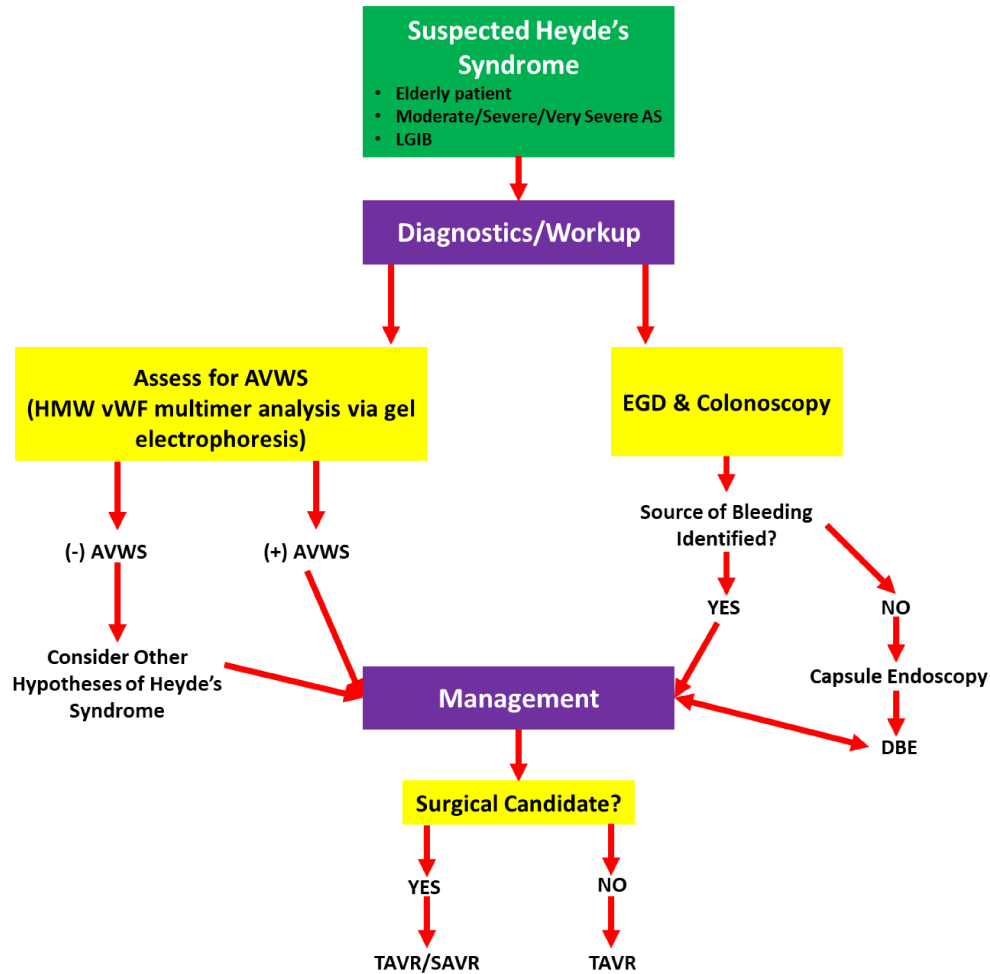


**Supplemental Fig. 1** Flowchart Detailing the Workup and Management of Suspected Heyde's Syndrome.



AS: Aortic Stenosis; LGIB: Lower Gastrointestinal Bleeding; AVWS: Acquired Von Willebrand Syndrome; HMW: High Molecular Weight; vWF: Von Willebrand Factor; EGD: Esophagogastroduodenoscopy; DBE: Double Balloon Enteroscopy; TAVR: Transcatheter Aortic Valve Replacement; SAVR: Surgical Aortic Valve Replacement

Appendix A**9 June 21****PubMed/MEDLINE (191)**

(aortic stenosis OR aortic valve stenosis OR "Aortic Valve Stenosis"[Mesh]) AND (gastrointestinal bleeding OR gastrointestinal hemorrhage OR hematochezia OR melena OR "gastrointestinal hemorrhage"[MeSH Terms] OR angiodysplasia OR "angiodysplasia"[MeSH Terms]) AND ("von Willebrand"[tiab] OR "von Willebrand Diseases"[Mesh]) OR ("heyde"[tiab])

**9 June 21****CINAHL - EBSCO (51)**

Search	Statement	Results
#10	#6 OR #9	51
#9	#7 OR #8	46
#8	AB heyde	20
#7	TI heyde	38
#6	#1 AND #2 AND #5	12
#5	#3 OR #4	1,618
#4	AB "von Willebrand"	1,371
#3	TI "von Willebrand"	552
#2	"gastrointestinal bleeding" OR "gastrointestinal hemorrhage" OR hematochezia OR melena OR angiodysplasia OR angiodysplasias	9,937

#1	“aortic stenosis” OR “aortic valve stenosis” OR “aortic valve stenoses”	7,156
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**10 June 21**

**Web of Science (Core Collection, Biological Abstracts, SciELO Citation Index),  
Timespan=all years (329)**

Search	Statement	Results
#10	#9 OR #8	329
#9	#5 AND #2 AND #1	111
#8	#7 OR #6	259
#7	TI=("heyde syndrome" OR "heyde's syndrome" OR "heyde-*") NOT TI=heyde theor*	184
#6	AB=("heyde syndrome" OR "heyde's syndrome" OR "heyde-*") NOT AB=heyde theor*	141
#5	#4 OR #3	24,453
#4	AB=("von Willebrand")	19,763
#3	TI=("von Willebrand")	11,115
#2	TS=("gastrointestinal bleeding" OR "gastrointestinal hemorrhage" OR hematochezia OR melena OR angiodysplasia OR angiodysplasias)	49,989
#1	TS=("aortic stenosis" OR "aortic valve stenosis" OR "aortic valve stenoses")	37,407

**10 June 21**

**Google Scholar (23)**

allintitle: heyde syndrome case OR report OR series

[searched “allintitle” to avoid pulling author last names]

## Appendix B

### **Inclusion Criteria:**

- Sex: Males and Females
- Age:  $\geq 18$  years old
- Valve Type: Natural
- Definition of Heyde's Syndrome:
  - Diagnosis of Heyde's Syndrome explicitly stated **&**
  - Evidence of aortic stenosis (based on echocardiogram/cardiac catheterization but not physical exam alone) **&**
  - GI bleed (hematochezia, melena, hematemesis) and/or angiodysplasia/AVMs but not iron deficiency anemia alone
- Language: English
- Study Design: Case reports/series and letter to the editor

### **Exclusion Criteria:**

- Age:  $< 18$  years old
- Valve Type: Mechanical, prosthetic
- Definition of Heyde's Syndrome: Cases presenting like Heyde's Syndrome but involve LVAD, ECMO, HOCM, mitral valve conditions rather than aortic stenosis
- Language: Any language other than English
- Study Design:
  - Prospective/retrospective cohort study
  - Case-control study
  - Reviews that do not include a case report/series
  - Randomized controlled trial
  - Abstract/conference proceedings