

Center for Gastrointestinal Health

Atlas Diagnostic Imaging

Medications and Contrast Media Guidelines

For CT and MRI

In adults

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I. Introduction

Contrast medium is a substance used to increase the contrast of structures or fluid within the body and it is used to improve medical imaging. Several forms of contrast media can be administered intravenously, intra-arterially, or intraluminally. Contrast media is considered a pharmaceutical agent and like all other pharmaceuticals, contrast media is not completely devoid of risk. The purpose of this guideline is to assist the radiologist in managing the small but real risks inherent in the use of contrast media.

II. Patient selection

Before any administration of contrast media, the radiologist should consider the following:

1. Assessment of risk versus benefit of the contrast administration.
2. Alternate imaging that would provide the same or better diagnostic information without need for contrast media.
3. Valid clinical indication for contrast administration.

III. Intravenous iodinated contrast media

The following parameters will be used when assessing patients requiring IV iodinated contrast media:

1. All patients with a serum creatinine (regardless of age) which measures **< 2.0 mg/dL (and/or eGFR > 40)** are eligible for intravenous iodinated contrast administration.
2. All patients **>60 years old** require a serum creatinine (and/or eGFR) performed within the last **30 days**.
3. Patients **< 60 years old**, scheduled for a routine intravascular study, and do not have one or more risk factors (listed below), do not require a baseline serum creatinine (and/or eGFR) determination before iodinated contrast medium administration.
4. Administration of intravenous contrast with serum creatinine **≥ 2.0 mg/dL (and/or eGFR ≤ 40)** requires radiologist's approval. The radiologist should consider discussing the risks of contrast-induced nephropathy (see page 12) with a member of the patients care team which may include the requesting physician or physician extender prior to approval of contrast administration.

Risk factors that may require serum creatinine determination:

Chemotherapy: All patients receiving nephrotoxic chemotherapy require a serum creatinine performed since the last dose.

Metformin: Patients on metformin require serum creatinine and estimated GFR. See page 11 for more information.

Renal disease: Patients with a history of significant renal disease (e.g. may result in impaired renal function), nephrectomy, kidney transplant, or recognized upward trend in creatinine may have a point of care creatinine performed at the discretion of the nurse, technologist, radiologist, or ordering provider

Hypertension requiring medical therapy, diabetes mellitus, or gout:

patients with one or more of these three conditions may have a point of care creatinine performed at the discretion of the nurse, technologist, radiologist, or ordering provider.

Recent intravenous contrast: All patients who have received IV iodinated contrast in the last 24 hours require approval by a radiologist for additional intravenous contrast media administration and may have a point of care creatinine performed at the discretion of the nurse, technologist, radiologist, or ordering provider.

Emergency patients

The ordering physician can choose to bypass screening in an emergency and have IV contrast administered without screening. This screening process bypass must be documented by the nurse, technologist or the ordering physician. The ordering physician's name must be included in the documentation.

Gadolinium:

Eligibility criteria for administration of intravenous gadolinium are described in the section on Gadolinium guidelines (see page 13)

Specific conditions:

Allergic-like reactions and premedication protocols—see pages 6-8.

Renal insufficiency—see page 12.

Pregnancy or breast-feeding—see page 17-18.

Sickle cell disease/trait: There is no evidence for any clinically significant risk, particularly after the injection of low-osmolality contrast media (1, 2).

Myasthenia Gravis: This condition may be a relative contraindication for intravenous administration of iodinated contrast media (1, 3).

Thyroid Disease:

a. Hyperthyroidism: patients with hyperthyroidism or other thyroid disease can potentially experience iodine-provoked delayed hyperthyroidism. This effect may appear 4-6 weeks after IV administration of iodinated contrast media. This condition is usually self-limited. However, patients with history of hyperthyroidism should follow-up with their endocrinologists after receiving iodinated contrast media (1, 4).

b. Thyroid carcinoma: Iodinated contrast media may interfere with both diagnostic scintigraphy and radio-iodine treatment. Therapeutic uptake of I¹³¹ radioiodine therapy may be decreased substantially after iodinated contrast injection. Patients are required to wait a minimum of 4 weeks (preferably 6 weeks) after receiving intravenous iodinated contrast administration, before undergoing either I¹²³ diagnostic scintigraphy or I¹³¹ radioiodine therapy (1, 4).

Multiple myeloma: Paraproteinemias, such as multiple myeloma, has previously been considered a risk factor for developing contrast-induced nephropathy after high-osmolality contrast media. However, there are no data predicting risk with the use of low-osmolality or iso-osmolality intravenous contrast media (1). Patients with multiple myeloma may receive iodinated IV low-osmolality or iso-osmolality contrast if they are adequately hydrated and not significantly hypercalcemic (5).

Pheochromocytoma: Previous investigations have shown no clinical effect from injection of low osmolality contrast media in patients with pheochromocytoma (6). However, direct injection of high or low osmolality contrast media into the adrenal or renal artery is to be avoided because of the risk of causing a hypertensive crisis (1)

Dialysis: Most low-osmolality iodinated contrast media are not protein-bound, have relatively low molecular weights, and are readily cleared by dialysis. Unless a usually large volume of contrast medium is administered, or there is substantial underlying cardiac dysfunction, there is no need for urgent dialysis after IV contrast administration(1).

