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Incidence, Presentation, and Natural History of Adrenal Hemorrhage: An Institutional Analysis

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Short Title: Incidence and Natural History of Adrenal Hemorrhage

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Author Contributions: MNS, CEG, MJC, and MTC contributed to conception and design of the study. MNS and KAS acquired and compiled the data, which was analyzed by MNS and CEG. MNS, ALW, and CEG drafted the article, with content

37 analysis and critical revisions by REG, MJC, and MTC. All authors provided final
38 approval of the article in its current form.

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48 **Abstract:**

49 Background

50 Adrenal hemorrhage (AH) can occur due to multiple etiologies with variable
51 radiographic appearance, often indistinguishable from underlying adrenal
52 neoplasms. There is a lack of AH literature and evidence-based guidelines. Our
53 study aimed to understand the prevalence and etiology of AH, follow-up, and
54 incidence of underlying neoplasm.

55

56 Materials and Methods

57 An institutional database was queried from 1/2006 to 10/2021 for patients with AH
58 on imaging, excluding patients with known malignancies, adrenal masses, or prior
59 adrenal surgery. Demographics, medical history, hematoma size, laterality,
60 biochemical evaluation, intervention, and additional imaging were reviewed.

61

62 Results

63 Of 490,301 imaging reports queried, 530 (0.11%) with AH met inclusion criteria.

64 Most imaging (n=485, 91.5%) was performed during trauma evaluation. Two
65 patients underwent dedicated intervention at presentation. Interval imaging was
66 performed in 114 (21.5%) patients at a median of 2.6 (IQR 0.99-13.4) months, with
67 resolution (n=84, 73.7%) or decreased size of AH (n=21, 18.4%) in most patients.

68 Only 10 patients (1.9%) saw an outpatient provider in our system to address AH or
69 evaluate for underlying mass, and 9 (1.7%) underwent biochemical screening.

70 Thirteen patients (11% of 118 patients with any follow-up) had evidence of an

71 adrenal mass, confirmed on serial imaging (n=10) or adrenalectomy (n=3). Scans
72 performed for non-trauma indications were significantly more likely to have an
73 underlying mass (n=6/26 (23.1%)) than those performed for trauma evaluation
74 (n=7/92 (7.6%), p=0.04).

75

76

77 Conclusion

78 AH is a rare finding associated with an increased rate of underlying adrenal mass
79 compared to adrenal incidentaloma, particularly when unrelated to trauma. Most
80 AH resolves spontaneously without intervention. Follow-up imaging at 6 months
81 can help distinguish mass-associated AH from simple hemorrhage.

82

83

84 **Keywords:** adrenal hemorrhage, adrenal hematoma, adrenal bleed, adrenal
85 incidentaloma, adrenal tumor, adrenal mass

86

87 **Introduction**

88 Adrenal hemorrhage (AH) can occur as a result of trauma, as well as non-
89 traumatic etiologies including infection, coagulopathy, severe stress, or an
90 underlying neoplasm.¹⁻³ While bilateral AH can be life-threatening and lead to
91 acute adrenal insufficiency,³ unilateral AH is often clinically silent. With the
92 increasing use of diagnostic imaging, particularly computed tomography (CT),
93 identification has become more frequent.⁴⁻⁶ The radiographic appearance of AH
94 can vary widely and is often indistinguishable from adrenal neoplasms or
95 concurrent AH within an underlying neoplasm.⁷ Determining the underlying
96 etiology of AH is important in order to diagnose possible primary or metastatic
97 adrenal neoplasm.^{5,8-10} One literature review found that 83% (110/133) of patients
98 with spontaneous AH had an adrenal neoplasm, of which 48% were
99 pheochromocytoma and 20% were malignant lesions including adrenocortical
100 carcinoma or metastatic disease.¹¹

101 While there are standardized guidelines on the evaluation and management
102 of adrenal incidentalomas,¹²⁻¹⁴ there is a lack of research and evidence-based
103 guidelines on AH. Given the paucity of data on this topic, we performed a
104 retrospective analysis of AH at our institution. Our study aimed to determine the
105 prevalence and etiology of AH, the rate of follow-up, the incidence and type of
106 underlying neoplasm, and patient outcomes.

