UCONN JOHN DEMPSEY HOSPITAL DEPARTMENT OF PHARMACY M E M O R A N D U M

To: JDH Staff

From: Pharmacy Department RE: IV Benzodiazepine Shortage

Date: July 19, 2022

Situation: The use of lorazepam and diazepam for IM and IV injection needs to be limited.

<u>Background:</u> There is a low supply of IV benzodiazepines due to the ongoing shortage of lorazepam and diazepam injection.

<u>Assessment:</u> Alternative benzodiazepines and other agents need to be used to lower the usage of lorazepam and diazepam injection while there is a shortage.

<u>Recommendation:</u> The following are recommendations being made in response to the benzodiazepine injection shortage. Patient-specific clinical assessment should be used when determining alternative medications.

Supply:

 Reserve a supply of lorazepam injection for behavioral health/emergency departments and diazepam injection for seizure treatment

Alternative Benzodiazepines:

- When IV benzodiazepines are necessary, utilize alternative injectable benzodiazepines. There
 are no direct dose conversions between agents so consider their pharmacokinetic profiles to
 determine dosing (reference table attached below for your reference).
 - Midazolam infusions will be benzodiazepine injection of choice in ICU
- Utilize oral benzodiazepines whenever possible
 - o Including CMO patients
 - Oral lorazepam and diazepam can be used at equivalent doses to IV formulations
- For seizures, benzodiazepines are 1st line.
 - o IV/IM lorazepam remains first line while supply is available
 - In units where IV/IM midazolam and diazepam are available for use, IM midazolam 10mg is the preferred alternative agent.
- Use alternative benzodiazepines for anxiety/sedation
 - Antipsychotics can also be used
 - Multiple agents are available in PO, IM and IV formulations

Alcohol Withdrawal:

- Use adjunctive agents with scheduled benzodiazepines to decrease PRN benzodiazepine needs
 - Gabapentin taper at start of therapy
 - 300-400mg tid x 3 days, then 300mg-400mg bid x 1 day, then d/c
 - Phenobarbital in ICU setting
 - Dexmedetomidine infusion in the ICU setting
 - Haloperidol
 - Oral benzodiazepines such as chlordiazepoxide
 - Oral diazepam/lorazepam as symptom-triggered PRN in mild-moderate withdrawal
 - IV diazepam is reserved for symptom-triggered management in select units (ED, UT2, UT1)

Please note patient specific pharmacokinetic considerations should be considered when converting doses for benzodiazepines (see tables below). For any related questions, contact your floor pharmacist or the Pharmacy Department at x7627

Table 1. Pharmacokinetics of Injectable Benzodiazepines7-11

Agent	Onset of Action (min) Intravenous	Onset of Action (min) Intramuscular	Duration of Action (hours) Intravenous	Duration of Action (hours) Intramuscular	Half-life (hours)	Active Metabolites
Diazepam	1 to 5	slow and erratic absorption	0.3 to 0.5	slow and erratic absorption	20 to 120	Yes
Lorazepam	5 to 20	15 to 30	6 to 8	6 to 8	8 to 15	No
Midazolam	1 to 5	5 to 15	≤ 2a	2a	3 to 11	Yes

a The pharmacologic effect of midazolam may last up to 6 hours in some patients.

Benzodiazepine	Onset of	Half-	Active
	Action/	life	metabolites?
	Time to		
	Peak		
Alprazolam (PO)	1-2 hours	12	No
		hours	
Chlordiazepoxide	1 hour	5-30	Yes
(PO)		hours	
Clonazepam	20-40	17-60	No
(PO)	minutes	hours	
Diazepam (PO,	PO: 15-	PO:	Yes
IM, IV)	30	44-48	
	minutes	hours	
	IM:	IM:	
	unreliable	60-72	
	IV: 1	hours	
	minute	IV:	
		33-45	
		hours	
Midazolam (PO,	PO: 10-	IM:	Yes
IM, IV)	20	up to	
	minutes	6	
	IM: 15	hours	
	minutes	IV:	
	IV: 1-5	less	
	minutes	than	
		2	
		hours	
Oxazepam (PO)	3 hours	8	No
		hours	
Temazepam	1-2 hours	4-18	No
(PO)		hours	