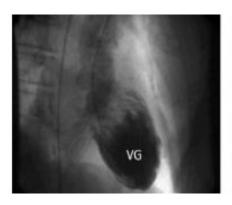
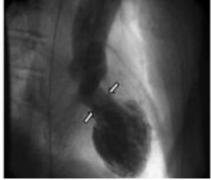
# Takotsubo Syndrome: Prediction of hospitalization outcomes

Python Machine Learning Project
July 10th 2021





Daisuke KUWABARA Nesrine BENANTEUR

# **OUTLINE**

Introduction: Reminders of basic heart physiology to understand the context

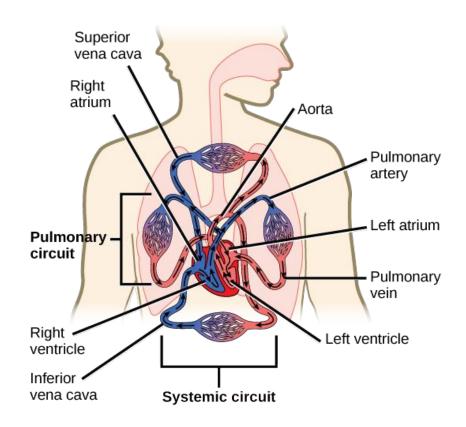
**Takotsubo Syndrome** 

Description of the cohorts and the variables used for prediction

**Algorithms predictions** 

**Conclusion: Perspectives and improvements** 

#### GENERAL HEART PHYISOLOGY TO UNDERSTAND THE CONTEXT

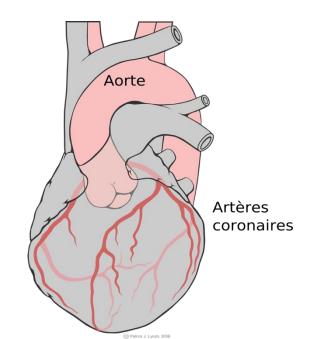


The heart is made of specialized cardiac muscle tissue called **myocardium**.

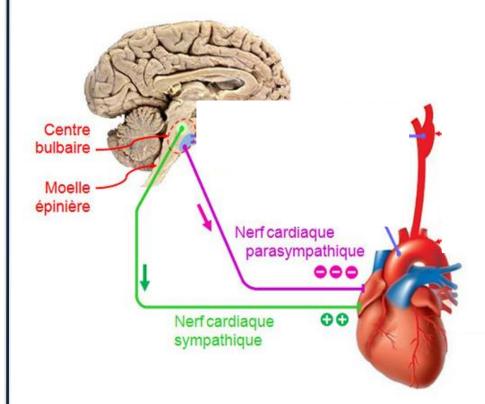
It is made of specific cells called cardiomyocytes.

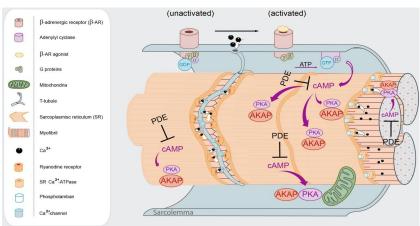
Circulatory system: network consisting of blood, blood vessels, and the heart.

This network supplies tissues in the body with oxygen and other nutrients, transports hormones, and removes unnecessary waste products.



