Online Supplemental Material

Online supplemental table A. Example quotes related to problem detection: cues that 'flagged' an initial concern and helped the participant notice a renal-drug issue.

THEME	Example(s) from the Incidents					
Vigilance	Medications: "I review meds for each patient towards beginning of patient visit, in part to look					
	for drug-disease interactions and red-flag meds that require renal dosing" [MD]					
	Patient status/renal function: "First thing in morning, I looked for acute changes from overnight.					
	I look at all my patient's renal functions throughout the morning. I want to catch it [any issues]					
	before the 10 am dose" [PharmD], "Intentionally check for patient's renal function during prep					
	for patient appointment" [MD]					
	Specific monitoring: "If patient is on tenofovir, we do an annual urinalysis to look for protein in					
	urine. It's just a screening test that the Department of Health recommends and we do it when					
	they're on the tenofovir" [MD]					
Alert	"I was not aware that this patient had a decreased CrCl –until [the EHR] basically beat me over					
	the head with it, which I appreciated" [PharmD]					
	"I entered the drug valganciclovir, [EHR alert] popped up and said this person has a CrCI [32.5]",					
Desire	"[EHK] caught me" [MD]					
Dosing	buomg Gabapentin Tib is a nerty dose [IVID]					
Abnormal	"It came in for routing visit in April 2012 and did routing labs. That was the first time that I					
Abriormai	really noted that it [SCr] was apparent [1, 5], and over the desired [SCr] for continuation for					
	metformin" [MD]					
value	"Pt's electrolytes being off were another marker that his kidneys were failing K+ was 7.9" [MD]					
Trends in	"Noticed that SCr had now almost doubled 0.9 to 1.6. Meets criteria for acute renal failure"					
laboratory	[PharmD]					
value						
Nephrotoxic	"Not concerned that other meds the patient is taking might be the cause; losartan is 'notorious'					
medication	for causing renal problems. 'Kind of medicine' losartan is, they're called ARBs and ACEs, and					
	those tend to cause injury to the kidneythe creatinine to go up. That's something that					
	happens because of the way that the drug works" [NP]					
New	"Patient had been taking bunch of anti-inflammatories for acute pain 2 weeks prior to					
medication	appointment, thought these were the culprit since had recently started these meds" [MD]					
	"When I saw enoxaparin, I immediately started thinking, was it the right drug for the patient,					
	was it the right route? Was it the right dose?" [PharmD]					
Renally	"Gabapentin is a notorious med that is renally cleared/eliminated so it 'always jumps out'" [MD]					
cleared						
medication						
Risk factor(s)	"Elevated SCr, plus pt being 90 years old tells me patient has CKD" [MD]					
	"So the dehydration, on top of that medicine [Losartan], can cause a problem [renal decline]"					
	"Pt had several reasons for having elevated SCr: patient in OR night before, had a history of					
	nephrectomy [one kidney removed] learned from rounding team, so his creatinine, was					
	potentially going to be elevated anyway, then he for whatever reason was hypotensive as well"					
Subcognant	[Pridimu] ("Dt had standed anti inflammatories, but PUN: 22: SCr: 2.1: both warss [then befare]. Use!th					
dotoction	rt nau stopped anti-innaminatories, but BUN: 33; SCr: 2.1; both Worse [than before]. Health					
uelection						

Tipped off	"When reviewing last renal note, I just happened upon this statement that was in all caps that
	said 'DO NOT SEND HIM HOME ON AN ACE OR AN ARB'" [MD]
	"[Physician] fellow told me SCr was 1.9 and we looked up eGFR. Fellow mentioned patient's
	renal function is getting worse" [MD]

Abbreviations. ACE(s): Angiotensin converting enzyme inhibitors; ARB(s): Angiotensin receptor blockers; BUN: Blood Urea Nitrogen; CKD: Chronic Kidney Disease; eGFR: estimated Glomerular Filtration Rate; EHR: Electronic Health Record; OR: operating room; SCr: Serum Creatinine; TID: three times daily

Participant's type in brackets. MD: physician; NP: nurse practitioner; PharmD: pharmacist

Online supplemental table B. Example quotes related to additional information gathering to respond to the incident

