



Enhanced clinical efficiency in radiology with protocol optimization & right choice of iodinated contrast media

Speaker Name: Dr Saritha Nair

Designation: Regional Medical Advisor, Bayer Radiology

Date: 25 May 2022





Disclaimer

- All rights reserved. No part of these slides may be reproduced, distributed, or transmitted in any form or by any means, without the prior written permission.
- Use of these slides is permitted for scientific and educational presentations only
- Author, reviewer and presenter shall not have any liability with respect to any damages caused by the use of these slides or parts thereof
- Bayer does not accept responsibility and shall have no liability for miscommunication of these data if they are changed in any way
- The data contained within this slide deck do not support or recommend the use of Ultravist, Gadovist & Primovist in any countries or indications in which it is not approved
- The safety of current and future patients is of utmost importance to Bayer. Capturing as many of these side effects, however rare they may be in absolute terms, from worldwide sources. Please report all adverse events to the following email address : drugsafety.hk@bayer.com



Number of CT scans is on the rise¹



- From 2007 to 2017, the number of CT examinations rose by an average of 56%.¹
- Use of iodinated contrast media (ICM) is growing accordingly.²
- In Korea alone, it is estimated that more than 4 million CT scans involving contrast use are performed each year.²
- Low-osmolar contrast media (LOCM) are the most widely used class of CT contrast media.³

1. OECD (2019), "CT exams, 2007 and 2017 (or nearest year)", in Health care activities, OECD Publishing, Paris, <https://doi.org/10.1787/da83b311-en>.
2. Cha MJ, et al. Hypersensitivity Reactions to Iodinated Contrast Media: A Multicenter Study of 196 081 Patients. *Radiology*. 2019. Oct;293(1):117-124. doi:10.1148/radiol.2019190485.
3. Becker, C.R. Selecting a contrast medium for MDCT investigation. *Eur Radiol Suppl* 16, D33–D37 (2006). <https://doi.org/10.1007/s10406-006-0185-2>



Confidence with Ultravist



> To diagnose with confidence, it is important to be familiar with the comprehensive data on image quality, patient comfort and safety of contrast media.



Image Quality



Patient Comfort



General Safety



Renal Safety



Delayed Skin Reactions



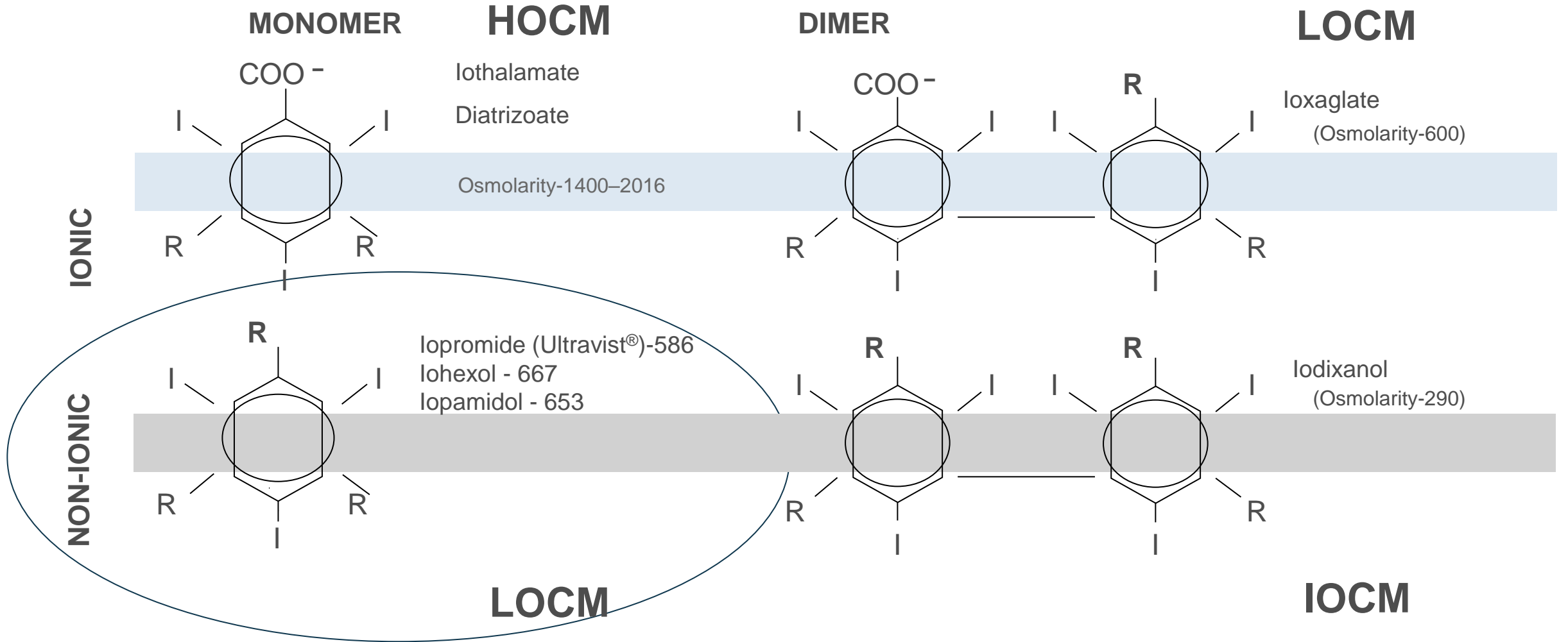
Cardiac Safety



Thyroid Safety



Structure, Ionicity & Osmolarity* of Iodinated Contrast Media



Adapted from- Richard Solomon, "Contrast Media: Are There Differences in Nephrotoxicity among Contrast Media?", *BioMed Research International*, vol. 2014, Article ID 934947, 8 pages, 2014. <https://doi.org/10.1155/2014/934947>



Physico-chemical properties of Iodinated Contrast Media



COMPOUND	OSMOLALITY*	VISCOSITY 20 °C	37 °C
Ionic CM			
Diatrizoate	1,502 ± 16	9.3 ± 0.1	4.9 ± 0.0
Ioxaglate 320	584 ± 10	16.2 ± 1.0	7.8 ± 0.4
Non ionic CM			
Iopamidol	640 ± 6	8.4 ± 0.1	4.5 ± 0.0
Iohexol	667 ± 8	10.9 ± 0.2	5.7 ± 0.1
Iopromide	586 ± 5	8.9 ± 0.1	4.7 ± 0.1
Iomeprol	521 ± 24	8.1 ± 0.7	4.5 ± 0.4
Ioversol	661 ± 3	10.1 ± 1.2	5.2 ± 0.6
Iopentol	683 ± 4	12.9 ± 0.2	6.5 ± 0.1
Iotrolan	294 ± 3	17.5 ± 0.0	8.5 ± 0.0
Iodixanol	290	25,4	11.1

* Osmolality (mOsm/kg H₂O) for 300 mg Iodine / mL, 37 °C
(mean ± standard deviation, as far as data are available)