# GENERAL HEART PHYISOLOGY TO UNDERSTAND THE CONTEXT



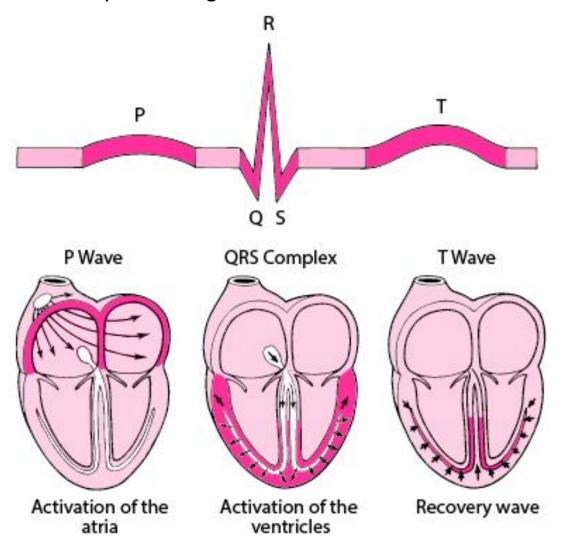


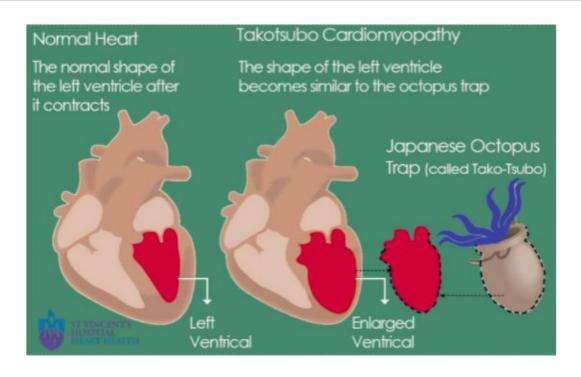
How does the heart beat?

- Nerve conduction: sympathic cardiac nerve (increases the heartbeat) or parasympathic cardiac nerve (decreases the heartbeat).
- Biochemicals: catecholamines (adrenaline, noradrenaline) are released to increases the contraction of the cardiomyocytes.

# **GENERAL HEART PHYISOLOGY TO UNDERSTAND THE CONTEXT**

▶ The heart rate can be analyzed through an exam called ECG.

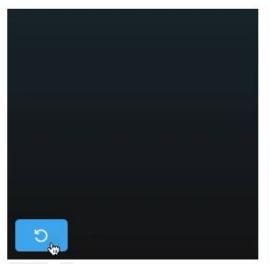


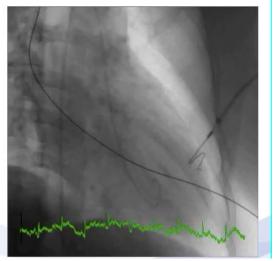


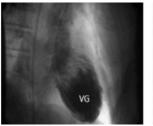
Takotsubo cardiomyopathy is a transient weakening of the left ventricle, the heart's main pumping chamber.
Usually: result of severe

emotional or physical

stress.

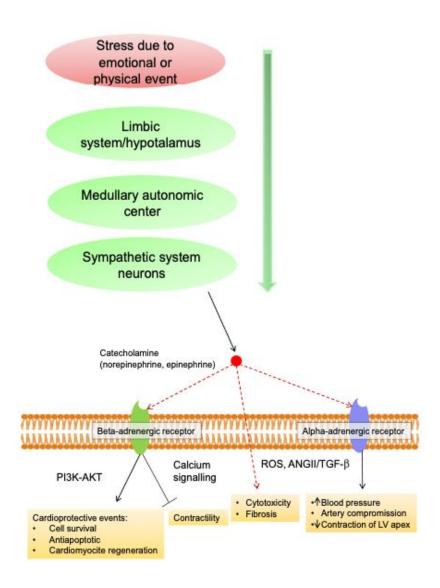








# **Possible causes:**

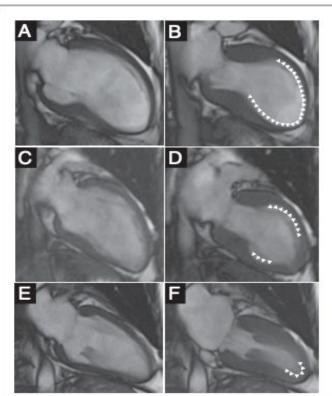


Lots of hypothesis today, but articles seem to all incriminate a catecholamine toxicity.

High prevalence in middle-aged women seem to show that estrogen deficiency might predispose them to the disease.

# <u>Criterion for Takotsubo diagnosis:</u> (Mayo Clinic, 2004)

1. Transient hypokinesis, akinesis, or dyskinesis of the left ventricular mid-segments with or without apical involvement; the regional wall motion abnormalities extend beyond a single epicardial vascular distribution; a stressful trigger is often, but not always present.



A, C, E: Diastole – heart relaxing and filling up with blood.

B, D, F: Systole – heart contraction

Diversity of left ventricle contraction patterns

# <u>Criterion for Takotsubo diagnosis:</u> (Mayo Clinic, 2004)

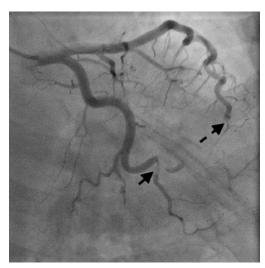
- 1. Transient hypokinesis, akinesis, or dyskinesis of the left ventricular mid-segments with or without apical involvement; the regional wall motion abnormalities extend beyond a single epicardial vascular distribution; a stressful trigger is often, but not always present.
- 2. Absence of obstructive coronary disease or angiographic evidence of acute plaque rupture.