I. Allergic-like reactions to contrast media

A history of prior allergy-like reaction to contrast media is associated with up to a five-fold increased likelihood of the patient experiencing a subsequent reaction. Patients who describe an "allergy" to contrast media or other substances (food, medication) should be questioned further to clarify the type and severity of the "allergy" or reaction. A patient with a history of anaphylaxis or life-threatening reaction to contrast media should not receive contrast media, regardless of prep. The patient records should be reviewed to see that this allergy is appropriately documented.

Please see Appendix A for additional information regarding allergic-like reactions to contrast media.

II. Categorization: Acute adverse events can be categorized as either allergic-like or physiologic, and classified into three categories: mild, moderate, or severe (1).

Mild:

Allergic-like: limited urticarial/pruritus, cutaneous edema, "itchy"/
"scratchy" throat, nasal congestion, sneezing, rhinorrhea

Physiologic: limited nausea/vomiting, transient flushing, headache,
dizziness, mild hypertension, vasovagal reaction that resolves spontaneously

Moderate:

Allergic-like: diffuse urticarial/pruritus; diffuse erythema (stable vital signs),
facial edema or throat hoarseness without dyspnea, wheezing with no
hypoxia

Physiologic: protracted nausea/vomiting, isolated chest pain, vasovagal
reaction requiring treatment

Severe:

Allergic-like: diffuse or facial edema with dyspnea, diffuse erythema with
hypotension, laryngeal edema with stridor and/or hypoxia,
wheezing/bronchospasm with hypoxia, anaphylactic shock (hypotension +
tachycardia)

Physiologic: vasovagal resistant to treatment, arrhythmia, seizures,
hypertensive emergency, cardiovascular collapse

I. Premedication guidelines

Steroid prep needed (if no prep given, consult radiologist; if prep given, do not consult radiologist):

- a. Prior allergic-like contrast reaction (mild, moderate) to the same class of contrast media agent (iodinated / gadolinium)
- b. Actively asthmatic (e.g., in the ER with active asthma exacerbation)
- c. Prior mild breakthrough reaction. Note: an individual radiologist may elect to not perform the examination in this setting. This decision should be coordinated such as during ordering/scheduling process.

Steroid prep not needed (may scan patient without consulting radiologist):

- a. Physiologic reaction to contrast media such as nausea or vomiting
- b. Seafood or shellfish allergy (mild or moderate)
- c. Allergy to a different type of contrast agent (e.g., allergy to gadolinium if receiving iodinated contrast or allergy to iodinated contrast if receiving gadolinium)

- Steroid prep possibly needed (consult radiologist if no prep given):**
- a. One or more allergies to any substance(s) except for the same type of contrast material (which would necessitate a steroid prep, see 1a above) and also excluding seafood or shellfish (see 2b above)
 - b. Asthma

Contrast media administration is relatively contraindicated (consult radiologist):

- a. Prior severe reaction to any substance (***active angioedema, laryngeal edema, anaphylactic shock***) including any type of contrast media agent (iodinated or gadolinium); if such a history is present, please consult a radiologist
- b. Prior moderate or severe breakthrough reaction. For prior mild breakthrough reaction, contrast media may be administered after a steroid prep (see 1c above)

Miscellaneous:

- a. If an alternate prep has been given, consult the radiologist. If all scheduled doses of a full steroid prep (12-13 hours) have been given but the timing is off by 2-3 hours, no consultation is required and the scan can be performed. The initial dose of steroids must precede contrast material administration by not less than 4 hours.
- b. Using Visipaque (iodixanol) may somewhat reduce the risk of a contrast reaction in patients with a history of allergic-like reaction to iodinated contrast, but the data supporting this is weak and the practice is not required.

II. Elective Prep: 13-Hour Prep (Greenberger Protocol)

1. Adults:

Prednisone 50 mg (oral) q 6 hours x 3 doses starting 13 hours prior to scan:
13 hours + 7 hours + 1 hour prior to scan

Optional: Benadryl 50 mg maximum dose 1 hr prior to exam

Steroid Equivalences

Decadron (dexamethasone) 8 mg X 3 doses, IV or oral

OR

Solu-Cortef (hydrocortisone) 200 mg X 3 doses, IV or oral (IV preferred)

OR

Solu-Medrol (Methylprednisolone) 40 mg x 3 doses, IV or oral (IV preferred)

Total Doses required for full strength prep prior to contrast media

50 x 3	= 150 mg Prednisone
8 x 3	= 24 mg Decadron (Dexamethasone)
200 x 3	= 600 mg Solu-Cortef (Hydrocortisone)
40 x 3	= 120 mg Solu-Medrol (Methylprednisolone)

III. Emergency Prep: 4-hour Prep

Use the following prep when you do not have 13 hours to follow the Greenberger protocol listed above.