107

108 **Methods**

109 This study is a retrospective cohort analysis of patients with evidence of AH

110 on imaging performed from January 2006 to October 2021 at our institution, a

111 quaternary care academic health center with high volume endocrine surgery

112 practice and level one trauma center. This study was approved by our institutional

113 review board, which allowed for waiver of informed consent. An institutional

114 database was queried for any radiology report containing the words “adrenal” and

115 “hematoma” or “hemorrhage” within three words of each other. This method was

116 chosen based on our group’s previous experience with our institutional

117 database,^{15,16} and a relatively specific query was chosen due to the relatively high

118 frequency of the words “hematoma” and “hemorrhage” in the database. A

119 secondary screening of all identified reports was performed manually to confirm

120 the read suggested adrenal hemorrhage or hematoma. Any inpatient or

121 outpatient cross-sectional CT imaging that visualized the adrenal glands, including

122 chest, abdominal, or spinal CT was included. If multiple studies were performed

123 within 24 hours, with a different scan ruling out AH, that patient was excluded. To

124 focus this study on patients with “de novo” adrenal hemorrhage, patients with

125 prior known adrenal masses, history of adrenal surgery, and known malignancy

126 were also excluded, as these factors change the index of suspicion for adrenal
127 mass, and therefore clinical decision-making.

128 Electronic charts of identified patients were reviewed for data including
129 patient demographics and medical history, size of hematoma, laterality,
130 biochemical evaluation, and follow-up, including intervention and additional
131 imaging. Interval imaging was defined as additional cross-sectional imaging (CT or
132 MRI) that included the adrenal glands performed at least 21 days after index
133 imaging. This 21-day period was chosen based on our aim to capture scans that
134 were outside of the immediate surveillance period (e.g. for immediate surveillance
135 of traumatic hemorrhage) and that we thought would be able to capture evolution
136 of hemorrhage. For patients with more than one additional scan after the finding
137 of AH, we collected data on the first scan after the 21-day interval and the most
138 recent applicable study in our system. Using the hospital's picture archiving and
139 communicating system (PACS), interval imaging without documented resolution of
140 hemorrhage underwent dedicated review by an attending radiologist with adrenal
141 expertise for evidence of underlying mass.

142 All data was collected and recorded using Research Electronic Data Capture
143 (REDCap).¹⁷ STATA software (StataCorp LLC, College Station, TX, USA) was used to
144 calculate all statistics. Interval data between groups was analyzed using the
145 Wilcoxon rank sum test, and categorical data was analyzed with Pearson's chi

square analysis. Fisher's exact test was used if more than 20% of an analysis table's cells had expected frequencies of less than five. Significance was set at $P < 0.05$. Time to resolution data was generated using Kaplan-Meier survival analysis via Prism 9 (GraphPad, San Diego, CA, USA). Only patients with interval imaging were included in this analysis. Scans with hemorrhage resolution were counted as having a resolution "event," while interval scans without demonstrated resolution were considered a censored observation.

Results

Index Imaging and Presentation

A total of 490,301 imaging reports were queried from our database, yielding 778 possible scans from our search criteria. Secondary manual screening confirmed a total of 579 reports describing AH. After excluding 17 patients with known adrenal masses on prior imaging, 12 patients with prior adrenal surgery, 19 patients with known history of malignancy, and one patient with same-day repeat contrast-enhanced imaging clearly ruling out hemorrhage, 530 (0.11%) scans reported evidence of AH. Imaging was predominately contrast-enhanced CT abdomen with or without pelvis ($n = 500$, 94.3%, Supplementary Table 1). Most scans ($n=485$, 91.5%) were performed during trauma evaluation, predominately following motor vehicle collision ($n=292$, 55.0% of trauma scans) or pedestrian or

bicycle struck by a motorized vehicle (n=90, 17.0% of trauma scans). Of the 213 scans that included measurements, the median maximum diameter of AH was 3cm, with an interquartile range (IQR) of 2.3 - 3.7cm. Full cohort characteristics are described in Table 1.