THEME	Example(s) from the Incidents				
Logistics	"I service rejected it back to the provider but it, once again it was a resident and they were here				
	once a week so they didn't see it till the next week and then it actually kind of became this huge				
	snowball effect because it was delayed initially because a resident was involved in ordering the				
	medication" [PharmD]				
	"I do like to keep the file because I feel like sometimes the doctors don't address things				
	appropriately and I have to go back and make sure that they have and in this situation it worked				
	out so" [PharmD]				
	"It was getting to be about the time patient was going to show up; therefore must move quickly				
	to avoid patient receiving higher dose" [PharmD]				
Medication	"For fenofibrate, the cutoff dose is 60. [Below CrCl of 60 fenofibrate needs to be adjusted]				
	looked up appropriate dose in a reference" [PharmD]				
	"Colchicine isn't removed by dialysis source of cue is that the reference says to wait 2 weeks				
	before giving the medication, again" [PharmD]				
	"Pt was taking 2 Aleve BID, 4 Advil BID" [MD]				
Renal	"If CrCl was 20-40, that would match patient's current dose of Piperacillin/tazobactam IV, but				
function	patient's CrCl was below 20. I used CrCl calculator and found CrCl was 18.6 mL/min" [PharmD]				
	"Given patient on dialysis, I estimated GFR to be 10-15 mL/min" [PharmD]				
	"I calculate eCrCL for the older pts. 80, 90 [year old], frail little guys" [PharmD]				
	"Usually look at the last 3-4 labs" [PharmD]				
	"Calculate CrCl because when you look up drug information like in Micromedex or LexiComp for				
	the package insert, when they do talk about renal adjustment, it usually goes off a CrCl, not so				
	much like a SCr, or a eGFR. CrCl is 'like the standard'" [PharmD]				
Renal patho-	"[Order extra labs to test my hypothesis] ordered a urine protein and creatinine, spot urine				
physiology	creatinine, urine eosinophils since anti-inflammatories can cause interstitial nephritis" [MD]				
Risk factor(s)	"Right radical' in notes indicates the patient's whole kidney was removed" [PharmD]				
A					

Abbreviations. BID: twice daily; CrCI: Creatinine Clearance; eCrCI: estimated Creatinine Clearance GFR: Glomerular Filtration Rate; IV: Intravenous

Participant's type in brackets. MD: physician; NP: nurse practitioner; PharmD: pharmacist

Online supplemental table C. Example quotes related to actions taken to help address the incident, including the participants' rationale for their approach.

THEME	Example(s) from the Incidents						
Actions related to treatment interventions							
Avoid risky	"Avoid NSAIDs as an alternative because they are hard on kidneys and this is a dialysis						
medications	patient" [PharmD]						
	"[The] only other fibrate available is gemfibrozil, but we can't use that because can't use						
	gemfibrozil in patients with renal failure" [MD]						
Continue	"I left piperacillin/tazobactam dose the same because [patient] was not at that CrCl cutoff						
medication	yet. Cutoff is CrCl of 20 and patient's [estimated] CrCl was 26" [PharmD]						
	"There aren't a lot of alternative medications for gabapentin to treat peripheral						
	neuropathy – duloxetine or pregabalin. For the alternatives that exist, they require special						
	approval, have to 'jump through hoops'" [MD]						
	"Tenofovir is the backbone of almost all of our regimens and so if you lose it then you're						
	kind of limited by what alternatives you have to treat people with so you have a high						
	threshold for switching that medication out." [MD]						
Medication	"Patient's SCr [is] worse, and CrCl [is] now below the cutoff for Piperacillin/tazobactam so						
dose	[I] adjusted Piperacillin/tazobactam. Reduced it from 3.375 grams every 6 hours to 2.25						
selection	grams every 6 hours" [PharmD]						
	"[I wanted to] avoid splitting the [colchicine] tablet. Patient may not do it right or may get						
	confused. Seek simple dose for patient: easier for patient to take one dose than split pill						
	and take at intervals" [PharmD]						
Restart	"We noticed on the day of discharge he [his blood pressure] started trending up a little						
medication	bit. It was in the 145 over 70 range so it was a good time to restart the lisinopril" [NP]						
Stop	"His renal function was [previously] stable but then when it had that bump when he went						
medication	into renal failure on [date] and had acute kidney injury, I decided that it wasn't worth [it						
	to keep patient on tenofovir containing combo pilij. Wasn't worth [risk of further kidney						
	damage, to keep patient on tenolovirj [IVID] "Cince lab results did not support anti-inflammaterias as source. [1] asked nationt to stop						
	Since lab results did not support anti-innaminatories as cause, [i] asked patient to stop						
	"Diperacillin/tazobactam was stopped because the patient had no medication indication						
	for it" [PharmD]						
Switch	"In order to switch [to abacavir] you have to do a genetic test [HI & B5701] -To make sure						
medication	that he doesn't have a hypersensitivity thankfully he didn't to the new regimen and that						
medication	came back negative so he was ready then to start on [it]" [MD]						
	"Since 3 days and nothing growing, recommended switching antibiotic and de-escalating"						
	[PharmD]						
Non-drug	"I gave him some heel wedges for his shoes which is another non-operative treatment for						
treatment	osteoarthritis of the knee" [MD]						
Actions not related to treatment interventions							
Contingency	"I thought about calcium channel blocker, amlodipine, as an alternative med for						
plan	HCTZ/Lisinopril since it doesn't cause renal impairment as much, but blood pressure was						
	normal and decided to watch blood pressure instead and see if it trends up without						
	meds" [MD]						

