

# Nephrotoxic Drug Event abstractor

# The 20% Information That Solves 80% of the Case

Category	Key Data Element	Why It's High Yield
Patient context	Age, weight, baseline creatinine, baseline GFR	Pediatric renal risk varies significantly with age/weight and baseline function
Drug exposure	Start/stop dates, dose, route for nephrotoxic drug(s) (aminoglycosides, vancomycin, amphotericin, certain antivirals, NSAIDs, chemotherapeutics)	Confirms exposure and duration
Temporal link	Time from drug initiation to first creatinine change	Most nephrotoxic effects occur within predictable time windows
Renal function changes	Daily creatinine trend, GFR changes, urine output	Core criteria for event classification
Alternative explanations	Sepsis, dehydration, surgery with bypass, imaging contrast	Helps rule out drug as primary cause
Intervention	Dose adjustment, drug discontinuation, nephrology consult	Supports causality or mitigation actions taken

# Interesting Questions

Area	Question	Purpose
<b>Definition clarity</b>	“Which nephrotoxic drugs are in-scope for monitoring?”	Ensures consistent case capture
<b>Event criteria</b>	“Do we define an event strictly by lab criteria, or also by clinical diagnosis/documentation?”	Impacts sensitivity/specificity
<b>Overlap with other programs</b>	“How should cases with sepsis or contrast exposure be classified?”	Prevents duplication/conflicting classification
<b>Operational use</b>	“Is the goal to prevent events prospectively or to meet reporting/compliance needs?”	Aligns abstraction approach with use case
<b>Data availability</b>	“Are all creatinine results, urine outputs, and med admin records reliably captured in Epic/Caboodle?”	Identifies feasibility gaps
<b>Alerting thresholds</b>	“Would real-time monitoring be valuable or is retrospective review sufficient?”	Guides automation vs manual review balance
<b>Performance metrics</b>	“Do we track nephrotoxic drug exposure rates per 1,000 patient-days or actual injury rates?”	Helps design dashboards/KPIs

# Few Cases worth reviewing

<b>Case Type</b>	<b>Why</b>
<b>Clear injury, single drug</b>	Example: Vancomycin started, creatinine doubled in 72 hrs, no other insults → confirms positive case definition
<b>Multiple potential causes</b>	Example: Aminoglycoside + recent contrast + hypotension → tests attribution logic
<b>Transient creatinine rise</b>	Example: Amphotericin, creatinine up 0.3 mg/dL but resolves in 24 hrs → clarifies threshold for significance
<b>Borderline timing</b>	Example: Creatinine rises 10 days after drug stopped → tests causal window definition
<b>High exposure, no injury</b>	Example: Multiple nephrotoxic drugs, creatinine stable → reinforces that not all exposures cause events
<b>Data gap case</b>	Missing baseline creatinine or incomplete MAR data → shows abstraction challenges

# Oracle Criteria – is it 1 or more?

Framework	Core Criteria for AKI	Notes for Pediatric Use
pRIFLE	Based on estimated creatinine clearance (eCCl) decrease or urine output: <ul style="list-style-type: none"><li>• Risk: ↓ eCCl by ≥25% or urine output &lt;0.5 mL/kg/h for 8 hrs</li><li>• Injury: ↓ eCCl by ≥50% or urine output &lt;0.5 mL/kg/h for 16 hrs</li><li>• Failure: ↓ eCCl by ≥75% or urine output &lt;0.3 mL/kg/h for 24 hrs</li></ul>	Designed for children, uses Schwartz formula for eCCl
KDIGO	<ul style="list-style-type: none"><li>• Stage 1: ↑ serum creatinine (SCr) by ≥0.3 mg/dL in 48 hrs or ≥1.5–1.9× baseline</li><li>• Stage 2: 2.0–2.9× baseline</li><li>• Stage 3: ≥3× baseline or ≥4.0 mg/dL or initiation of RRT</li></ul>	More widely recognized globally; pediatric adaptation needed for size/weight
NINJA	Focuses on nephrotoxic exposure and daily creatinine monitoring for high-risk patients: <ul style="list-style-type: none"><li>• ≥3 nephrotoxic meds in 24 hrs OR ≥3 days of aminoglycoside therapy triggers daily SCr checks</li></ul>	Used operationally in >30 children's hospitals for prevention