Image Quality



Image quality is key



- High-quality images simplify interpretation and are essential for correct diagnosis and treatment planning.
- The key parameter for a high imaging quality differs for organ studies and imaging of vessels
 - Key parameter for organ studies: Total Iodine Load (TIL)¹
 - Key parameter in CT angiography: Iodine Delivery Rate (IDR)²
- Other aspects to be considered when optimizing protocols are:
 - The patient's body weight³
 - Peak pressure²
 - Viscosity of the contrast medium (CM)²
 - Type of injector used to inject the CM⁴

1. Bae KT. Intravenous contrast medium administration and scan timing at CT: considerations and approaches. *Radiology*. 2010;256(1):32–61. doi: 10.1148/radiol.10090908

2. Rengo M, et al. Impact of iodine concentration and iodine delivery rate on contrast enhancement in coronary CT angiography: a randomized multicenter trial (CT-CON). *Eur Radiol*. 2019 Nov;29(11):6109-6118. doi: 10.1007/s00330-019-06196-7 doi: 10.1007/s00330-019-06196-7

3. Hendriks BMF, et al. Individually tailored contrast enhancement in CT pulmonary angiography. *Br J Radiol* 2016; 89: 20150850. doi: 10.1259/bjr.20150850

4. Chaya A, et al. Piston-Based vs Peristaltic Pump-Based CT Injector Systems. *Radiol Technol*. 2019 Mar;90(4):344-352. PMID: 30886031



Peak pressure and image quality

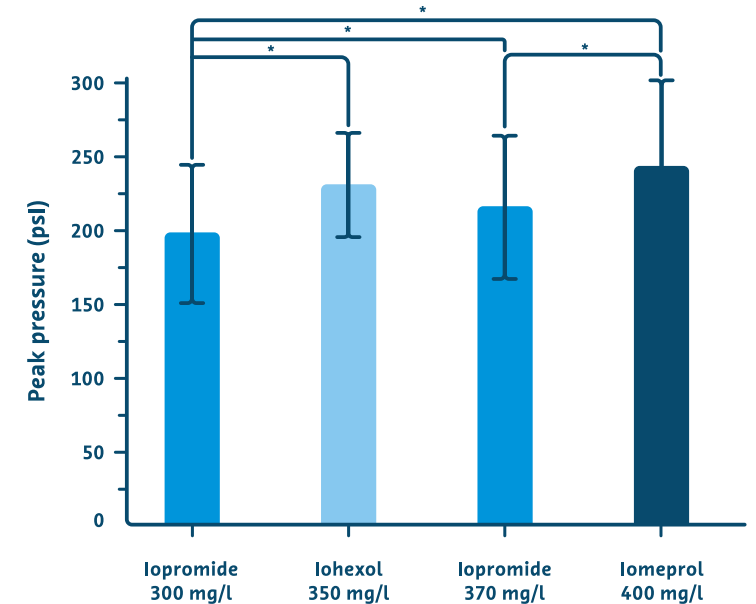


- CT-CON demonstrated that higher iodine concentrated contrast media with simultaneously increased viscosity result in higher peak pressures.¹
- Lower viscosity is known to be beneficial in terms of injection pressure.^{2,3}
- High viscosity complicates the contrast administration at higher flow rates.⁴



High peak pressures should be avoided to achieve optimal image quality.¹

A low viscosity contrast media is beneficial for the higher flow rates required for CTA examinations.^{1,5}



Peak pressure (psi)** 197.4 ± 47.7 229.8 ± 35.7 216.1 ± 46.1 243.7 ± 58.7

Influence of viscosity on peak pressures for different contrast media.¹

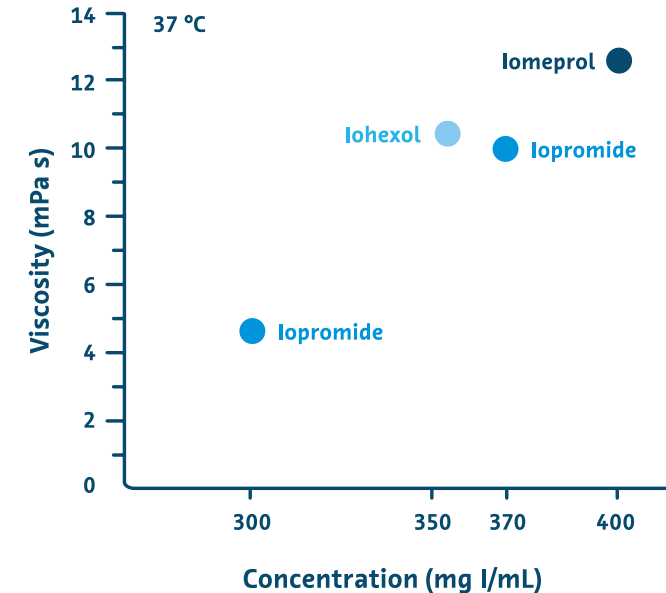
1. Rengo M, et al. Impact of iodine concentration and iodine delivery rate on contrast enhancement in coronary CT angiography: a randomized multicenter trial (CT-CON). Eur Radiol. 2019 Nov;29(11):6109-6118. doi: 10.1007/s00330-019-06196-7
2. Bae KT. Intravenous contrast medium administration and scan timing at CT: considerations and approaches. Radiology. 2010;256(1):32-61. doi: 10.1148/radiol.10090908
3. Kok M, et al. Patient Comfort During Contrast Media Injection in Coronary Computed Tomographic Angiography Using Varying Contrast Media Concentrations and Flow Rates: Results From the EICAR Trial. Invest Radiol. 2016 Dec;51(12):810-815. doi: 10.1097/RLI.0000000000000284
4. Faggioni L, Gabelloni M. Iodine Concentration and Optimization in Computed Tomography Angiography: Current Issues. Invest Radiol. 2016 Dec;51(12):816-822. doi: 10.1097/RLI.0000000000000283 Review. Dec;51(12):810-815.
5. Mihl C, et al. Intravascular enhancement with identical iodine delivery rate using different iodine contrast media in a circulation phantom. Invest Radiol. 2013;48:813-818. doi: 10.1097/RLI.0b013e31829979e8



Ultravist[®] – a good choice to apply IDR



- Both Ultravist[®] 300 and 370 have a relatively low viscosity in relation to their iodine concentration which allows for lower peak pressures at identical flow rates.¹
- Despite having a lower iodine concentration, the viscosity of Iohexol 350 is higher than that of Ultravist[®] (Iopromide) 370.*,¹



Relation of CT-CON investigated contrast media concentrations to their viscosity at 37° Celsius as stated in the respective prescribing information²⁻⁴

* numerically, not statistically significant

1. Rengo M, et al. Impact of iodine concentration and iodine delivery rate on contrast enhancement in coronary CT angiography: a randomized multicenter trial (CT-CON). Eur Radiol. 2019 Nov;29(11):6109-6118. doi: 10.1007/s00330-019-06196-7
2. Iopromide prescribing information. <http://www.mhra.gov.uk/spc-pil/?subsName=IOPROMIDE&pageID=SecondLevel>. Date of access: December 2019.
3. Iomeprol prescribing information. https://imaging.bracco.com/sites/braccoimaging.com/files/technica_sheet_pdf/de-de-2018-11-15-spc-Imeron.pdf. Date of access: December 2019.
4. Iohexol prescribing information. https://www.accessdata.fda.gov/drugsatfda_docs/label/2018/018956s101lbl.pdf. Date of access: December 2019.