Two patients underwent dedicated intervention at the time of presentation. One patient, who presented as a trauma with multiple stab wounds, had down trending hemoglobin and active extravasation on CT into a hematoma of the left adrenal gland. Supraceliac angiography also demonstrated contrast extravasation in the region of the left adrenal gland, which was embolized. The patient stabilized and was discharged home after a 7-day hospitalization. No interval imaging was obtained. The second patient had a 9.8cm left adrenal hematoma (no prior trauma, presumed spontaneous) and developed fever and leukocytosis concerning for infection. This patient underwent percutaneous aspiration, with cytology demonstrating "hemorrhagic fibrinous material with histiocytes and neutrophils." The patient was discharged the day after the procedure, and follow-up CT 5 months later showed a 2.7cm evolving hematoma with peripheral calcification.

Follow-up

A total of 114 (21.5%) patients had interval imaging at our institution at least 21 days

186 after index imaging. Initial follow-up imaging was performed at a median of 2.6
187 (IQR 0.99-13.4) months after the index scan. The majority of scans were abdominal
188 (+/- pelvis) CT with contrast (n = 94, 82.5%), and only two patients (1.8%) received
189 dedicated adrenal-protocol CTs (Supplementary Table 1). Forty-six patients (8.9% of
190 total cohort) had more than one follow-up scan, with their most recent imaging at
191 a median of 15.9 (IQR 4.3 – 100.9) months after the index scan. The longest interval
192 was 14.6 years. Patients with non-trauma index scans were more likely to have
193 interval imaging (23/45 , 51.1% vs. 91/485, 18.8%, $p < 0.01$), and AH measuring
194 larger than 3cm were more likely to receive interval imaging than those 3cm or
195 smaller (36/102, 35.3% vs. 23/111, 20.7%; $p = 0.02$).

196 On follow-up imaging, 84 patients (73.7%) demonstrated resolution of
197 hemorrhage at a median of 8.3 (IQR 2.5 – 30.7) months after the index scan. Figure
198 1 demonstrates our time to resolution Kaplan-Meier plot. A representative
199 example of index imaging and resolution is shown in Figures 2a and 2b,
200 respectively. An additional 21 patients (18.4%) showed decreased size of
201 hemorrhage at a median of 1.5 (IQR 1.1-3.6) months following the index scan.
202 There was no difference in resolution or decreased size between patients with AH
203 larger than 3cm and those 3cm or smaller (33/36, 91.7% vs. 18/23, 78.3%, $p=0.14$).

204 Very few patients (10/530, 1.9%) had a note in our system by an outpatient
205 provider that addressed the hemorrhage or possibility of adrenal mass for follow-

up evaluation. These visits were distributed amongst multi-disciplinary providers: surgeon (4 patients), primary care provider, including family medicine or internal medicine practitioner (4 patients), and endocrinologist (2 patients). Only 9 patients (1.7%) had blood and/or urine tests performed for evaluation of adrenal function, with 4 performed during the acute presentation and 5 performed at outpatient follow-up. Only one patient had clear evidence of abnormal function with markedly elevated plasma metanephrines; this patient ultimately had a confirmed pheochromocytoma on pathology following surgical resection.

Patients were more likely to have a follow-up visit or imaging if the radiologist noted concern for an underlying mass on their index imaging (41/118 (34.8%) vs. 77/412 (18.7%), $p < 0.01$) or if a recommendation for additional evaluation was explicitly stated in the report (27/74 (36.5%) vs. 91/456 (20.0%), $p < 0.01$).

Confirmed Adrenal Masses

Thirteen patients (11.0% of the 118 patients with any clinical or imaging follow-up) had confirmed evidence of an adrenal mass, either on serial imaging (n=10) or at adrenalectomy (n=3). One patient was found to have an adrenal adenoma contralateral to the side of AH. Imaging characteristics and biochemical

workup (when applicable) of patients with confirmed masses are described in Table 2.