Adults:

Solu-Medrol (Methylprednisolone) 60 mg IV Q 4 hours x 2 doses prior to contrast administration (the first dose is given 4 hours prior to the scan and the second dose is given before the patient is put on the CT table)

Benadryl (Diphenhydramine) 25-50 mg IV one-hour prior to scan (per Radiologist)

I. Contrast Extravasation

Risk Factors:

- Non-communicative patients: non-English speaking and unconscious
- Small peripheral veins (hands and feet)
- Injection of an older IV line
- Multiple attempts at IV access
- Abnormality in limb to be injected (trauma, lymphedema, etc.)
- Higher injection rates (4-5 mL/sec)

Sequela of Extravasations:

Iodinated contrast media is toxic to surrounding tissues/skin resulting in an acute local inflammatory response. The vast majority of patients recover with no significant injury.

Possible significant injuries include:

1. Compartment syndrome: more likely to occur with large volumes or injection in a small tight space (i.e. ventral or dorsal surface of wrist)
2. Skin ulceration/blister/tissue necrosis

Actions/Treatment:

Call referring physician to notify of event and severity.

Evaluate and observe patient in the Department of Radiology as appropriate depending on amount and severity of extravasation

Physical Examination

- Confirm pulses in affected limb
- Assess skin color/sensation and monitor for change by comparing to unaffected limb
- Elevate affected extremity
- Apply cold compresses

Indications for Surgical Consult if:

- Increasing pain over 1-2 hours
- Increasing pain on passive stretch of flexor or extensor tendons
- Skin blistering
- Altered tissue perfusion by decrease in capillary refill
- Change in sensation or decrease in sensation distal to site of extravasation

Provide the patient with radiology contact information before release and confirm that patient is aware of the need to contact the facility if she/he develops:

- Significant residual pain or increasing pain
- Skin changing (including blistering)
- Hardness or drop in temperature sensation at extravasation site
- Change in sensation site of extravasation

Document extravasation as incident report

I. Metformin Guidelines (1)

Metformin (see Appendix C for brand names) is an oral anti-hyperglycemic agent used primarily to treat insulin resistant diabetes mellitus. The most significant adverse effect of metformin therapy is the potential for the development of metformin-associated lactic acidosis. Iodinated contrast media is not an independent risk factor for patients taking metformin but is a concern only if post-contrast acute kidney injury should develop.

C **Metformin and Iodine-based Contrast Agents:**

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Category 1: In patients with no evidence of acute kidney injury and with eGFR ≥ 45 mL/min/1.73m², there is no need to discontinue metformin either prior to or following the intravenous administration of iodinated contrast media, nor is there an obligatory need to reassess the patient's renal function following the test or procedure.

Category 2: In patients taking metformin who are known to have acute kidney injury or chronic kidney disease (eGFR < 45), or are undergoing arterial catheter studies that might result in emboli (atheromatous or other) to the renal arteries, metformin should be temporarily discontinued at the time of or prior to the procedure, and withheld 48 hours subsequent to the procedure and reinstated only after renal function has been re-evaluated and found to be normal. Patients should be given written information to contact their PCP (primary care provider).

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Metformin and IV Gadolinium: It is not necessary to discontinue metformin prior to contrast medium administration when the amount of gadolinium-based contrast material administered is in the usual dose range of 0.1 to 0.3 mmol per kg of body weight.

I. Renal Insufficiency and Contrast Induced Nephropathy

Contrast-induced nephropathy (CIN) is a specific term to which is used describe a type of post-contrast acute kidney injury (PC-AKI) where the deterioration in renal function is directly attributed to the administration of contrast media.

The most important risk factor for CIN is pre-existing severe renal insufficiency. A threshold of eGFR of $< 30 \text{ mL/min/1.73m}^2$ has been proposed as the limit below which intravenous contrast administration becomes a risk factor for developing PC-AKI and CIN(1).

II. Acute Kidney Injury Network (AKIN) definition of acute kidney injury:

1. Absolute serum creatinine increase $\geq 0.3 \text{ mg/dL}$.
2. A percentage increase in serum creatinine $\geq 50\%$ (≥ 1.5 -fold above baseline).
3. Urine output reduced to $\leq 0.5 \text{ mL/kg/hour}$ for at least 6 hours.

III. Clinical course: The clinical course of CIN or PC-AKI depends on the baseline renal function, coexisting risk factors, degree of hydration, and other factors. However, the usual course consists of a transient asymptomatic elevation in serum creatinine. Serum creatinine usually begins to rise within 24 hours of IV iodinated contrast administration, peaks at 4 days, and often returns to baseline within 7 to 10 days. It is unusual for patients to develop permanent renal dysfunction.

IV. Treatment: The treatment of CIN or PC-AKI is largely supportive. The major preventative action to mitigate the risk of CIN is to provide intravenous volume expansion (see hydration protocol, page 21). One possible protocol would be 0.9% saline at 100 mL/hr, beginning 6-12 hours before and continuing 4-12 hours after iodinated contrast administration. This protocol is only practical in the inpatient setting.