Concentration to Viscosity Ratio

- Concentration and viscosity are two properties of contrast media & its ratio is important for an evaluation of a contrast medium
- Hence, the concentration to viscosity ratio (CVR) has been proposed in the InnoVatE study,2020, as a new parameter
- This study assessed the impact of injection system technologies and CM viscosity on achievable iodine delivery rates and vascular enhancement.
- It concluded that- the tested piston-based injection systems (like Centargo) combined with low viscosity contrast media provide higher achievable IDRs and higher peak vascular enhancement than the tested peristaltic-based injectors.

$$\frac{\text{Concentration}}{\text{Viscosity}} = \text{CVR}$$

mg I / mL
cP
mg I / mL / cP



Concentration to Viscosity Ratio

$$\begin{array}{ccc}
 \text{Concentration} & & \text{Viscosity} & & \text{CVR} \\
 \mathbf{370} & \div & \mathbf{17.10} & = & \mathbf{21.6} \\
 \text{370 milligrams of} & & \text{Measured viscosity in} & & \text{grams of} \\
 \text{iodide per milliliter} & & \text{centipoise} & & \text{iodine} \\
 \text{(mgI/mL)} & & \text{(cP)} & & \text{per second}
 \end{array}$$

InnoVatE study measured the viscosity of iodinated contrast media at room temperature. In addition, viscosity as published by the respective manufacturer is also provided.

To calculate the CVR, the concentration e.g. 370 is divided by viscosity (UV 370= 17.10), leading to a CVR of 21.6

Generic	Concentration (mgI/mL)	Published Viscosity (cP)*	Measured Viscosity (cP)**	Concentration / Viscosity Ratio (mgI/mL/cP)***	Concentration / Viscosity Ratio (mgI/mL/cP) at 37°C****
Iopromide	300	9.2	7.64	39.3	61.2
Iodixanol	320	26.6	21.10	15.2	27.1
Iohexol	350	20.4	18.70	18.7	33.7
Iopromide	370	22.0	17.10	21.6	37.0
Iomeprol	400	27.5	23.00	17.4	31.7

- * Official data from manufacturers at 20°C
- ** Measured data using Brookfield DV-II+ Pro Viscometer at tested temperature of 21.5°C
- *** Determined using measured contrast media viscosity
- **** Calculated from manufacturer reported viscosities at 37°C



General Safety



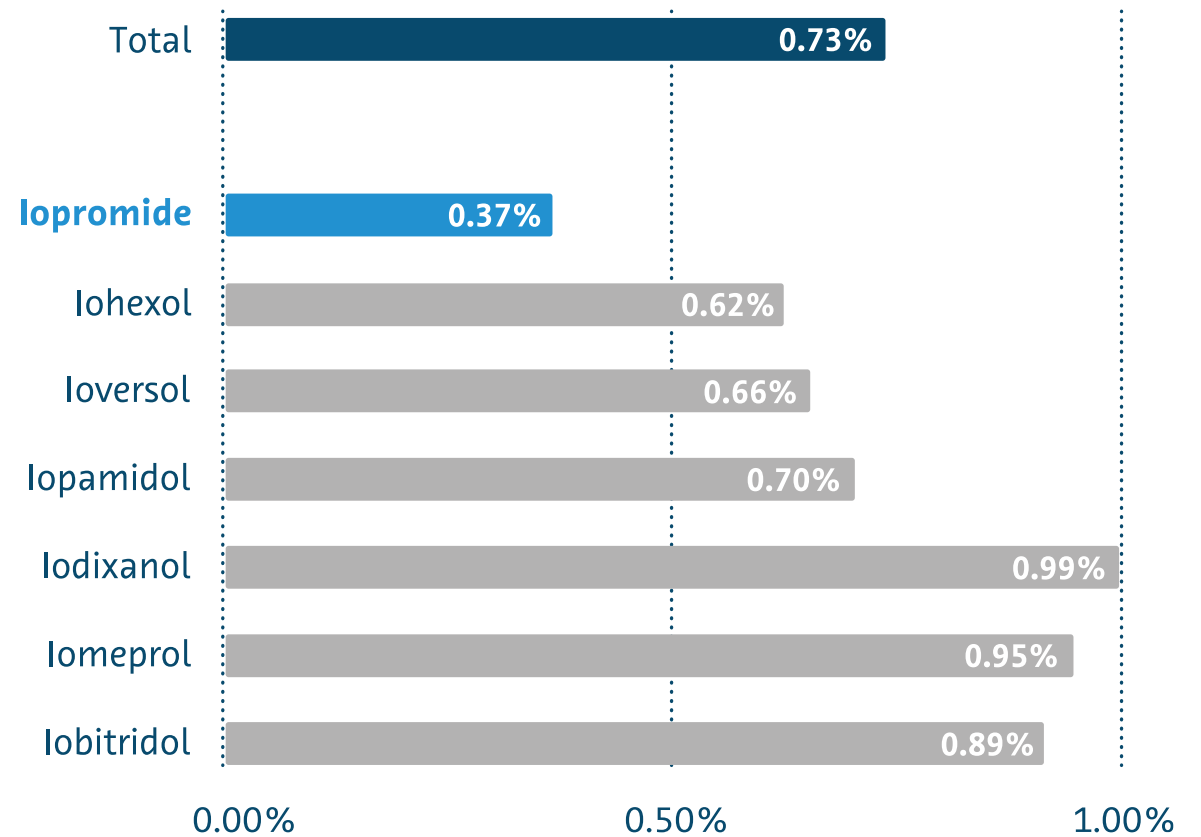
Ultravist® has a low prevalence of HSRs¹



- A 2019 published Korean nationwide registry study, sponsored by the Ministry of Food and Drug Safety as a multi-center study (March 2017 and October 2017) from seven tertiary referral hospitals in Korea including 196,081 patients and seven ICM examined:²
 - Prevalence
 - Risk factors
 - And preventive measures for HSRs



Iopromide (Ultravist®) showed the lowest prevalence of HSRs in this study.



Prevalence of hypersensitivity reactions in the Korean Registry Study by Cha et al.²

1. Endrikat J et al. Risk of Hypersensitivity Reactions to Iopromide After Intra-Arterial Versus Intravenous Administration: A Nested Case-Control Analysis of 133,331 Patients. Invest Radiol. 2020 Jan;55(1):38-44. doi: 10.1097/RLI.0000000000000611.