Normal coronary arteries on the left side of the heart



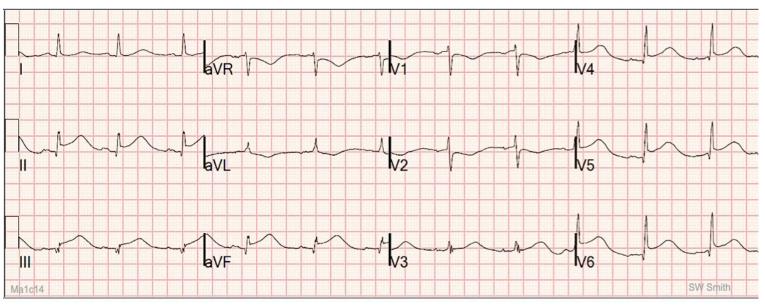
Normal coronary arteries on the right side of the heart

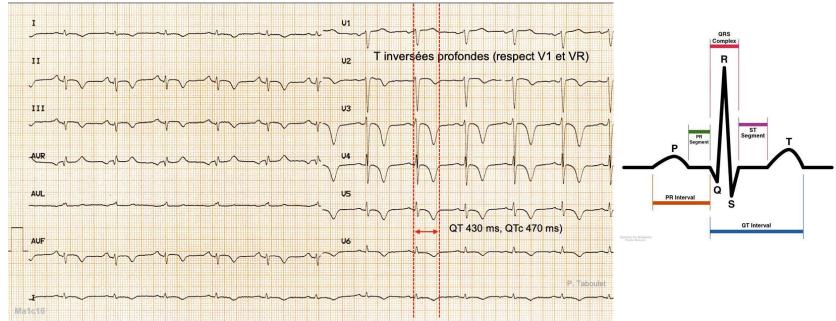


Coronary arteries obstructed

# <u>Criterion for Takotsubo diagnosis:</u> (Mayo Clinic, 2004)

- 1. Transient hypokinesis, akinesis, or dyskinesis of the left ventricular mid-segments with or without apical involvement; the regional wall motion abnormalities extend beyond a single epicardial vascular distribution; a stressful trigger is often, but not always present.
- 2. Absence of obstructive coronary disease or angiographic evidence of acute plaque rupture.
- 3. New electrocardiographic abnormalities (either ST-segment elevation and/or T-wave inversion) or modest elevation in cardiac troponin.





# <u>Criterion for Takotsubo diagnosis:</u> (Mayo Clinic, 2004)

- 1. Transient hypokinesis, akinesis, or dyskinesis of the left ventricular mid-segments with or without apical involvement; the regional wall motion abnormalities extend beyond a single epicardial vascular distribution; a stressful trigger is often, but not always present.
- 2. Absence of obstructive coronary disease or angiographic evidence of acute plaque rupture.
- 3. New electrocardiographic abnormalities (either ST-segment elevation and/or T-wave inversion) or modest elevation in cardiac troponin.
- 4. Absence of: a. Pheochromocytoma



## b. Myocarditis



# What happens during a Takotsubo event?

Left ventricle pumps
blood less efficiently

Amount of blood going out
of the heart decreases
significantly (LVEF >)

Organs receive less blood
and suffer from the lack of
oxygen and nutrients.

# Why chest pains?

- Coronary arteries receive a lot less blood than usual.
- → Cardiomyocytes aren't fed in oxygen and nutrients, and die.

# **Consequences?**

- Cardiogenic shock: the blood pumped into the whole body cannot meet the other organs' needs, provoking damages to the liver, kidneys from lack of oxygen, which can be permanent.
- Rhythmic abnormalities
- Thrombus due to residual blood in the left ventricle.
- Death

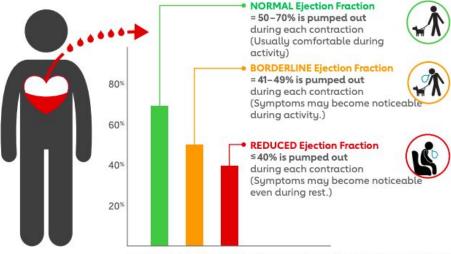
# **TAKOTSUBO SYNDROME: PROBLEMATIC**

# **Project goals:**

- Predict the in-hospital complications
- Predict the kind of heart failure to anticipate the medical care needed.



