

Timely Information for Providers in South Carolina

An outreach service for Medicaid providers to help identify and prevent potential gaps in evidence-based care, as well as detect fraud, abuse, overuse or inappropriate use.

SCRIPTS Tips for Pharmacists

Optimizing Data Entry and Clinical Use

https://www.schealthviz.sc.edu/tipsc-1

SCROLL DOWN TO



MARCH 2024

TO CREATE A PATIENT REQUEST GO TO: <u>http://southcarolina.pmpaware.net</u>

- Click <MENU>, then <PATIENT REQUEST>
- Input required fields: FIRST NAME, LAST NAME, DOB (mm/dd/yyyy)
- Scroll down and click <SEARCH>

TO QUICKLY VIEW A REPORT

Patient, Test, 33F Refine Search

Date of Birth: Recent Address: 10/24/1988 View Linked Records (3)

Integrated Patient Records Total (3)	11.0	one administration details rst responders and major ilities	SEE THE FOLLOWING SECTIONS			
Incident Date	Medication Given	Dosage	Administered By	Zip code of Administration		
12/12/2021	Naloxone (Narcan)	1 mg	University Hospital	29401		
12/12/2021	Naloxone	1 Each	EMS	29407		
06/20/2021	Naloxone	1 Each	EMS	29407		

Prescriptions		Review controlled substances (CII-CIV) dispensed										
		to patient (default displays 2 years)										
08/22/2018	08/22/2018	3	Tramadol Hcl 50 Mg Tablet	60.00	30	Te Doc	20180822	App (1119)	0/0	10.00 MME	Private Pay	SC
04/15/2017	02/15/2017	2	Alprazolam 0.5 Mg Tablet	100.00	7	Bo Tes	305860	Gut (1119)	2/2	14.29 LME	Comm Ins	SC
03/15/2017	02/15/2017	2	Alprazolam 0.5 Mg Tablet	100.00	7	Bo Tes	305860	Gut (1119)	1/2	14.29 LME	Comm Ins	SC
)3/15/2017	02/15/2017	2	Oxycodone-Acetaminophen 5-325	100.00	7	Bo Tes	305862	Gut (1119)	1/0	107.14 MME	Comm Ins	SC
2/15/2017	02/15/2017	2	Alprazolam 0.5 M			Bo Tes	305860	Gut (1119)	0/2	14.29 t nE	Comm Ins	SC
2/15/2017	02/15/2017	2		y impacts		Bo Tes	305861	Gut (1119)	1/0	107.14 MME	Comm Ins	SC
2/02/2017	02/01/2017	1	Oxycodone-Aceta SCRIPTS	S Narx rep	orts;	Da Tes	4455	Dav (0000)	0/0	15.00 N ME	Comm Ins	SC
1/26/2017	01/26/2017	1	Oxycodone Hd () flip to backside Oxycodone Hd () for quick tips			AI Tes	3344	Car (8506)				SC
/26/2016	12/26/2016	1				AI Tes	2233	Car (8506)	Review calculated Morphine Milligram			SC
/26/2016	11/26/2016	1	Oxycontin Er 80 N	Ca Tes	1122	Car (5555)	SC					
0/31/2016	10/31/2016	1	Oxycodone-Acetaminophen 5-325	60.00	30	Da Tes	5566	Dav (0000)	Eau	ivalent (MM	E) per	SC
Disclaimer										for each ind		< 1
									pres	cription		

Morphine Milligram Equivalent Prescribed Over Time



Click desired time frame to view MME per day over time (*default is last 30 days*)





CONSIDERATIONS FOR REVIEWING SOUTH CAROLINA PRESCRIPTION A SCRIPTS Narx Report (also called a DHEC or PMP report) is one tool to help review and

WHAT IF:

APPARENTLY GOOD RESULTS (1 PHARMACY, 1 OPIOID PRESCRIBER)¹

- Does it match patient profile and personal presentation in the pharmacy (e.g., interactions, conversations)?
- Consider non-adherence behaviors not captured in results (e.g., binging, running out early).