I. Gadolinium Contrast Use and Nephrogenic Sclerosing Fibrosis

The association of nephrogenic sclerosing fibrosis (NSF) and gadolinium is well documented. However, the precise mechanism of the relationship is controversial and incompletely understood. One hypothesis is that the development of NSF is related to the release of gadolinium ions from the chelates in gadolinium-based contrast agents (GBCA), in patients with decreased renal function and prolonged clearance time of GBCA from the bloodstream. Our gadolinium guidelines are therefore directed at identifying patients who are at risk for NSF.

II. Gadolinium Guidelines

Background

This document is a guideline for administration of intravenous (IV) contrast agents during MRI in particular patient populations (particularly individuals with compromised renal function) for the Department of Radiology, Duke University Medical Center. The choices of whether to administer a contrast agent and the type of agent should ultimately be guided by patient clinical needs. Any MRI area in which IV contrast agents (or other medications) may be administered must have a supervising physician who is prepared for the evaluation and treatment of idiosyncratic and allergic reactions. Such measures and policies are not described in detail in this document.

Note that the following guidelines assume stable renal function over a period of several months. If there is any suggestion that the patient may have sustained an acute kidney injury, a point of care (POC) estimated glomerular filtration rate (eGFR) should be measured within 24 hours of the scan, and the decision regarding contrast media administration made in light of the most recent eGFR value and its trend over time. The presence of an acute kidney injury in and of itself, regardless of the eGFR, may represent a relative contraindication to the administration of gadolinium-based contrast agents (GBCA).

MRI contrast agents available at Duke

Gadolinium-based agents (GBCAs):

Eovist (gadoxetate disodium)

Gadavist (gadobutrol)

Magnevist (gadopentetate dimeglumine)

MultiHance (gadobenate dimeglumine)

ProHance (gadoteridol)

Non-Gadolinium-based agent (off-label use as contrast agent):

Feraheme (ferumoxytol)

Patients Requiring a Serum eGFR Measurement

- 1) All patients >60 years old; eGFR within last 30 days.
- 2) All patients receiving nephrotoxic chemotherapy; eGFR since the last dose of nephrotoxic chemotherapy.
- 3) Any patient with a history of renal disease, nephrectomy (complete or partial), kidney transplant, recognized downward trend in eGFR or upward trend in creatinine; eGFR within last 30 days or since last kidney-related intervention/event.

Categorization of patient renal function*

- eGFR > 60 mL/min/1.73m²: Normal function (**category 0**)
- eGFR 40-59 mL/min/1.73m²: Normal to mildly impaired (**category I**)
- eGFR 30-39 mL/min/1.73m²: Moderately impaired, borderline (**category II**)
- eGFR < 30 mL/min/1.73m²: Severely impaired (**category III**)
(note: patients on chronic dialysis are also considered to fall within **category III** regardless of eGFR)

*Note that the above categorization system is for internal use only, and does not correspond to the National Kidney Foundation system for staging chronic kidney disease.

Guidelines

Category 0 (eGFR ≥ 60 mL/min/1.73m², measured within 3 months): Normal renal function. GBCAs may be administered up to standard dosages.

Category I (eGFR between 40 and 59 mL/min/1.73m², measured within 1 month): Normal to mildly impaired renal function. Risk of developing nephrogenic systemic fibrosis (NSF) from GBCA administration in this patient population is extremely low. GBCAs may be administered up to standard dosages.

Category II (eGFR between 30 and 39 mL/min/1.73m²): Borderline renal function. Importantly, it has been shown that eGFR measurements can fluctuate from day to day. As a result, these patients must have an eGFR rechecked within 24 hours of potential GBCA administration; this will be done via POC iSTAT test when the patient arrives in MRI for their scan if no other qualifying eGFR measurement is available. Those with an eGFR ≥ 30 mL/min/1.73m² within 24 hours of GBCA administration are managed as **category I** renal function. Those with eGFR < 30 mL/min/1.73m² are managed as **category III**. If it is not possible to check the eGFR within 24 hours of MRI, then these patients are considered **category III** for purposes of potential GBCA administration.

Category III (eGFR < 30 mL/min/1.73m²): Severely impaired renal function. Note that since such severe renal insufficiency is typically chronic, a recent (within 3 months) eGFR measurement is only required if a provider has reason to believe that the patient's renal function may have improved. Administration of ANY GBCA is relatively contraindicated in this patient population. Alternative techniques, such as non-MRI investigations, non-contrast MRI, or administration of an iron-based agent (ferumoxytol)

should be considered. In certain scenarios, where the future risk of development of NSF is outweighed by the need for GBCA administration, it may be reasonable to administer a GBCA. The following precautions must be observed:

