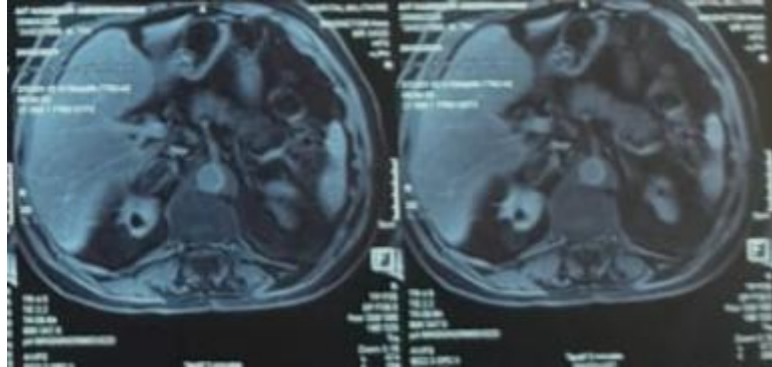
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Magnetic resonance imaging (MRI) of the adrenal gland refines the diagnosis, revealing a
74 nodular formation occupying the entire right adrenal gland, with low T2 signal intensity, T1
75 isosignal, with signal drop on phase opposition sequences, site of peripheral diffusion
76 restriction with progressive peripheral enhancement, measuring 21x36mm.

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77 Left adrenal thickening with a maximum thickness of 7.5 mm, showing low signal intensity
78 on all sequences and calcifications; conclusion: low-fat right adrenal adenoma against a
79 background of bilateral adrenal granulomatosis (Figure 2).

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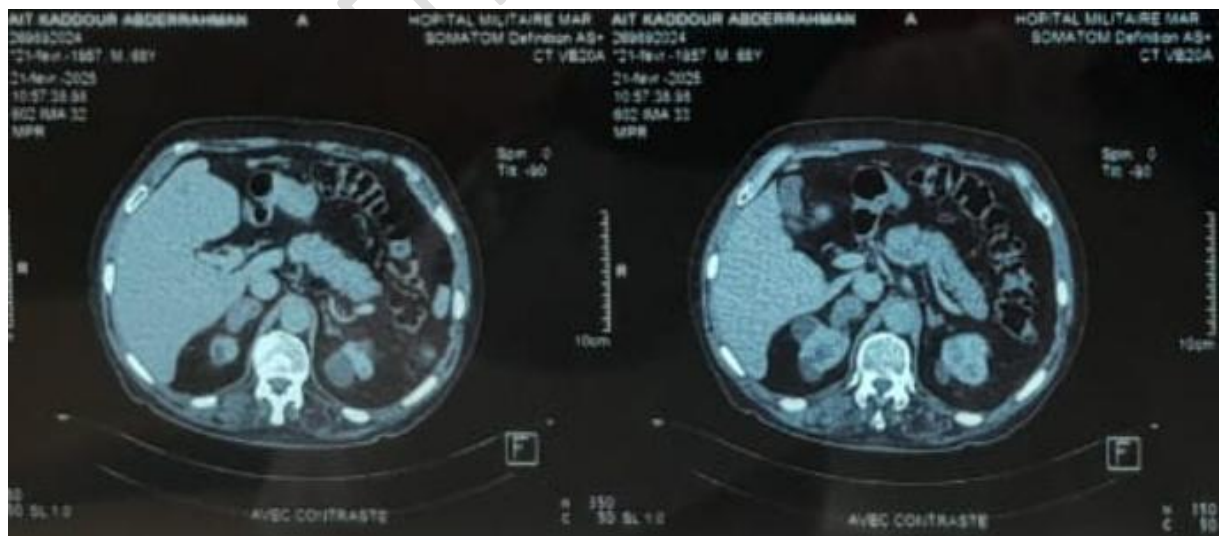
Figure 2: Adrenal MRI showing a nodular formation occupying the entire right adrenal gland, with low T2 signal, T1 isosignal, measuring 21x36mm, and left adrenal thickening with a maximum thickness of 7.5mm with low signal intensity on all sequences, showing calcifications and concluding a low-fat right adrenal adenoma against a background of bilateral adrenal granulomatosis.

