# Hand Foot Skin Reaction and Hand Foot Syndrome: What's the Difference?

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## **Hand Foot Skin Reaction**

## Pathophysiology<sup>1</sup>

- Not fully understood
- Likely a combination of mechanisms
- Vascular endothelial growth factor inhibition + platelet-derived growth factor receptors inhibition

### Causes<sup>2</sup>

- Multi-targeted kinase inhibitors (MKI) to varying degrees
- Sorafenib
- Regorafenib

## • Symptoms<sup>2</sup>

- Hyperkeratotic lesions or calluses
- Blisters
- Inability to complete activities of daily living due to associated pain

## **Prevention and Treatment**

## • Prevention<sup>2</sup>

- Daily skin care
- Monitor for calluses
- Exfoliate calluses with a pumice stone
- Avoid hot water
- Wear nonrestrictive thick cotton socks
- Thick moisturizing cream or 20-40% urea to protect skin barrier

## • Treatment<sup>2</sup>

- Urea 20-40% if not already using
- Pain management
- NSAIDs
- GABA analogs
- Opioids, if necessary
- Dose reduction of MKI, depending on grade
- Topical anti-infectives
- May need temporary or permanent break

## Picture 1: Hand Foot Syndrome<sup>3</sup>



Picture 2: Hand Foot Skin Reaction<sup>4</sup>



#### References

- McLellan B, Ciardiella F, Lacouture ME, Segaert S, Van Cutsem E. Regorafenib-associated hand-foot skin reaction: practical advice on diagnosis, prevention, and management. *Ann Oncol.* 2015;26(10): 2017-2026.
   DOI: 10.1093/annonc/mdv.244.
   Lacouture ME, Wu S, Robert C, et al. Evolving strategies for the management of hand-foot skin reaction associated with the multitargeted kinase inhibitors sorafenib and sunitinib. *Oncologist*. 2008;13(9):1001-1011. DOI:10.1634/theoperalogist. 2008-0131.
- NSW Government. Hand-foot syndrome (HFS) palmar-plantar erythrodysaesthesia (PPE) associated with chemotherapeutic agents. Accessed March 17, 2022. https://www.eviq.org.au/clinical-resources/side-effect-and-toxicity-management/hair-skin-and-nails/1416-hand-foot-syndrome-associated-with-chemothera#assessment
- 10.1007/s40257-016-0197-1.
  5) Kwakman JJM, Elshot YS, Punt CJA, Koopman M. Management of cytotoxic chemotherapy-induced hand-foot syndrome. *Oncol Rev*. 2020; 14(1): 442. DOI: 10.4081/oncol.2020.442.

## **Hand Foot Syndrome**

## Pathophysiology<sup>5</sup>

- Different for each class medication
- 5-FU & capecitabine metabolites cause WBC infiltration, dilated blood vessels, and edema
- Liposomal doxorubicin concentrates in hands and feet, activates chemokine signaling pathways, and causes keratinocyte apoptosis

#### • Causes<sup>5</sup>

- 5-Fluorouracil
- Capecitabine
- Liposomal doxorubicin

## • Symptoms<sup>5</sup>

- Neuropathy in hands and feet
- Redness
- Scaling of skin
- Swelling
- Pain

## Prevention and Treatment

## • Prevention<sup>5</sup>

- Daily skin care
- Monitoring signs and symptoms
- Thick moisturizing cream such as CeraVe
- Wear thick, nonrestrictive socks
- Avoid hot water
- Decrease friction to hands and feet
- Avoid items, such as pumice stones

## • Treatment<sup>5</sup>

- Continue prevention efforts
- Urea 20-40%
- Pain management primarily with NSAIDs
- Dose reduction of anti-cancer therapy
- Adjustment of treatment cycles
- One on/one off for 5-FU-based regimens instead of two on/one off