- 1) A discussion of the risks and benefits must be undertaken with the referring attending physician, and a note must be placed in the medical record by the referring attending physician to document that the benefits of the examination are believed to outweigh the risks.
- 2) **For patients with eGFR < 15 mL/min/1.73 m²:** A discussion of the risks and benefits must be undertaken with the patient (or designee) by a physician member of the Radiology department or the referring clinical team. The patient (or designee) must provide written informed consent, and the scanned into PACS or MaestroCare. If the consent document is obtained by the referring team but documentation cannot be verified at the time of MRI examination, written consent must be re-obtained in MRI prior to GBCA administration.
- 3) Prior to GBCA administration, the non-contrast portion of the examination must be evaluated by a Radiologist to determine whether GBCA administration is necessary.
- 4) Following the above, if deemed necessary, GBCA may be administered at the lowest reasonable dose that is expected to yield a diagnostic examination and answer the clinical question, as determined by the supervising Radiologist. The choice of GBCA will depend upon the clinical situation, however agents with favorable safety profiles (MultiHance, ProHance, or Gadavist) should be used whenever possible. Magnevist is contraindicated for patients in this category of renal function, in concordance with the Food and Drug Administration's guidelines.
- 5) The Radiologist reporting the MRI result must clearly summarize the steps taken above in their report, including verification of informed consent (if applicable) and the note from the referring physician.

While dialysis following GBCA administration has not shown clear benefit, the risks and benefits of dialysis should be considered following GBCA administration to patients who receive dialysis.

eGFR Quick Reference				
	Category 0	Category I	Category II	Category III
Prior eGFR	eGFR ≥ 60	40 ≤ eGFR < 59	30 ≤ eGFR < 39	eGFR < 30 or chronic dialysis
Time eGFR is valid	3 months	1 month	≤ 24 hours	Any time (most recent eGFR)
Action	Normal dose GBCA	Normal dose GBCA	eGFR must be within 24 hours; if contraindicated most recent eGFR ≥ 30, manage as category I; otherwise manage as category III	GBCA relatively needed, follow steps 1-4 above

Ferumoxytol

Ferumoxytol is an iron-based agent with no theoretical increased risk of NSF. It is a bloodpool agent and has shown utility in vascular MRI. Use of ferumoxytol for non-vascular indications should be avoided due to its high cost and the uncertainty of incremental benefit from its use. The final determination regarding its suitability for use as a contrast agent is the responsibility of the individual divisions/protocoling physicians.

Pregnancy

Gadolinium chelates may accumulate in the amniotic fluid, and the potential effects on the pregnancy are unknown. As a result, GBCA administration is contraindicated in pregnant or potentially pregnant patients. If it is felt necessary to administer a GBCA to such a patient, this process is managed identically to that described for patients with category III renal dysfunction, as above.

Breast-feeding mothers

Given the tiny amount of GBCA excreted in breast milk, the even smaller amount expected to be absorbed by a breast-feeding infant's GI tract, and the absence of any evidence of toxicity to breast-feeding infants in the literature, administration of GBCA at typical doses is considered safe in breast-feeding mothers. If a breast-feeding mother remains concerned about potential ill effects, she may express and discard breast milk ("pump and dump") for 24 hours following GBCA administration to further reduce the already small risks associated with GBCA administration.

Reference

ACR Manual on Contrast Media v10.1

<http://www.acr.org/~media/37D84428BF1D4E1B9A3A2918DA9E27A3.pdf>

I. Pregnant or Breast-feeding patient

1. Pregnant or Potentially Pregnant patients (1)

Iodinated contrast agents

Diagnostic iodinated contrast media have been shown to cross the human placenta and enter the fetus when given in the usual clinical doses. In-vivo tests in animals have shown no evidence of either mutagenic or teratogenic effects with low-osmolar contrast media. However, no well-controlled studies of the teratogenic effects of these media in pregnant women have been performed.

For those patients who are known to be pregnant or may be pregnant and for whom iodinated IV or (or internal) contrast enhancement is most appropriate for performance of the CT examination, there is no need to get signed, informed consent to use contrast media.

The data does not demonstrate mutagenic effects, fetal thyroid dysfunction or other biological effects, including renal insufficiency. According to the ACR manual on contrast media, Version 10.1 (2015), "given that there are no available data to suggest any potential harm to the fetus from exposure to iodinated contrast medium by maternal IV or intra-arterial injection, we cannot recommend routine screening for pregnancy prior to contrast media use." This recommendation is also supported by the FDA classification of most iodinated contrast as Category B medications. Please also see departmental policy on use of ionizing radiation in pregnant or potentially pregnant patients.

Gadolinium based contrast agents

Gadolinium chelates may accumulate in the amniotic fluid, and the potential effects on the pregnancy are unknown. As a result, GBCA administration is contraindicated in pregnant or potentially pregnant patients. If it is felt necessary to administer a GBCA to such a patient, this process is managed identically to that described for patients with category III renal dysfunction, as above, in the previous section on **Gadolinium Contrast Use**.

2 Breast Feeding Patients (1)

Iodinated contrast agents

The available literature on the excretion into breast milk of iodinated contrast media and the gastrointestinal absorption of these agents from breast milk is very limited. However, several studies have shown that the expected dose of contrast medium absorbed by an infant from ingested breast milk is extremely low. Therefore, it is safe for the mother and infant to continue breast-feeding after receiving iodinated contrast agents. An informed decision to temporarily stop breast-feeding should be left up to the mother after these facts are communicated. If the mother remains concerned about any potential ill effects to the infant, she may abstain from breast-feeding from the time of contrast administration for a period of 12-24 hours. There is no value to stop breast-feeding beyond 24 hours. The mother should be told to express and discard breast milk from both breasts during that period.

