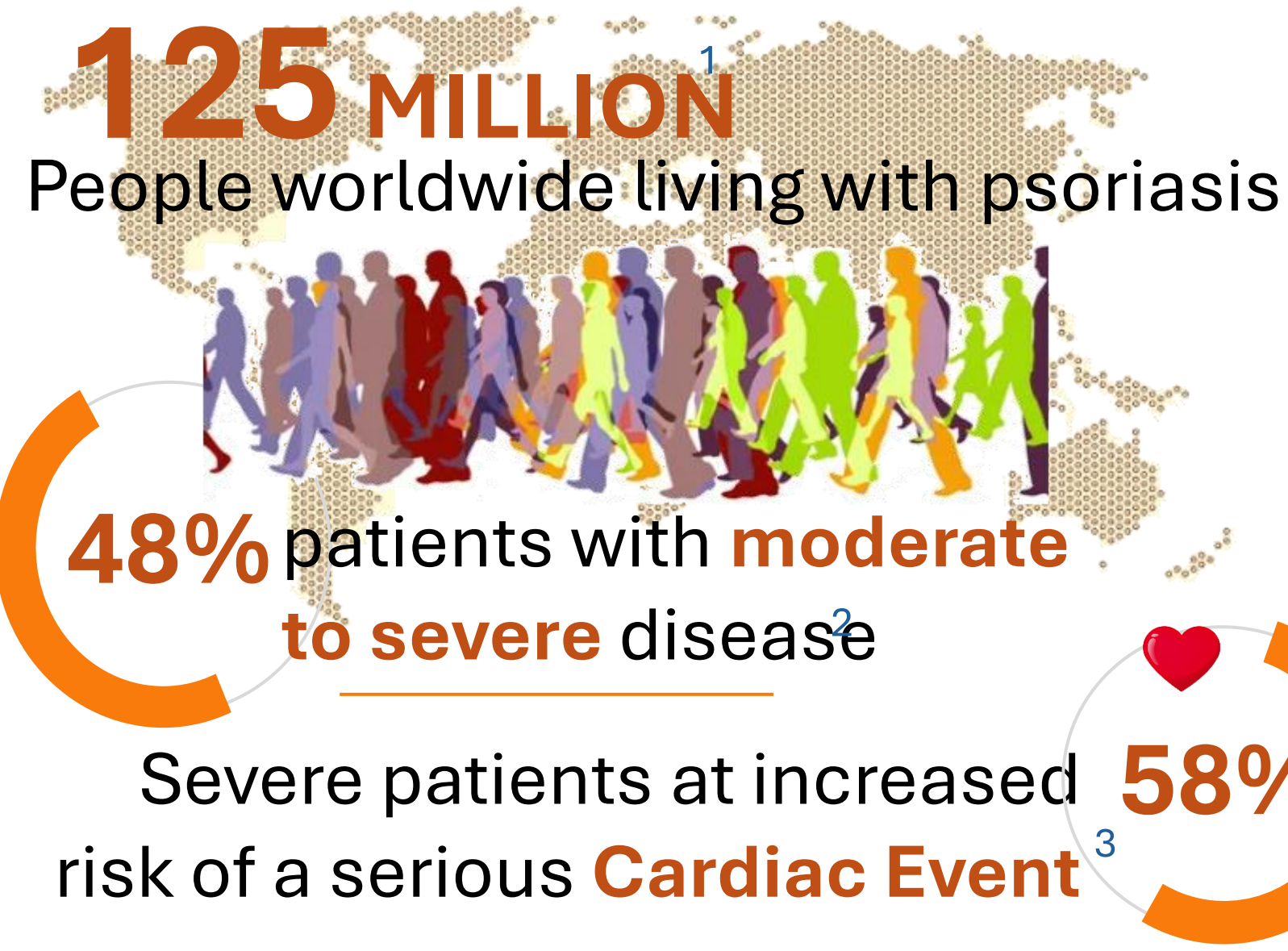


Bridging the Gap: Exploring Cardiovascular Risk with Modern Biotherapies in Psoriasis

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INTRODUCTION



- The treatment for psoriasis has undergone a paradigm shift with the advent of modern biotherapies. These **targeted agents have revolutionized the management of moderate to severe psoriasis**. However, concerns have also emerged regarding their potential impact on cardiovascular health.
- This scoping review aims to **synthesize** the current evidence, **fill research gaps**, and **provide valuable insights** into the evolving role of biological agents in managing psoriasis.

METHODS

- Study design:** Follow PRISMA guideline
- Databases:** PubMed, ScienceDirect, Cochrane Library, Wiley Online Library.
- Keywords:** psoriasis, psoriasis arthritis, TNF- α , cardiovascular events
- Eligible criteria:** Publications between 2013 and 2023, English only. Articles focused on psoriasis adult patients undergoing biotherapies and cardiovascular events.