The final pathology of the three resected glands demonstrated one adrenocortical carcinoma, one pheochromocytoma, and one ganglioneuroma. The first patient was a 44 year-old man who presented with acute flank pain and emesis, found to have a 13.4cm left-sided AH with high suspicion for underlying mass. He was followed outpatient by endocrine surgery with normal biochemical evaluation and repeat imaging 6 weeks after initial presentation that showed resolving hematoma posteriorly but enhancement anteriorly that was concerning for underlying mass of at least 6cm. He underwent open adrenalectomy 2 months after initial presentation, with pathology confirming adrenal cortical carcinoma with oncocytic features. The second patient was a 43 year-old woman who presented with right flank pain, headache, and hypertension and was found to have a 3cm heterogeneous right adrenal mass with surrounding fat stranding. Biochemical evaluation was significant for elevated plasma and 24-hour urine metanephrine and normetanephrine, consistent with pheochromocytoma. She was started on alpha blockade with doxazosin, increasing doses over two weeks, then underwent laparoscopic adrenalectomy approximately one month after initial presentation. Pathology confirmed a pheochromocytoma, Pheochromocytoma of the Adrenal gland Scaled Score (PASS) = 2. The third patient was a 12-year-old girl

245 who presented with acute left upper quadrant pain, nausea, and emesis, and was
246 found to have a ruptured ovarian cyst and 5.4cm left adrenal hemorrhage vs. mass
247 on ultrasound and subsequent CT. MRI was performed two days after admission,
248 consistent with indeterminate, mildly enhancing adrenal mass. She was followed
249 by pediatric surgery, and biochemical evaluation demonstrated normal urine
250 cortisol and metanephrines. Resection was recommended based on size of mass,
251 and she underwent laparoscopic left adrenalectomy 3 weeks after presentation.
252 Final pathology was consistent with a ganglioneuroma.

253 The subset of patients with follow-up evaluation, with and without
254 underlying mass, is described in Table 3. Patients with index scans performed for
255 non-trauma indications were significantly more likely to have an underlying mass
256 ($n=6/26$ (23.1%)) than those performed for trauma evaluation ($n=7/92$ (7.6%),
257 $p=0.04$). Patients with an underlying mass were also more likely be non-Hispanic or
258 Latino ($p = 0.01$) and to have a left-sided AH ($p = 0.04$). The majority of patients with
259 an underlying mass were noted to have concern for a mass on their index imaging
260 and were recommended to undergo additional evaluation (76.9%, 10/13). This was
261 significantly higher than those without an underlying mass who were noted to
262 have concern for mass (29.5%, (31/105)) or were recommended to undergo
263 additional imaging (16.2% (17/105), $p<0.01$ for both.

264

265 Discussion

266 In this study, we aimed to determine the prevalence and etiology of AH, the
267 rate of follow up, the incidence and type of underlying mass, and patient outcomes
268 at our high-volume trauma and endocrine surgery center. AH was a rare
269 phenomenon that was present in 0.1% of CT scans performed over a nearly
270 sixteen-year interval. Our cohort consisted predominately of trauma-associated AH
271 (91.5%), which correlates with the most common etiology of AH reported in the
272 literature,¹⁸ as well as our institution's position as a high-volume trauma center.
273 The rate of underlying mass in our cohort overall was low at 2.3%, but this was
274 limited by a low rate of follow-up, including imaging and/or outpatient clinical
275 evaluation, of 22.3%. Focusing on only patients with follow-up, the rate of
276 underlying mass was 11%. Patients with AH identified on a non-trauma-related
277 index scan were significantly more likely to have an underlying mass than those
278 with trauma-related AH (23% vs. 8%). Notably, both rates are higher than the
279 recently-reported prevalence of adrenal incidentaloma in the general population,
280 0.5-1.4%.^{19,20} While most identified masses were consistent with adenoma or
281 presumed benign due to stability on serial imaging, our cohort included an
282 adrenocortical carcinoma, which is associated with significant mortality, and a
283 pheochromocytoma, which is associated with significant clinical morbidity, risk of
284 malignancy, and is often heritable.¹⁴