Gadolinium based contrast agents

Given the tiny amount of GBCA excreted in breast milk, the even smaller amount expected to be absorbed by a breast-feeding infant's GI tract, and the absence of any evidence of toxicity to breast-feeding infants in the literature, administration of GBCA at typical doses is considered safe in breast-feeding mothers. If a breast-feeding mother remains concerned about potential ill effects, she may express and discard breast milk ("pump and dump") for 24 hours following GBCA administration to further reduce the already small risks associated with GBCA administration.

Emergency patients

The ordering physician can choose to bypass screening in an emergency and have IV contrast administered without screening. This screening process bypass must be documented by the nurse, technologist or the ordering physician. The ordering physician's name must be included in the documentation.

Use of Serum Creatinine to evaluate renal function

1. Glomerular Filtration Rate (GFR) or Creatinine Clearance (CrCl) is calculated using the appropriate calculation equation depending upon serum creatinine testing method:

a. Serum Creatinine use:

i. No calculation for patients less than 1 year of age.

ii. Use serum creatinine to evaluate renal function; should be < 0.54mg/dL.

ii. For patients 1 to 16 years old

Use the "Bedside Schwartz Equation":

$(0.413 \times \text{Height}) \div \text{SCr}$

Age = age in years

SCr = serum creatinine concentration in mg/dL

Height = height/length in centimeters

I. Hydration protocol

The major preventive action to mitigate the risk of contrast-induced nephropathy is to provide intravenous volume expansion prior to contrast medium administration. The ideal infusion rate and volume is unknown, but isotonic fluids are preferred (Lactated Ringer's or 0.9% normal saline).

IV Hydration:

Patients who need IV hydration prior to and after contrast administration should make arrangements to have this performed by the referring physician in his/her clinic. We cannot provide intravenous or oral hydration to patients in the radiology department.

PO Hydration:

Oral hydration can be utilized but it is considered less effective than intravenous hydration.

I. Non-vascular Contrast Media

Contrast media may be administered into the body through the gastrointestinal tract, genitourinary tract, cutaneous fistulae, lymphatics, and intrathecal space. Adverse reactions to non-vascular contrast agents are rare; however, the appropriate management of contrast media in this setting is described in this section.

II. Barium sulfate: Barium contrast agents are frequently used for outpatient conventional fluoroscopic gastrointestinal studies and as an oral agent for some abdominal/pelvic CT and MRI scans (Redicat, Volumen).

1. **Complications:**

- a. Allergic-like reactions to Barium are exceedingly rare. If a patient reports a history of allergic-like reaction to Barium, then an alternate intraluminal agent (Isovue or Gastrografin) may be substituted. Alternatively, if the reported reaction is mild, the patient may undergo a standard steroid pre-medication (see pages 6-8).
- b. Leakage into the pleural space, mediastinum, or peritoneal cavity: Barium leakage can lead to mediastinitis or peritonitis and is contraindicated in situations where extraluminal leakage is possible.
- c. Aspiration: While Barium is generally inert when aspirated, large volume aspiration can lead to inflammation or pneumonia and therefore should be avoided in patients at risk for aspiration.

III. Iodinated contrast media: Water-soluble iodinated contrast media agents which are specifically designed for enteric opacification can be used for certain indications. These include, but not limited to, suspected bowel perforation, leak, or to confirm feeding tube position.

1. **Complications:**

- a. High-osmolar contrast media agents, e.g. Gastrografin: these agents are hypertonic and if aspirated can cause a life-threatening pulmonary edema and pneumonitis. These agents are contraindicated in patients at risk for aspiration. In these patients, low-osmolar contrast media agents, e.g. Isovue, should be substituted.
- b. Allergic-like reactions to luminal administration of iodinated contrast media are rare. Nonetheless, the potential for systemic absorption of iodinated contrast media exists. Therefore, patients with a history of allergic like reaction to contrast media should be treated the same as if receiving intravenous dosing and undergo steroid pre-medication if appropriate (see pages 6-8).

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Appendix A

Frequently Asked Questions Regarding Pretreatment in Pediatric and Adult Patients with a History of Allergy-like Reactions to Iodinated Contrast

1. If a patient has a prior reaction (mild/moderate) to contrast media, should we generally begin a pre-treatment regimen?

Yes, a pretreatment regimen with steroids should be prescribed.

2. If a patient has a mild or moderate allergy to shellfish, should we begin a pretreatment regimen? No.

3. If a patient has multiple allergies but no documented reaction to iodinated contrast media, should we begin a pretreatment regimen?

This is a controversial area. In general, no, but a strong history of allergies should increase awareness of the risk for reaction. In most cases, there is no need to avoid injection of contrast media in a patient with multiple mild or moderate allergies.