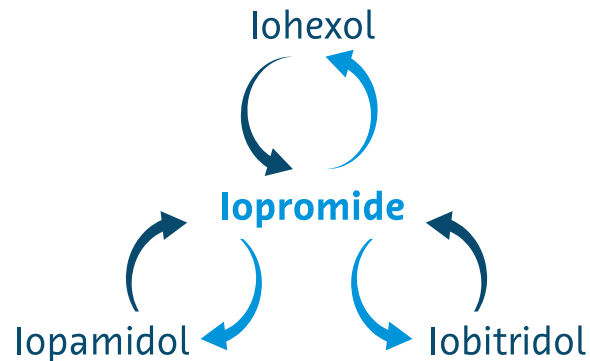
2. Cha MJ, et al. Hypersensitivity Reactions to Iodinated Contrast Media: A Multicenter Study of 196 081 Patients. Radiology. 2019. Oct;293(1):117-124. doi:10.1148/radiol.2019190485



Choosing the right contrast agent



- Guidelines suggest that changing ICM is helpful to reduce HSR recurrence^{1,2}
- In 2018 Park et al. evaluated different exchange combinations of ICM³:
 - It was concluded that the optimal choice could be individualized according to the culprit agent
 - Five favorable combinations of substitutes depending on culprit agent were identified
 - Ultravist[®] is suggested in 3 of these 5 pairs that showed a reduced risk for HSR recurrence³

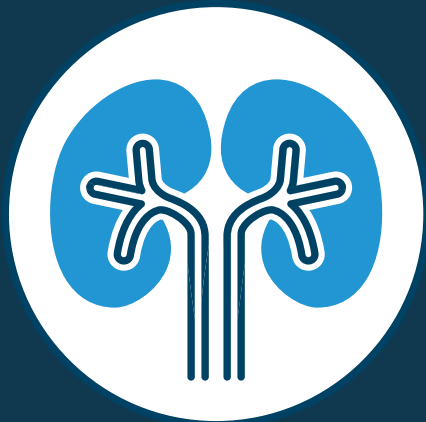


Combination of Contrast Media	Recurrence	Odds ratio*	95% Confidence Interval	P Value	Odds Ratio*	95% Confidence Interval	P Value
Iohexol / Iopamidol	24/110 (21.8)	0.470	0.247, 0.892	.021	1.863	0.995, 3.487	.052
Iopamidol / Iopromide	11/58 (19.0)	0.366	0.163, 0.820	.015	1.466	0.654, 3.289	.353
Iohexol / Iopromide	37/268 (13.8)	0.402	0.227, 0.712	.002	1.353	0.788, 2.321	.273
Iobitridol / Iohexol	32/261 (12.3)	0.245	0.127, 0.474	<.0001	0.887	0.468, 1.680	.713
Iobitridol / Iopromide	18/169 (10.7)	0.296	0.129, 0.682	.004	1.048	0.462, 2.374	.911
Iobitridol / Iopamidol	7/17 (41.2)	0.942	0.340, 2.608	.909	4.175	1.545, 11.281	.005
Iobitridol / Ioversol	7/21 (33.3)	0.890	0.321, 2.466	.823	3.310	1.210, 9.056	.020
Iopromide / Ioversol	1/2 (50.0)	1.344	0.083, 21.898	.835	5.961	0.369, 96.259	.208
Iohexol / Ioversol	5/12 (41.7)	0.876	0.237, 3.232	.842	3.585	0.978, 13.137	.054
Iohexol / Iomeprol	3/13 (23.1)	0.709	0.163, 3.084	.647	1.937	0.437, 8.591	.384
Iomeprol / Iopamidol	4/19 (21.1)	0.472	0.144, 1.551	.216	1.804	0.551, 5.913	.330
Iopamidol / Ioversol	2/10 (20.0)	0.188	0.023, 1.569	.123	0.779	0.094, 6.454	.817
Iomeprol / Iopromide	2/11 (18.2)	0.612	0.108, 3.451	.578	1.877	0.321, 10.980	.485
Iobitridol / Iomeprol	0/8 (0)	...	NA	NA	...
Iomeprol / Ioversol	0/1 (0)	...	NA	NA	...

* Compared with exposure to same CM.+ Compared with exposure to different CM.

HSR recurrence and odds ratios for different substitute combinations.³

1. ACR Committee on Drugs and Contrast Media -ACR Manual on Contrast Media. 2018; Version 10.3. Available at: https://www.acr.org/-/media/ACR/Files/Clinical-Resources/Contrast_Media.pdf.
 2. European Society of Urogenital Radiology: ESUR Guidelines on Contrast Media (version 10.0) 2019: URL: http://www.esur.org/fileadmin/content/2019/ESUR_Guidelines_10.0_Final_Version.pdf
 3. Park SJ, et al. Immediate Mild Reactions to CT with Iodinated Contrast Media: Strategy of Contrast Media Readministration without Corticosteroids. Radiology. 2018 Sep;288(3):710-716.doi: 10.1148/radiol.2018172524.



Renal Safety



Guideline recommendations regarding renal safety



➤ Recommendations on the choice of iodinated contrast medium do not state a preference for IOCM over LOCM regarding renal safety.

Scientific Association	Recommendations
American College of Radiology (2020 update) ¹	“Studies have failed to establish a clear advantage of IV iso-osmolality iodixanol over IV LOCM with regard to PC-AKI or CIN . A 2009 meta-analysis (Heinrich et al.,2009) using data pooled from 25 trials found no difference in the rate of PC-AKI between iodixanol and low osmolality agents after intravenous administration .”
American College of Radiology and the National Kidney Foundation (2020) ²	“ There are no clinically relevant differences in CI-AKI risk between iso-osmolality and low-osmolality iodinated contrast media .”
ESUR Contrast Media Safety Committee (ESUR V10.0) ³	For all patients use low- or iso-osmolar contrast media . Use the lowest dose of contrast medium consistent with a diagnostic result.
Canadian Association of Radiologist (Owen RJ et al. 2014) ⁴	“Larger studies and meta-analyses revealed no significant difference between iodixanol and most low-osmolar contrast media . [...] Currently, the Canadian Association of Radiologists recommends the use of iso- or low-osmolar media in patients with GFR <45 mL/min in intravenous administration and GFR <60 ml/min at intraarterial administration.”
The Renal Association, British Cardiovascular and Intervention Society and The Royal College of Radiologists (2013 Update) ⁵	“Currently there is only one type of iso-osmolar media which has failed to demonstrate any clear benefit compared to different low-osmolar media in preventing CI-AKI ”
American College of Cardiology Foundation/Society for Cardiovascular Angiography and Interventions (Bashore TM et al. 2012) ⁶	“[The volume of] contrast media should be minimized, and low-osmolar or iso-osmolar contrast media should be used ”
European Society of Cardiology (2018 Update) ⁷	“Use of low-osmolar or iso-osmolar contrast media is recommended for patients with moderate or severe CKD”

1. ACR Committee on Drugs and Contrast Media -ACR Manual on Contrast Media. Version 2020. Available at: https://www.acr.org/-/media/ACR/Files/Clinical-Resources/Contrast_Media.pdf

2. Davenport M., et al. Use of Intravenous Iodinated Contrast Media in Patients with Kidney Disease: Consensus Statements from the American College of Radiology and the National Kidney Foundation. Radiology 2020. 294:660–668. 10.1148/radiol.2019192094.