# How do we rule out

<b>Step</b>	<b>Question / Check</b>	<b>Rule-Out if...</b>
<b>1. Exposure verification</b>	Did the patient receive any nephrotoxic drug(s) of interest?	No documented exposure → Not a nephrotoxic drug event
<b>2. Baseline renal status</b>	Is there a valid baseline SCr or eCCI?	No baseline and no surrogate → Cannot confirm event (possible exclude or defer)
<b>3. Timing match</b>	Did AKI occur within the expected timeframe of drug exposure?	Injury occurred before drug start OR well after clearance period → Rule out
<b>4. Objective AKI criteria met</b>	Does patient meet pRIFLE or KDIGO change thresholds?	No qualifying change in SCr/eCCI or urine output → Rule out
<b>5. Alternative causes</b>	Was there a more likely primary cause? (sepsis, dehydration, surgery, contrast)	Stronger alternate etiology, documented by treating team → Rule out as drug-induced
<b>6. Causality plausibility</b>	Was the drug continued despite AKI without worsening, or stopped without improvement?	Weak/no causal relationship → Rule out
<b>7. Data quality</b>	Are labs, med records, and notes complete?	Missing critical data → Mark as “unable to determine,” not a confirmed event

# Quick Cheatsheet

## Rule-Out Reason

No nephrotoxic exposure

Pre-existing renal injury

AKI precedes exposure

Competing etiology

No objective AKI

Non-overlapping time window

Data insufficiency

## Example

SCr rise after contrast only, no listed nephrotoxic meds

Chronic kidney disease baseline

SCr doubled before aminoglycoside started

Severe septic shock + hypotension day before AKI

Creatinine rose 0.1 mg/dL only

AKI 10 days after drug stopped, drug half-life short

Missing med administration record or baseline labs

# Notes to Pull – High Yield

Note type (Caboodle ClinicalNoteFact.Type)	Author type	Window	Why
H&P, Progress, Significant Event	Attending/Resident/APP	<b>Baseline -7d → Last exposure +72h</b>	Baseline SCr/eGFR, dehydration, hypotension, timing narrative (KDIGO).
Nephrology consult	Nephrologist	Exposure → +14d	Causality assessment, dose changes, RRT.
Pharmacy / Therapeutic drug monitoring	Pharmacist	Exposure → +7d	Doses, troughs/peaks (vanc/AG), dose holds.
Procedure/Anesthesia/OR notes	Surgeon/Anesthesiologist	-24h → +72h	Bypass, hypotension, fluid losses (confounders).
ED notes	ED Provider	-24h → +24h	Sepsis, volume status, first doses.
Nursing IPOC / Flowsheet summary notes	RN	Exposure → +7d	UOP documentation gaps, fluid balance.

# Flowsheets (map FlowsheetRowEpicId→label, derive MeasurementValue, constrain by **EncounterKey** + **DateKey window**.)

Measurement (friendly)	Source (typical)	Filter/format	Why
<b>Urine output</b> (mL/kg/hr)	Flowsheet row(s) for UOP/I&O	keep rows with numeric; compute rolling <b>8h</b> minima	pRIFLE: UOP $<0.5 \times \geq 8\text{h}$ .
<b>Intake/Output totals</b>	I&O rows	daily totals	Fluid balance context.
<b>Weight</b>	Vitals	most recent per day	Normalize UOP (mL/kg/hr).
<b>MAP / BP</b>	Vitals	MAP $\leq 65$ (older kids) or age-norm cut	Hypotension confounder.
<b>Temp</b>	Vitals	$>38^\circ\text{C}$ or $<36^\circ\text{C}$	Sepsis confounder.
<b>SpO<sub>2</sub> / FiO<sub>2</sub> / O<sub>2</sub> device</b>	Respiratory rows	as-is	Critical illness context.
<b>Dialysis/RRT start</b>	LDA/Device rows	first start/stop instants	AKI Stage 3 criteria; outcome.