# How much blood is pumped out?



It is also possible to have a diagnosis of heart failure with a seemingly normal (or preserved) ejection fraction of greater than or equal to 50%.

# Who?

Takotsubo patients between 2015 and 2021 from the CHU de Toulouse.

# What?

Variables selected by reading research articles on the short and long-term prognosis or diagnosis of the disease.

# How?

Reading the patients records and extracting the data patient by patient.

	âge	Date hospit init		lomme	▼ poids	▼ tai	lle ▼	IMC ( kg/ ci = ATCD dépression/ =	ATCD psychiatrique	patho neurologiques	*	HTA 🔻	Dyslipidém =	Tabac 🔻	Diabète =	IRC	AVC/AIT	ATCD Canc ▼	Cano
3/5/1955	(	6 5/	3/2015		1	65	1,78	20,52		0	1	0	0	1	0		0 0	0	. 0
1/6/1964		7 3/	2/2016		0	58	1,6	22,66	)	0	0	1	0	1	0		0 0	0	
30/6/1940		1 10/	2/2016		0	50	1,53	21,36	)	0	0	1	0	0	1		0 0	0	
20/10/1962		8 25/	2/2016		1	60	1,78	18,94	)	0	1	0	0	1	0		0 0	0	
19/8/1939		1 1/	3/2016		0	70	1,56	28,76	)	0	0	1	0	0	0		0 0	1	
6/11/1929		1 28/	2/2016		0	57	1,54	24,03		0	1	1	0	0	0		0 0	0	
15/2/1935		6 6/	4/2016		0	65		(	)	0	0	1	0	0	1		0 0	0	
20/4/1944	7	7 20/	4/2016		0	80	1,56	32,87	)	0	1	1	0	0	1	1	0 1	1	9
29/5/1991		0 24/	4/2016		0	63	1,58	25,24	)	0	0	0	0	0	0		0 0	0	
3/9/1958		2 2/	5/2016		1	55	1,68	19,49	)	1	0	0	1	1	1		0 0	0	
11/5/1928		3 18/	5/2016		0	55	1,55	22,89	)	0	0	0	0	1	0		0 0	0	
12/3/1949		2 2/	6/2016		0	80	1,55	33,30	)	0	0	1	1	0	0		0 0	0	
11/8/1939		1 3/	6/2016	,	0	98	1,69	34,31	)	0	0	1	1	1	1		0 0	0	
12/10/1934		6 13/	7/2016		0	67	1,7	23,18	)	0	1	1	0	0	0		0 1	1	
27/6/1949	7	2 23/	7/2016		0	65	1,69	22,76		0	0	1	0	0	0		0 0	0	
12/3/1931		0 23/	7/2016		1	73	1,7	25,26	)	0	0	0	0	0	0		0 0	0	
27/7/1955		5 26/	7/2016		0	82	1,67	29,40		0	0	1	0	1	0		0 0	0	
30/7/1955		5 17/	8/2016		0	58	1,63	21,83	)	0	0	1	0	1	1	8	0 0	0	9
26/5/1933		8 20/	9/2016		0	57	1,55	23,73	)	0	0	1	0	0	1		0 0	0	
31/12/1941		9 27/	9/2016		0	74	1,5	32,89	)	0	0	1	1	0	0		0 0	0	
27/6/1952		9 5/1	0/2016		0	58	1,6	22,66		0	0	1	0	0	0		0 0	0	
1/7/1935		6 12/1	0/2016		0	60	1,58	24,03	)	0	0	1	1	0	0		0 0	0	
19/1/1944		7 19/1	0/2016		0	55	1,55	22,89	)	0	0	1	1	0	0		0 0	0	
20/2/1943		8 13/1	1/2016		0	70	1,76	22,60	)	0	0	0	0	0	1		0 0	0	7
31/1/1939		2 29/1	1/2016		0	68	1,64	25,28		0	1	1	0	0	0		0 1	0	
15/1/1949		2 28/1	2/2016		1	70	1,73	23,39	)	0	0	0	0	1	0		0 0	0	
27/12/1944		6 4/	1/2017		0	49	1,6	19,14	)	0	0	0	0	1	0		0 0	0	
3/10/1942		8 20/	1/2017		0	60	1,54	25,30		0	0	0	0	0	0		0 1	0	
3/5/1973	4	8 26/	1/2017		1	84	1,86	24,28	)	0	0	0	0	0	0		0 0	0	
20/9/1998	- 2	2 27/	1/2017		0	59	1,6	23,05	)	0	0	0	0	0	0		0 0	0	
5/12/1946	- 1	4 8/	2/2017		0	75	1,58	30,04	)	0	0	1	1	0	0		0 0	1	
23/7/1931		9 27/	2/2017		0	63	1,53	26,91	)	0	0	1	0	0	0		0 0	0	
16/2/1949		2 27/	2/2017		0	67	1,59	26,50	)	0	0	0	1	1	0		0 0	1	
31/12/1949		1 14/	5/2017		0	64	1,69	22,41	)	0	1	0	1	1	0	4	0 0	0	9
11/6/1959	(	2 11/	6/2017		0	65	1,68	23,03		0	0	0	1	0	0		0 0	0	
19/4/1924		7 14/	6/2017		0	53	1,6	20,70	)	0	0	_r_ 1	0	0	0		0 0	0	
8/2/1930		1 20/	6/2017		0	60	1,57	24,34	)	0	0	0	0	0	0		0 0	1	
20/10/1043			6/2017		^	E7	16	22.27		n	4	0	0	^	0	1	0 1	^	$\overline{}$

