

Neutropenia following Kidney Transplant

I. PURPOSE

Identification and treatment plan for neutropenia after receiving a kidney transplant.

II. SETTING

Both inpatient and outpatient

III. POLICY

DEFINITIONS:

Neutropenia based on ANC count/mm3:

Mild: 1000-1500 Moderate: 500-1499 Severe: ANC < 500 Agranulocytosis: < 100

Leukopenia and neutropenia are common after kidney transplant.

Causes:

- Medications: MMF, tacrolimus, SMX, Ganciclovir, valganciclovir, acyclovir, azathioprine, Rituximab.
- Thymoglobulin in the first 2 weeks.
- Other medications: Cephalosporins, vancomycin, Oseltamivir, Semisynthetic penicillin, leflunomide, furosemide, propranolol, ACEI, Dapsone
- Infections: particular attention CMV, and EBV.
- Nutritional deficiency (short gut and malabsorption): vitamin B12, folic acid, copper, ceruloplasmin, methylmalonic acid and homocysteine
 - In the first 2 weeks after transplant leukopenia is most likely secondary to induction with thyomoglobulin and is expected to improve without intervention.
 - Dapsone can cause severe neutropenia.

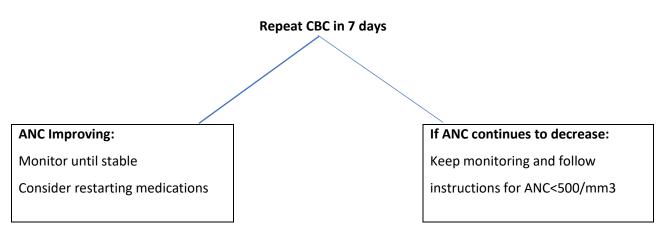
Procedure/Management:

- Admit if unwell or febrile with severe (ANC <500) and moderate neutropenia.
- Screen for viral infections: CMV, EBV viral loads and others based on clinical presentation.
- For patients at high risk of nutritional deficiencies consider checking the above vitamin and trace element levels if neutropenia persists
- ANC <500/mm3
- Check CMV viral load
- GCSF 5mcg/kg SQ for 3-5 days

- If response but neutropenia recurs, or if no response, stop/hold TMP-SMX (consider substituting with alternative in first 6 months: Dapsone or Atovaquone)
- If no response, reduce MMF, Azathioprine dose by 25%-50% Hold mTORI containing regimens
- Hold valcyte ulness CMV viral load positive or if high risk CMV status, monitor CMV viral load
- Repeat CBC 1-2 days after GCSF completion and then 1 week later to assess For ANC >500/mm3: monitor CBC weekly until neutropenia stabilizes

ANC 500-1000/mm3

- CMV viral load negative:
 - First 6 months after transplant: replace SMX with: pentamidine, atovaquone,
 Dapsone if not G6PD deficient.
 - More than 6 months after transplant: Hold SMX
- Hold valcyte, monitor CMV viral load



If immunosuppressive does are reduced, monitor DSA and Allosure closely.

IV. RESPONSIBILITY

Maha Haddad, MD Pediatric Nephrology

V. REFERENCES

See below

VI. REVIEWED BY

- Lavjay Butani, MD
- Arun Kale, MD
- Stephanie Nguyen, MD
- Gia Oh, MD

- Machi Kaneko, MD

VII. REVIEWED DATE and REVIEW CYCLE: 6/24/2024

References:

- 1. Baradaran H, Hashem Zadeh A, Dashti-Khavidaki S, Laki B. Management of drug-induced neutropenia, thrombocytopenia, and anaemia after solid organ transplantation: A comprehensive review. J Clin Pharm Ther. 2022;47(12):1895-912.
- 2. Harshman LA, Williams R, Engen RM. Neutropenia in pediatric solid organ transplant. Pediatr Transplant. 2022;26(8):e14378.
- 3. Ingold L, Halter J, Martinez M, Amico P, Wehmeier C, Hirt-Minkowski P, et al. Short- and long-term impact of neutropenia within the first year after kidney transplantation. Transpl Int. 2021;34(10):1875-85.
- 4. Jarasvaraparn C, Choudhury S, Rusch C, Nadler M, Liss KHH, Stoll J, et al. Characteristics, risk factors, and outcomes of neutropenia after liver or kidney transplantation in children. Pediatr Transplant. 2022;26(1):e14131.
- 5. Keles M, Yildirim R, Uyanik A, Turkmen M, Bilen Y, Aydinli B, et al. Neutropenia related to valacyclovir and valganciclovir in 2 renal transplant patients and treatment with granulocyte colony stimulating factor: a case report. Exp Clin Transplant. 2010;8(2):181-3.
- 6. Regev-Sadeh S, Borovitz Y, Steinberg-Shemer O, Gilad O, Shoham S, Yacobovich J. Cytopenias in pediatric kidney transplant recipients: preceding factors and clinical consequences. Pediatr Nephrol. 2023;38(10):3445-54.
- 7. Varnell CD, Fukuda T, Kirby CL, Martin LJ, Warshaw BL, Patel HP, et al. Mycophenolate mofetil-related leukopenia in children and young adults following kidney transplantation: Influence of genes and drugs. Pediatr Transplant. 2017;21(7).

Medical Legal Disclaimer:

Welcome to the UC Davis Health, Department of Pediatrics, Clinical Practice Guidelines Website. All health and health-related information contained within the Site is intended chiefly for use as a resource by the Department's clinical staff and trainees in the course and scope of their approved functions/activities (although it may be accessible by others via the internet). This Site is not intended to be used as a substitute for the exercise of independent professional judgment. These clinical pathways are intended to be a guide for practitioners and may need to be adapted for each specific patient based on the practitioner's professional judgment, consideration of any unique circumstances, the needs of each patient and their family, and/or the availability of various resources at the health care institution where the patient is located. Efforts are made to ensure that the material within this Site is accurate and timely but is provided without warranty for quality or accuracy. The Regents of the University of California; University of California, Davis; University of California, Davis, Health nor any other contributing author is responsible for any errors or omissions in any information provided or the results obtained from the use of such information. Some pages within this Site, for the convenience of users, are linked to or may refer to websites not managed by UC Davis Health. UC Davis Health does not control or take responsibility for the content of these websites, and the views and opinions of the documents in this Site do not imply endorsement or credibility of the service, information or product offered through the linked sites by UC Davis Health. UC Davis Health provides limited personal permission to use the Site. This Site is limited in that you may not:

- Use, download or print material from this site for commercial use such as selling, creating course packets, or posting information on another website.
- Change or delete propriety notices from material downloaded or printed from it. · Post or transmit any unlawful, threatening, libelous, defamatory, obscene, scandalous, inflammatory, pornographic, or profane material, any propriety information belonging to others or any material that could be deemed as or encourage criminal activity, give rise to civil liability, or otherwise violate the law.
- Use the Site in a manner contrary to any applicable law.

You should assume that everything you see or read on this Site is copyrighted by University of California or others unless otherwise noted. You may download information from this Site as long as it is not used for commercial purposes, and you retain the proprietary notices. You may not use, modify, make multiple copies, or distribute or transmit the contents of this Site for public or commercial purposes without the express consent of UC Davis Health.