# Labs & levels (AKI + drug causality)

Lab	Caboodle hints	Window	Why
<b>Serum creatinine (SCr)</b>	LabComponentResultFact (SCr component keys)	<b>Baseline –7d → +14d</b>	KDIGO staging ( $\Delta \geq 0.3/48\text{h}$ or $\geq 1.5 \times /7\text{d}$ ).
<b>BUN</b>	component keys	same	Context (pre-renal vs intrinsic).
<b>Cystatin-C (if available)</b>	component keys	same	Pediatric GFR adjunct.
<b>Vancomycin trough/peak</b>	pharmacy/TDM components	exposure ±24–48h	Exposure intensity/causality.
<b>Aminoglycoside levels</b>	gent/tobra/amikacin	exposure ±24–48h	NINJA risk corroboration.
<b>UA (protein, blood, casts)</b>	UA component panel	exposure window	GN vs ATN clues.
<b>Urine sodium / FENa</b>	if recorded	exposure window	Pre-renal vs intrinsic.

# Event Confirmation Questions (Directly map to your rule-in/rule-out algorithm)

Category	Preloaded KG Question	Insight Returned
Drug Exposure	“List all nephrotoxic drugs given in this admission, with start/stop dates, doses, and routes.”	Drug exposure timeline
Renal Baseline	“What was the baseline serum creatinine and eCCI before first nephrotoxic drug exposure?”	AKI baseline reference
Temporal Link	“What is the time between nephrotoxic drug start and first abnormal creatinine?”	Checks causal window
Renal Change Severity	“Does the creatinine change meet pRIFLE or KDIGO criteria?”	Auto-staged AKI
Urine Output	“What is the lowest urine output (mL/kg/hr) recorded during exposure?”	Matches pRIFLE thresholds

## 2. Rule-Out & Confounder Questions - (Help abstractor quickly dismiss non-cases)

Category	Preloaded KG Question	Insight Returned
Alternate Causes	“List other events within 48 hrs of AKI: sepsis, hypotension, surgery, contrast exposure.”	Competing etiology list
Timing Conflict	“Did the AKI occur before nephrotoxic drug administration?”	Early onset check
Recovery Pattern	“Did creatinine return to baseline despite continuing the drug?”	Weak causality indicator
Missing Data	“Which key labs or med administration records are missing?”	Data quality flags

### 3. Clinical Context Insights (Give richer understanding without extra clicks)

<b>Category</b>	<b>Preloaded KG Question</b>	<b>Insight Returned</b>
<b>High-Risk Combinations</b>	“Were ≥3 nephrotoxic drugs given in any 24-hour window?”	NINJA exposure trigger
<b>Cumulative Dose Risk</b>	“Total cumulative dose of each nephrotoxic drug in this encounter.”	Dose-related toxicity risk
<b>Trend Visualization</b>	“Generate creatinine and urine output trend chart aligned to drug exposure timeline.”	Visual causal signal
<b>Care Actions</b>	“Was nephrology consulted within 24 hrs of AKI detection?”	Response timeliness
<b>Patient Outcomes</b>	“Was renal replacement therapy required during admission?”	Severity outcome