3. European Society of Urogenital Radiology: ESUR Guidelines on Contrast Media (version 10.0) 2019: URL: http://www.esur.org/fileadmin/content/2019/ESUR_Guidelines_10.0_Final_Version.pdf

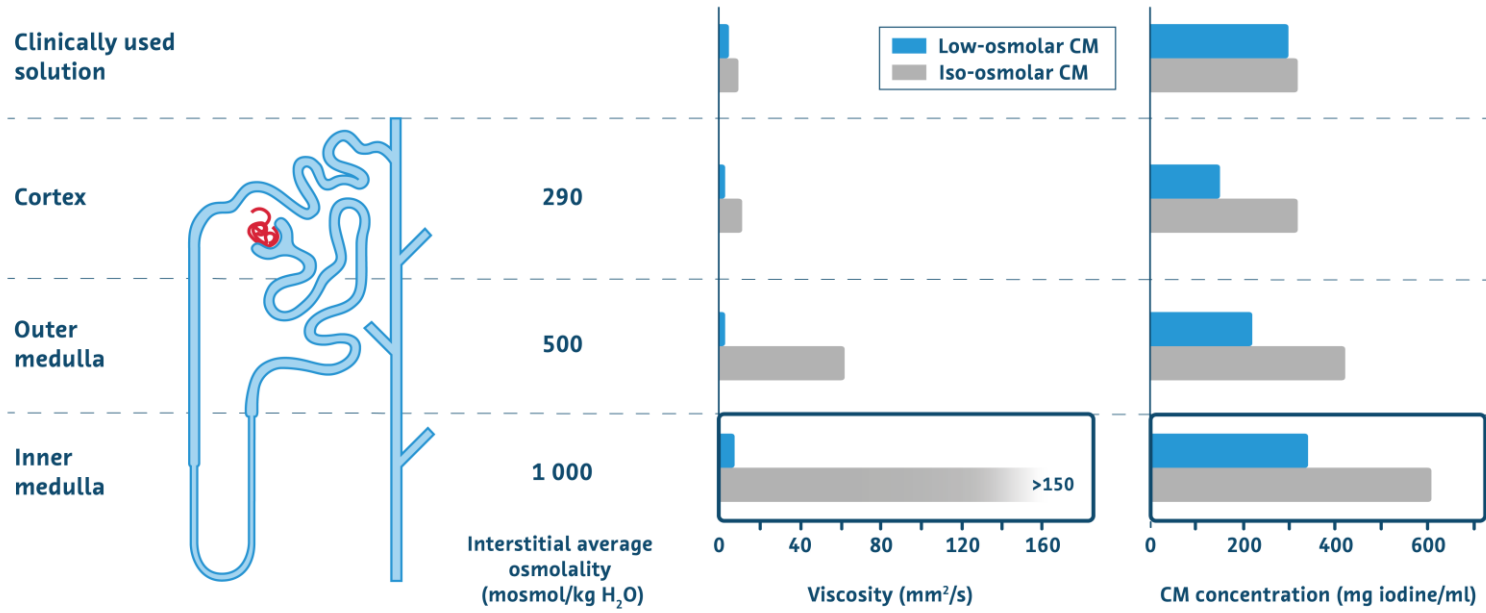
4. Owen RJ, et al. Canadian Association of Radiologists consensus guidelines for the prevention of contrast-induced nephropathy: update 2012. Can Assoc Radiol J. 2014 May;65(2):96-105. doi: 10.1016/j.carj.2012.11.002

5. The Renal Association, British Cardiovascular Intervention Society and The Royal College of Radiologists, Prevention of Contrast Induced Acute Kidney Injury (CI-AKI) In Adult Patients

6. Bashore TM, et al. ACCF Task Force Members. 2012 American College of Cardiology Foundation/Society for Cardiovascular Angiography and Interventions expert consensus document on cardiac catheterization laboratory standards update: A report of the American College of Cardiology Foundation Task Force on Expert Consensus documents developed in collaboration with the Society of Thoracic Surgeons and Society for Vascular Medicine. J Am Coll Cardiol. 2012 Jun 12;59(24):2221-305. doi: 10.1016/j.jacc.2012.02.010

7. Neumann F.-J., et al., 2018 ESC/EACTS Guidelines on myocardial revascularization, European Heart Journal, 2019; 40, 87–165 doi:10.1093/eurheartj/ehy394

Viscosity level of contrast media significantly differs in different layers of the kidney

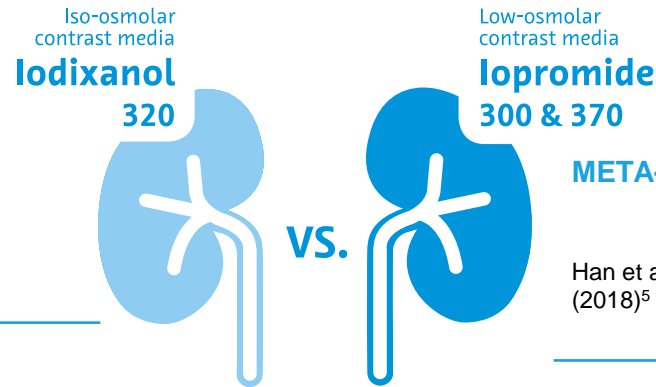


- At 1,000 mosmol/kg H₂O, the level of viscosity for IOCM markedly exceeds the maximum value which a viscometer can measure.
- In contrast, the viscosity level of LOCM only slightly increased compared to the initial solution.

- At 1,000 mosmol/kg H₂O, the concentration of low-osmolar CM is slightly elevated.
- In contrast, the concentration of iso-osmolar CM is about twice as high as that of the original solutions.

1. Fähring M et al. Understanding and preventing contrast-induced acute kidney injury. Nat Rev Nephrol 13, 169–180 (2017). <https://doi.org/10.1038/nrneph.2016.196>

Scientific and clinical evidence on renal safety



INDIVIDUAL COMPARISON IOPROMIDE VS. IODIXANOL*

Chen et al. (2012) ¹	Iodixanol 320 (N=284) Iopromide (Ultravist®) 370 (N=278)	NON-INFERIOR (p<0.001)
Bolognese et al. (2012) ²	Iodixanol 320 (N=236) Iopromide (Ultravist®) 370 (N=239)	NON-INFERIOR (p<0.0002)
Shin et al. (2011) ³	Iodixanol 320 (N=215) Iopromide (Ultravist®) 300 (N=205)	NO SIGNIFICANT DIFFERENCE (p=0.394)
Juergens et al. (2009) ⁴	Iodixanol 320 (N=91) Iopromide (Ultravist®) 370 (N=100)	NO SIGNIFICANT DIFFERENCE (p=0.56)

