### Sarcomatoid renal cell carcinoma

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October 15, 2020

**RAD 4001** 

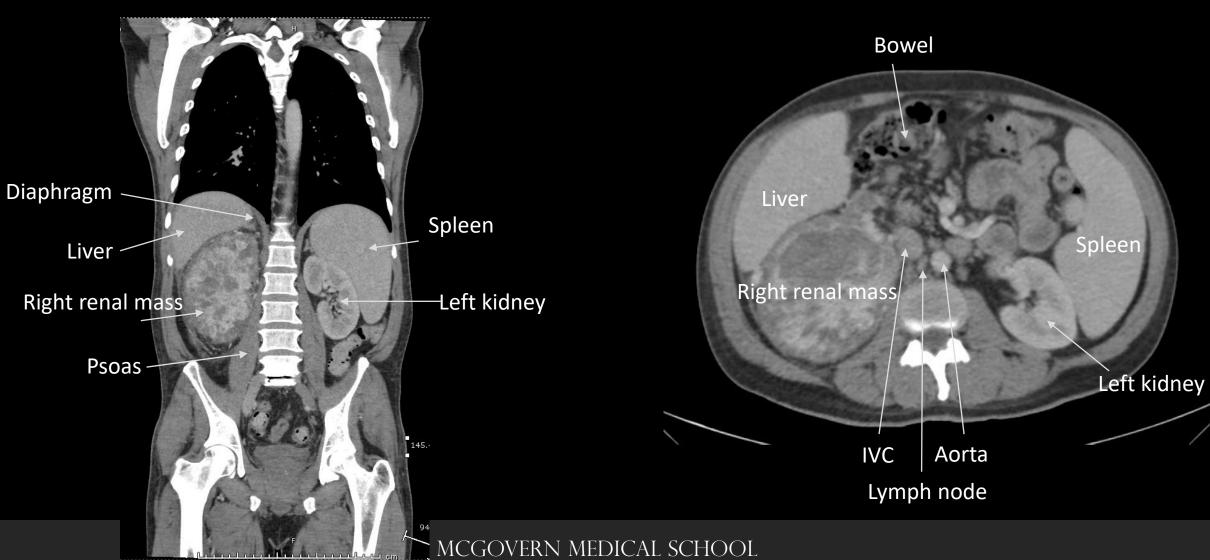
Julia Talley, M.D. and Joseph Hasapes, M.D.



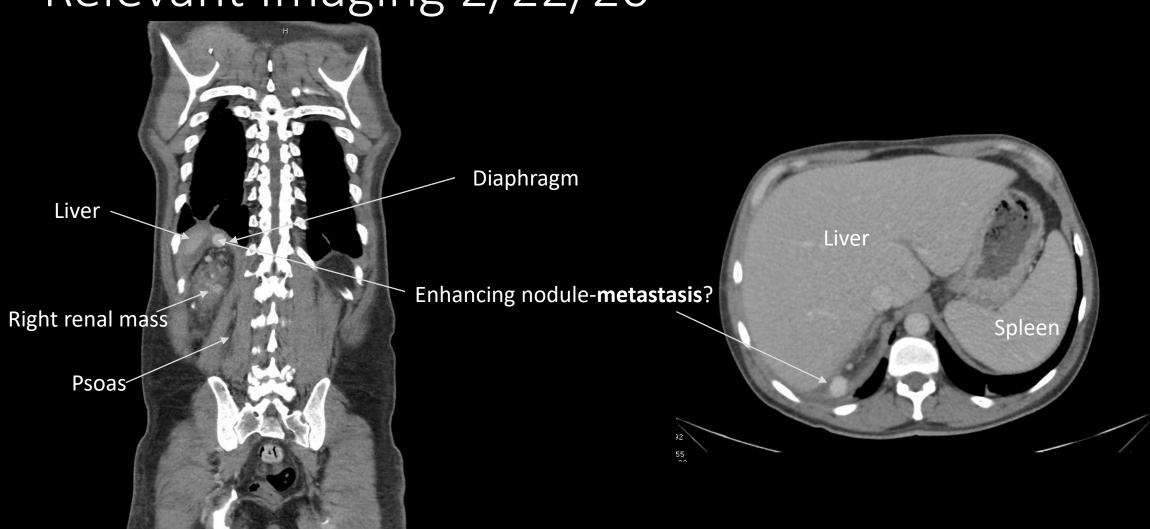
#### Clinical History

- 2/2020: 51 year-old man presents to ED with right flank pain & hematuria
  - PMH: T2DM, HTN, GERD
  - SH: ~30 pack year history, rare EtOH use
  - ROS: Unintentional weight loss (20 lbs in 6 mos); hematuria, dysuria
  - Hgb 4.4
  - Physical exam: TTP in RUQ and R CVA, firm mass palpated R side/flank
  - CT C/A/P ordered