WHAT IF:

TOTAL MORPHINE MILLIGRAM EQUIVALENTS PER DAY (MME/day)² SUGGESTS CONCERN FOR ADVERSE EVENTS OR OVERDOSE^{3,4}

- More recent guidelines recommend additional precautions when prescribing ≥ 50 MME/day including offering naloxone^{4,5} along with overdose prevention education to patients and caregivers.
- For patients already on higher opioid daily doses, risks and benefits of continuing or tapering opioid dose must be carefully weighed. Guidelines do not recommend abrupt tapering or discontinuation of opioids.

All homes that contain opioids carry some risk of overdose for the patient and others.

WHAT IF:

OVERDOSE RISK SCORE (ORS)⁶ SUGGESTS CONCERN FOR ADVERSE EVENTS OR OVERDOSE

- ORS indicates the risk of unintentional overdose death; the odds begin to increase sharply at a score of 200.⁶
- Look at the list of Key Contributing Factors⁷ to help identify potential patient risks.
- Use the ORS⁷ as a signal to review the complete patient profile and speak with patient and family to identify factors not captured in SCRIPTS that can influence unintentional overdose (e.g., depression, respiratory conditions).

The ORS alone is NOT sufficient information to decide to dispense or refuse to dispense medication.

WHAT IF:

COMBINATION OF OPIOID AND OTHER CONTROLLED SUBSTANCE(S), ESPECIALLY BENZODIAZEPINES^{8,9}

- Pain guidelines concur benzodiazepines and opioids are high risk combinations, especially in the elderly; many recommend against combination unless clearly indicated.
- Encourage patients to make sure all providers know they are on an opioid AND benzodiazepine; it is important for providers to monitor for respiratory depression if the benefits of the combination outweighs the risk.
- Check and offer naloxone^{4,5} along with overdose prevention education.

For more information on tapering opioids and/or benzodiazepines visit: <u>https://bit.ly/opioids_benzos</u>

WHAT IF:

OPIOID-ACETAMINOPHEN COMBINATION PRODUCT

- Counsel patient on risk of exceeding 4000 mg total daily acetaminophen dose. Consider 3000 mg total daily dose, especially if elevated liver function tests, known liver impairment, or older age; limit use to 2000 mg total daily dose in patients with alcohol use disorder or taking warfarin.
- Counsel patient to avoid alcohol and medications with alcohol (e.g., cough syrups) when taking acetaminophen unless provider has given different instructions.
- Remind patients about "hidden" acetaminophen and alcohol found in some medications and over-the-counter (OTC) allergy, cold, and sleep products (e.g., night time pain relievers).

Instructing patients and caregivers on the proper use of naloxone and taking time to have them demonstrate technique is one way to show you care



confirm a patient's controlled substance (CII - CIV) medication use and history

WHAT IF:

POTENTIAL ABERRANT BEHAVIOR (2 OR MORE PHARMACIES, 2 OR MORE OPIOID PRESCRIBERS)³

- Does it match patient interactions with you and staff or feedback from patient, family, and friends?
- Recognize the multiple reasons for possible inappropriate opioid use:
 - □ **ADDICTION** often characterized by behaviors that may include loss of control over drug use, craving, compulsive use, and continued use despite harm to health or relationships (See table at right)

Physical dependence and tolerance are normal physiologic adaptations to extended opioid therapy and are NOT the same as addiction.

- PHYSICAL DEPENDENCE biologic adaptation to drug that results in abstinence syndrome (signs and symptoms of withdrawal) upon cessation, rapid dose reduction and/or administration of antagonist
- □ **TOLERANCE** a physiologic state of reduced effect over time from regular drug exposure in which increased dosage is needed to produce specific effect (increase in dose and no increase in effect may mean opioid is ineffective)
- □ **HYPERALGESIA** increase in pain sensitivity that can be seen with rapid opioid dose escalation or high opioid dose (consider if increase in pain with increase in dose)
- □ **PSEUDO-ADDICTION** aberrant drug-related behaviors driven by uncontrolled pain (*relief seeking vs drug seeking*) that are reduced by improved pain control
- □ **OTHER PSYCHIATRIC ILLNESSES** such as anxiety, depression, PTSD, "chemical coping" (knowingly or unknowingly taking medications to decrease or numb negative emotions)
- DIVERSION moving medications from legal/ medically indicated users to illegal/unauthorized users