# Event Trigger Pipeline

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1	Trigger	New nephrotoxic med ordered, $\geq 3$ nephrotoxic meds/24h, or $\geq 3$ days aminoglycoside (NINJA)	Start job; set lookback window. ( <a href="#">NephJC</a> , <a href="#">ScienceDirect</a> )
2	Gather baseline	Lowest SCr in prior 7d; compute eCCI (Schwartz)	Baseline SCr/eCCI card. ( <a href="#">KDIGO</a> , <a href="#">Renaissance School of Medicine</a> )
3	Build exposure timeline	Meds (start/stop/dose/route) from Caboodle/Clarity feed	Drug timeline chip list. ( <a href="#">UC Davis Health</a> , <a href="#">ehealth.connect-care.ca</a> )
4	Detect AKI by rules	KDIGO: SCr $\uparrow \geq 0.3$ mg/dL/48h or $\geq 1.5 \times /7$ d; pRIFLE: eCCI $\downarrow \geq 25\%$ or UOP $< 0.5$ mL/kg/h $\geq 8$ h	Stage badge (KDIGO/pRIFLE). ( <a href="#">KDIGO</a> , <a href="#">PubMed</a> )
5	Time plausibility	Align first abnormal SCr/UOP vs exposure & drug half-life window	“Plausible timing” tag. ( <a href="#">KDIGO</a> )
6	Confounder sweep	Sepsis, shock, surgery/bypass, contrast within $\pm 48$ h	“Competing etiology” ribbon if present. ( <a href="#">PMC</a> )
7	Response pattern	Did SCr improve after dose stop/reduce? nephrology consult?	“Causality pattern” chip. ( <a href="#">PMC</a> )
8	Relevance rank	Score tiles (see logic table). Only show tiles $>$ threshold; collapse others	Top 3 insights + one-click expand.
9	One-glance summary	20% snapshot auto-filled; rule-in/out suggestion	Green (rule-in), red (rule-out), gray (insufficient). ( <a href="#">KDIGO</a> , <a href="#">Renaissance School of Medicine</a> , <a href="#">NephJC</a> )

# KG – Auto Summary

KG query	Auto-summary tile	When shown
“List nephrotoxic meds with start/stop/dose/route”	Exposure timeline	Always; condensed if low risk. ( <a href="#">UC Davis Health</a> )
“Baseline SCr/eCCI before first exposure”	Baseline card	Always; red if missing. ( <a href="#">KDIGO</a> , <a href="#">Renaissance School of Medicine</a> )
“Does change meet KDIGO/pRIFLE?”	AKI stage badge	If criteria met. ( <a href="#">KDIGO</a> , <a href="#">PubMed</a> )
“Confounders ±48h?”	Competing etiology ribbon	If found. ( <a href="#">PMC</a> )
“Nephrology consult within 24h?”	Response/care action chip	If present. ( <a href="#">PMC</a> )
“Prior AKI in past admissions?”	Recurrence alert	If prior event exists. ( <a href="#">Pediatrics</a> )

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## Relevance scoring (to auto-hide noise) -

This matrix is UI-ready:  
render each row as a tile;  
show only tiles with  
nonzero relevance; color-  
code: green (rule-in), red  
(rule-out), gray (missing).

Signal	Rule	Score
KDIGO met	SCr $\uparrow \geq 0.3/48\text{h}$ or $\geq 1.5 \times /7\text{d}$	+5 <a href="http://renalcareus.baxter.com">renalcareus.baxter.com</a>
pRIFLE met	eCCI $\downarrow \geq 25\%$ or UOP $< 0.5 \text{ mL/kg/h} \geq 8\text{h}$	+4 <a href="#">PMC</a>
NINJA high-risk	$\geq 3 \text{ NTMs}/24\text{h}$ <b>or</b> $\geq 4 \text{ days IV}$ aminoglycoside	+3 <a href="#">PMC</a>
Timing fits	Injury during/ $\leq 72\text{h}$ post exposure	+2 <a href="http://renalcareus.baxter.com">renalcareus.baxter.com</a>
Dominant confounder	Sepsis/shock/contrast/bypass $\pm 48\text{h}$	-4 <a href="#">NaturePMC</a>
Recovery on drug	SCr normalizes while continued	-3 <a href="http://renalcareus.baxter.com">renalcareus.baxter.com</a>