285 Due to the heterogeneous appearance of AH on imaging, a single study is
286 often insufficient to differentiate AH from adrenal neoplasms or coexisting AH and
287 neoplasm. Most CT examinations performed for trauma or abdominal pain are
288 acquired during a single phase after IV contrast administration, and differentiation
289 between high density blood and an enhancing mass can be challenging or
290 impossible. Comparison to prior and follow-up imaging can evaluate the expected
291 evolution of AH, including decrease in size and attenuation, liquification,
292 calcification, and often complete resolution over time.^{6,7} In this retrospective study,
293 74% of patients with follow-up imaging demonstrated complete resolution of AH at
294 a median of 8.3 (IQR 2.5 – 30.7) months. Based on this data, our practice in our
295 endocrine surgery team is now to recommend follow-up imaging with an “adrenal
296 protocol” CT approximately 6 months after a finding of AH. Our adrenal protocol
297 specifies an upper abdominal scan with fixed kVp of 120 and slices of 1-3mm with
298 an unenhanced phase, IV contrast injection with 100mL of Omnipaque-350 (GE
299 Healthcare, Chicago, IL) at a rate of 2.5 mL/second, portal venous phase 70
300 seconds after contrast injection, and a 15-minute delayed phase. Interval imaging
301 that is suspicious for an underlying mass is then treated according to guidelines
302 for adrenal incidentalomas with imaging characterization and biochemical
303 evaluation.^{13,14} Imaging consistent with resolving hemorrhage is again followed at
304 6-month intervals until complete resolution is demonstrated. A shorter interval for

305 repeat imaging is recommended if the imaging findings have any features that
306 raise concern for an adrenocortical carcinoma, which can present with
307 intratumoral bleeding, particularly central areas of necrosis and hemorrhage, with
308 contrast retention on delayed contrast-enhanced CT.²¹⁻²³ Though patients with
309 known malignancies were not evaluated in this study, the adrenal gland is a
310 common site for metastases, particularly in melanoma and lung, renal cell, breast,
311 and colon cancer.¹¹ Patients with AH should be asked about history of malignancy,
312 and positive history should raise suspicion for metastatic lesions and shorter
313 interval imaging to evaluate for growth.

314 The characteristics of our trauma-related cohort support previous analyses
315 of trauma populations, indicating that AH is relatively rare, most commonly seen
316 as the result of blunt trauma such as motor vehicle collisions, predominately right-
317 sided, and associated with other injuries.^{4,24-27} Dedicated adrenal acute intervention
318 was rare (2/530, 0.4%) and performed only in the setting of secondary infection of
319 a hematoma or penetrating trauma to the adrenal gland resulting in persistent
320 active bleeding. Though clinically significant, isolated persistent hemorrhage of the
321 adrenal is uncommon, both due to the frequency of coexisting injuries requiring
322 intervention and the typically self-limited nature of AH, adrenal angiography and
323 embolization is an effective minimally-invasive alternative to surgical exploration,
324 as demonstrated in our patient.^{28,29}

325 Our study had multiple limitations. As a retrospective cohort analysis, this
326 study was unable to examine the true natural progression of an adrenal
327 hematoma over time and was limited to often unrelated follow-up imaging at
328 nonstandard time intervals. As our institution is a high-volume level one trauma
329 center, trauma-associated index imaging was predominant in our cohort, with a
330 relatively small volume of non-trauma AH patients. This cohort was also selected
331 based on radiology reports, with no dedicated overreads of the index imaging.
332 Though all reports were signed by an attending radiologist per institutional
333 protocol, variations or mistakes in reporting may have led to selection bias.

