

CRISIS IN THE CORTEX

Hajare Iraqi, MD¹, Maral Koolian, MD², Jill Pancer, MD³, Teresa Cafaro, MD²

¹ Department of Medicine, McGill University, Montreal, Quebec, Canada

² Division of Internal Medicine, Jewish General Hospital, McGill University, Montreal, QC, Canada; Center for Clinical Epidemiology and Community Studies, Lady Davis Institute, Jewish General Hospital, Montreal, QC, Canada; Department of Medicine, Jewish General Hospital, McGill University, Montreal, QC, Canada.

³ Division of Endocrinology and Metabolism, Jewish General Hospital, Montreal, Quebec, Canada



Hôpital général juif
Jewish General Hospital



McGill

CASE PRESENTATION

RECORDS	PROGRESS NOTES
	ID: 64F
	CHIEF CONCERN: "generalized weakness"
	PAST MEDICAL HISTORY:
	1. IgG MM s/p auto-SCT (2018) in PR, on Lenalidomide 5mg DIE
	2. Paroxysmal Afib (CHADS 0), on Apixaban 5mg BID
	3. Previous pericarditis (2021)
	4. Polyarthritits NYD (2021)
	5. Migraines with aura
	HISTORY OF PRESENT ILLNESS: Two-week history of sudden-onset fatigue, anorexia, and vomiting. No abdominal pain, infectious symptoms, B symptoms, or sick contacts/travel.
	Two previous ER visits with impression of viral gastroenteritis and possible community-acquired pneumonia (treated with amoxicillin-clavulanate/doxycycline).
	PHYSICAL EXAM: BP 105/71 HR 106 RR 18 SpO2 97% RA T 36.6
	Appeared fatigued, but non-toxic.
	Examination otherwise non-contributory.

INVESTIGATIONS:

CBC: **Hb 102** ← **110** (MCV 91.8) **Plt 92** ← **167** ← **116** **WBC 3.5** ← 5.2

CHEM: Cr 82 **Na 130** K 4.8 HCO3 22 (AG 13) Mg 0.78

LFT: Bili 12 ALT < 5 ALP 89 Albumin 31

COAG: **INR 1.2** ← **2.1**

VBG: pH 7.37/pCO2 42/HCO3 24 Lactate 1.3

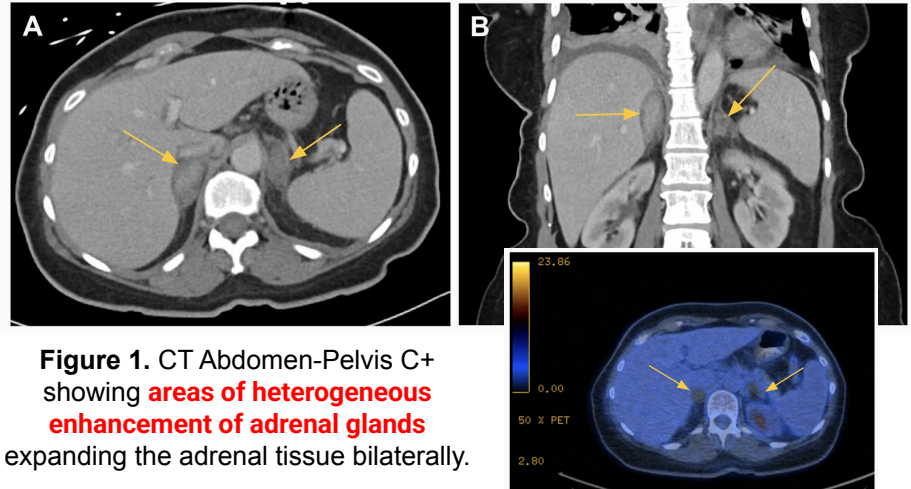


Figure 1. CT Abdomen-Pelvis C+ showing **areas of heterogeneous enhancement of adrenal glands** expanding the adrenal tissue bilaterally.

Figure 2. PET scan with **spontaneously hyperdense bilateral adrenal glands with absent FDG uptake centrally** and rim of minimal activity on the left, SUV 3.5.



DIAGNOSIS

A.M. CORTISOL 61 nmol/L

ADRENAL INSUFFICIENCY SECONDARY TO BILATERAL ADRENAL HEMORRHAGE

Laboratory Test	Result (Reference Range)
Endocrine	
Early morning serum cortisol	61 nmol/L (172 - 497 nmol/L)
Adrenocorticotrophic hormone	13.8 pmol/L (1.6 - 13.9 pmol/L)
21-hydroxylase antibody	5 ISR (0 - 35 ISR)
Coagulation	
INR	2.1 (0.8 - 1.2)
APTT	25.4 s (24.6 - 37.6 s)
Fibrinogen	8.0 g/L (2.0 - 4.5 g/L)
Haptoglobin	3.06 g/L (0.30 - 2.00 g/L)
Antiphospholipid Antibody Screen	
Lupus anticoagulant normalized ratio	1.14 (0.00 - 1.20)
Anticardiolipin antibody IgG	< 9 GPL u/mL (0-21 GPL u/mL)
Anticardiolipin antibody IgM	< 9 MPL u/mL (0-21 MPL u/mL)
B2-glycoprotein	< 1.4 U/mL (0.0 - 12.0 U/mL)

Table 1. Results of laboratory investigations for the etiology of bilateral adrenal hemorrhage.