## Relevant Imaging 2/22/20



## Relevant Imaging 2/22/20



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#### Differential Diagnosis

- Renal cell carcinoma
- Oncocytoma
- Medullary carcinoma
- Collecting duct carcinoma
- Transitional cell carcinoma
- Angiomyolipoma
- Lymphoma
- Metastasis

#### Discussion

- Large, solid renal masses with enhancement  $\rightarrow$  RCC until proven otherwise
- As mass size increases:
  - Risk of malignancy increases
  - Likelihood of aggressive pathology increases
- Classic clinical presentation of RCC: flank pain, hematuria, palpable flank mass (5-10% of cases)
- Median age dx: 66. M>F
- Long-term dialysis: 3-6x risk compared to normal population
- Risk factors: smoking, obesity, petroleum exposure, ionizing radiation, etc.
- Tissue required for formal diagnosis—also informs treatment and prognosis
- 5-year OS:
  - >50% if no distant mets
  - <10% with distant metastases

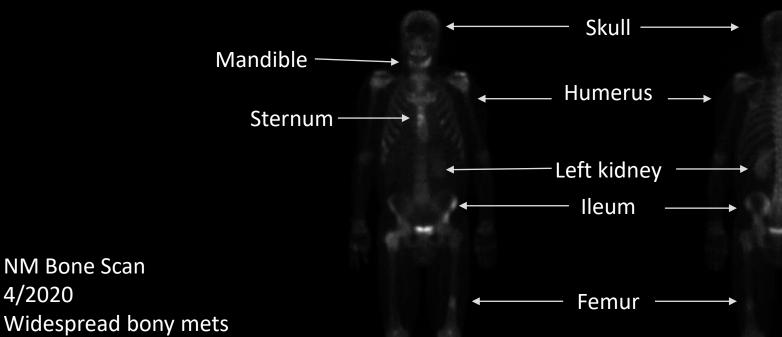
#### Clinical course

- 3/2020: Right radical nephrectomy, adrenalectomy, RPLND, resection of right diaphragmatic nodule, diaphragm repair
  - Pathology: clear cell renal cell carcinoma (WHO grade 4) with sarcomatoid morphology
    - 16.5cm invading in vessel, renal sinus, perirenal fat, and beyond Gerota's fascia. No adrenal involvement. 0/6 LN. Peritoneal biopsy negative. Diaphragmatic nodule positive for RCC → pT4N0M1
- 4/2020: Widely metastatic disease identified [NM bone scan]
- 5/2020: Ipilimumab and nivolumab started
- 7/2020: CT CAP with disease progression → switched to cabozantinib and nivolumab

#### Relevant imaging

NM Bone Scan

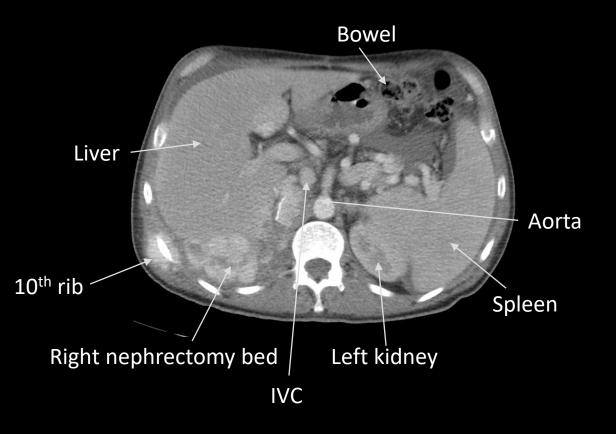
4/2020

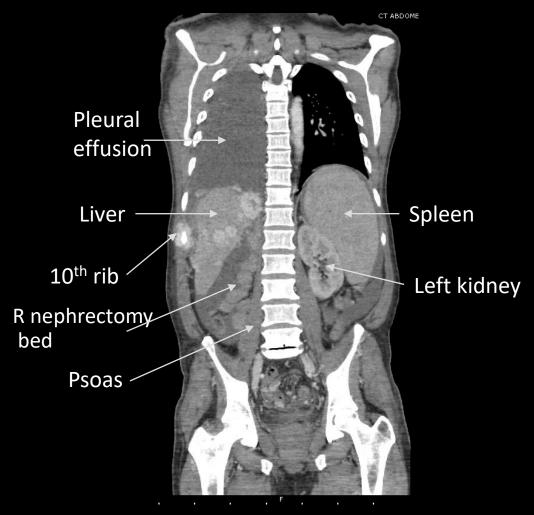


→ 5/2020: Ipilimumab and nivolumab started (3 cycles)