#### **DEMOGRAPHICS**

Age, BMI, Sex.

# **MEDICAL HISTORY**

- Depression or Anxiety
- Psychiatric disorders
- Neurological diseases
- Hypertension
- Dyslipidemia
- Smoking
- Diabetes
- Chronic Renal Failure
- Stroke or Transient Ischemic Attack (TIA)
- Cancer history or active Cancer
- Chronic Obstructive Pulmonary Disease (COPD) or asthma
- Alcoholism
- Cardiac diseases history

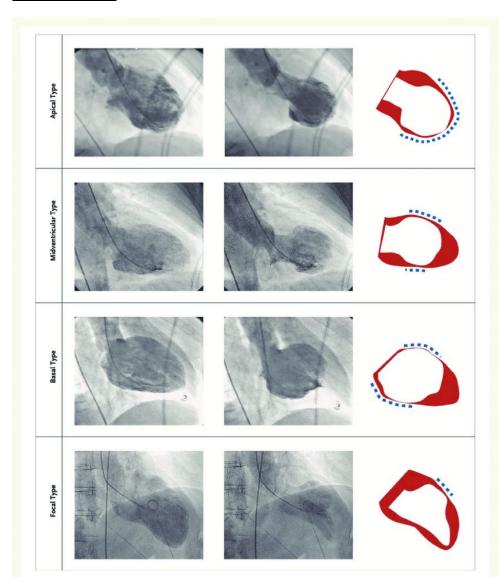
## **STRESS FACTOR**

- Emotional
- Physical

# TREATMENTS BEFORE/AFTER EVENT

- Beta Blockers
- Angiotensin Converting Enzyme Inhibitor
- Angiotensin II receptor blockers
- Aspirin
- Antiplatelets drugs
- Oral anticoagulation drugs
- Statins
- Anti-depressants and anxiolytics

# **ANATOMY**



Apical type

Medioventricular type

Basal type

Focal type

#### **HAEMODYNAMICS**

- Left Ventricle Ejection Fraction (LVEF)
- ECG abnormalities

## **BIOMARKERS**

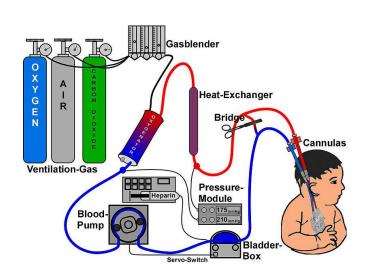
- Troponin T (entry and peak)
- NT pro-BNP
- CRP

# **CORONAROGRAPHY**

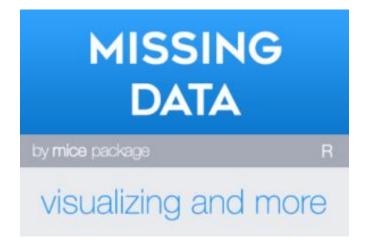
- Harm
- Healthy

# **IN-HOSPITAL COMPLICATIONS**

- Heart Failure
- Right Ventricle Harm
- Ventricular arrhythmia
- Left ventricle thrombus
- Cardiogenic shock
- ECMO (Extracorporeal membrane oxygenation)
- Death



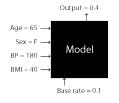
# **ALGORITHM PREDICTION**



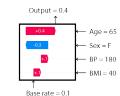












# **CONLUSION AND PERSPECTIVES**

- Difficulty of data collection: precision
- Clinical features mostly how about genetics, epigenetics, proteomics...?
- Difficulty in data processing: dropping data when we already have just a few?
- ICM codes  $\rightarrow$  make the algorithm more precise by distinguishing the physical/emotional stresses.
- Contradiction of adopting the latest technologies.