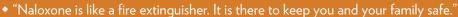
CONCERNING BEHAVIORS FOR ADDICTION

- Requests for increases in opioid dose
- Requests for specific opioid by name, "brand name only" or allergic to all but the desired opioid
- Overwhelming focus on opioids during visits instead of underlying disease process
- O Multiple office contacts regarding opioids
- O Unwilling to follow through with recommended therapy/ referrals (e.g., physical therapy)
- O Running out early due to unsanctioned dose escalation
- O Resistance to change therapy despite harm or negative consequences (e.g., over-sedation); unwilling to consider non-opioid therapy
- O Concurrent alcohol or substance abuse
- O Deterioration in function at home and work
- O Opposition to monitoring (e.g., pill counts, UDT)
- O Three or more requests for early refills
- O Multiple "lost," "spilled," or "stolen" opioid prescriptions
- O Multiple sources for opioids
- Illegal activities forging prescriptions, selling opioid prescriptions
- O Overdose

Familiarity with educating patients about use of naloxone⁵ can put you and the patient more at ease and potentially improve communication and outcomes (just like with inhalers and other medical devices).

Consider sharing that:

- "Prescribing naloxone is like prescribing epinephrine (Epi-pen®) to someone with a food allergy."
- "It is there to keep you safe in case something accidentally happens."



For more information about naloxone, including formulation-specific administration instructions, go to http://opirescue.com/rescue

⁶ An ORS of 000 is an error reading meaning no ORS could be calculated.



Adapted with permission: Boston University SCOPE of Pain Program www.scopeofpain.com

¹ Not all dispensed opioids require reporting to SCRIPTS, such as methadone dispensed from Opioid Treatment Programs (i.e., 'methadone clinics') or < 48-hour supply from emergency department.

² Morphine Milligram Equivalents (MME) is a mathematical conversion that standardizes risk evaluation of the different opioids.

 $^{^{3}}$ Increased risk of opioid overdose-related death has been associated with 4+ opioid prescriptions, 4+ pharmacies, or total MME/day \geq 100.

⁴ Multiple guidelines agree on co-prescribing naloxone to anyone currently on chronic opioids in ANY ONE of the following higher risk groups: opioid dose ≥ 50 MME/day; concomitant benzodiazepine and opioid use; history of opioid overdose or substance use disorder; respiratory conditions (e.g., COPD, sleep apnea); mental health condition(s); excessive alcohol use; AND previously on chronic opioids with a lost tolerance to previous dose and at risk of resuming that dose using prescription or illicit drugs (e.g., opioid taper underway, recent release from prison or detoxification facility).

⁵ Multiple formulations and dosages of opioid overdose antidotes (e.g., naloxone, nalmefene) are now available; evidence suggests high-dose version of naloxone (8 mg) does not improve survival rates compared to 4 mg.

⁷ Key Contributing Factors displayed and used in the ORS calculation include: greater than 6 opioid prescriptions dispensed; benzodiazepine-narcotics overlap; # of high-risk prescriptions dispensed; # of pharmacies where opioids/sedatives dispensed; total days supply of short-acting opioid prescriptions; and # of overlapping opioid/sedative dispensations.

⁸Benzodiazepines and opioid medication labelings carry black box warnings highlighting the risks associated with concomitant use.

⁹Lorazepam Milligram Equivalent (LME) values in SCRIPTS offer one way to compare sedative hypnotic medications for dose-related risk considerations.

DATA ENTRY TIPS THAT MAY IMPROVE SCRIPTS NARX REPORTS

Data entry impacts SCRIPTS Narx reports for everyone – affecting timely refills and the ability to identify clinical concerns. Basic suggestions to prevent multiple patient records for the same patient are to carefully enter:

- ✓ Legal FIRST and LAST NAME as it appears on driver's license (avoid nicknames and middle names)
- ✓ Date of Birth
- ✓ FULL Address (utilize United States postal service [USPS] address standards to help minimize creation of multiple PMP records for an individual patient). To look up USPS approved abbreviations, go to https://pe.usps.com/text/pub28/28c2_001.htm
- ✓ AVOID special characters (to help minimize data rejection or creation of multiple records for an individual patient)

Other suggestions to help minimize issues with refills and evaluating SCRIPTS Narx reports are to be mindful of:

- ✓ "Days Supply" especially for as-needed prescriptions (can affect calculated daily MMEs, refill dates, and patient care)
- ✓ "Fill Date" (can potentially affect timing for refills)

Appreciate that accurate data entry into any field is key; even data entered into non-essential fields may have the potential to affect algorithms and data uploading to SCRIPTS.