334 Finally, the rate of follow-up in this study was low, at 22%, which limited our
335 evaluation of these patients' AH and clinical course. One factor related to this low
336 rate may be the high proportion of trauma patients, who may have been acutely
337 evaluated at our trauma center and had follow-up elsewhere. An institution-wide
338 prospective study of AH is warranted to better evaluate the natural history of
339 hemorrhage and true rate of underlying masses. Notably, patients were more
340 likely to have a follow-up outpatient visit or imaging if their index imaging report
341 specified concern for an underlying mass or explicitly recommended additional
342 evaluation. This again highlights the effectiveness of clear recommendations in
343 radiology reports and suggests that standardized report macros may be useful in
344 improving follow-up of AH. This intervention has been previously demonstrated by

our group in increasing adherence to clinical guidelines for evaluation and follow-up of adrenal incidentalomas.³⁰

Conclusions

AH is a rare imaging finding associated with an increased rate of underlying adrenal mass, particularly when identified on imaging unrelated to trauma. Most AH resolves spontaneously without intervention. Follow-up imaging can evaluate for signs of hemorrhage evolution to distinguish simple hemorrhage from mass-associated hemorrhage and should be considered at an interval of approximately 6 months from index imaging.

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Disclosures

MTC is a consultant for Corcept Therapeutics. The remaining authors report no proprietary or commercial interest in any product mentioned or concept discussed in this article.

365

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367

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467 **TABLES**

468 **Table 1:** Characteristics of patients with adrenal hemorrhage on imaging
 469 performed for trauma and non-trauma indications. Percentages are out of column
 470 header denominator, except as noted. AH= adrenal hemorrhage

Patient Characteristic	Full Cohort (n= 530)		Trauma (n= 485)		Non-Trauma (n=45)		P-value
Age, years, median (IQR)	36	(17-53)	34	(17-51)	44	(32-62)	<0.01
Male, n (%)	347	(65.5)	331	(68.3)	16	(35.6)	<0.01
Race, n (%)							0.2
White	196	(37.0)	176	(36.3)	20	(44.4)	
Black	33	(6.4)	28	(5.8)	5	(11.1)	
Asian	22	(4.2)	22	(4.5)	0	(0)	
American Indian or Alaska Native	6	(1.1)	6	(1.2)	0	(0)	
Native Hawaiian or Other Pacific Islander	6	(1.1)	4	(0.8)	2	(4.4)	
More than one race	4	(0.8)	3	(0.6)	1	(2.2)	
Unknown/Not Reported/Missing	263	(49.6)	246	(50.7)	17	(37.8)	
Ethnicity, n (%)							0.8
Not Hispanic or Latino	273	(51.5)	248	(51.1)	25	(55.6)	
Hispanic or Latino	106	(20.0)	98	(20.2)	8	(17.8)	
Unknown/Not Reported/Missing	151	(28.5)	139	(28.7)	12	(26.7)	
BMI, kg/m², median (IQR)	27.3	(24.0-31.7)	27.2	(24.0-31.6)	29.0	(25.1-34.9)	0.3
Antihypertensive medication, n (% of patients with known medications)*	47	(11.0)	31	(6.4)	16	(35.6)	<0.01
Number of antihypertensives, median (range)	2	(1-4)	2	(1-4)	2	(1-4)	
Anticoagulation medication, n (%)**	15	(3.5)	6	(1.2)	9	(20.0)	<0.01
Antiplatelet medication, n (%)***	17	(3.9)	13	(2.7)	4	(8.9)	0.04
Trauma-related index imaging, n (%)	485	(91.5)					
Motor vehicle collision, n (% of trauma scans)			292	(60.2)			
Pedestrian or bicycle struck by motorized vehicle			90	(18.6)			
Fall from above ground level			42	(8.7)			
Fall from ground level			14	(2.9)			
Assault with knife			3	(0.6)			
Assault other (fight, bludgeoning, etc.)			7	(1.4)			
Gunshot			3	(0.6)			
Other/unknown			34	(7.0)			
Other injuries on scan, n (%)			434	(89.5)			
Non-trauma imaging indications, n (%)	45	(8.5)					