ETIOLOGIES OF ADRENAL HEMORRHAGE

- Trauma
- Neoplasm (malignant > benign)
- Sepsis, Waterhouse-Friderichsen syndrome
- Infections (e.g. COVID-19)
- Inherited coagulopathy
- Acquired coagulopathy: **anticoagulation**, antiplatelets, antiphospholipid syndrome, DIC, HIT, VITT, etc.
- Peripartum
- Iatrogenic
- Idiopathic

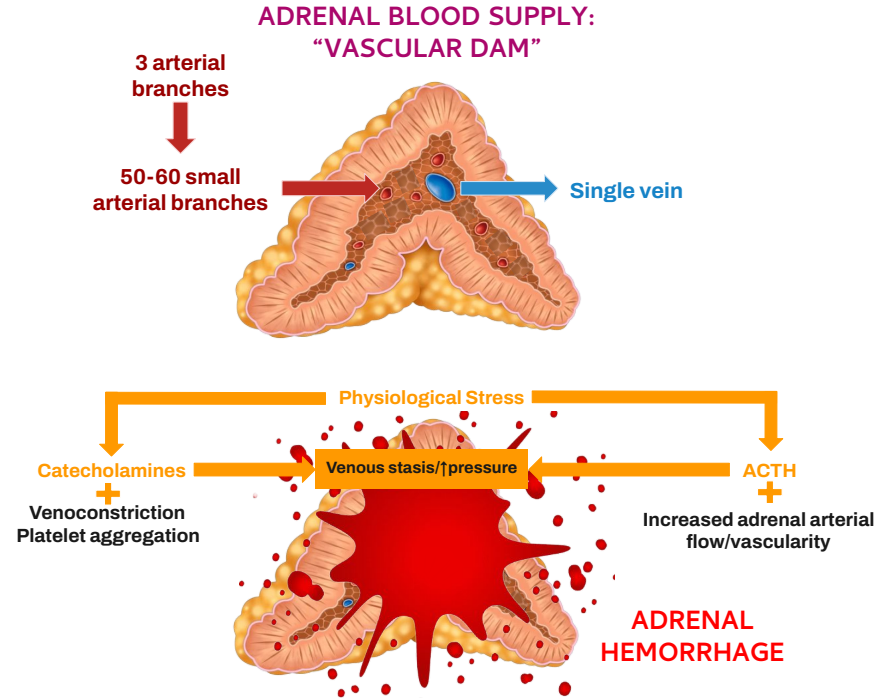
TREATMENT: parenteral corticosteroids initiated with rapid improvement, switched to oral regimen at discharge (no indication for mineralocorticoid replacement), anticoagulation held temporarily.

ADRENAL HEMORRHAGE

KEY POINTS

Adrenal hemorrhage is a **potentially life-threatening complication** that is often overlooked due to its **vague, non-specific presentation**.

Wang et al. (2024) retrospective study of 193 patients with adrenal hemorrhage found that compared to patients diagnosed after trauma, shock, or post-operatively, **“anticoagulation was noted to be a risk factor among patients with spontaneous adrenal hemorrhage (26.4% vs 6.9%; $p < 0.002$)”**



CONSIDER THE ADRENAL GLANDS AS POTENTIAL ATYPICAL BLEEDING SITES IN PATIENTS WITH BLEEDING DIATHESIS OR RECEIVING ANTICOAGULANTS TO ENSURE TIMELY IDENTIFICATION, TREATMENT, AND PREVENTION OF DECOMPENSATION.