4. If a patient has well-controlled asthma but no documented reaction to iodinated contrast media, should we begin a pretreatment regimen?

No, patients with a history of well-controlled asthma do not require premedication. Patients who are actively symptomatic (i.e., currently using their inhaler) from their asthma should be premedicated or an alternate test should be considered.

5. If a patient has a remote history of an allergic reaction to contrast media, but has had intervening, uneventful, contrast enhanced scans without a prep, should they receive a pretreatment regimen with steroids?

No, in general, but these patients remain at risk for an adverse reaction. This is a controversial area.

6. If a patient with a prior contrast media reaction has undergone a steroid prep, what is their risk for a reaction?

The risk is lowered but a breakthrough reaction may occur.

7. If a patient has had a severe and life-threatening reaction to contrast media (such as anaphylaxis) should we pretreat them?

We should avoid administration of iodinated contrast media to such patients. An alternative imaging procedure should be considered. If a contrast enhanced scan is deemed absolutely necessary, a steroid prep should be given.

8. If a patient has had a steroid prep that differs from the Duke steroid prep, should we cancel the study and reschedule with a Duke prep?

No. There are several appropriate steroid preps that have been recommended. It is not clear whether one prep is advantageous over another. In these scenarios, consult the radiologist.

If all scheduled doses of a full steroid prep (12-13 hours) have been given but the timing is off by 2-3 hours, no consultation is required and the scan can be performed.

9. If a patient has had a prior breakthrough reaction, should they be prevented from having another contrast-enhanced exam?

The answer depends on the severity of the reaction.

Prior moderate or severe breakthrough reaction (anaphylaxis, laryngeal edema, hypotension): The patient should in general not be exposed to the same class of contrast (iodinated / gadolinium), regardless of premedication

Prior mild breakthrough reaction (itching, rash, hives): The patient can safely receive contrast if pre-medicated. An individual radiologist may elect to not perform the examination in this setting. This decision should be coordinated such as during ordering/scheduling process.

10. What is the likelihood that a patient will have a severe contrast reaction?

Patients with no risk factors have a risk of 4 in 10,000

Patients with a history of a prior contrast reaction have a risk of 18 in 10,000

Patients with a history of asthma have a reported risk of 23 in 10,000, but this data is likely somewhat skewed.

Patients with a history of a prior contrast reaction that are pre-medicated have a theoretical risk that is similar to someone with no risk factors. This has not been confirmed experimentally.

11. If someone has a reaction history to iodinated contrast, should they be pre-medicated before gadolinium contrast exposure (or vice versa)? No.

12. What is the definition of a mild, moderate, and severe reaction? Are all adverse events considered contrast media reactions?

The definitions are not set in stone, but these are the most commonly published categories:

a. Mild (prep needed) - Rash, scattered hives, mild facial swelling, sneezing, cough, nasal stuffiness

b. Moderate (prep needed) - Mild laryngeal edema ("scratchy throat", hoarseness), without dyspnea, wheezing without hypoxia, diffuse urticaria, significant facial swelling

c. Severe (avoid test altogether; if no other option, prep needed) - Severe respiratory distress, moderate or severe laryngeal edema, cardiopulmonary arrest, anaphylactic shock (hypotension and tachycardia)

d. Not an allergic-like reaction (no prep needed) - Nausea, flushing, vomiting, sensation of warmth

13. What is the minimum length of an effective emergency prep?

At least four hours are required for efficacy. The data supporting this are scant, but it is known that 1 and 2 hour steroid preps are ineffective. In cases where the clinical urgency demands it (e.g., trauma, aortic dissection, etc.), emergency preps shorter than 4 hours may be used at the Radiologist's discretion.

14. What do we do for a patient on chronic steroids?

There is no data on this. We recommend the following: If a patient is getting more than the equivalent of 150 mg prednisone daily, no additional steroids are needed; they may be managed as though they had received a standard steroid prep. If the patient is getting less than 150 mg prednisone equivalents daily, supplement the daily dose with enough additional steroids to reach 150 mg. When doing this, discuss the situation with the clinical service so they understand our recommendation, consent to the altered steroid dose, and avoid overdosing the patient.

Appendix B

Basic Management of Adverse Reactions to Contrast Media

Adults:

ALL ADVERSE EVENTS

1. Stay calm
2. Obtain vital signs
3. Get help
4. Communicate with the patient (before and after treatment)
5. At conclusion, call referring physician
6. Document an incident report to be added to the medical record

URTICARIA:

1. No treatment needed in most cases
2. If severe: Diphenhydramine (Benadryl) 25-50 mg PO,IM,or IV 3.
Patients receiving Benadryl need a driver

LARYNGEAL EDEMA:

1. Call a code blue
2. Oxygen 10L/min by facemask
3. ADULT Epi Kit: 0.3 mL **IM** (1:1000 sol) repeat every 5-15 min up to 3 times. Available on Omnicell medication cart. **IM** is the preferred initial treatment route.
4. Epinephrine 1-3 mL **IV**(1:10,000 sol); inject slowly up to 10 mL
5. Do not intubate. Use bag-mask ventilation if needed