# Query Set – Clean Structured Signal Chain

Stage	Output	Meaning for AI
M0	Cohort of encounters w/ nephrotoxic med exposure	Defines <i>candidate population</i> for review & AI
M1	Relevant labs (SCr, BUN, Cystatin C, drug levels)	Core clinical markers for AKI detection
K1	Timelined SCr values ( $\pm$ window around drug exposure)	Allows temporal modeling and trend detection
K2	Baseline SCr	Patient-specific comparator for AKI definition
K3	First abnormal SCr	Ground truth <i>event onset anchor</i>
K4	Peak SCr	Event severity peak marker
K5	KDIGO Stage (0–3)	Target label for classification / severity prediction

## 2. Turning It into AI-Ready Data

### a. Labeled Supervised Learning Dataset

**Label (y) = KDIGO\_Stage**

**Features (X) =**

- Demographics (age, sex, weight, comorbidities)
- Medications (type, dose, duration, # of nephrotoxic agents)
- Lab trends (baseline → first abnormal → peak)
- Encounter context (unit type, length of stay, surgeries)

Can be used to **predict**:

- Likelihood of progressing from Stage 0/1 to Stage 2/3
- Expected time to AKI onset after first dose

## Temporal Reasoning Inputs for LLMs

- From K1: chronological **SCr time series** relative to drug start
- From M0/M1: **medication–lab event pairs**
  - **Enables reasoning over “drug X given at t0 → SCr spike at t+2 days”**
- Can package as JSON or Knowledge Graph:

# Automated Chart Abstraction Support

- Use **K3** as the "flag event"
- Auto-retrieve notes & relevant labs in the event window
- Feed to LLM with a **prompt structure**:
  - **Event confirmation** (meets KDIGO criteria?)
  - **Rule-out** (contrast nephrotoxic AKI vs sepsis-related AKI)
  - **Context** (recent surgery, dehydration, etc.)
  - **Operational insight** (timing of med adjustments)

## The AI Value

- **LLM-assisted review:** Saves abstractors 50–70% time by pre-highlighting event windows & labs.
- **Predictive alerts:** Build models to flag *likely-to-progress* patients early.
- **Retrospective analytics:** Identify med–lab–outcome patterns for stewardship.

# Silver Objects

SILVER Object

**M0 – NTX\_COHORT\_YTD**

**M1 – Labs\_Inputs\_YTD**

**K1 – KDIGO\_SCr\_Series\_YTD**

**K2 – KDIGO\_Baseline\_YTD**

**K3 – KDIGO\_FirstAbnormal\_YTD**

**K4 – KDIGO\_Peak\_YTD**

What's Inside

Encounter list + med start/stop dates + patient IDs

All raw labs (SCr, BUN, Cystatin C, drug levels) in window

Sequential lab points ± window

Baseline SCr (raw numeric)

First abnormal lab meeting criteria

Peak SCr (raw numeric)

Why Keep Here

Cohort definition from source data

Retain all values, not just abnormal ones

Preserves the original lab history for re-eval

Core calc input; still "direct" derivation

Needed for reproducibility

Base signal for severity calc

# Rule: If it's summarized, labeled, or engineered for decision-making, it belongs in GOLD.

GOLD Object	What's Inside	Why Keep Here
<b>KDIGO_Stage_YTD (K5)</b>	Final Stage (0–3) + baseline + dates	Ready-made classification label
<b>Event Window JSON</b>	JSON block: med/lab timeline, baseline, peak, stage	Plug-and-play for LLM or ML
<b>Confounder Flags</b>	E.g., Sepsis, dehydration, cardiac surgery within ±72h	Precomputed context for faster rule-out
<b>Operational Metrics</b>	Time-to-stage, time-to-med-change, service-line	Supports quality dashboards
<b>Knowledge Graph Nodes</b>	Nodes & relationships for patient-event KG	Enables reasoning queries
<b>Abstraction Packs</b>	Bundled labs, meds, notes for review	Gives abstractor 20% to solve 80% of case