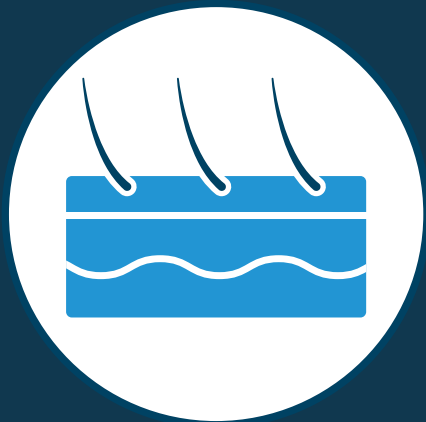
META-ANALYSES LOCM VS. IODIXANOL

Han et al. (2018) ⁵	Diabetic patients, 12 trials Iodixanol 320 (N=575) LOCM (N=525)	NO SIGNIFICANT DIFFERENCE Subgroup analysis: Significant difference between iohexol and iodixanol
From et al. (2010) ⁶	36 trials Iodixanol 320 (N=3,672) LOCM (N=3,494)	NO SIGNIFICANT DIFFERENCE Subgroup analysis: Significant difference between iohexol and iodixanol
Heinrich et al. (2009) ⁷	25 trials Iodixanol (N=1,701) LOCM (N=1,569)	NO SIGNIFICANT DIFFERENCE Subgroup analysis: Significant difference between iohexol and iodixanol



No significant difference in scientific and clinical evidence regarding renal safety between IOCM and LOCM.

- Chen Y et al. Renal tolerability of iopromide and iodixanol in 562 renally impaired patients undergoing cardiac catheterisation: the DIRECT study.
- Bolognese L, et al. Impact of iso-osmolar versus low-osmolar contrast agents on contrast-induced nephropathy and tissue reperfusion in unselected patients with ST-segment elevation myocardial infarction undergoing primary percutaneous coronary intervention. Am J Cardiol. 2012 Jan 1;109(1):67-74. doi: 10.1016/j.amjcard.2011.08.006
- Shin DH, et al. Comparison of contrast-induced nephrotoxicity of iodixanol and iopromide in patients with renal insufficiency undergoing coronary angiography. Am J Cardiol. 2011 Jul 15;108(2):189-94. doi: 10.1016/j.amjcard.2011.03.019
- Juergens CP, et al. Nephrotoxic effects of iodixanol and iopromide in patients with abnormal renal function receiving N-acetylcysteine and hydration before coronary angiography and intervention: a randomized trial. Intern Med J. 2009 Jan;39(1):25-31. doi: 10.1111/j.1445-5994.2008.01675.x
- Han XF, et al. Contrast-induced nephropathy in patients with diabetes mellitus between iso- and low-osmolar contrast media: A meta-analysis of full-text prospective, randomized controlled trials. PLoS One. 2018 Mar 20;13(3). doi: 10.1371/journal.pone.0194330
- From AM, et al. Iodixanol versus low-osmolar contrast media for prevention of contrast induced nephropathy: meta-analysis of randomized, controlled trials. Circ Cardiovasc Interv. 2010 Aug;3(4):351-8. doi: 10.1161/CIRCINTERVENTIONS.109.917070
- Heinrich MC, et al. Nephrotoxicity of iso-osmolar iodixanol compared with nonionic low-osmolar contrast media: meta-analysis of randomized controlled trials. Radiology. 2009; 250(1):68-86. doi: 10.1148/radiol.2501080833



Delayed Skin Reactions



Late Adverse Reactions -ICM

Definition: A late adverse reaction to intravascular ICM is defined as a reaction which occurs 1 h (30-60mins)^A to 1 week after CM injection.^E

- Majority occurring between 3 hours and 2 days^A

ESUR (V.10.0) & ACR (V.11)

Reactions^E

- **Skin reactions** similar in type to other drug induced eruptions occur.
 - Maculopapular rashes, erythema, swelling and pruritus are most common.
 - Most skin reactions are mild to moderate and self-limiting.
- A variety of late symptoms (e.g., nausea, vomiting, headache, musculoskeletal pains, fever) have been described following CM, but many are not related to the CM.

Risk factors for skin reactions^E

- Previous late CM reaction
- Interleukin-2 treatment
- **Use of non-ionic dimers**

Incidence^A

- 0.5% to 14% (Delayed allergic-like reactions)
- A prospective study of 258 individuals receiving intravenous iohexol demonstrated a delayed reaction rate of **14.3%** compared to **2.5%** in a control group undergoing imaging without intravascular contrast material. (Loh et al., 2010)

Note: Late skin reactions of the type which occur after ICM have not been described after GBCA and ultrasound contrast media.

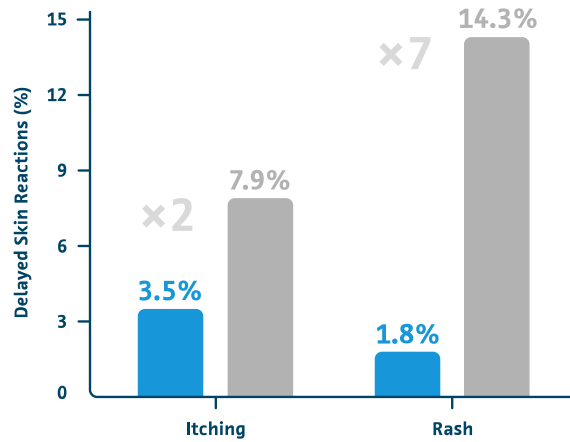
ICM: iodine-based contrast media
GBCA: gadolinium-based contrast agent
CM: Contrast Medium

^EESUR Guidelines on Contrast Media Version 10.0; 2018

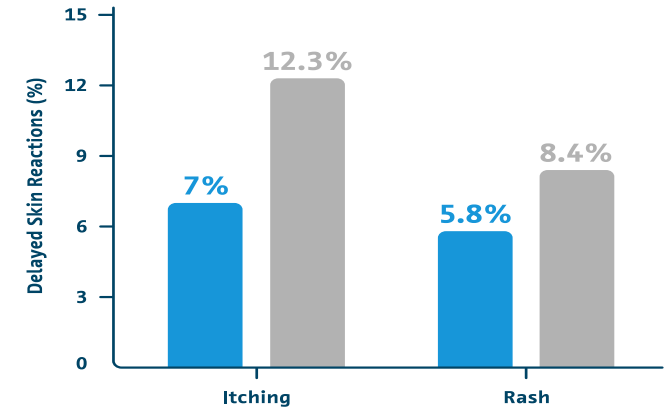
^AACR Manual on Contrast Media Version 11; 2020