#### **BIBLIOGRAPHY AND SOURCES**

#### **ARTICLES:**

Templin, C., Ghadri, J. R., Diekmann, J., Napp, L. C., Bataiosu, D. R., Jaguszewski, M., Cammann, V. L., Sarcon, A., Geyer, V., Neumann, C. A., Seifert, B., Hellermann, J., Schwyzer, M., Eisenhardt, K., Jenewein, J., Franke, J., Katus, H. A., Burgdorf, C., Schunkert, H., ... Lüscher, T. F. (2015). Clinical Features and Outcomes of Takotsubo (Stress) Cardiomyopathy. New England Journal of Medicine, 373(10), 929–938.

Bloom, M. W., Greenberg, B., Jaarsma, T., Januzzi, J. L., Lam, C. S. P., Maggioni, A. P., Trochu, J.-N., & Butler, J. (2017). Heart failure with reduced ejection fraction. Nature Reviews Disease Primers, 3(1).

Sharkey, S. W., Windenburg, D. C., Lesser, J. R., Maron, M. S., Hauser, R. G., Lesser, J. N., Haas, T. S., Hodges, J. S., & Maron, B. J. (2010). Natural History and Expansive Clinical Profile of Stress (Tako-Tsubo) Cardiomyopathy. Journal of the American College of Cardiology, 55(4), 333–341.

Ono, R., & Falcão, L. M. (2016). Takotsubo cardiomyopathy systematic review: Pathophysiologic process, clinical presentation and diagnostic approach to Takotsubo cardiomyopathy. International Journal of Cardiology, 209, 196–205.

Ghadri, J. R., Kato, K., Cammann, V. L., Gili, S., Jurisic, S., Di Vece, D., Candreva, A., Ding, K. J., Micek, J., Szawan, K. A., Bacchi, B., Bianchi, R., Levinson, R. A., Wischnewsky, M., Seifert, B., Schlossbauer, S. A., Citro, R., Bossone, E., Münzel, T., ... Templin, C. (2018). Long-Term Prognosis of Patients With Takotsubo Syndrome. Journal of the American College of Cardiology, 72(8), 874–882.

Veillet-Chowdhury, M., Hassan, S. F., & Stergiopoulos, K. (2014). Takotsubo cardiomyopathy: A review. Acute Cardiac Care, 16(1), 15–22.

Limongelli, G., D'Alessandro, R., Masarone, D., Maddaloni, V., Vriz, O., Minisini, R., Citro, R., Calabrò, P., Russo, M. G., Calabrò, R., Pacileo, G., Bossone, E., & Elliott, P. M. (2013). Takotsubo Cardiomyopathy. Heart Failure Clinics, 9(2), 207–216.

Nguyen, T. H., Neil, C. J., Sverdlov, A. L., Mahadavan, G., Chirkov, Y. Y., Kucia, A. M., Stansborough, J., Beltrame, J. F., Selvanayagam, J. B., Zeitz, C. J., Struthers, A. D., Frenneaux, M. P., & Horowitz, J. D. (2011). N-Terminal Pro-Brain Natriuretic Protein Levels in Takotsubo Cardiomyopathy. The American Journal of Cardiology, 108(9), 1316–1321.

Scantlebury, D. C., & Prasad, A. (2014). Diagnosis of Takotsubo Cardiomyopathy. Circulation Journal, 78(9), 2129–2139.

#### **WEBSITES**

 $\underline{https://www.health.harvard.edu/heart-health/takotsubo-cardiomyopathy-broken-heart-syndrome}$ 

http://www.remede.org/librairie-medicale/pdf/e9782224030766.pdf

https://www.svhhearthealth.com.au/conditions/takotsubo-cardiomyopathy

https://www.heartfoundation.org.nz/your-heart/heart-tests/coronary-angiography

https://www.vidal.fr/actualites/22737-syndrome-du-c-ur-brise-de-takotsubo-publication-d-un-consensus-d-experts-mais-des-questions-demeurent.html

https://www.takotsubo-registry.com/takotsubo-score.html

http://www.mednuc.net/wp-content/uploads/2017/05/Takotsubo-et-ph%C3%A9ochromocytome.pdf

https://www.uofmhealth.org/health-library/tx4091abc