Abdominal pain, n (% of non-trauma scans)					16	(35.6)	
Post-procedure					14	(31.1)	
Concern for bleeding					10	(22.2)	
Other					22	(48.9)	
Right-sided AH, n (%)	367	(69.3)	352	(72.6)	15	(33.3)	<0.01
Left-sided AH	129	(24.3)	106	(21.9)	23	(51.1)	<0.01
Bilateral AH	34	(6.4)	27	(5.6)	7	(15.6)	<0.01
Largest dimension of AH, cm, median (IQR)^	3.0	(2.3-3.7)	3.0	(2.3-3.7)	3.1	(2.3-4.2)	0.7
Concern for adrenal mass noted on index scan, n (%)	118	(22.3)	105	(21.7)	13	(28.9)	0.3
Recommendation for additional evaluation or imaging	74	(14.0)	62	(12.8)	12	(26.7)	0.01
Follow-up scan performed	114	(21.5)	91	(18.8)	23	(51.1)	<0.01

471 *n=101 unknown/missing
472 **n=99 unknown/missing
473 ***n=98 unknown/missing
474 ^ n=272 missing
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477 **Table 2:** Characteristics of cohort with confirmed adrenal masses on imaging or
 478 adrenalectomy

Patient	Age	Sex	Indication for index scan	Laterality of hemorrhage	Size on index scan	Imaging Characteristics	Outcome
1	44	M	Trauma	Right	4.2cm	Adenoma	Stable on serial imaging, normal 24h urine cortisol, no intervention
2	57	M	Trauma	Right	not measured	Myelolipoma	3.3 cm on recent imaging, no intervention
3	53	F	Non-trauma	Left	not measured	Indeterminate mass	Patient died from septic shock 3 months after index imaging
4	43	F	Trauma	Left	2.6cm	Adenoma	Stable on 10-year follow-up imaging, no biochemical evaluation
5	56	M	Trauma	Left	4.1cm	Indeterminate	Stable over 4 years on imaging (contrast-enhanced only), no biochemical evaluation, lost to follow-up
6	58	M	Trauma	Right	3.4cm	Adenoma with hemorrhage	Serial imaging with initial decrease size due to resolving hemorrhage; follow-up MRI confirmed underlying adenoma
7	53	F	Trauma	Left	1.7cm	Indeterminate	Stable over 2 years on imaging (contrast-enhanced only), no biochemical evaluation
8	12	F	Non-trauma	Left	5.4cm	Indeterminate	Adrenalectomy; pathology consistent with ganglioneuroma
9	58	F	Non-trauma	Left	3.6cm	Adenoma with hemorrhage	Smaller on 6-week follow-up imaging, no biochemical evaluation, lost to follow-up
10	44	M	Non-trauma	Left	13.4cm	Indeterminate	Adrenalectomy; pathology consistent with adrenocortical carcinoma
11	43	F	Non-trauma	Right	3.2cm	Indeterminate	Adrenalectomy; pathology consistent with pheochromocytoma
12	69	F	Non-trauma	Left	2.9cm	Indeterminate	Decrease in size over 2 years on imaging, no biochemical evaluation
13	36	M	Trauma	Right	Right: 3.7cm Left: 1cm	Right: hematoma Left: adenoma	Right hematoma resolved on 1 year imaging; small left-sided adenoma stable, biochemical evaluation of plasma metanephrines only - normal

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Table 3: Characteristics of patients with follow-up with and without confirmed adrenal mass. Percentages are out of column header denominator, except as noted. AH= adrenal hemorrhage