ANGIOEDEMA (DIFFUSE ERYTHEMA AND HYPOTENSION)

1. Call a code blue
2. Oxygen 10 L/min by facemask
3. ADULT Epi Kit: 0.3 mL **IM** (1:1000 sol) repeat every 5-15 min up to 3 times. Available on Omnicell medication cart. **IM** is the preferred initial treatment route. Note that with profound hypotension, decreased peripheral perfusion may limit IM absorption.
4. Epinephrine 1-3 mL **IV** (1:10,000 sol); inject slowly up to 10 mL
5. Do not intubate. Use bag-mask ventilation if needed.
6. Isotonic IV fluids (normal saline or Lactated Ringer's)
 - One or more liters wide open
7. Raise legs 60 degrees
8. Remove compression if present

MILD /MODERATE BRONCHOSPASM

1. Oxygen 10L/min by face mask
2. Albuterol 2-3 puffs with spacer (If spacer available)
3. Do not intubate. Use bag-mask ventilation if needed.

MODERATE/SEVERE BRONCHOSPASM:

1. Call a code blue
2. Oxygen 10L/min by facemask
3. ADULT Epi Kit: 0.3 mL **IM** (1:1000 sol) repeat every 5-15 min up to 3 times.
Available on Omnicell medication cart. **IM** is the preferred initial treatment route.
Epinephrine 1-3 mL **IV**(1:10,000 sol); inject slowly up to 10 mL
4. Do not intubate. Use bag-mask ventilation if needed
- 5.

HYPOTENSION WITH TACHYCARDIA (ANAPHYLACTIC SHOCK):

1. Call a code blue
2. Oxygen 10L/min by facemask
3. ADULT Epi Kit: 0.3 mL **IM** (1:1000 sol) repeat every 5-15 min up to 3 times.
Available on Omnicell medication cart. **IM** is the preferred initial treatment route. Note that with profound hypotension, decreased peripheral perfusion may limit IM absorption
4. Epinephrine 1-3 mL **IV**(1:10,000 sol); inject slowly up to 10 mL
5. Do not intubate. Use bag-mask ventilation if needed
6. Isotonic IV fluids (normal saline or Lactated Ringer's)
 - One or more liters wide open
7. Raise legs 60 degrees
8. Remove compression if present

HYPOTENSION WITH BRADYCARDIA • VASOVAGAL REACTION

1. Oxygen 10 L/min by face mask
2. Raise legs 60 degrees
3. Remove compression
4. Medications not normally needed
5. If persistent:
Isotonic IV fluids (normal saline or Lactated Ringer's)
One or more liters wide open
6. Atropine 0.6-1.0 mg IV slowly into running fluids, repeat up to 3 mg total
7. Call a code if persistent hypotension (115)

HYPERTENSION, SEVERE (Diastolic BP > 12 mmHg; systolic BP > 200 mm Hg):

1. Oxygen 10 L/min by face mask
2. Labetalol 20 mg IV; administer slowly over 2 mins; double dose every
3. 10 min (e.g. 40 mg 10 min later, 80 mg 10 min after that)
4. Watch for iatrogenic bradycardia or heart block
If labetalol is not available:
 1. Furosemide (Lasix) 20-40 mg IV slowly over 2 min
 2. Nitroglycerine 0.4 mg SL; repeat every 5-10 min

PULMONARY EDEMA:

1. Oxygen 10L/min by facemask
2. Elevate head
3. Stop IV fluids
4. Furosemide (Lasix) 20-40 mg IV slowly over 2 min

EPI-KIT Guidelines

The **IM administration will be given as the first option for all patients (even with existing lines) with anaphylactic or potentially anaphylactic reactions.** The Epi- Kit can be found in CT drug box and can only be used by the nursing staff after healthcare provider's approval

Anaphylaxis Epinephrine Kits:

Intramuscular injection is the standard route for treatment of anaphylaxis following contrast administration

The code cart will remain as is; code dose epinephrine (1:10,000) should be pulled from the drug box

Appendix C

I. Oral Hypoglycemic Agents with Metformin

Brand Name	Generic
Glucophage (XR); Glumetza; Riomet; Fortamet	Metformin
Glucovance	Metformin; Glyburide
Metaglip	Metformin; Glipizide
Actoplus Met	Metformin; Pioglitazone
Avandamet	Metformin; Rosiglitazone
Janumet	Metformin; Sitagliptin

II. Oral Hypoglycemic Agents without Metformin

Brand Name	Generic
Tolinase	Tolazamide
Tolbutamide	Tolbutamide
Diabinese	Chlorpropamide
Glucotrol (XL)	Glipizide
Amaryl	Glimepiride
DiaBeta; Glynase; Micronase	Glyburide
Glyset	Miglitol
Precose	Acarbose
Avandia	Rosiglitazone
Actos	Pioglitazone
Starlix	Nateglinide
Prandin	Repaglinide
Avandaryl	Rosiglitazone; Glimepiride
Duetact	Pioglitazone; Glimepiride
Januvia	Sita