RESULTS

Association of Biological therapies and Cardiac risks in Psoriatic Patients

	26-52 weeks	Authors, years	1-5 years	Authors, years	≥ 5 years	Authors, years
Composed Biological agents						
Cardiac Events (CEs)			Compared to topical agents No elevated risks ⁵ (HR= 1.0, 0.4-2.7)	Carretero et al., 2015	Compared to biologic-naive Reduced risks ¹⁰ (HR= 0.8, 0.70-0.95)**	Lee et al., 2018
Major CEs	Compared to baseline Decreased coronary inflammation ($p < 0.001$)***, suggesting reduced risks ⁴	Elnabawi et al., 2019	Compared to mild disease No elevated risks ⁶ (HR= 0.9, 0.55-1.36)	Hong et al., 2021		
			Compared to topical agents No elevated risks ⁷ (IR/100 PY= 0.8, 0.41-1.31)	Reich et al., 2015	Compared to placebo No elevated risks ¹¹ (OR=1.5, 0.34-6.24)	Rungapiromnan et al., 2017
			Compared to other therapies No elevated risks ⁸ (HR= 0.6, 0.30-1.10)	Ahlehoff et al., 2015		
CE deaths			Compared to topical agents Decreased risks ⁶ (HR=0.5, 0.29-0.74)**	Hong et al., 2021		
			Compared to other therapies Reduced risks ⁹ (HR= 0.5, 0.17-1.38)*	Ahlehoff et al., 2013		
Tumor necrosis factor-α inhibitors (TNF-α)						
Cardiac Events (CEs)	Compared to baseline A significant increase CFR***, suggest a decreased risk ¹² .	Pisericco et al., 2016			Compared to biologic-naive Reduced risks ⁹ (HR= 0.9, 0.76-0.95)**	Lee et al., 2018
Major CEs	Compared to non-PsO patients A significant improvement in TAPSE and right ventricular free wall peak systolic velocity, suggest a decreased risk ¹³ ***	Herédi et al., 2016				
	Compared to topical/photo therapies Having a 5%** reduction in non-calcified plaque burden, suggests a reduced risk ⁷ .	Elnabawi et al., 2019	Compared to methotrexate Reduced risks ¹⁴ (HR= 0.6, 0.45-0.67)***	Wu et al., 2017	Compared to placebo No protected risk ¹⁰ (OR= 0.7, 0.10-4.63)	Rungapiromnan et al., 2017
Myocardial infarction			Compared to other therapies Reduced risks ⁹ (HR= 0.5, 0.22-0.98)*	Ahlehoff et al., 2015	Compared to photo therapy Reduced risks ¹⁶ (H=0.8, 0.60-0.99)*	Wu et al., 2018
			Compared to methotrexate Reduced risks ¹⁴ (HR= 0.5, 0.34-0.71)***	Wu et al., 2017		
Stroke			Compared to topical agents Significantly lower ¹⁵ : IR: 4.9 vs 12.3**	Shaaban et al., 2018		
			Compared to methotrexate Reduced risks ¹⁴ (HR= 0.6, 0.42-0.71)***	Wu et al., 2017		
Angina			Compared to methotrexate Reduced risks ¹⁴ (HR=0.6, 0.41-0.82)**	Wu et al., 2017		

In the review, 1,380,483 psoriasis patients were studied across 155 publications. **Psoriasis can be effectively treated with biologic medicines with a reduction in cardiovascular hazard risks**, particularly:

- TNF- α show reduced cardiac-related risk biomarkers and lower hazard risks for major cardiac events (CEs), myocardial infarction (MI), stroke, and angina, when compared to non-biologic anti-psoriatic medications and even without systemic anti-psoriatic therapy.
- Anti-IL-17 and Anti-IL-12/23 medications exhibit potential cardioprotective effects against coronary artery disease. However, alternative studies suggest conflicting evidence regarding a potential association between these therapies and an elevated risk of major cardiovascular events.
- Anti-IL-23 therapies show no discernible effect on major cardiovascular events or heart failure in short-term studies.
- Further research is needed for comprehensive understanding.

	≤ 26 weeks	Authors, years	26-52 weeks	Authors, years	1-5 years	Authors, years
Anti-Interleukin-17 inhibitor (Anti-IL-17)						
Major CEs	Compared to placebo No protected risk ¹⁰ (OR= 1.0, 0.09-11.09)	Rungapiromnan et al., 2017	Compared to topical/photo therapies Having a 12%** reduction in non-calcified plaque burden; suggests a reduced risk ⁷ .	Elnabawi et al., 2019	Compared to TNF α Increased risks ¹⁸ (weighted HR= 1.9, 1.2-3.0)***	Vegas et al., 2022
Anti-Interleukin-12/23 inhibitor (Anti-IL-12/23)						
Major CEs	Compared to control Increased risks ¹⁷ (OR=4.2, 1.07- 16.75)**	Tzellos et al., 2013	Compared to other therapies ⁸ No elevated effect (HR=1.5, 0.47-4.94)	Ahlehoff et al., 2015		
			Compared to topical/photo therapies Having a 2%** reduction in non-calcified plaque burden, suggests reduced risks ⁷ .	Elnabawi et al., 2019	Compared to TNF- α Increased risks ¹⁸ (weighted HR=2.0, 1.3-3.0)***	Vegas et al., 2022
Anti-Interleukin-23 inhibitor (Anti-IL-23)						
Major CEs	Compared to placebo No effect ¹⁹	Champs et al., 2019				
Heart failure	Compared to placebo No effect ¹⁹	Champs et al., 2019				

CONCLUSIONS

- TNF- α inhibitors demonstrate **consistent cardiovascular benefits**, including reduced incidence rates of major CEs.
- IL 12/23 inhibitors, IL-17 inhibitors, and IL-23 inhibitors exhibit conflicting findings underscore the complexity of these treatments' effects, necessitating further research to comprehensively elucidate their long-term cardiovascular impact
- These findings emphasize the importance of individualized treatment approaches in optimizing psoriasis management strategies, weighing both efficacy and cardiovascular safety considerations.

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