Delayed skin reactions with LOCMs vs. IOCMs

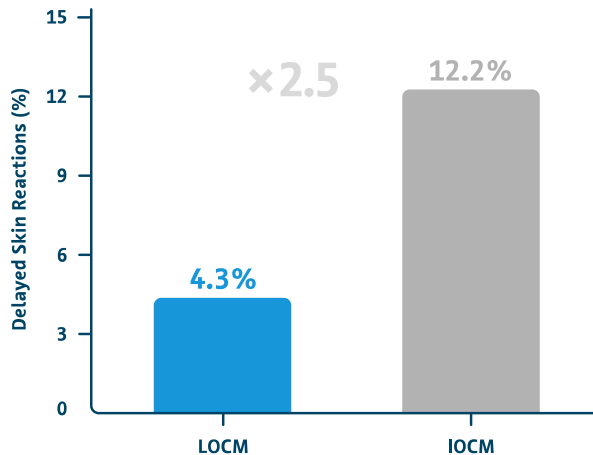


Delayed skin reactions are less frequent with LOCMs than with IOCMs.^{1,2,3}

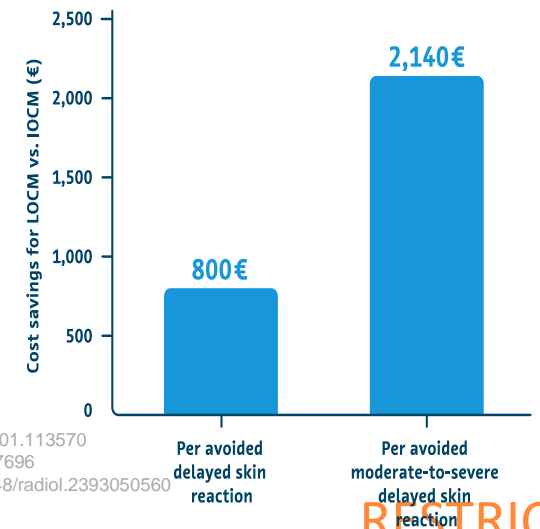


Ultravist® (Iopromide) Iotrolan

Reactions to CM may cause additional costs such as concomitant medication, healthcare personnel or hospitalization⁴



LOCMs like Ultravist® have less additional costs due to delayed skin reactions than IOCMs⁴



1. Sutton AG, et al. Early and late reactions after the use of iopamidol 340, ioxaglate 320, and iodixanol 320 in cardiac catheterization. Am Heart J. 2001 Apr;141(4):677-83. doi: 10.1067/mhj.2001.113570
 2. Gharekhanloo F and Torabian S, Comparison of allergic adverse effects and contrast enhancement between iodixanol and iopromide. Iran J Radiol. 2012;9(2):63-66. doi: 10.5812/iranjradiol.7696
 3. Schild HH, et al. Adverse events after unenhanced and monomeric and dimeric contrast-enhanced CT: a prospective randomized controlled trial. Radiology 2006; 240(1): 56 – 64. doi: 10.1148/radiol.2393050560
 4. Arana E and Catalá-López F, Cost-effectiveness of iodinated contrast media for CT scanning in Spain: A decision-based analysis. Imaging Med. 2012 4(2), 193–99



Clinical Updates on ACR 2021 Guidelines





ACR 2021 Guidelines Update Summary

Changes	Details
Addition of Chapter 5- Fasting Prior to Intravascular Contrast Media Administration	<ul style="list-style-type: none"> Given the potential for negative consequences due to fasting and a lack of evidence that supports the need for fasting, <u>fasting is not required prior to routine intravascular contrast material administration.</u> However, for patients receiving conscious sedation, anesthesia guidelines should be consulted
Chapter 10- Change in terminologies	<p>Post-contrast acute kidney injury (PCAKI) IS NOW Contrast-associated acute kidney injury (CA-AKI)</p> <p>Contrast-induced nephropathy (CIN) IS NOW Contrast-induced acute kidney injury (CI-AKI)</p>
Chapter 10- Volume Expansion protocol recommendations to prevent CA-AKI made more concrete	<ul style="list-style-type: none"> Isotonic fluid such as 0.9% normal saline (NS) is preferred. Typical prophylaxis regimens begin 1 hour prior to the exam and continue 3-12 hours after. Typical doses may be fixed volume (e.g., 500 mL NS) before and after or weight-based volumes (1-3mL/kg per hour) The ideal infusion rate and volume is unknown
Chapter 10- Addition of Indications & Contraindications for volume expansion to prevent CA-AKI	<p>Indications-</p> <ul style="list-style-type: none"> Patients who have AKI or severe CKD with an eGFR less than 30 mL/min/1.73m², although the risks of volume expansion (i.e., heart failure or other hypervolemic conditions) should be considered before initiation. Considered on an individual basis for high-risk circumstances (e.g., numerous risk factors, recent AKI, borderline eGFR) in patients with an eGFR of 30-44 mL/min./1.73 m² at the discretion of the ordering provider <p>Contraindication-</p> <ul style="list-style-type: none"> General population of patients with stable eGFR greater than or equal to 30 mL/min 1.73 m² or patients on chronic dialysis.

CM: Contrast Medium
 CT: Computed Tomography
 MRI: Magnetic resonance Imaging
 LOCM: Low Osmolar Contrast Medium
 IOCM: Iso Osmolar Contrast medium



ACR 2021 Guidelines Update Summary contd.....

Changes	Details
Changes in Chapter 10- Use of N-acetylcysteine & Sodium Bicarbonate for prevention of CA-AKI	<ul style="list-style-type: none"> Recent randomized trial showed that <u>N-acetylcysteine</u> was no more effective than placebo at preventing CA-AKI for intra-arterial iodinated contrast media administration and is therefore <u>not recommended</u> for intravenous contrast media prophylaxis <u>Bicarbonate</u> is likely similar to normal saline for the prevention of CA-AKI, but it is <u>not preferred</u> due to the additional requirement for pharmacist compounding.
Renal Dialysis Patients and the Use of Iodinated Contrast Medium	<ul style="list-style-type: none"> Patients undergoing dialysis who make more than 1-2 cups of urine/day (100 mL) should be considered <u>nonanuric and treated as high-risk patients</u> similar to patients with AKI or eGFR less than 30 mL/min/1.73m² who are not undergoing hemodialysis. Patients should not <u>have acute dialysis nor continuous renal replacement therapy initiated or alter their schedule solely based on iodinated contrast media administration regardless of renal function</u> due to the risks, costs and lack of benefit .
Chapter 16- NSF- Identifying patients at-risk of NSF	<ul style="list-style-type: none"> <u>Addition of-</u> History of CKD or prior history of AKI to the list <u>Removal of-</u> History of hypertension requiring medical therapy from the list <u>Changed to Optional-</u> History of diabetes mellitus
Chapter 16- Calculating eGFR	<ul style="list-style-type: none"> Methods of calculating eGFR are in flux as efforts are underway to <u>remove race from clinical calculators</u>.
Chapter 16- Changes made to Additional Specific Recommendations for Specific Groups of Patients- Patients with end-stage renal disease on chronic dialysis	<ul style="list-style-type: none"> The ACR & NKF recommend that in patients who are already receiving dialysis, if feasible, <u>elective GBCA-enhanced MRI examinations be performed before regularly scheduled dialysis</u>. Due to the risks of catheter placement and infection, the possibility of worsening kidney function in patients with AKI and CKD, and the perceived very low risk of NSF from group II and III GBCM agents, <u>dialysis should not be initiated or altered in patients receiving a group II GBCM</u>



*An exciting
opportunity to share
rare & interesting
cases from your
practice!!!*