More relevant imaging 7/2020

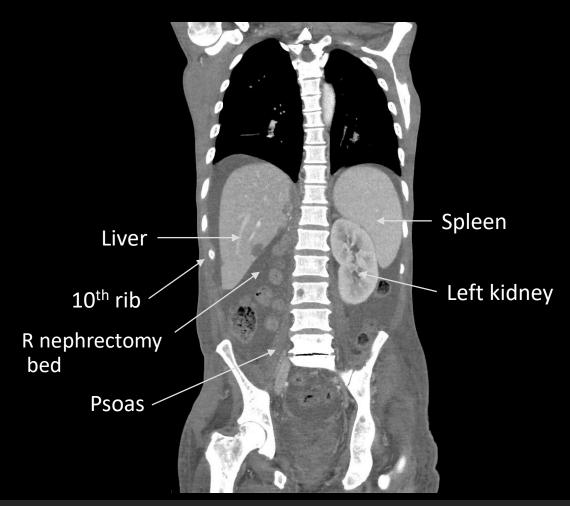
[s/p 3 cycles ipi+nivo]





### More relevant imaging 9/2020

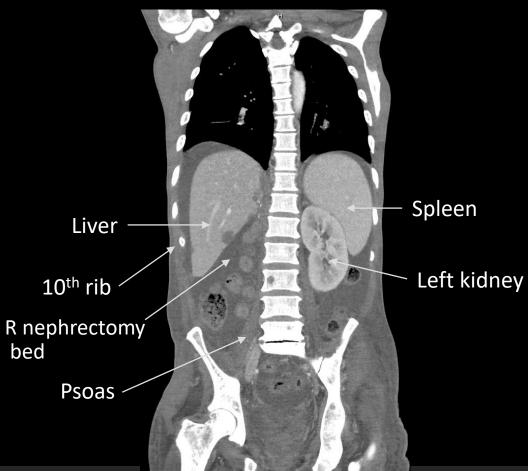
[s/p 3 cycles nivo+cabozantinib]



## More relevant imaging

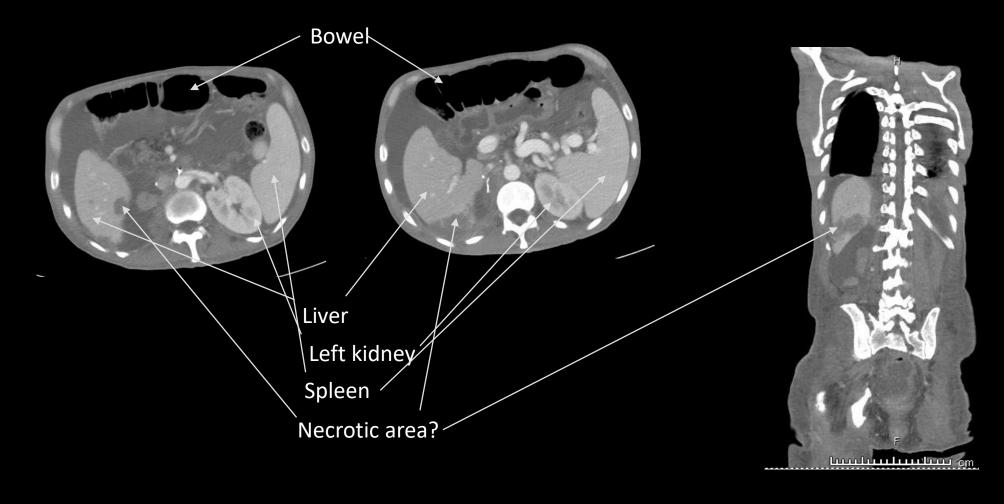
lpi + nivo 7/2020 Pleural effusion -Liver Spleen 10<sup>th</sup> rib Left kidney R nephrectomy bed Psoas -

Cabo + nivo 9/2020



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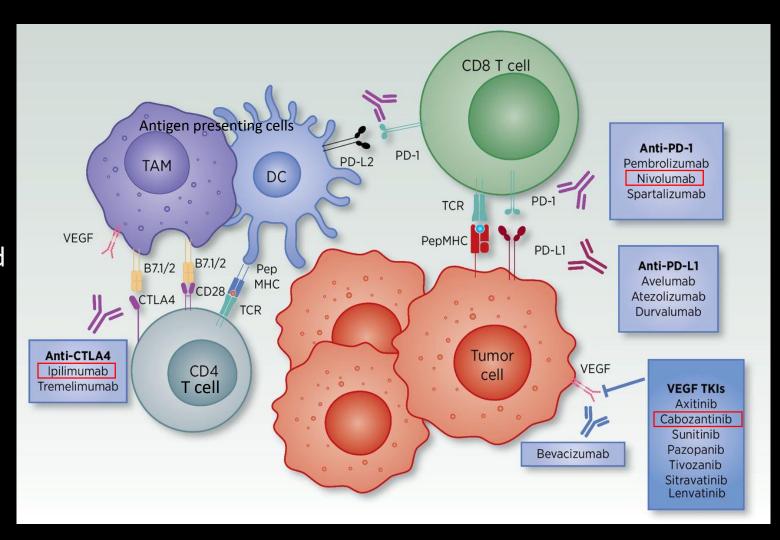
## More relevant imaging 9/30/20