Patient Characteristic	Full Cohort (n=118)		No mass (n= 105)		Mass (n=13)		P-value
Age, years, median (IQR)	40.5	(29-53)	40	(27-53)	53	(43-57)	0.06
Male, n (%)	78	(66.1)	72	(68.6)	6	(46.2)	0.1
Race, n (%)							0.07
White	55	(46.4)	46	(43.8)	9	(69.2)	
Black	9	(7.6)	6	(5.7)	3	(23.1)	
Asian	0	(0)	0	(0)	0	(0)	
American Indian or Alaska Native	2	(1.7)	2	(1.9)	0	(0)	
Native Hawaiian or Other Pacific Islander	4	(3.4)	4	(3.8)	0	(0)	
More than one race	1	(0.9)	1	(1.0)	0	(0)	
Unknown/Not Reported/Missing	47	(39.8)	46	(43.8)	1	(7.7)	
Ethnicity, n (%)							0.01
Not Hispanic or Latino	70	(59.3)	58	(55.2)	12	(92.3)	
Hispanic or Latino	27	(22.9)	27	(25.7)	0	(0)	
Unknown/Not Reported/Missing	21	(17.8)	20	(19.0)	1	(7.7)	
BMI, kg/m², median (IQR)	28.3	(24.4-33.9)	27.4	(24.4-33.9)	29.6	(25.1-34.4)	0.5
Antihypertensive medication, n (% of patients with known medications)*	19	(16.1)	14	(13.3)	5	(38.5)	0.1
Number of antihypertensives, median (range)	2	(1-4)	2	(1-4)	2	(2-4)	
Anticoagulation medication, n (%)**	9	(7.6)	9	(8.6)	0	(0)	0.3
Antiplatelet medication, n (%)***	5	(4.2)	5	(4.8)	0	(0)	1.0
Trauma-related index imaging, n (%)	92	(78.0)	85	(81.0)	7	(53.9)	0.04
Other injuries on scan, n (%)	89	(75.4)	83	(79.1)	6	(46.2)	0.6
Laterality, n (%)							0.04
Right-sided AH	68	(57.6)	63	(60.0)	5	(38.5)	
Left-sided AH	37	(31.4)	29	(27.6)	8	(61.5)	
Bilateral AH	13	(11.0)	13	(12.4)	0	(0)	
Largest dimension of AH, cm, median (IQR)^	3.3	(2.4-4.4)	3.3	(2.4-4.5)	3.5	(2.8-4.2)	0.7
Concern for adrenal mass noted on index scan, n (%)	41	(34.6)	31	(29.5)	10	(76.9)	<0.01
Recommendation for additional evaluation or imaging	27	(22.9)	17	(16.2)	10	(76.9)	<0.01

*n=26 unknown/missing

**n=26 unknown/missing

***n=26 unknown/missing

489 ^ n=55 missing
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492 **Supplementary Table 1: Imaging modalities**

Type of Imaging	Index Scan (n = 530)		Initial Follow-up (n = 114)		Most recent (n = 46)	
CT abdomen (+/- pelvis) with contrast, n (%)	500	(94.3)	94	(82.5)	38	(82.6)
CT abdomen (+/- pelvis) without contrast, n (%)	10	(1.9)	6	(5.3)	0	(0)
CT chest with contrast, n (%)	12	(2.3)	9	(7.9)	4	(8.7)
CT chest without contrast, n (%)	3	(0.01)	0	(0)	0	(0)
CT spine, n (%)	5	(4.2)	N/A*		N/A	
Adrenal protocol CT, n (%)	0	(0)	2	(1.8)	0	(0)
MRI (chest or abdomen) with contrast, n (%)	N/A		3	(2.6)	3	(6.5)
MRI (chest or abdomen) without contrast, n (%)	N/A		0	(0)	1	(2.2)

493 *CT spine imaging was excluded on interval imaging as insufficient for adrenal
 494 assessment

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Figures

Figure 1: Kaplan-Meier estimator curve of resolution probability. Timing of interval imaging showing complete resolution of adrenal hemorrhage or hematoma is plotted on the X axis, with the Y axis demonstrating the probability of resolution at the given time. Number at risk at corresponding time points are listed in table below. Figure has been truncated at 100 months, when number at risk is 10.

Figure2 A: Axial slice of abdominal computed tomography imaging demonstrating a 3.8cm right adrenal gland concerning for hematoma or hemorrhage. **B.** Axial slice of abdominal computed tomography performed 5 months after part A demonstrating normal-appearing right adrenal gland with complete resolution of hematoma.