88

89 The thoraco-abdominal-pelvic CT scan shows, in addition to adrenal involvement, a retractile
90 focus of condensation in the right posterior basal pleural space associated with scattered
91 pulmonary nodules and micronodules (Figure 3).

92 The hypothalamic-pituitary MRI shows no abnormalities.

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Figure 3: Thoraco-abdominal-pelvic CT scan shows, in addition to adrenal involvement, a retractile focus of condensation in the right posterior basal pleural space associated with nodules and

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97 After these images pointed to an inflammatory and infectious process, suggestive of

98 granulomatosis, a GeneXpert test was performed on sputum samples and came back positive
99 for Mycobacterium tuberculosis, leading to a diagnosis of pulmonary tuberculosis.

100

101 Treatment consisted of substitution with injectable and then oral hydrocortisone,
102 fludrocortisone and anti-tuberculosis therapy, with favourable results: clinical improvement
103 (good general condition and normal blood pressure) and biological improvement (correction
104 of hydroelectrolytic disorders: sodium level at 136 mmol/l and potassium level at 3.80
105 mm/l).

106

107 Discussion

108 Addison's disease was first described by Thomas Addison in 1855. He demonstrated the
109 destruction of the bilateral adrenal glands by tuberculosis [5]. The disease often develops
110 over several years after an initial active or latent tuberculosis infection, predominantly in
111 adults [3].

112 Classically, adrenal tuberculosis manifests as signs of slow adrenal insufficiency after
113 destruction of more than 90% of the cortex. The advent of modern imaging has made it
114 possible to identify atypical radiological forms with pseudotumour masses that may suggest
115 metastases, low-fat adenomas, or other rare neoplasms. [2,3]

116 In our case, the absence of known infection and initial pulmonary signs illustrates the
117 characteristic clinical latency of tuberculous adrenal disease.

118 Dissemination occurs via the bloodstream, and adrenal destruction is due to granulomatous
119 infiltration followed by fibrosis and calcification [6].

120

121 Radiologically, the natural history of adrenal tuberculosis has three stages of progression [7]
122 (Table 1).

Phase	CT scan appearance	Significance
1. Acute	Homogeneous bilateral adrenal hypertrophy, hypodense, sometimes with peripheral enhancement	Active inflammation
2. Subacute	Heterogeneous nodular masses, sometimes pseudotumoral, with necrotic foci	Parenchymal destruction
3. Chronic	Calcifications, atrophy or fibrous masses	Sequelae scars

123 **Table 1:** The three radiological stages of adrenal tuberculosis

124 Atypical forms combine nodular, calcified and pseudotumoral lesions, making differential
125 diagnosis difficult with bilateral metastases, adrenal lymphoma and atypical adenomas or
126 low-fat myelolipomas [8].

127 The classic features of adrenal MRI are illustrated in Table 2 [7, 9].

Phase MRI / Imaging Interpretation / Appearance

Acute	Hyposignal T1 / Hypersignal T2	Caseous necrosis
Chronic	Hyposignal T2, Atrophy, Calcifications	Fibrosis, sequelae

128 Table 2: Classic MRI features of the different phases of adrenal tuberculosis.

129 The radiological features described in our observation, particularly the bilateral,
130 heterogeneous lesions, low washout and calcifications, correspond to a subacute or chronic
131 form, but atypical due to the presence of a pseudo-tumoural nodular appearance.