REFERENCE LIST

AMA Opioid Task Force. Help save lives: co-prescribe naloxone to patients at risk of overdose. Chicago: American Medical Association; August 2017. Accessed March 7, 2024. Available from: https://www.end-opioid-epidemic.org/wp-content/ uploads/2017/08/AMA-Opioid-Task-Force-naloxone-one-pager-updated-August-2017-FINAL.pdf

An Act to Amend the Code of Laws of South Carolina, 1976, by Adding Section 44-53-361 so as to Require Prescribers to Offer a Prescription for Naloxone Hydrochloride or Other Approved Drug to a Patient Under Certain Circumstances and for Other Purposes, S1.A3,C53,T44 (SC 2021). April 26, 2021. Accessed March 7, 2024. Available from: https://www. scstatehouse.gov/sess124_2021-2022/prever/571_20210420.htm

Arizona Department of Health Services. Arizona Opioid Prescribing Guidelines. 2018. Accessed March 6, 2024. Available from: https://www.azdhs.gov/documents/audiences/clinicians/clinical-guidelines-recommendations/prescribing-guidelines.pdf

Bamboo Health 2023. Narx Care Support Center. About Overdose Risk Scores. Accessed March 6, 2024. Available from: https://narxcare.zendesk.com/hc/en-us/categories/4409656763411-About-Overdose-Risk-Scores

Bamboo Health: New Version of the Unintentional Overdose Risk Score Model; December 4, 2023. Internal Document

Bamboo Health 2023. Understanding Narxcare and The New Score Tile. Accessed March 6, 2024. Available from: https:// bamboohealth.com/wp-content/uploads/2023/10/NarxCare-Fact-Sheet_No-Subhead_FINAL-V09.11.2023.pdf

Calculating total daily dose of opioids for safer dosage. Centers for Disease Control and Prevention website. Updated August 28, 2019. Accessed March 6, 2024. Available from: https://www.cdc.gov/opioids/providers/prescribing/pdf/calculating-total-daily-dose.pdf

Coffin PO, Behar E, Rowe C, et al. Nonrandomized intervention study of naloxone coprescription for primary care patients receiving long-term opioid therapy for pain. Ann Intern Med. 2016;165(4):245-252. doi:10.7326/M15-2771

Crawford C. CDC Warns of Misapplication of Its Opioid Guideline. American Academy of Family Physicians website. May 9, 2019. Accessed March 7, 2024. https://www.aafp.org/news/health-of-the-public/20190509cdcopioidgdln.html

Dowell D, Ragan KR, Jones CM, Baldwin GT, Chou R. CDC Clinical Practice Guideline for Prescribing Opioids for Pain — United States, 2022. MMWR Recomm Rep 2022;71(No. RR-3):1–95. DOI: http://dx.doi.org/10.15585/mmwr.rr7103a1

FDA requires strong warnings for opioid analgesics, prescription opioid cough products, and benzodiazepine labeling related to serious risks and death from combined use. U.S. Food and Drug Administration website. August 31, 2016. Accessed March 6, 2024. https://www.fda.gov/news-events/press-announcements/fda-requires-strong-warnings-opioid-analgesics-prescription-opioid-cough-products-and-benzodiazepine

Gifford JD, Anderson JE, Baley JM, et al. Guidelines for the Chronic Use of Opioid Analgesics. Federation of State Medical Boards website. April 2017. Accessed June 3, 2020. https://www.fsmb.org/siteassets/advocacy/policies/opioid_guidelines_as_adopted_april-2017_final.pdf

Gwira Baumblatt JA, Wiedeman C, Dunn JR, Schaffner W, Paulozzi LJ, Jones TF. High-risk use by patients prescribed opioids for pain and its role in overdose deaths. JAMA Intern Med. 2014;174(5):796-801. doi:10.1001/jamaint-ennmed.2013.12711