	"There were other medications [piperacillin/tazobactam] on his list [besides famotidine]						
	that if it [SCr] had gotten worse would have needed to be adjusted, but I just waited						
	because I was trying to see whether it would get better or worse" [PharmD]						
	"If the patient is more at risk for bleeding and complications then I'll err on the side of						
	caution and adjust the dose [of enoxaparin], if they aren't, then I will follow the package						
	insert and not adjust the dose" [PharmD]						
Counsel	"I told the patient 'look, based on your renal function, you probably shouldn't continue						
patient	with [naproxen]'" [MD]						
	"I told patient I think we should reduce the dose of gabapentin because it can cause						
	grogginess. [I] explain[ed] there is a risk of becoming comatose, unarousable, risk of						
	extreme grogginess and coma." [MD]						
Document in	"I view alerted the provider via an addendum to one of my notes. Goals were: to let them						
EHR	know patient should be on 48 mg fenofibrate (lower dose), to make provider aware of the						
	poor kidney function, so provider can also go back and look at my [original] note [about						
	lowering fenofibrate dose]" [PharmD]						
Follow up	"I would have checked patient's trough next on the 10 th or 11 th , but since pts SCr changed						
	[and kidneys are unlikely to be clearing the vancomycin] I checked the trough earlier in						
	the week [1-2 days earlier than expected] on the 9 th " [PharmD]						
	"So this time I just looked at his labs, because I knew from the day before that it						
	[Piperacillin/tazobactam] was probably going to be a problem" [PharmD]						
	"Schedule patient's follow-up appointment sooner than usual – 2 months instead of 3 or						
	4 - so patient can see dietician and we can check how patient is doing without the						
	fenofibrate" [MD]						
Weigh risk	"I don't think there are any circumstances with the nitrofurantoin and this patient where						
verses	the benefits of the med would outweigh the risks. There are other antibiotics we could						
benefit	give the patient" [PharmD]						
Weigh act	"I deferred to patient's PCP because: patient had a history of tendinitis, I didn't know the						
verses	patient personally, wanted to see if levofloxacin dose should be decreased, given patients						
recommend	decreased kidney function" [PharmD]						
	"[I] changed the gabapentin order myself [rather than relying on original prescriber]						
	because: I get the sense gabapentin is one of those meds that's frequently prescribed and						
	not as much attention [monitoring] paid to it" [MD]						

<u>Abbreviations</u>. CrCl: Creatinine Clearance; EHR: electronic health record; ER: emergency room; HCTZ: Hydrochlorothiazide; HLA: human leukocyte antigen; NSAIDs: Non-Steroidal Anti-Inflammatory Drugs; PCP: Primary Care Physician; SCr: Serum Creatinine; TID: Three Times Daily

Participant's type in brackets. MD: physician; NP: nurse practitioner; PharmD: pharmacist

Online Supplemental Table D. Template for the decision requirements table used in this investigation. Column headings were selected from among those commonly used in decision requirements tables^{1,2} and then adapted for this study as needed. Time expenditures in the last two rows were captured during the interview whenever possible.

Decision or	Why	Cue (Source)	Strategies	Potential	Design Seeds				
Judgment made by	Difficult?	Cue: salient	Used to make	Errors	Design ideas				
the expert; often end	Reasons	information or	the decision	Errors	that may				
in 'ing'	difficult for	signals applied	and the	that could	improve				
	the expert	when making a	actions that	occur or	healthcare				
	or could be	decision	were taken	that non-	systems, care				
	difficult for	(Source: where	to address	experts	coordination,				
	those with	information came	the decision	may tend	IT designs,				
	less	from that was used		to make	etc.				
	experience	to make the							
		decision)							
1. [decision point]									
2. [decision point]									
[rows added to accommodate the number of decision points in the case]									
Time spent looking up information in references:									

Total time spent resolving issue:

REFERENCES

- 1. Militello LG, Klein G. Decision-centered design. *The Oxford handbook of cognitive engineering*. 2013:261-271.
- 2. Crandall B, Klein GA, Hoffman RR. *Working minds: A practitioner's guide to cognitive task analysis.* Mit Press; 2006.