#### Treatment

- Frontline tx RCC: immunotherapy (dual checkpoint inhibition)
  - CheckMate 214 study ipi + nivo vs. sunitinib: improved ORR, OS (Motzer et al, NEJM, 2018)
- Cabozanitib + nivo
  - CheckMate 9ER study: cabozantinib + nivo vs. sunitinib (Choueiri et al, J Ann Onc, 2020): superior PFS, OS, ORR



## Final Diagnosis

Metastatic sarcomatoid renal cell carcinoma responsive to second line treatment

#### ACR appropriateness Criteria

ariant 4: Gross hematuria. Initial imaging.			
Procedure	Appropriateness Category	Relative Radiation Level	
CTU without and with IV contrast	Usually Appropriate	<del>ଡ</del> ଼	
MRU without and with IV contrast	Usually Appropriate	0	
CT abdomen and pelvis without and with IV contrast	May Be Appropriate	***	
MRI abdomen and pelvis without and with IV contrast	May Be Appropriate	0	
MRI abdomen and pelvis without IV contrast	May Be Appropriate	0	
US kidneys and bladder retroperitoneal	May Be Appropriate	0	
CT abdomen and pelvis with IV contrast	May Be Appropriate	999 999	
CT abdomen and pelvis without IV contrast	May Be Appropriate		
Radiography abdomen and pelvis (KUB)	Usually Not Appropriate		
Arteriography kidney	Usually Not Appropriate	<b>⊕⊕⊕</b>	
Radiography intravenous urography	Usually Not Appropriate	***	

#### American College of Radiology ACR Appropriateness Criteria®

Clinical Condition: Renal Cell Carcinoma Staging

Radiologic Procedure	Rating	Comments	RRL*
CT abdomen without and with IV contrast	9	This procedure is complementary to x-ray chest.	***
X-ray chest	8	This procedure is complementary to CT.	<b>⊕</b>
MRI abdomen without and with IV contrast	8	This procedure is an alternative to CT.	0
CT abdomen with IV contrast	7	This procedure is an alternative to CT without and with contrast.	<del>ବଳ</del> ନ
CT chest without IV contrast	6		<b>\$\$\$</b>
CT chest with IV contrast	6		<b>\$\$\$</b>
CT abdomen and pelvis with IV contrast	5		<b>⊕⊕⊕</b>
CT abdomen and pelvis without and with IV contrast	5	This procedure may be appropriate but there was disagreement among panel members on the appropriateness rating as defined by the panel's median rating.	***
MRI abdomen without IV contrast	5		0
Bone scan whole body	5		<b>⊕⊕⊕</b>
MRI head without and with IV contrast	4		0
CT head with IV contrast	4		***
CT abdomen and pelvis without IV contrast	3		ବଳକ
CT chest without and with IV contrast	3		<b>⊕⊕⊕</b>
MRI head without IV contrast	3		0
CT head without IV contrast	3		888
CT head without and with IV contrast	3		<b>⊕⊕⊕</b>
US abdomen	3		0
FDG-PET/CT skull base to mid-thigh	3		***
CT abdomen without IV contrast	2		⊕⊕⊕
Rating Scale: 1,2,3 Usually not appropriate; 4,5,6 M	ay be appropriate; 7	,8,9 Usually appropriate	*Relative Radiation Level

#### Take Home Points

- Large renal mass? Think malignancy
- Imaging of metastatic disease is complex
- Marked response to therapy is possible-even in the presence of diffuse metastatic disease

#### References

- Ng, Chaan S., et al. "Renal cell carcinoma: diagnosis, staging, and surveillance." American Journal of Roentgenology 191.4 (2008): 1220-1232.
- Motzer, Robert J., et al. "Nivolumab plus ipilimumab versus sunitinib in advanced renal-cell carcinoma." *New England Journal of Medicine* (2018).
- Aggen, David H., Charles G. Drake, and Brian I. Rini. "Targeting PD-1 or PD-L1 in Metastatic Kidney Cancer: Combination Therapy in the First-Line Setting." Clinical Cancer Research 26.9 (2020): 2087-2095.
- Choueiri, T., et al. "Nivolumab+ cabozantinib vs sunitinib in first-line treatment for advanced renal cell carcinoma: First results from the randomized phase III CheckMate 9ER trial: 6960\_PR." Annals of Oncology 31 (2020).

# Questions?



**The University of Texas Health Science Center at Houston**  Medical School