Gebauer MG, Nyfort-Hansen K, Henschke PJ, Gallus AS. Warfarin and acetaminophen interaction. Pharmacotherapy. 2003;23(1):109-112. doi:10.1592/phco.23.1.109.31913

Häuser W, Morlion B, Vowles KE, et al. European* clinical practice recommendations on opioids for chronic noncancer pair - Part 1: Role of opioids in the management of chronic noncancer pain. Eur J Pain. 2021 May;25(5):949-968. doi: 10.1002/ ejp.1736. Epub 2021 Mar 2. PMID: 33655607; PMCID: PMC8248186

Hurstak EE, Kushel M, Chang J, et al. The risks of opioid treatment: perspectives of primary care practitioners and patients from safety-net clinics. Subst Abus. 2017;38(2):213-221. doi:10.1080/08897077.2017.1296524

Interagency Guideline on Prescribing Opioids for Pain. Washington State Agency Medical Directors' Group website. June 2015. Accessed March 6, 2024. http://www.agencymeddirectors.wa.gov/Files/2015AMDGOpioidGuideline.pdf

Larson AM, Polson J, Fontana RJ, et al. Acetaminophen-induced acute liver failure: results of a United States multicenter, prospective study. Hepatology. 2005;42(6):1364-1372. doi:10.1002/hep.20948

LiverTox: Clinical and Research Information on Drug-Induced Liver Injury. Bethesda (MD): National Institute of Diabetes and Digestive and Kidney Diseases; 2012. Acetaminophen. Updated January 28, 2016. Available from: https://www.ncbi. nlm.nih.gov/books/NBK548162/

Naloxone Can Save a Life. Charleston: tipSC; April 2020. Accessed March 6, 2024. Available from: https://schealthviz. sc.edu/Data/Sites/1/media/downloads/tipsc_resources/tipsc_mailer_v10_hyperlinks_%2002%20naloxone%20sp%20ed.pdl

NaloxoneSavesSC.org. c2017. Accessed March 7, 2024. Available from: http://naloxonesavessc.org/

Naloxone: the opioid reversal drug that saves lives. Washington: Department of Health and Human Services (US); Accessed March 7, 2024. Available from: https://www.hhs.gov/system/files/naloxone-coprescribing-guidance.pdf

Pain, Palliative Care, and Addiction Special Interest Group (PPCA SIG). Let's talk about naloxone – it saves lives. Washington: American Pharmacists Association: Accessed March 7, 2024 p. Available from: https://georgiagerontologysociety.org/ wp-content/uploads/2019/08/Lets TalkAboutNaloxone-1.pdf

Payne ER, Stancliff S, Rowe K, Christie JA, Dailey MW. Comparison of Administration of 8-Milligram and 4-Milligram Intranasal Naloxone by Law Enforcement During Response to Suspected Opioid Overdose - New York, March 2022-August 2023. MMWR Morb Mortal Wkly Rep. 2024 Feb 8;73(5):110-113. doi: 10.15585/mmwr.mm7305a4. PMID: 38329911; PMCID: PMC10861201

PMP AWARxE[™] South Carolina Prescription Drug Monitoring Program Data Submission Guide for Dispensers Version 2.1. South Carolina Department of Health and Environmental Control website. January 2020. Accessed May 20, 2020. https://www.scdhec.gov/sites/default/files/media/document/sc-pmp-data-submission-dispenser-guide-Jan2020.pdf

PMP AWARXE^{III} South Carolina Prescription Drug Monitoring Program Requester User Support Manual Version 2.3. South Carolina Department of Health and Environmental Control website. February 2020. Accessed June 3, 2020. Available from: https://dib1sdx6nwlphm.cloudfront.net/aware/sc_aws_prod/narxcare_user_guide.pdf

PrescribeToPrevent.org. San Francisco: PrescribeToPrevent.org; c2022. Accessed February 21, 2024. Available from: https://prescribetoprevent.org

Rich RLC Jr. Prescribing Opioids for Chronic Pain: Unintended Consequences of the 2016 CDC Guideline. Am Fam Physician. 2020 Apr 15;101(8):458-459. PMID: 32293841