Abbreviated Product Information - Ultravist

Ultravist® (iopromide) Prescribing Information (Refer to package insert before prescribing) **Presentation:** Intravascular injections of nonionic iopromide in strengths of 300mg and 370mg of iodine/ml. Intravascular injections of nonionic iopromide in strengths of 300mg and 370mg of iodine/ml. Indications: For diagnostic use only. Delineation of the vascular and renal systems and of body cavities **Posology and administration:** Adults Intravenous urography: minimum doses: Ultravist 370: 0.8ml/kg body weight; Ultravist 300: 1ml/kg body weight. Children **Intravenous urography:** see package insert. **Adults Computed tomography:** Cranial CT: Ultravist 300: 1-max, 2ml/kg body weight; Ultravist 370: 1-max, 1.5ml/kg body weight. Whole-body CT: Dosage and administration rate depend on investigation and scanner. **Adults Angiography:** depends on age, weight, cardiac output, general condition, clinical problem, examination technique and the nature and volume of the vascular region to be investigated. (see package insert). **Paediatric population:** young infants (age < 1 year) and especially newborns are susceptible to electrolyte imbalance and haemodynamic alterations. Care should be taken regarding the dose of contrast medium to be given, the technical performance of the radiological procedure and the patient status. **Renal impairment:** to reduce the risk of additional contrast media-induced renal impairment in patients with pre-existing renal impairment, the minimum possible dose should be used (see package insert). **Hepatic impairment:** no dosage adjustment is necessary. **Elderly:** possibility of reduced renal function should be considered. **Contra-indications:** Uncontrolled thyrotoxicosis. **Warnings and precautions:** Can be associated with anaphylactoid/hypersensitivity reactions, ensure preparedness for institution of emergency measures. Allergy-like reactions from mild to severe possible, mostly within 30 min, but delayed reactions (hours to days) may occur. Particularly careful risk/benefit judgement required for patients with: known hypersensitivity to Ultravist or its excipients; previous reaction to any contrast medium or; history of bronchial asthma or allergic disorders (increased risk). Pre-medicate with corticosteroids if necessary. To minimise risk: administer Ultravist to recumbent patients; observe patients closely for 15 minutes and keep them in hospital for at least one hour after the last injection. Patients on beta-blockers may be resistant to the effects of beta agonists. If severe reaction occurs, patients with cardiovascular disease are more susceptible to serious or fatal outcomes. Caution in patients with: known/suspected hyperthyroidism or goitre, monitor thyroid function in neonates exposed via mother or during neonatal period. Caution in patients with cerebral arteriosclerosis, pulmonary emphysema, poor general health, renal insufficiency, dehydration, diabetes mellitus, multiple myeloma/ paraproteinaemia, repetitive and/ or large doses of Ultravist. Nephrotoxicity may occur or rarely acute renal failure. Ensure adequate hydration of patients; correct water or electrolyte imbalances before administration. With cardiac or severe coronary artery disease, increased risk of haemodynamic changes or arrhythmia. Intravascular injection may precipitate pulmonary oedema in patients with heart failure. Increased risk of neurological complications in patients with seizure history or CNS disorders. Caution in patients with reduced seizure threshold. May aggravate the symptoms of myasthenia gravis. Flush intravascular catheters frequently with physiological saline (if possible with addition of heparin) and minimise procedure length to minimise procedure-related thromboembolism risk. Patients with pheochromocytoma may be at increased risk of developing a hypertensive crisis. Minimise excitement, anxiety and pain. Do not use in myelography. Sensitivity testing is not recommended. **Interactions:** Consider interruption of biguanides treatment prior to Ultravist administration as a precaution against development of lactic acidosis. Prevalence of delayed reactions higher in patients who have received interleukin-2. Diagnosis and treatment of thyroid disorders with thyrotropic radioisotopes may be impeded for up to several weeks due to reduced radioisotope uptake. Pregnancy and lactation: Adequate and well-controlled studies in pregnant women have not been conducted. Safety for nursed infants has not been investigated. **Effects on ability to drive and use machines:** Driving or operating machinery is not advisable for 30 minutes after the last injection. **Undesirable effects:** Common: dizziness, headache, dysgeusia, blurred/disturbed vision, chest pain/ discomfort, hypertension, vasodilatation, vomiting, nausea, pain, injection site reactions (e.g. oedema, soft tissue injury post extravasation), feeling hot. Uncommon: Hypersensitivity/anaphylactoid reactions (anaphylactoid shock, respiratory arrest, bronchospasm, laryngeal/pharyngeal/face oedema, tongue oedema, laryngeal/pharyngeal spasm, asthma, conjunctivitis, lacrimation, sneezing, cough, mucosal oedema, rhinitis, hoarseness, throat irritation, urticaria, pruritus, angioedema), vasovagal reactions, confusional state, restlessness, paraesthesia/ hypoaesthesia, somnolence, arrhythmia, hypotension, dyspnea, abdominal pain, oedema. Rare: Anxiety, cardiac arrest, myocardial ischemia, palpitations. Frequency not known: Thyrotoxic crisis, thyroid disorder, coma, cerebral ischaemia/infarction, stroke, brain oedema, convulsion, transient cortical blindness, loss of consciousness, agitation, amnesia, tremor, speech disorders, paresis/paralysis, hearing disorders, myocardial infarction, cardiac failure, bradycardia, tachycardia, cyanosis, shock, thromboembolic events, vasospasm, pulmonary edema, respiratory insufficiency, aspiration, dysphagia, salivary gland enlargement, diarrhoea, bullous conditions (e.g. Stevens-Johnson's or Lyell syndrome), rash, erythema, hyperhidrosis, compartment syndrome in case of extravasation, renal impairment, acute renal failure, malaise, chills, pallor, body temperature fluctuation. *These adverse reactions may have a fatal or life-threatening outcome and are considered the most serious adverse drug reactions. Prescribers should consult the SmPC in relation to other side effects. **Overdose:** Symptoms may include fluid and electrolyte imbalance, renal failure, cardiovascular and pulmonary complications. Monitoring of fluids, electrolytes and renal function recommended in case of intravascular overdosage. Treatment of overdose should be directed towards the support of vital functions. Ultravist is dialysable. **Incompatibilities:** Because of possible precipitation, X-ray contrast media and prophylactic agents must not be injected as mixed solutions. **Special Precautions for Storage:** Protect from light and X-rays. **Date of revision of text:** April 2017. Please note: for current prescribing information refer to the package insert and/or contact your local Bayer HealthCare organization. **Bayer HealthCare Ltd.** 14th Floor, Oxford House, Taikoo Place, 979 King's Road, Quarry Bay, Hong Kong.



THANK YOU!





Polling Question:

What do scientific associations like American College of Radiology(ACR), NKF(National Kidney Foundation), and likewise, recommend on the choice of iodinated contrast medium between Low Osmolar Contrast Medium (LOCM) and Isosmolar Contrast Medium(IOCM), regarding renal safety.

- A. The LOCMs are better than IOCM
- B. There are no clinically relevant differences in CI-AKI risk between iso-osmolality and low-osmolality iodinated contrast media
- C. The IOCM is better than LOCMs
- D. None of the above