REFERENCES

1. Badawy M, Gaballah AH, Ganeshan D, Abdelalziz A, Remer EM, Alsabbagh M, et al. Adrenal hemorrhage and hemorrhagic masses; diagnostic workup and imaging findings. *The British Journal of Radiology*. 2021 Nov 1;94(1127). DOI:10.1259/bjr.20210753
2. Elhassan YS, Ronchi CL, Wujewickrama P, Baldeweg SE. Approach to the patient with adrenal hemorrhage. *J Clin Endocrinol Metab*. 2022 Nov 21;108 (4):995-1006. DOI:10.1210/clinem/dgac672
3. Watts NB, Tindall GT. Rapid Assessment of Corticotropin Reserve After Pituitary Surgery. *JAMA*. 1988 Feb 5;259(5):708–8. DOI:10.1001/jama.1988.03720050044021
4. Charmandari E, Chrousos GP. Adrenal Insufficiency. *Encyclopedia of Endocrine Diseases*. 2004;75–80. DOI:10.1016/b0-12-475570-4/00029-9
5. Bornstein SR, Allolio B, Arlt W, Barthel A, Don-Wauchope A, Hammer GD, et al. Diagnosis and Treatment of Primary Adrenal Insufficiency: an Endocrine Society Clinical Practice Guideline. *The Journal of Clinical Endocrinology & Metabolism*. 2016 Feb;101(2):364–89. DOI:10.1210/jc.2015-1710
6. Alidoost M, Soomro R, Gubeladze A, Morabia A, Holland S, Asif A, et al. Rivaroxaban Related Bilateral Adrenal Hemorrhage: A Rare Complications of Direct Oral Anticoagulants – A Case Reports. *American Journal of Case Reports*. 2019 Nov 2;20:1607–11. DOI:10.12659/AJCR.917780
7. Aloini ME, Manella S, Biondo I, Maggio R, Roberto G, Ricci F, et al. Bilateral adrenal hemorrhage: learning notes from clinical practice and literature review. *Frontiers in Endocrinology*. 2023 Nov 3;14. DOI:10.3389/fendo.2023.1233710
8. Al-Rawi S, Seidahmed M, Emam SS, Othman ES. A 65-Year-Old Man with Bilateral Adrenal Hemorrhage Following Prophylaxis for Postoperative Deep Vein Thrombosis with Rivaroxaban. *American Journal of Case Reports*. 2023 Jul 19;24. DOI:10.12659/AJCR.939816
9. Arosemena MA, Rodriguez A, Hasini Ediriweera. Bilateral adrenal haemorrhage secondary to rivaroxaban in a patient with antiphospholipid syndrome. *BMJ Case Reports*. 2020 Jul 1;13(7):e234947–7. DOI:10.1136/bcr-2020-234947
10. Comuth W, Christiansen JJ, Bloch-Münster AM, Husted S. Bilateral adrenal gland hemorrhage in a patient treated with rivaroxaban. *Blood Coagulation & Fibrinolysis*. 2017 Jan;28(1):102–4. DOI:10.1097/MBC.0000000000000541
11. Elhassan YS. Insights on Adrenal Hemorrhage. *Mayo Clinic Proceedings*. 2024 Mar 1;99(3):355–6. DOI:10.1016/j.mayocp.2024.01.010
12. Ly BA, Quintero L. Adrenal Insufficiency from Unilateral Adrenal Hemorrhage in a Patient on Rivaroxaban Thromboprophylaxis. *AACE Clinical Case Reports*. 2018 Oct 5;5(1):e70–2. DOI:10.4158/accr-2018-0340
13. Nasser B, Bughrra MS, Alakhras H, Nasser Z, Jameel OF. Unilateral Adrenal Hemorrhage: A Rare Complication of Anticoagulant Use. *Cureus*. 2022 Jun 10;14(6):e25821. DOI:10.7759/cureus.25821
14. Sanford Z, Nanjundappa A, Annie FH, Embrey S. Adrenal Hemorrhage in a Patient Anticoagulated with Apixaban with Antiphospholipid Syndrome. *Cureus*. 2019 Jul 9; 11 (7): e5108. DOI:10.7759/cureus.5108
15. Sheklabadi E, Sharifi Y, Tabarraee M, Tamehrizadeh SS, Rabiee P, Hadaegh F. Adrenal hemorrhage following direct oral anticoagulant (DOAC) therapy: two case reports and literature review. *Thrombosis Journal*. 2022 Jul 5;20(1). DOI:10.1186/s12959-022-00397-9
16. Tan JW, Shukla A, Hu JR, Majumdar SK. Spontaneous Adrenal Hemorrhage with Mild Hypoadrenalism in a Patient Anticoagulated with Apixaban for Antiphospholipid Syndrome: A Case Report and Literature Review. *Case Reports in Endocrinology*. 2022 Nov 30;2022:1–6. DOI:10.1155/2022/6538800
17. Wang TN, Padmanaban V, Bashian EJ, Davis HW, Kirsch MJ, Phay JE, Miller BS, Hackett CE, Dedhia PH. Clinical characteristics and outcomes of adrenal hemorrhage. Surgery. 2024 Jul;176(1):76-81. doi: 10.1016/j.surg.2024.03.004. Epub 2024 Apr 8. PMID: 38594100.

CONFLICT DISCLOSURES

	Co-author	Conflict disclosures
1	Hajare Iraqi	No conflict of interest to disclose.
2	Maral Koolian	No conflict of interest to disclose.
3	Jill Pancer	No conflict of interest to disclose.
4	Teresa Cafaro	No conflict of interest to disclose.

CREDITS: This presentation template was created by Slidesgo, including icons by Flaticon, and infographics & images by Freepik.