Royal College of Physicians. Chronic pain (primary and secondary) in over 16s: assessment of all chronic pain and management of chronic primary pain. London (UK): NICE; April 7, 2021. Accessed March 20, 2023. Available from: https://www. nice.org.uk/guidance/ng193/chapter/Recommendations#managing-chronic-primary-pain

San Francisco Department of Public Health. Naloxone for Opioid Safety: A provider's guide to prescribing naloxone to patients who use opioids. January 2015. Accessed March 7, 2024. Available from: https://www.chcf.org/wp-content/uploads/2017/12/PDF-NaloxoneOpioidSafetyProviders.pdf

SCRIPTS (PDMP Reports) Quick Tips and Tricks. tipSC. October 2022. Accessed March 6, 2022. Available from: https:// schealthviz.sc.edu/Data/Sites/1/media/downloads/tipsc_resources/scripts_tips_tricks.pdf

SO.S. for Safer Opioid Prescribing, Charleston: tipSC; August 2020. Accessed March 17, 2023. Available from: https:// schealthviz.sc.edu/Data/Sites/1/media/downloads/tipsc_resources/SOS%20for%20Safer%20Opioid%20Prescribing%20 2020%20with%20hyperlinks.pdf

South Carolina Department of Health and Environmental Control. Columbia (SC): South Carolina Department of Health and Environmental Control; c2018. Prescription Monitoring; 2022. Accessed March 6, 2024. Available from: https://scdhec.gov/healthcare-quality/drug-control-register-verify/prescription-monitoring

Systematic Review on Opioid Treatments for Chronic Pain: Surveillance Report 3. Rockville (MD): AHRQ; March 16, 2022. Accessed March 17, 2023. Available from: https://effectivehealthcare.ahrq.gov/sites/default/files/product/pdf/opi-oid-chronic-pain-surveillance-report-3.pdf

Talem Health[™]. Continuing Pharmacy Education. Data Quality in Prescription Monitoring Programs. Accessed March 12, 2024. Available from: https://ce.talemhealth.com/a/HLQIJS

Tylenol® for Healthcare Professionals. Adult Dosing Charts. Accessed March 13, 2024. Available from: https://www.tylenolprofessional.com/adult-dosage

VA/DoD Clinical Practice Guideline. (2022). Use of Opioids in the Management of Chronic Pain Work Group. Washington, DC: U.S. Government Printing Office

VA PBM Academic Detailing Service. Acute pain management: meeting the challenges. A VA Clinician's Guide. Washington (DC) Department of Veteran Affairs, Department of Defense: July 2017. Accessed March 7, 2024. Available from: https://www.pbm.va.gov/PBM/AcademicDetailingService/Documents/508/IB10-998PAIN-Provider-AcutePainProvider-EducationalGuide_508Ready.pdf

U.S. Food and Drug Administration. Silver Spring (MD): FDA drug safety communication: prescription acetaminophen products to be limited to 325 mg per dosage unit; boxed warning will highlight potential for severe liver failure. January 11, 2011; updated February 7, 2018. Accessed March 7, 2024. Available from: https://www.fda.gov/drugs/drug-safety--and-availability/fda-drug-safety--communication-prescription-acetaminophen-products-be-limited-325-mg-dosage-unit

U.S Food and Drug Administration. Silver Spring (MD): Benzodiazepines and opioid medication labelings carry black box warnings highlighting the risks associated with concomitant use 2016 Aug 31. Accessed March 7, 2024. Available from: https://www.fda.gov/drugs/information-drug-class/new-safety-measures-announced-opioid-analgesics-prescription-opioid-cough-products-and#:~:text=FDA%20is%20warning%20patients%20and,breathing%2C%20coma%2C%20and%20 death

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The information contained in this summary is intended to assist pharmacists and pharmacy technicians in the management of chronic non-cancer pain in adults in the primary care setting. This information is advisory only and is not intended to replace sound clinical judgement, nor should it be regarded as a substitute for individualized diagnosis and treatment. Special considerations are needed when treating some populations with certain conditions (such as respiratory/sleep disorders; cardiac, liver and renal impairment; debility; addiction; and pregnancy/breast-feeding).