132 Recent studies confirm that the pseudo-tumoural phase of adrenal tuberculosis is often
133 confused with metastasis or a low-fat adenoma:

134 In 2012, Yang et al. compared 25 cases of adrenal tuberculosis and 30 metastases: the
135 tuberculous lesions had a significantly lower absolute washout (<50%) and a heterogeneous
136 T2 signal, similar to our case. [3]

137 In 2014, Guo et al. showed that the presence of bilateral calcifications is highly suggestive,
138 but late. Our patient already had a calcified left adrenal gland, indicating an advanced stage.
139 [10]

140 The originality of our case lies in several factors: the patient's advanced age, atypical adrenal
141 imaging simulating an adenoma, and microbiological confirmation of pulmonary tuberculosis.
142 It demonstrates the need to maintain tuberculosis as a differential diagnosis for adrenal
143 masses, even in the absence of a history of tuberculosis.

145 Conclusion

146 Adrenal tuberculosis, although rare, remains a cause to consider in any case of adrenal
147 insufficiency associated with heterogeneous bilateral lesions, particularly in countries where
148 tuberculosis is endemic.

149 Computed tomography and MRI play an important role in the diagnosis of adrenal
150 tuberculosis. Understanding the imaging characteristics of adrenal tuberculosis is essential
151 for guiding diagnosis and initiating prompt and essential treatment of Addison's disease
152 secondary to adrenal tuberculosis.

153 An integrated clinical, hormonal, radiological and microbiological approach is essential. The
154 prognosis depends on rapid diagnosis, appropriate hormone replacement therapy and
155 appropriate anti-tuberculosis treatment.

156 Références bibliographiques

- 157 1. Huang Y, Zhang Y, Wang H, Zhang N. Primary bilateral adrenal tuberculosis
158 associated with Addison's disease: a case report. Urol Case Rep. 2024;56:102837.
159 doi:10.1016/j.eucr.2024.102837
160 2. Zhang XC, Yang ZG, Li Y, Min PQ, Guo YK, Deng YP, Dong ZH. Addison's disease due

- 161 to adrenal tuberculosis: MRI features. *Abdom Imaging*. 2008;33:689–694.
162 doi:10.1007/s00261-007-9352-8
- 163 3. Yang ZG, Guo YK, Li Y, Deng YP, Ma ES. Differentiation between tuberculosis and
164 metastases in the adrenal glands: evaluation with contrast-enhanced CT. *EurRadiol*.
165 2012;22(9):2104–2111.
- 166 4. Rezgui A, Ben Fredj F, Mzabi A, Karmani M, Laouani C. Multifocal tuberculosis in
167 immunocompetent individuals. *Pan Afr Med J*. 4 May 2016;24:13. doi:
168 10.11604/pamj.2016.24.13.6030. PMID: 27583077; PMCID: PMC4992387.
- 169 5. Tran NQ, et al. Bilateral adrenal masses due to tuberculosis: how to diagnose
170 without extra-adrenal tuberculosis. *Endocrinol Diabetes Metab Case Rep*.
171 2021;2021:21-0093. doi:10.1530/EDM-21-0093
- 172 6. Huang YC, Chen WY, Chen CL, et al. Evaluation of primary adrenal insufficiency
173 secondary to tuberculous adrenalitis with computed tomography and magnetic
174 resonance imaging: current status. *World J Radiol*. 2015;7(10):336–342.
- 175 7. Guo YK, Yang ZG, Li Y, et al. Imaging features of adrenal tuberculosis: comparison
176 with primary adrenocortical carcinoma and metastases. *EurRadiol*. 2007;17(7):1765–
177 1775.
- 178 8. Song JH, Chaudhry FS, Mayo-Smith WW. The incidental adrenal mass on CT:
179 prevalence and interpretation. *AJR Am J Roentgenol*. 2008;190(5):1163–1168.
- 180 9. Al-Mamari A, Balkhair A, Gujjar A, et al. A case of disseminated tuberculosis with
181 adrenal insufficiency. *Sultan Qaboos Univ Med J*. 2009;9(3):324–327.
- 182 10. Guo YK, Yang ZG, Li Y, et al. Addison's disease due to adrenal tuberculosis: contrast-
183 enhanced CT features. *Abdom Imaging*. 2014;39(5):1069–1077.